

# **Dossier zur Nutzenbewertung gemäß § 35a SGB V**

*Sparsentan (Filspari™)*

Vifor Pharma Deutschland GmbH

## **Separater Anhang 4-G**

*Erwachsene Patienten mit primärer Immunglobulin-A-Nephropathie (IgAN) mit einer Ausscheidung von Eiweiß im Urin von  $\geq 1,0$  g/Tag (oder einem Protein/Kreatinin-Quotienten im Urin von  $\geq 0,75$  g/g)*

**Medizinischer Nutzen und  
medizinischer Zusatznutzen,  
Patientengruppen mit therapeutisch  
bedeutsamem Zusatznutzen**

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PT2RPR_FSLM: Percent change from baseline in UP/C by subgroup _____	1339
PT2RPR_FSHM: Course of UP/C (percentage) by subgroup _____	1356
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PF2URFT_FSKM: Time to complete remission by subgroup _____	1514
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PT2DT_FSTM: Time to death by subgroup _____	1697
PF2DT_FSKM: Time to death by subgroup _____	1698
PT2CKD_FSIM: Patients entering CKD stage 4 or 5 by subgroup _____	1699
PT2CKDT_FSTM: Time to CKD stage 4 or 5 by subgroup _____	1711
PF2CKDT_FSKM: Time to CKD stage 4 or 5 by subgroup _____	1723
PT2MIS_FSIM: Patients with systemic immunosuppressive medication by subgroup _____	1725
PT2MIST_FSTM: Time to systemic immunosuppressive medication by subgroup _____	1737
PF2MIST_FSKM: Time to systemic immunosuppressive medication by subgroup _____	1747
PT2MIK_FSIM: Patients with systemic immunosuppressive renal medication by subgroup _____	1750
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PF2VSIT_FSKM: Time to increase in EQ-5D VAS by at least 15 points by subgroup _____	1885
PT2VSDT_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup _____	1887
PF2VSDT_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup _____	1899
PT2KBUC_FSHM: KDQOL: Course of burden of kidney disease by subgroup _____	1905
PT2KBUC_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup _____	1946
PF2KBUC_FSGM: KDQOL: Change from baseline in burden of kidney disease by subgroup _____	1962
PT2KBUIT_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup _____	1963
PF2KBUIT_FSKM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup _____	1975
PT2KBUDT_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup _____	1976
PF2KBUDT_FSKM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup _____	1988
PT2KEFC_FSHM: KDQOL: Course of effect of kidney disease by subgroup _____	1989
PT2KEFC_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup _____	2020
PF2KEFC_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup _____	2046
PT2KEFIT_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup _____	2056
PF2KEFIT_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup _____	2067
PT2KEFDT_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup _____	2074
PF2KEFDT_FSKM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup _____	2086
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PF2KSYC_FSGM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup _____	2147
PT2KSYIT_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup _____	2149
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PT2KSYDT_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup _____	2162
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PT2KPSC_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup _____	2207
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PT2KPSIT_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup _____	2234
PF2KPSIT_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup _____	2246
PT2KPSDT_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup _____	2260
PF2KPSDT_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup _____	2272
PT2KMSC_FSHM: KDQOL-SF12: Course of MCS by subgroup _____	2279
PT2KMSC_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup _____	2310
PF2KMSC_FSGM: KDQOL-SF12: Change from baseline in MCS by subgroup _____	2336
PT2KMSIT_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup _____	2337
PF2KMSIT_FSKM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup _____	2349
PT2KMSDT_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup _____	2350
PF2KMSDT_FSKM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup _____	2362
PT2A_SSIM: Incidence of TEAEs during double-blind period by subgroup _____	2363
PT2AN_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup _____	2375
PT2AC_SSIM: Incidence of severe TEAEs during double-blind period by subgroup _____	2387
PT2AS_SSIM: Incidence of serious TEAEs during double-blind period by subgroup _____	2399
PT2AT_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup _____	2411
PT2AD_SSIM: Incidence of fatal TEAEs during double-blind period by subgroup _____	2423
PT2AA_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup _____	2424
PT2AAN_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup _____	2436
PT2AAC_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup _____	2448
PT2AAS_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup _____	2460
PT2A_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups _____	2472
PT2AC_SSSM: Incidence of severe TEAEs during double-blind period by SOC/PT by subgroups _____	2496
PT2AS_SSSM: Incidence of serious TEAEs during double-blind period by SOC/PT by subgroups _____	2497
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PT2AEFN_SSIM: Incidence of AESI abnormal liver function during double-blind period – non-severe by subgroup _____	2510
PT2AEFC_SSIM: Incidence of AESI abnormal liver function during double-blind period – severe by subgroup _____	2522

PT2AEFS_SSIM: Incidence of AESI abnormal liver function during double-blind period – serious by subgroup	2523
PT2AEV_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup	2524
PT2AEVN_SSIM: Incidence of AESI COVID-19 during double-blind period – non-severe by subgroup	2536
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PT2AECC_SSIM: Incidence of AESI cardiovascular system during double-blind period – severe by subgroup	2585
PT2AECS_SSIM: Incidence of AESI cardiovascular system during double-blind period – serious by subgroup	2586
PT2AEH_SSIM: Incidence of AESI hypotension during double-blind period by subgroup	2587
PT2AEHN_SSIM: Incidence of AESI hypotension during double-blind period – non-severe by subgroup	2599
PT2AEHC_SSIM: Incidence of AESI hypotension during double-blind period – severe by subgroup	2611
PT2AEHS_SSIM: Incidence of AESI hypotension during double-blind period – serious by subgroup	2612
PT2AEL_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup	2613
PT2AELN_SSIM: Incidence of AESI hepatic disorders during double-blind period – non-severe by subgroup	2625
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PT2AELS_SSIM: Incidence of AESI hepatic disorders during double-blind period – serious by subgroup	2638
PT2AEP_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup	2639
PT2AEPN_SSIM: Incidence of AESI acute pancreatitis during double-blind period – non-severe by subgroup	2651
PT2AEP_C_SSIM: Incidence of AESI acute pancreatitis during double-blind period – severe by subgroup	2663
PT2AEP_S_SSIM: Incidence of AESI acute pancreatitis during double-blind period – serious by subgroup	2664
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PT2AEOC_SSIM: Incidence of AESI fluid retention during double-blind period – severe by subgroup	2689
PT2AEOS_SSIM: Incidence of AESI fluid retention during double-blind period – serious by subgroup	2690
PT2AEA_SSIM: Incidence of AESI anemia during double-blind period by subgroup	2691
PT2AEAN_SSIM: Incidence of AESI anemia during double-blind period – non-severe by subgroup	2703
PT2AEAC_SSIM: Incidence of AESI anemia during double-blind period – severe by subgroup	2715
PT2AEAS_SSIM: Incidence of AESI anemia during double-blind period – serious by subgroup	2716
PT2AEU_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup	2717

PT2AEUN_SSIM: Incidence of AESI hyperkalemia during double-blind period – non-severe by subgroup	2729
PT2AEUC_SSIM: Incidence of AESI hyperkalemia during double-blind period – severe by subgroup	2741
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PT2AEIC_SSIM: Incidence of AESI acute kidney injury during double-blind period – severe by subgroup	2793
PT2AEIS_SSIM: Incidence of AESI acute kidney injury during double-blind period – serious by subgroup	2794

Table PT1BTS\_FMA0: Treatment status double-blind period  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Treatment status	PROTECT	Sparsentan	Treatment completed	202	18 (8.9)
			Treatment discontinued	202	19 (9.4)
			Treatment ongoing	202	165 (81.7)
			Untreated patients	202	0 (0.0)
	Irbesartan		Treatment completed	202	15 (7.4)
			Treatment discontinued	202	29 (14.4)
			Treatment ongoing	202	158 (78.2)
			Untreated patients	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024

Table PT1BSX\_FMA0: Sex  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Sex	PROTECT	Sparsentan	Male	202	139 (68.8)
			Female	202	63 (31.2)
			Missing	202	0 (0.0)
		Irbesartan	Male	202	143 (70.8)
			Female	202	59 (29.2)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024



Table PT1BRC\_FMA0: Race  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Race	PROTECT	Sparsentan	White	202	130 (64.4)
			Black	202	1 (0.5)
			Asian	202	67 (33.2)
			Others	202	4 (2.0)
			Missing	202	0 (0.0)
		Irbesartan	White	202	142 (70.3)
			Black	202	3 (1.5)
			Asian	202	48 (23.8)
			Others	202	9 (4.5)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024

Table PT1BRG\_FMA0: Geographic region  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Geographic region	PROTECT	Sparsentan	North America	202	35 (17.3)
			Europe	202	98 (48.5)
			Asia Pacific	202	69 (34.2)
			Missing	202	0 (0.0)
		Irbesartan	North America	202	46 (22.8)
			Europe	202	115 (56.9)
			Asia Pacific	202	41 (20.3)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024

Table PT1BCO\_FMA0: Country  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Country	PROTECT	Sparsentan	Australia	202	19 (9.4)
			Belgium	202	9 (4.5)
			Croatia	202	3 (1.5)
			Czech Republic	202	4 (2.0)
			Estonia	202	1 (0.5)
			France	202	3 (1.5)
			Germany	202	9 (4.5)
			Hong Kong	202	14 (6.9)
			Italy	202	11 (5.4)
			Lithuania	202	7 (3.5)
			New Zealand	202	3 (1.5)
			Poland	202	7 (3.5)
			Portugal	202	8 (4.0)
			South Korea	202	24 (11.9)
			Spain	202	16 (7.9)
			Taiwan	202	9 (4.5)
			USA	202	35 (17.3)
			United Kingdom	202	20 (9.9)
	Irbesartan		Australia	202	12 (5.9)
			Belgium	202	3 (1.5)
			Croatia	202	8 (4.0)
			Czech Republic	202	9 (4.5)
			Estonia	202	6 (3.0)
			France	202	10 (5.0)
			Germany	202	7 (3.5)
			Hong Kong	202	9 (4.5)
			Italy	202	15 (7.4)
			Lithuania	202	4 (2.0)
			New Zealand	202	2 (1.0)
			Poland	202	6 (3.0)
			Portugal	202	4 (2.0)
			South Korea	202	14 (6.9)
			Spain	202	14 (6.9)
			Taiwan	202	4 (2.0)
USA	202	46 (22.8)			
United Kingdom	202	29 (14.4)			

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024

Table PT1BE1\_FMA0: Baseline eGFR - Group 1  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Baseline eGFR - Group 1	PROTECT	Sparsentan	< 60 mL/min/1.73 m**2	202	127 (62.9)
			60 to < 90 mL/min/1.73 m**2	202	49 (24.3)
			>= 90 mL/min/1.73 m**2	202	26 (12.9)
			Missing	202	0 (0.0)
	Irbesartan		< 60 mL/min/1.73 m**2	202	129 (63.9)
			60 to < 90 mL/min/1.73 m**2	202	48 (23.8)
			>= 90 mL/min/1.73 m**2	202	25 (12.4)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Results are presented in mL/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate.  
 Source Data: asl, created on: 26JAN2024

Table PT1BE2\_FMA0: Baseline eGFR - Group 2  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Baseline eGFR - Group 2	PROTECT	Sparsentan	< 45 mL/min/1.73 m**2	202	82 (40.6)
			45 to < 60 mL/min/1.73 m**2	202	45 (22.3)
			60 to < 90 mL/min/1.73 m**2	202	49 (24.3)
			>= 90 mL/min/1.73 m**2	202	26 (12.9)
			Missing	202	0 (0.0)
		Irbesartan	< 45 mL/min/1.73 m**2	202	80 (39.6)
			45 to < 60 mL/min/1.73 m**2	202	49 (24.3)
			60 to < 90 mL/min/1.73 m**2	202	48 (23.8)
			>= 90 mL/min/1.73 m**2	202	25 (12.4)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Results are presented in mL/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate.  
 Source Data: asl, created on: 26JAN2024

Table PT1BUR\_FMA0: Baseline UP/C  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Baseline UP/C	PROTECT	Sparsentan	<= 1.25 g/g	202	101 (50.0)
			> 1.25 g/g	202	101 (50.0)
			Missing	202	0 (0.0)
		Irbesartan	<= 1.25 g/g	202	104 (51.5)
			> 1.25 g/g	202	98 (48.5)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Results are presented in g/g. UP/C = Urine protein/creatinine ratio.  
 Source Data: asl, created on: 26JAN2024

Table PT1BPE\_FMA0: Baseline urine protein excretion  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Baseline urine protein excretion	PROTECT	Sparsentan	<= 1.75 g/day	202	98 (48.5)
			> 1.75 g/day	202	104 (51.5)
			Missing	202	0 (0.0)
		Irbesartan	<= 1.75 g/day	202	94 (46.5)
			> 1.75 g/day	202	108 (53.5)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Results are presented in g/day.  
 Source Data: asl, created on: 26JAN2024

Table PT1BAG\_FMB0: Age at informed consent  
 Full analysis Set

Variable	Study	Treatment	N	n	Mean (SD)	Min	Q25	Q50	Q75	Max
Age at informed consent	PROTECT	Sparsentan	202	202	46.56 (12.76)	18.0	37.00	47.00	57.00	73.0
		Irbesartan	202	202	45.43 (12.12)	19.0	36.00	46.00	55.00	76.0

N = total number of patients in analysis set. n = number evaluable values. SD = standard deviation. Q = quantile.  
 Results are presented in years.  
 Source Data: asl, created on: 26JAN2024



Table PT1BWI\_FMB0: Baseline BMI  
 Full analysis Set

Variable	Study	Treatment	N	n	Mean (SD)	Min	Q25	Q50	Q75	Max
Baseline BMI	PROTECT	Sparsentan	202	202	28.52 (5.21)	18.3	25.10	27.85	31.40	47.2
		Irbesartan	202	201	28.34 (5.66)	17.7	24.60	27.50	31.00	49.4

N = total number of patients in analysis set. n = number evaluable values. SD = standard deviation. Q = quantile.  
 Results are presented in kg/m\*\*2.  
 BMI = Body mass index.  
 Source Data: asl, created on: 26JAN2024

Table PT1BHA\_FMB0: Baseline HbA1c  
 Full analysis Set

Variable	Study	Treatment	N	n	Mean (SD)	Min	Q25	Q50	Q75	Max
Baseline HbA1c	PROTECT	Sparsentan	202	202	5.51 (0.57)	3.9	5.20	5.50	5.80	9.7
		Irbesartan	202	202	5.52 (0.59)	4.3	5.20	5.40	5.70	8.9

N = total number of patients in analysis set. n = number evaluable values. SD = standard deviation. Q = quantile.  
 Results are presented in %.  
 HbA1c = Hemoglobin Alc.  
 Source Data: asl, created on: 26JAN2024

Table PT1BE\_FMB0: Baseline eGFR  
 Full Analysis Set

Variable	Study	Treatment	N	n	Mean (SD)	Min	Q25	Q50	Q75	Max
Baseline eGFR	PROTECT	Sparsentan	202	202	56.86 (24.38)	24.0	38.00	50.00	71.00	128.0
		Irbesartan	202	202	57.08 (23.58)	26.0	39.00	50.00	70.00	123.0

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = total number of patients in analysis set. n = number evaluable values. SD = standard deviation. Q = quantile.  
 Results are presented in mL/min/1.73 m<sup>2</sup>. eGFR = estimated glomerular filtration rate.  
 Source Data: asl, created on: 26JAN2024

Table PT1BAD\_FMA0: Age at IgAN diagnosis - Group 1  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Age at IgAN diagnosis	PROTECT	Sparsentan	<= 18 years	202	9 (4.5)
			>18 to 40 years	202	102 (50.5)
			> 40 years	202	91 (45.0)
			Missing	202	0 (0.0)
		Irbesartan	<= 18 years	202	5 (2.5)
			>18 to 40 years	202	109 (54.0)
			> 40 years	202	88 (43.6)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Results are presented in years. IgAN = immunoglobulin A nephropathy  
 Source Data: asl, created on: 26JAN2024

Table PT1BRP\_FMA0: Time since renal biopsy - Group 1  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Time since renal biopsy	PROTECT	Sparsentan	<= 5 years	202	113 (55.9)
			> 5 years	202	89 (44.1)
			Missing	202	0 (0.0)
		Irbesartan	<= 5 years	202	127 (62.9)
			> 5 years	202	75 (37.1)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Results are presented in years.  
 Source Data: asl, created on: 26JAN2024

Table PT1BMI\_FMA0: Previous renal immunosuppressives  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Previous renal immunosuppressives	PROTECT	Sparsentan	Yes	202	10 (5.0)
			No	202	192 (95.0)
			Missing	202	0 (0.0)
		Irbesartan	Yes	202	11 (5.4)
			No	202	191 (94.6)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024

Table PT1BMR\_FMA0: RAAS inhibitors at Screening  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
RAAS inhibitors at Screening	PROTECT	Sparsentan	Yes	202	201 (99.5)
			No	202	1 (0.5)
			Missing	202	0 (0.0)
		Irbesartan	Yes	202	202 (100.0)
			No	202	0 (0.0)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 RAAS = Renin-angiotensin-aldosterone system.  
 Source Data: asl, created on: 26JAN2024

Table PT1BMRE\_FMA0: ACEI at MLD at Screening  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
ACEI at MLD at Screening	PROTECT	Sparsentan	Yes	202	51 (25.2)
			No	202	151 (74.8)
			Missing	202	0 (0.0)
		Irbesartan	Yes	202	53 (26.2)
			No	202	149 (73.8)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 ACEI = Angiotensin converting enzyme inhibitor. MLD = Maximum labeled dose.  
 Source Data: asl, created on: 26JAN2024



Table PT1BMRR\_FMA0: ARB at MLD at Screening  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
ARB at MLD at Screening	PROTECT	Sparsentan	Yes	202	85 (42.1)
			No	202	117 (57.9)
			Missing	202	0 (0.0)
		Irbesartan	Yes	202	76 (37.6)
			No	202	126 (62.4)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 ARB = Angiotensin II receptor blocker. MLD = Maximum labeled dose.  
 Source Data: asl, created on: 26JAN2024

Table PT1BMRB\_FMA0: ACEI and ARB at MLD at Screening  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
ACEI and ARB at MLD at Screening	PROTECT	Sparsentan	Yes	202	5 (2.5)
			No	202	197 (97.5)
			Missing	202	0 (0.0)
		Irbesartan	Yes	202	4 (2.0)
			No	202	198 (98.0)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 ACEI = Angiotensin converting enzyme inhibitor. ARB = Angiotensin II receptor blocker. MLD = Maximum labeled dose.  
 Source Data: asl, created on: 26JAN2024

Table PT1BMA\_FMA0: Antihypertensive medications at Baseline  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Antihypertensive medications at Baseline	PROTECT	Sparsentan	Yes	202	88 (43.6)
			No	202	114 (56.4)
			Missing	202	0 (0.0)
		Irbesartan	Yes	202	83 (41.1)
			No	202	119 (58.9)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024

Table PT1BML\_FMA0: Lipid-lowering medications at Baseline  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Lipid-lowering medications at Baseline	PROTECT	Sparsentan	Yes	202	112 (55.4)
			No	202	90 (44.6)
			Missing	202	0 (0.0)
		Irbesartan	Yes	202	111 (55.0)
			No	202	91 (45.0)
			Missing	202	0 (0.0)

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024

Table PT1BA\_FMB0: Age at IgAN diagnosis  
 Full Analysis Set

Variable	Study	Treatment	N	n	Mean (SD)	Min	Q25	Q50	Q75	Max
Age at IgAN diagnosis	PROTECT	Sparsentan	202	202	40.19 (13.35)	10.0	30.00	39.00	51.00	72.0
		Irbesartan	202	202	38.96 (12.38)	8.0	31.00	38.00	48.00	75.0

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = total number of patients in analysis set. n = number evaluable values. SD = standard deviation. Q = quantile.  
 Results are presented in years. IgAN = immunoglobulin A nephropathy  
 Source Data: asl, created on: 26JAN2024

Table PT1BR\_FMB0: Time since renal biopsy  
 Full Analysis Set

Variable	Study	Treatment	N	n	Mean (SD)	Min	Q25	Q50	Q75	Max
Times since renal biopsy	PROTECT	Sparsentan	202	202	6.41 (6.48)	0.0	1.00	4.00	10.00	33.0
		Irbesartan	202	202	6.37 (7.10)	0.0	1.00	4.00	10.00	36.0

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = total number of patients in analysis set. n = number evaluable values. SD = standard deviation. Q = quantile.  
 Results are presented in years.  
 Source Data: asl, created on: 26JAN2024

Table PT1BSS\_FMA0: Study status double-blind period  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Study status	PROTECT	Sparsentan	Completed	202	18 (8.9)
			Premature discontinuation	202	3 (1.5)
			Ongoing	202	181 (89.6)
		Irbesartan	Completed	202	14 (6.9)
			Premature discontinuation	202	11 (5.4)
			Ongoing	202	177 (87.6)

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024

Table PT1BSR\_FMA0: Reason for premature study discontinuation during double-blind period  
 Full analysis Set

Parameter	Study	Treatment	Category	N	n (%)
Reason premature study discontinuation	PROTECT	Sparsentan	No discontinuation	202	199 (98.5)
			Deaths	202	0 (0.0)
			Lost to follow-up	202	0 (0.0)
			PI decision	202	0 (0.0)
			Study terminated by sponsor	202	0 (0.0)
			Withdrawal of consent	202	3 (1.5)
		Irbesartan	No discontinuation	202	191 (94.6)
			Deaths	202	0 (0.0)
			Lost to follow-up	202	1 (0.5)
			PI decision	202	3 (1.5)
			Study terminated by sponsor	202	0 (0.0)
			Withdrawal of consent	202	7 (3.5)

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024



Table PT1BTR\_FMA0: Reason for premature treatment discontinuation during double-blind period  
 Full Analysis Set

Parameter	Study	Treatment	Category	N	n (%)	
Reason premature treatment discontinuation	PROTECT	Sparsentan	No discontinuation	202	183 (90.6)	
			Adverse event/adverse event of interest	202	14 (6.9)	
			Death	202	0 (0.0)	
			Diagnosis of class II-IV CHF	202	0 (0.0)	
			Hyperkalemia resistant to treatment	202	0 (0.0)	
			Lost to follow-up	202	0 (0.0)	
			Subject decision	202	3 (1.5)	
			Physician decision	202	0 (0.0)	
			Pregnancy	202	0 (0.0)	
			Protocol deviation	202	1 (0.5)	
			Receipt of kidney transplant or initiation of chronic dialysis	202	1 (0.5)	
			Site terminated by sponsor	202	0 (0.0)	
			Study terminated by sponsor	202	0 (0.0)	
			Other	202	0 (0.0)	
			Irbesartan		No discontinuation	202
		Adverse event/adverse event of interest			202	8 (4.0)
		Death			202	0 (0.0)
		Diagnosis of class II-IV CHF			202	0 (0.0)
		Hyperkalemia resistant to treatment			202	0 (0.0)
		Lost to follow-up			202	0 (0.0)
		Subject decision			202	12 (5.9)
		Physician decision			202	5 (2.5)
		Pregnancy			202	1 (0.5)
		Protocol deviation			202	1 (0.5)
		Receipt of kidney transplant or initiation of chronic dialysis	202	0 (0.0)		
Site terminated by sponsor	202	0 (0.0)				
Study terminated by sponsor	202	0 (0.0)				
Other	202	2 (1.0)				

N = total number of patients in analysis set. n = number of patients in category.  
 Source Data: asl, created on: 26JAN2024

Table PT1BDS\_FMB0: Study duration of double-blind period  
 Full analysis Set

Variable	Study	Treatment	N	n	Mean (SD)	Min	Q25	Q50	Q75	Max
Study duration	PROTECT	Sparsentan	202	202	68.66 (35.36)	9.9	34.71	77.07	99.57	119.1
		Irbesartan	202	202	67.90 (35.24)	0.6	34.43	75.79	97.00	122.9

$\bar{N}$  = total number of patients in analysis set. n = number evaluable values. SD = standard deviation. Q = quantile.  
 Results are presented in weeks.  
 Duration is defined as time from study day 1 to end of double-blind treatment (regular Week 114).  
 Source Data: asl, created on: 26JAN2024

Table PT1BDT\_FMB0: Treatment duration of double-blind period  
 Full analysis Set

Variable	Study	Treatment	N	n	Mean (SD)	Min	Q25	Q50	Q75	Max
Treatment duration	PROTECT	Sparsentan	202	202	64.29 (35.04)	0.1	32.57	73.43	95.71	115.9
		Irbesartan	202	202	61.27 (35.58)	0.1	27.00	60.86	93.57	114.9

$\overline{N}$  = total number of patients in analysis set. n = number evaluable values. SD = standard deviation. Q = quantile.  
 Results are presented in weeks.  
 Duration is defined as time from study day 1 to last study dose during double-blind period (regular Week 110).  
 Source Data: asl, created on: 26JAN2024

Table PT1RPR\_FML0: Percent change from baseline in UP/C  
 Full Analysis Set

Percent change from baseline in UP/C				Repeated measures analysis					
Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
				Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Week 6	Sparsentan	202	191 (94.6)	-39.47	(-44.52, -33.97)	0.652	(0.576, 0.737)	<0.001	*
	Irbesartan	202	191 (94.6)	-7.15	(-14.89, 1.29)				
Week 36	Sparsentan	202	139 (68.8)	-48.18	(-53.00, -42.88)	0.613	(0.534, 0.705)	<0.001	*
	Irbesartan	202	132 (65.3)	-15.52	(-23.46, -6.76)				
Week 58	Sparsentan	202	113 (55.9)	-46.34	(-51.83, -40.22)	0.632	(0.541, 0.739)	<0.001	*
	Irbesartan	202	99 (49.0)	-15.16	(-24.20, -5.03)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale.

LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. Only key visits up to Week 58 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate.

All observed values up to Week 58 are included in the model.

An increase reflects a worsening of the status of the patient. Results are presented in g/g. UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as a other common summarizing ratios (e.g. RR).

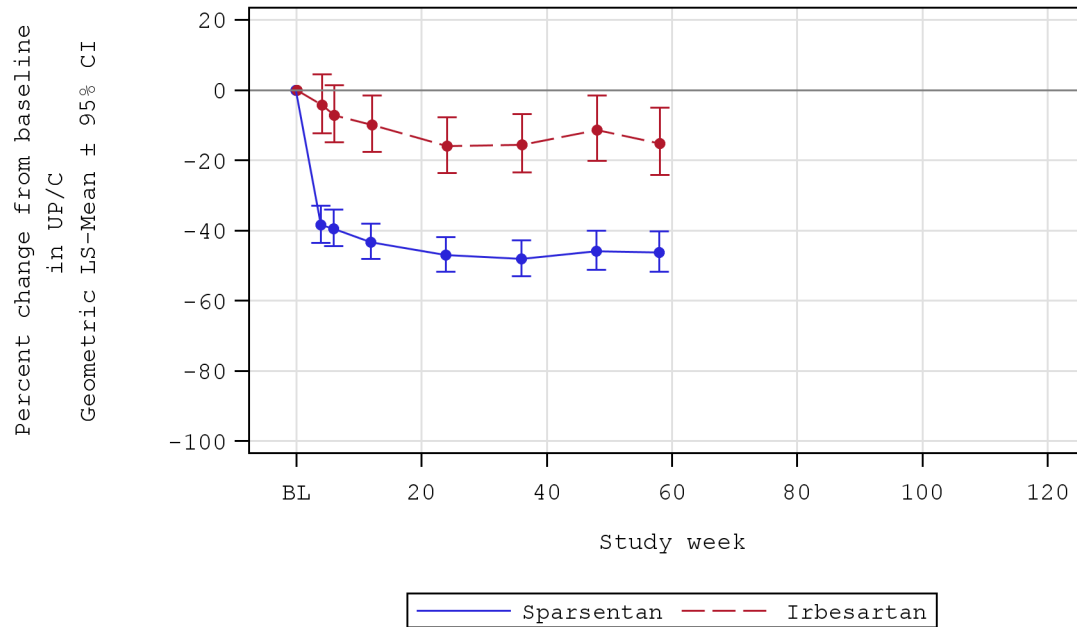
Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FMHO: Course of UP/C (percentage change)  
 Full Analysis Set

Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max
UP/C	Baseline	Sparsentan	202	202 (100.0)	1.43 (0.90)	0.1	0.78	1.25	1.82	7.0
		Irbesartan	202	202 (100.0)	1.44 (0.89)	0.2	0.88	1.23	1.72	6.9
	Week 6	Sparsentan	202	191 (94.6)	0.97 (0.80)	0.1	0.41	0.79	1.27	5.6
		Irbesartan	202	191 (94.6)	1.41 (0.93)	0.1	0.75	1.14	1.78	6.2
	Week 36	Sparsentan	202	139 (68.8)	0.97 (0.84)	0.1	0.36	0.70	1.37	4.4
		Irbesartan	202	132 (65.3)	1.34 (1.04)	0.1	0.63	1.03	1.73	6.6
	Week 58	Sparsentan	202	113 (55.9)	0.96 (0.79)	0.1	0.32	0.77	1.29	3.5
		Irbesartan	202	99 (49.0)	1.40 (1.08)	0.1	0.67	1.10	1.76	5.6
Percent change from baseline in UP/C	Week 6	Sparsentan	202	191 (94.6)	-28.26 (46.40)	-88.7	-60.81	-40.80	-7.96	256.2
		Irbesartan	202	191 (94.6)	5.28 (67.89)	-74.5	-29.48	-8.32	26.69	696.4
	Week 36	Sparsentan	202	139 (68.8)	-27.96 (54.16)	-92.4	-64.65	-43.61	-7.46	168.2
		Irbesartan	202	132 (65.3)	2.91 (80.60)	-83.9	-42.91	-9.96	27.88	581.8
	Week 58	Sparsentan	202	113 (55.9)	-26.68 (54.90)	-94.1	-67.15	-36.52	1.70	166.7
		Irbesartan	202	99 (49.0)	1.58 (79.68)	-93.2	-40.51	-10.95	17.70	451.6

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.  
 An increase reflects a worsening of the status of the patient. Results are presented in g/g. UP/C = urine protein/creatinine ratio.  
 Source Data: alb, created on: 19FEB2024

Figure PF1RPR\_FMG0: Percent change from baseline in UP/C  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	191	139	113
Irbesartan	191	132	99

An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data but original units are displayed. Number of available records are presented for key visits up to Week 58 only. UP/C = urine protein/creatinine ratio. Reference table: PT1RPR\_FML0

Table PT1UEC\_FMH0: Course of urine protein excretion  
 Full Analysis Set

Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Urine protein excretion	Baseline	Sparsentan	202	202 (100.0)	2.22 (1.65)	0.1	1.18	1.76	2.83	14.7		
		Irbesartan	202	202 (100.0)	2.16 (1.24)	0.5	1.33	1.81	2.60	7.5		
	Week 6	Sparsentan	202	192 (95.0)	1.49 (1.21)	0.2	0.67	1.09	1.98	6.5		
		Irbesartan	202	191 (94.6)	2.18 (1.59)	0.4	1.08	1.77	2.70	10.3		
	Week 36	Sparsentan	202	139 (68.8)	1.47 (1.54)	0.1	0.56	0.91	1.92	9.4		
		Irbesartan	202	133 (65.8)	2.12 (1.66)	0.2	1.01	1.67	2.68	9.2		
	Week 58	Sparsentan	202	113 (55.9)	1.45 (1.43)	0.1	0.57	1.03	1.73	7.5		
		Irbesartan	202	98 (48.5)	2.14 (1.71)	0.1	0.90	1.72	2.88	8.6		
	Change from baseline in urine protein excretion	Week 6	Sparsentan	202	192 (95.0)	-0.72 (1.42)	-13.0	-1.21	-0.67	-0.10	5.2	-0.54 [-0.75, -0.34]
			Irbesartan	202	191 (94.6)	0.01 (1.27)	-3.7	-0.63	-0.15	0.51	8.4	
Week 36		Sparsentan	202	139 (68.8)	-0.65 (1.26)	-4.2	-1.23	-0.67	-0.07	4.6	-0.43 [-0.67, -0.19]	
		Irbesartan	202	133 (65.8)	-0.07 (1.45)	-3.3	-0.92	-0.26	0.53	6.5		
Week 58		Sparsentan	202	113 (55.9)	-0.63 (1.29)	-4.4	-1.27	-0.62	0.02	3.3	-0.34 [-0.61, -0.06]	
		Irbesartan	202	98 (48.5)	-0.17 (1.41)	-2.7	-0.93	-0.38	0.25	6.7		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.  
 An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FMC0: Change from baseline in urine protein excretion  
 Full Analysis Set

Change from baseline in urine protein excretion				Repeated measures analysis					
Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Week 6	Sparsentan	202	192 (95.0)	-0.69 (0.09)	(-0.86, -0.51)	-0.67 (0.12)	(-0.91, -0.42)	<0.001 *	
	Irbesartan	202	191 (94.6)	-0.02 (0.09)	(-0.19, 0.15)				
Week 36	Sparsentan	202	139 (68.8)	-0.76 (0.10)	(-0.96, -0.56)	-0.70 (0.14)	(-0.98, -0.42)	<0.001 *	
	Irbesartan	202	133 (65.8)	-0.06 (0.10)	(-0.26, 0.14)				
Week 58	Sparsentan	202	113 (55.9)	-0.66 (0.11)	(-0.88, -0.44)	-0.57 (0.16)	(-0.89, -0.25)	<0.001 *	
	Irbesartan	202	98 (48.5)	-0.09 (0.12)	(-0.32, 0.14)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. Only key visits up to Week 58 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

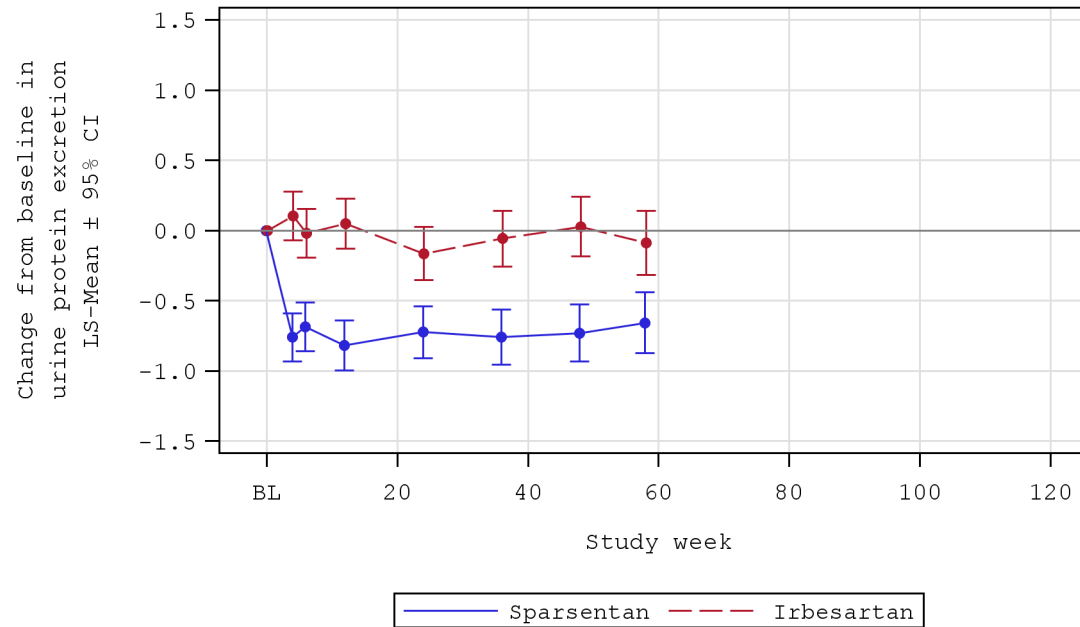
All observed values up to Week 58 are included in the model following a treatment policy approach.

An increase reflects a worsening of the status of the patient. Results are presented in g/day. A first order regressive covariance structure was used.

Source Data: alb, created on: 19FEB2024



Figure PF1UEC\_FMG0: Change from baseline in urine protein excretion  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	192	139	113
Irbesartan	191	133	98

Number of available records are presented for key visits up to Week 58 only. MMRM = Mixed model Repeated Measures. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. All observed values up to Week 58 are included in the model following a treatment policy approach.

An increase reflects a worsening of the status of the patient. Results are presented in g/day. Reference table: PT1UEC\_FMG0

Table PT1URP\_FMP0: Patients with partial remission  
 Full Analysis Set

Variable	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Partial remission	Week 6	Sparsentan	202	192 (95.0)	86 (44.8)	2.251 [1.627, 3.116]	3.267 [2.072, 5.149]	24.9 [15.3, 34.4]	<0.001 *
		Irbesartan	202	191 (94.6)	38 (19.9)				
	Week 36	Sparsentan	202	139 (68.8)	76 (54.7)	2.272 [1.621, 3.186]	3.808 [2.265, 6.399]	30.6 [18.9, 42.4]	<0.001 *
		Irbesartan	202	133 (65.8)	32 (24.1)				
	Week 58	Sparsentan	202	113 (55.9)	55 (48.7)	1.645 [1.148, 2.356]	2.256 [1.277, 3.987]	19.1 [5.2, 32.9]	0.005 *
		Irbesartan	202	98 (48.5)	29 (29.6)				

N = number of patients in analysis set. n = number of patients with event. nval = number of evaluable values.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 Source Data: alb, created on: 19FEB2024

Table PT1URPT\_FMT0: Time to partial remission  
 Full Analysis Set

Time to partial remission			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	145 (71.8)	6.9	(6.0, 12.3)	2.513	(1.929, 3.273)	<0.001 *
Irbesartan	202	94 (46.5)	59.6	(36.1, 114.1)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

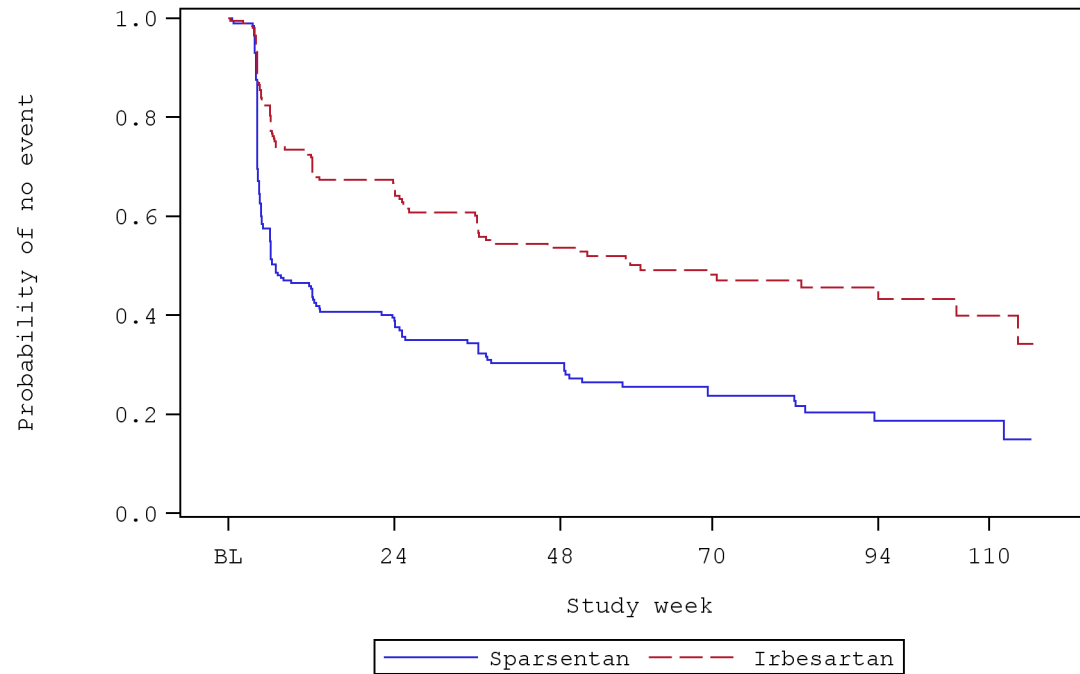
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

Partial remission means an urinary protein excretion of <1.0 g/day.

Source Data: alb\_tte, created on: 31MAY2024

Figure PF1URPT\_FMK0: Time to partial remission - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	62	42	25	11	6
Irbesartan	202	106	69	45	20	8

Partial remission means an urinary protein excretion of <1.0 g/day. No event means no partial remission.  
 Reference table: PT1URPT\_FMT0

Table PT1URF\_FMP0: Patients with complete remission  
 Full Analysis Set

Variable	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Complete remission	Week 6	Sparsentan	202	192 (95.0)	9 (4.7)	18.902 + [1.108, 322.477]	19.828 + [1.146, 343.135]	4.7 [1.2, 8.2]	0.004 *
		Irbesartan	202	191 (94.6)	0 (0.0)				
	Week 36	Sparsentan	202	139 (68.8)	14 (10.1)	2.679 [0.992, 7.232]	2.867 [1.003, 8.197]	6.3 [-0.4, 13.0]	0.056
		Irbesartan	202	133 (65.8)	5 (3.8)				
	Week 58	Sparsentan	202	113 (55.9)	19 (16.8)	2.746 [1.142, 6.602]	3.099 [1.185, 8.109]	10.7 [1.4, 20.0]	0.019 *
		Irbesartan	202	98 (48.5)	6 (6.1)				

N = number of patients in analysis set. n = number of patients with event. nval = number of evaluable values.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 Source Data: alb, created on: 19FEB2024

Table PT1URFT\_FMT0: Time to complete remission  
 Full Analysis Set

Time to complete remission			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	43 (21.3)	NE		2.672	(1.522, 4.692)	<0.001 *
Irbesartan	202	17 (8.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

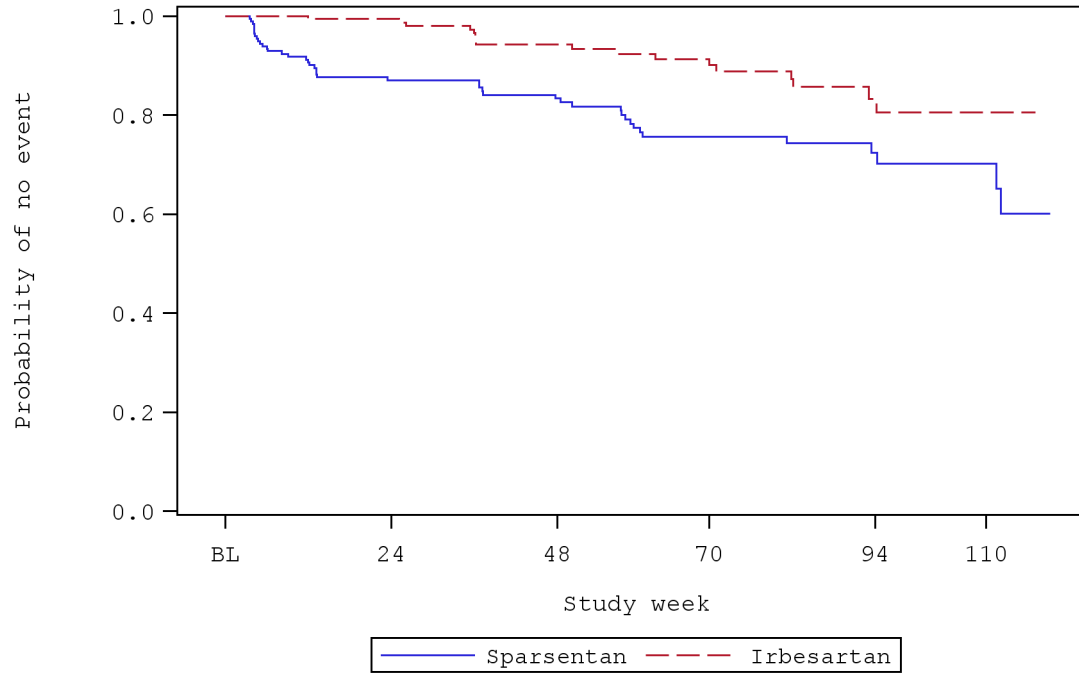
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

Complete remission means an urinary protein excretion of <0.3 g/day.

Source Data: alb\_tte, created on: 31MAY2024

Figure PF1URFT\_FMK0: Time to complete remission - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	134	110	78	36	16
Irbesartan	202	152	113	78	32	12

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 Reference table: PT1URFT\_FMT0

Table PT1GGC\_FMH0: Course of eGFR  
 Full Analysis Set

Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR	Baseline	Sparsentan	202	202 (100.0)	56.86 (24.38)	24.0	38.00	50.00	71.00	128.0		
		Irbesartan	202	202 (100.0)	57.08 (23.58)	26.0	39.00	50.00	70.00	123.0		
	Week 6	Sparsentan	202	193 (95.5)	55.92 (23.65)	21.0	36.00	52.00	70.00	121.0		
		Irbesartan	202	195 (96.5)	55.58 (23.39)	19.0	38.00	48.00	69.00	120.0		
	Week 36	Sparsentan	202	136 (67.3)	54.24 (23.12)	18.0	36.00	48.50	67.50	123.0		
		Irbesartan	202	133 (65.8)	50.59 (22.84)	7.0	34.00	44.00	62.00	125.0		
	Week 58	Sparsentan	202	110 (54.5)	53.73 (24.31)	13.0	36.00	47.00	69.00	133.0		
		Irbesartan	202	95 (47.0)	48.57 (22.12)	13.0	33.00	41.00	60.00	122.0		
	Change from baseline in eGFR	Week 6	Sparsentan	202	193 (95.5)	-1.13 (7.69)	-43.0	-4.00	-1.00	2.00	20.0	0.06 [-0.14, 0.25]
			Irbesartan	202	195 (96.5)	-1.53 (6.56)	-31.0	-5.00	-2.00	2.00	20.0	
		Week 36	Sparsentan	202	136 (67.3)	-3.84 (8.69)	-39.0	-8.00	-3.00	1.00	26.0	0.08 [-0.16, 0.32]
			Irbesartan	202	133 (65.8)	-4.49 (7.70)	-32.0	-8.00	-4.00	0.00	15.0	
Week 58		Sparsentan	202	110 (54.5)	-4.89 (8.95)	-34.0	-9.00	-4.00	0.00	24.0	0.16 [-0.11, 0.44]	
		Irbesartan	202	95 (47.0)	-6.27 (7.88)	-24.0	-12.00	-7.00	0.00	12.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.  
 A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m<sup>2</sup>. eGFR = estimated glomerular filtration rate.  
 Source Data: alb, created on: 19FEB2024



Table PT1GGC\_FMC0: Change from baseline in eGFR  
Full Analysis Set

Change from baseline in eGFR				Repeated measures analysis					
Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Week 6	Sparsentan	202	193 (95.5)	-1.12 (0.52)	(-2.13, -0.11)	0.38 (0.73)	(-1.04, 1.81)	0.598	
	Irbesartan	202	195 (96.5)	-1.50 (0.51)	(-2.51, -0.50)				
Week 36	Sparsentan	202	136 (67.3)	-3.70 (0.59)	(-4.87, -2.53)	0.85 (0.85)	(-0.81, 2.51)	0.315	
	Irbesartan	202	133 (65.8)	-4.55 (0.60)	(-5.73, -3.37)				
Week 58	Sparsentan	202	110 (54.5)	-4.61 (0.66)	(-5.91, -3.31)	1.55 (0.97)	(-0.35, 3.45)	0.110	
	Irbesartan	202	95 (47.0)	-6.16 (0.70)	(-7.54, -4.78)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. Only key visits up to Week 58 are displayed.

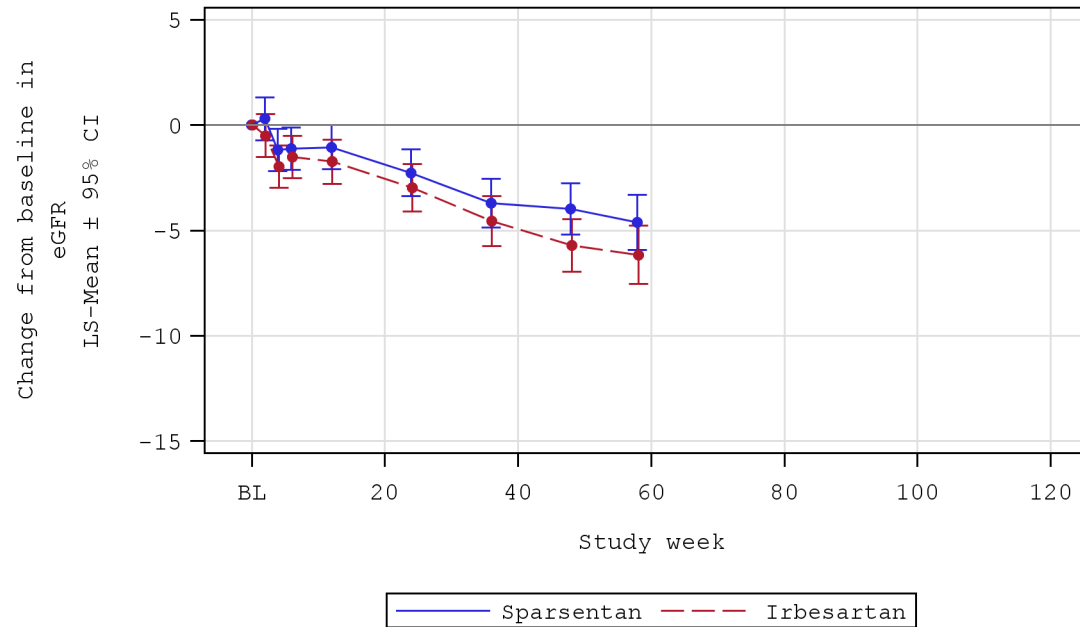
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

All observed values up to Week 58 are included in the model following a treatment policy approach.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m<sup>2</sup>. eGFR = estimated glomerular filtration rate. A first order regressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Figure PF1GGC\_FMG0: Change from baseline in eGFR  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	193	136	110
Irbesartan	195	133	95

Number of available records are presented for key visits up to Week 58 only. MMRM = Mixed model Repeated Measures. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. All observed values up to Week 58 are included in the model. A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m<sup>2</sup>. Reference table: PT1GGC\_FMC0

Table PT1GGA\_FMR0: Rate of change in eGFR (total slope) - Year 1  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1		Mixed Random Coefficient Model							
		Time	Treatment	N	Annualized Slope		Slope Difference		p-value
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline to Week 58	Sparsentan	202	-4.26 (0.89)	(-6.01, -2.52)	0.62 (1.27)	(-1.88, 3.11)	0.629		
	Irbesartan	202	-4.88 (0.91)	(-6.67, -3.09)					

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. MRCM = Mixed Random Coefficient Model.  
 An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject.  
 All observed post-baseline values up to Week 58 are included in the model. A first order autoregressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate.  
 Source Data: alb, created on: 19FEB2024

Table PT1GGLA\_FMR0: Rate of change in eGFR (chronic slope) - Year 1  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1		Mixed Random Coefficient Model					
		N	Annualized Slope		Slope Difference		
			LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time	Treatment						
Week 6 to Week 58	Sparsentan	202	-3.92 (0.94)	(-5.76, -2.08)	1.00 (1.34)	(-1.64, 3.63)	0.457
	Irbesartan	202	-4.92 (0.96)	(-6.80, -3.03)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. MRCM = Mixed Random Coefficient Model.  
 An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject.  
 All observed values up from Week 6 to Week 58 are included in the model. A first order autoregressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate.  
 Source Data: alb, created on: 19FEB2024

Table PT1MIS\_FMI0: Patients with systemic immunosuppressive medication  
 Full Analysis Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Systemic immunosuppressive medication	Double-blind period	Sparsentan	202	14 (6.9)	0.700 [0.364, 1.347]	0.678 [0.332, 1.382]	-3.0 [-8.9, 2.9]	0.370
		Irbesartan	202	20 (9.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aeff, created on: 28FEB2024

Table PT1MIST\_FMT0: Time to systemic immunosuppressive medication  
 Full Analysis Set

Time to systemic immunosuppressive medication			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	14 (6.9)	NE		0.661	(0.334, 1.309)	0.235
Irbesartan	202	20 (9.9)	NE				

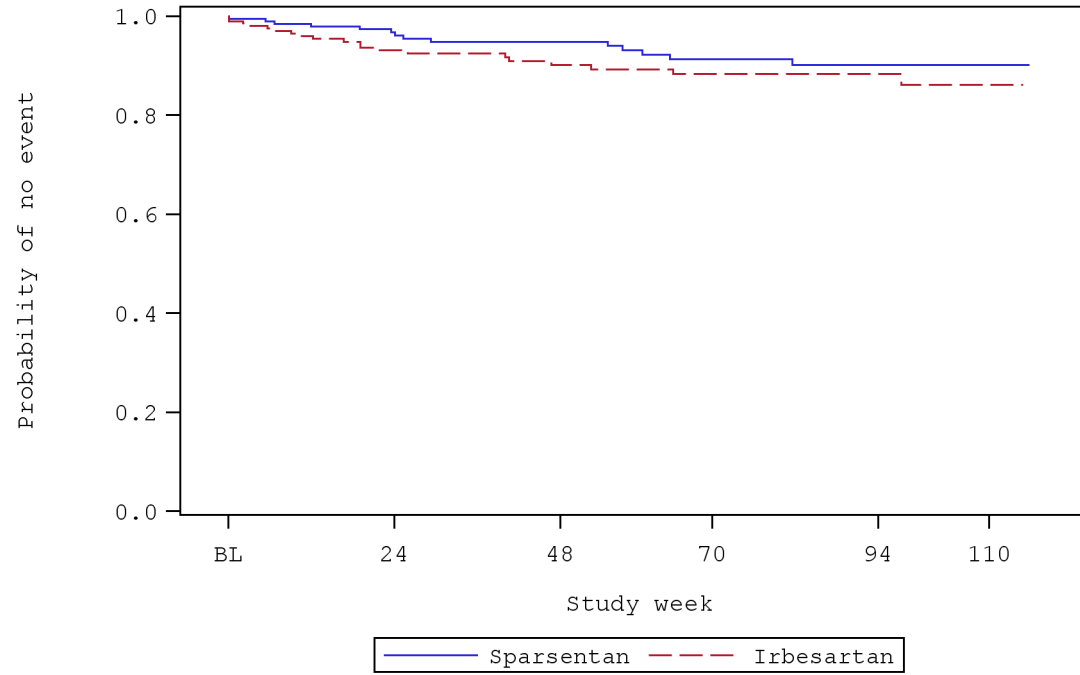
N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

Source Data: aeff\_tte, created on: 28FEB2024

Figure PF1MIST\_FMK0: Time to systemic immunosuppressive medication - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	156	121	97	50	16
Irbesartan	202	149	110	88	46	17

Reference table: PT1MIST\_FMT0

Table PT1MIK\_FMI0: Patients with systemic immunosuppressive renal medication  
 Full Analysis Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Systemic immunosuppressive renal medication (renal indication)	Double-blind period	Sparsentan	202	2 (1.0)	0.222 [0.049, 1.016]	0.214 [0.046, 1.005]	-3.5 [-7.1, 0.2]	0.062
		Irbesartan	202	9 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aeff, created on: 28FEB2024



Table PT1MIKT\_FMT0: Time to systemic immunosuppressive renal medication  
 Full Analysis Set

Time to systemic Immunosuppressive renal medication			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	2 (1.0)	NE		0.202	(0.044, 0.937)	0.041 *
Irbesartan	202	9 (4.5)	NE				

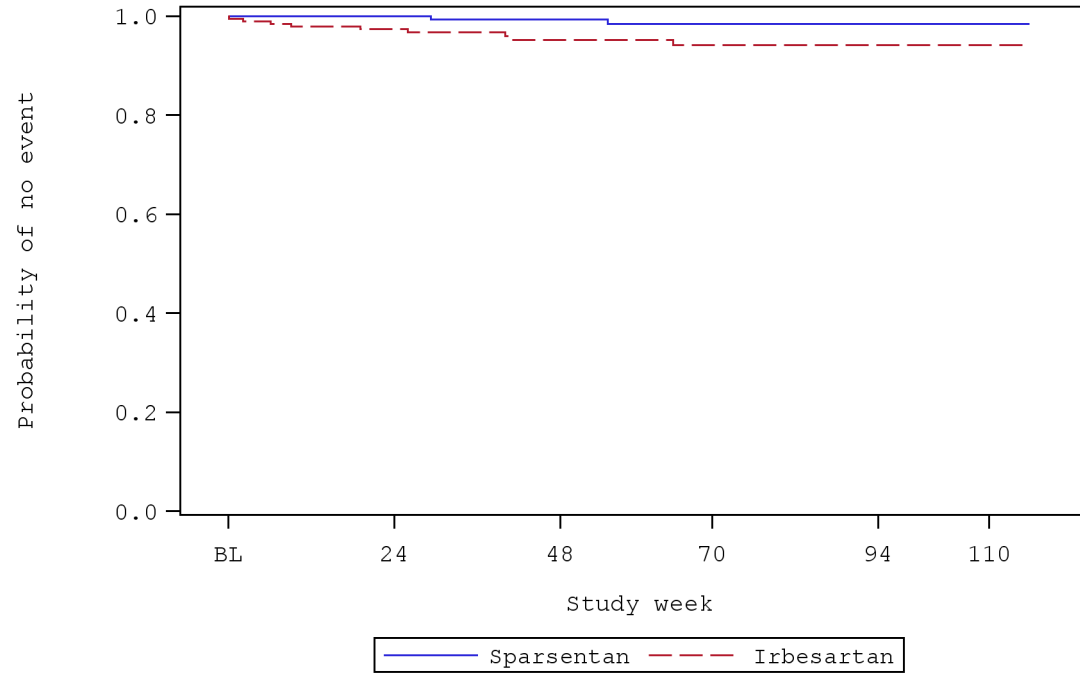
N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

Source Data: aeff\_tte, created on: 28FEB2024

Figure PF1MIKT\_FMK0: Time to systemic immunosuppressive renal medication - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	161	127	105	54	18
Irbesartan	202	154	114	92	48	19

Reference table: PT1MIKT\_FMT0

Table PT1HOS\_FMI0: Patients with hospitalizations  
 Full Analysis Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Hospitalizations	Double-blind period	Sparsentan	202	22 (10.9)	1.000 [0.572, 1.747]	1.000 [0.535, 1.870]	0.0 [-6.6, 6.6]	1.000
		Irbesartan	202	22 (10.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aeff, created on: 28FEB2024

Table PT1HNS\_FMNO: Number of hospitalizations  
 Full Analysis Set

Number of hospitalizations			Time at risk (years)	Number of events	Crude rate	Negative binomial model		
Treatment	N	nval (%)				Rate ratio	95% CI	p-value
Sparsentan	202	202 (100.0)	265.8	24	0.09	0.924	(0.520, 1.639)	0.786
Irbesartan	202	202 (100.0)	262.9	26	0.10			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
 NE = not evaluable.  
 Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
 A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset.  
 \* = significant treatment effect.  
 Source Data: aeff, created on: 28FEB2024

Table PT1HDS\_FMNO: Duration of hospitalizations  
 Full Analysis Set

Duration of hospitalizations			Negative binomial model					
Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Sparsentan	202	202 (100.0)	265.8	105	0.40	0.799	(0.307, 2.082)	0.647
Irbesartan	202	202 (100.0)	262.9	178	0.68			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
 NE = not evaluable.  
 Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
 A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset.  
 \* = significant treatment effect.  
 Source Data: aeff, created on: 28FEB2024

Table PT1HDS\_FMB0: Duration of hospitalizations  
 Full Analysis Set

Variable	Study	Treatment	N	n	Mean (SD)	Min	Q25	Q50	Q75	Max
Duration of hospitalizations	PROTECT	Sparsentan	202	202	0.52 (2.01)	0.0	0.00	0.00	0.00	18.0
		Irbesartan	202	202	0.88 (3.32)	0.0	0.00	0.00	0.00	30.0

N = total number of patients in analysis set. n = number evaluable values. SD = standard deviation. Q = quantile.  
 Results are presented in days.  
 Source Data: aeff, created on: 28FEB2024

Table PT1VSC\_FMH0: Course of EQ-5D VAS  
 Full Analysis Set

Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
EQ-5D VAS	Baseline	Sparsentan	202	191 (94.6)	80.48 (14.24)	7.0	75.00	81.00	90.00	100.0	
		Irbesartan	202	184 (91.1)	80.36 (14.21)	19.0	75.00	80.00	90.00	100.0	
	Week 24	Sparsentan	202	132 (65.3)	83.20 (10.84)	50.0	76.00	83.50	91.00	100.0	
		Irbesartan	202	115 (56.9)	80.93 (16.23)	4.0	77.00	81.00	91.00	100.0	
	Week 48	Sparsentan	202	113 (55.9)	82.11 (10.59)	49.0	79.00	82.00	90.00	100.0	
		Irbesartan	202	83 (41.1)	82.77 (12.72)	40.0	80.00	83.00	90.00	100.0	
Change from baseline Week 24 in EQ-5D VAS	Week 24	Sparsentan	202	132 (65.3)	3.20 (15.02)	-31.0	-4.50	1.00	10.00	75.0	0.16 [-0.09, 0.41]
		Irbesartan	202	115 (56.9)	0.57 (17.19)	-90.0	-5.00	0.00	10.00	52.0	
	Week 48	Sparsentan	202	113 (55.9)	2.84 (15.56)	-30.0	-4.00	1.00	9.00	87.0	0.06 [-0.23, 0.34]
		Irbesartan	202	83 (41.1)	2.07 (10.72)	-35.0	-1.00	2.00	9.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FMC0: Change from baseline in EQ-5D VAS  
 Full Analysis Set

Change from baseline in EQ-5D VAS				Repeated measures analysis					
Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Week 24	Sparsentan	202	132 (65.3)	3.19 (1.03)	(1.16, 5.21)	2.41 (1.51)	(-0.56, 5.38)	0.112	
	Irbesartan	202	115 (56.9)	0.78 (1.11)	(-1.40, 2.95)				
Week 48	Sparsentan	202	113 (55.9)	2.20 (1.11)	(0.02, 4.38)	-0.30 (1.70)	(-3.65, 3.04)	0.859	
	Irbesartan	202	83 (41.1)	2.50 (1.29)	(-0.03, 5.04)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

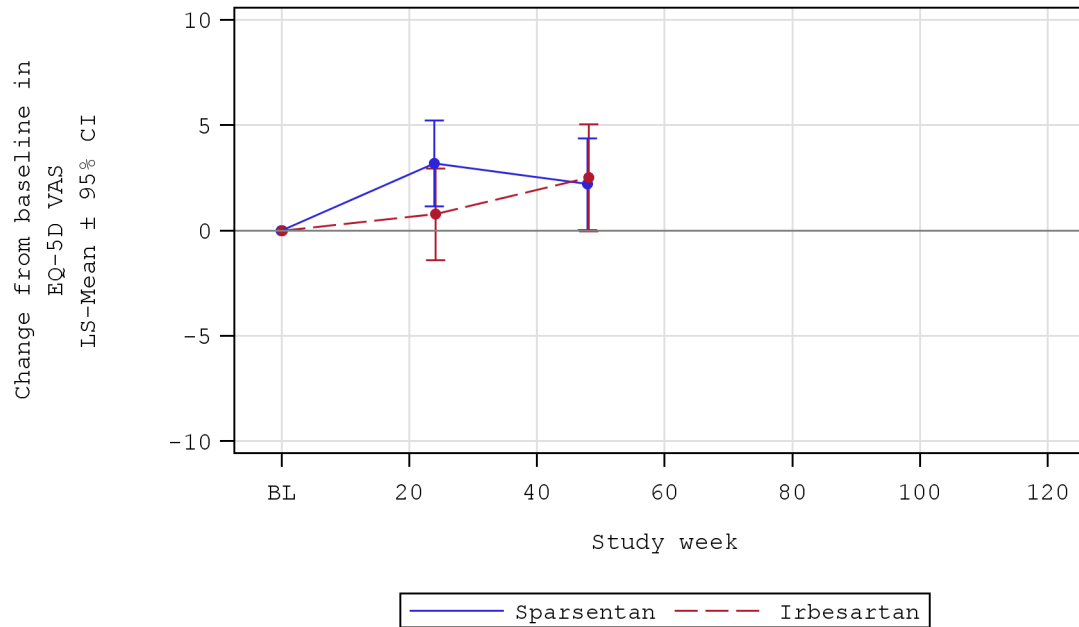
A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 04APR2024



Figure PF1VSC\_FMG0: Change from baseline in EQ-5D VAS  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	132	113
Irbesartan	115	83

Number of available records are presented per time-point. LS-Mean = Least square mean. 95% CI = 95% confidence interval.  
 A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1VSC\_FMG0

Table PT1VSIT\_FMT0: Time to increase in EQ-5D VAS by at least 15 points  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	29 (14.4)	NE		0.936	(0.539, 1.625)	0.813
Irbesartan	202	26 (12.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

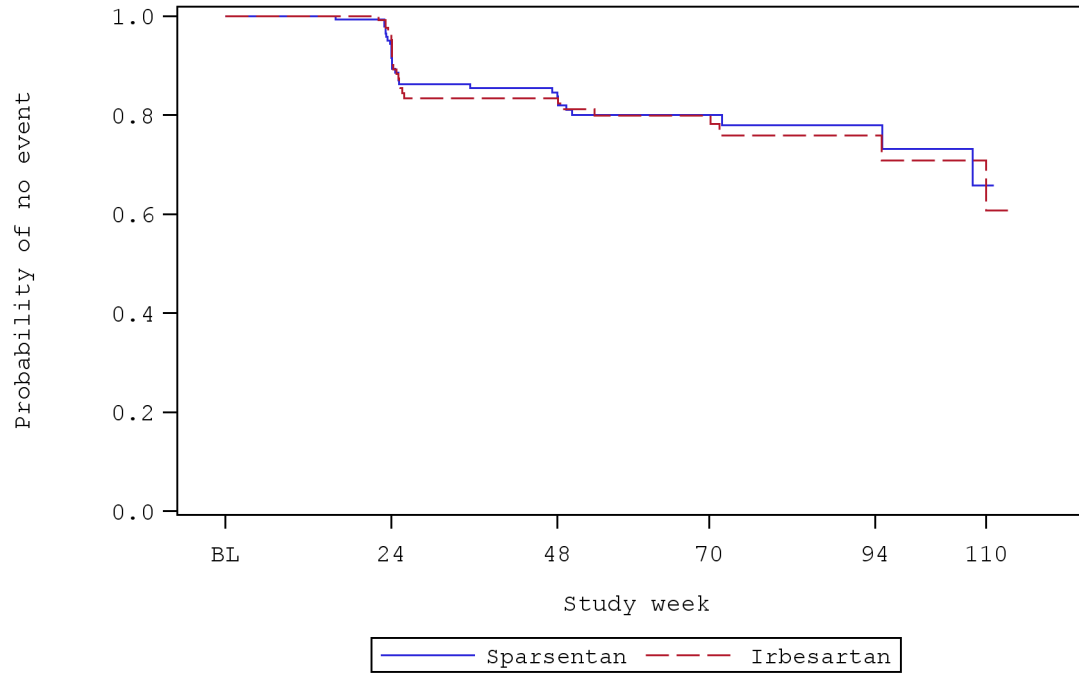
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

An increase reflects an improvement of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024

Figure PF1VSIT\_FMK0: Time to increase in EQ-5D VAS by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	133	98	69	29	8
Irbesartan	202	118	80	50	20	7

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1VSIT\_FMT0

Table PT1VSDT\_FMT0: Time to decrease in EQ-5D VAS by at least 15 points  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	26 (12.9)	112.1	(111.0, NE)	1.242	(0.660, 2.338)	0.502
Irbesartan	202	16 (7.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

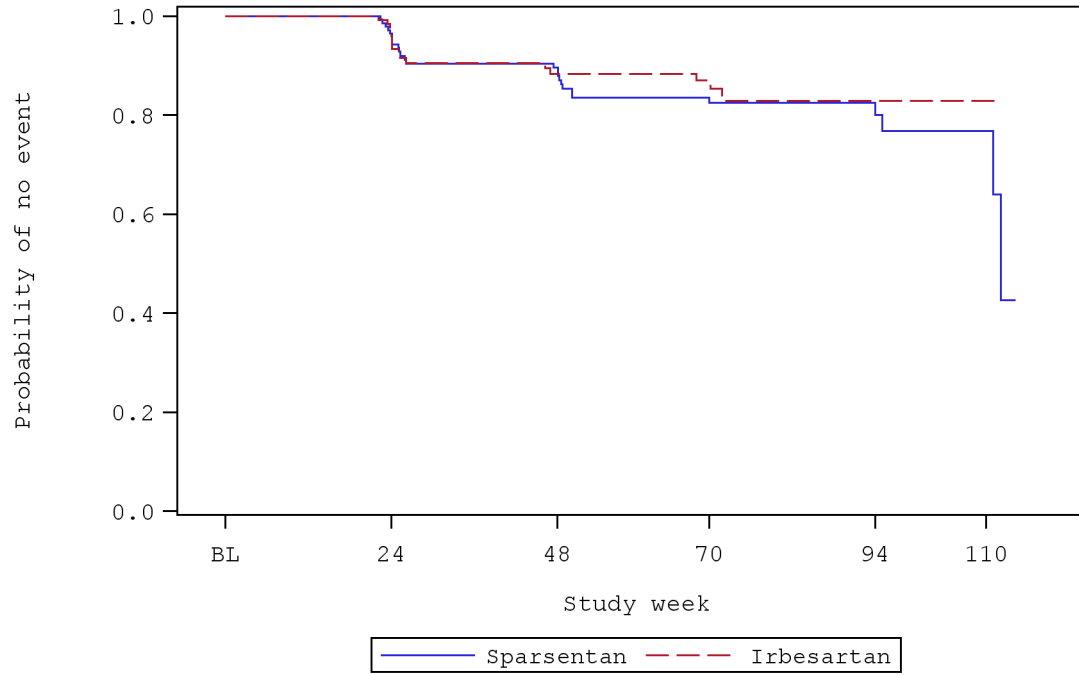
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024

Figure PF1VSDT\_FMK0: Time to decrease in EQ-5D VAS by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	137	109	79	35	13
Irbesartan	202	119	82	55	21	8

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1VSDT\_FMT0

Table PT1KBUC\_FMH0: KDQOL: Course of burden of kidney disease  
 Full Analysis Set

Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
KDQOL: burden of kidney disease	Baseline	Sparsentan	202	186 (92.1)	69.52 (25.46)	0.0	50.00	75.00	87.50	100.0		
		Irbesartan	202	179 (88.6)	76.85 (22.80)	6.3	62.50	81.25	93.75	100.0		
	Week 24	Sparsentan	202	125 (61.9)	75.55 (22.00)	12.5	68.75	81.25	93.75	100.0		
		Irbesartan	202	104 (51.5)	78.85 (22.14)	0.0	68.75	81.25	100.00	100.0		
	Week 48	Sparsentan	202	110 (54.5)	77.33 (20.92)	6.3	68.75	81.25	93.75	100.0		
		Irbesartan	202	77 (38.1)	80.76 (20.27)	6.3	75.00	81.25	100.00	100.0		
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	202	125 (61.9)	7.75 (22.94)	-81.3	0.00	6.25	18.75	75.0	0.38 [0.11, 0.64]
			Irbesartan	202	104 (51.5)	-0.06 (17.73)	-50.0	-12.50	0.00	6.25	56.3	
Week 48		Sparsentan	202	110 (54.5)	9.26 (20.44)	-62.5	0.00	6.25	25.00	62.5	0.34 [0.04, 0.63]	
		Irbesartan	202	77 (38.1)	2.84 (17.10)	-50.0	-6.25	0.00	12.50	56.3		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FMC0: KDQOL: Change from baseline in burden of kidney disease  
 Full Analysis Set

				Repeated measures analysis				
KDQOL: Change from baseline in burden of kidney disease				Change from Baseline		Treatment Difference		
Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Week 24	Sparsentan	202	125 (61.9)	5.48 (1.50)	(2.54, 8.42)	2.79 (2.24)	(-1.62, 7.21)	0.214
	Irbesartan	202	104 (51.5)	2.69 (1.65)	(-0.56, 5.93)			
Week 48	Sparsentan	202	110 (54.5)	7.31 (1.58)	(4.20, 10.42)	2.86 (2.48)	(-2.00, 7.73)	0.248
	Irbesartan	202	77 (38.1)	4.45 (1.89)	(0.74, 8.16)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

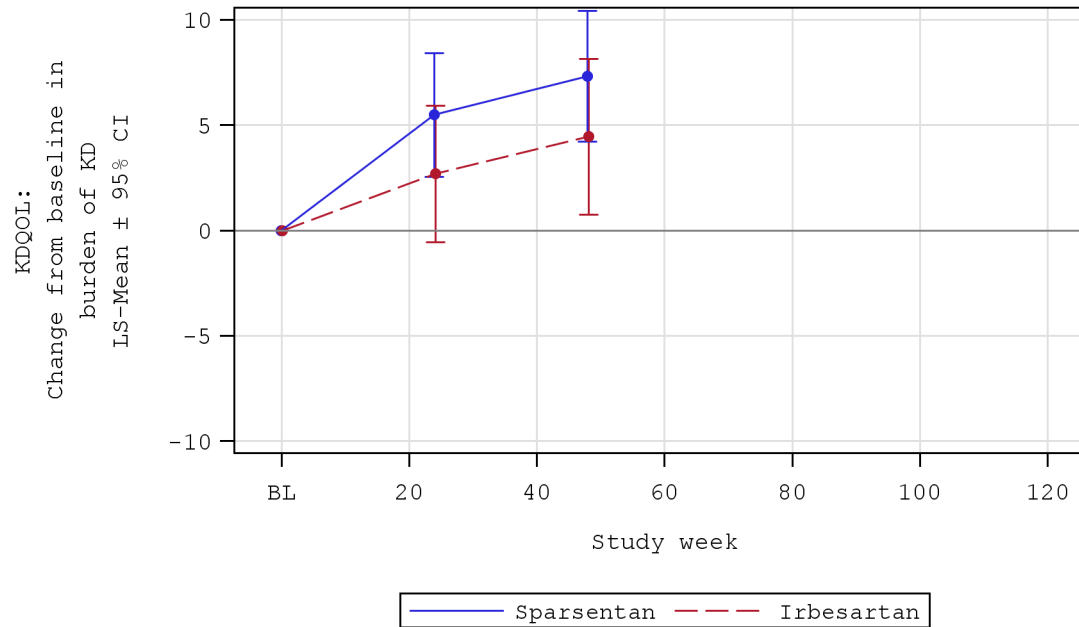
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 04APR2024

Figure PF1KBUC\_FMG0: KDQOL: Change from baseline in burden of kidney disease  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	125	110
Irbesartan	104	77

Number of available records are presented per time-point. LS-Mean = Least square mean. 95% CI = 95% confidence interval.  
 A decrease reflects a worsening of the status of the patient.  
 KD = kidney disease. Reference table: PT1KBUC\_FMG0



Table PT1KBUIT\_FMT0: KDQOL: Time to increase in burden of kidney disease by at least 15 points  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	61 (30.2)	75.3	(50.1, NE)	1.594	(1.013, 2.508)	0.044 *
Irbesartan	202	29 (14.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

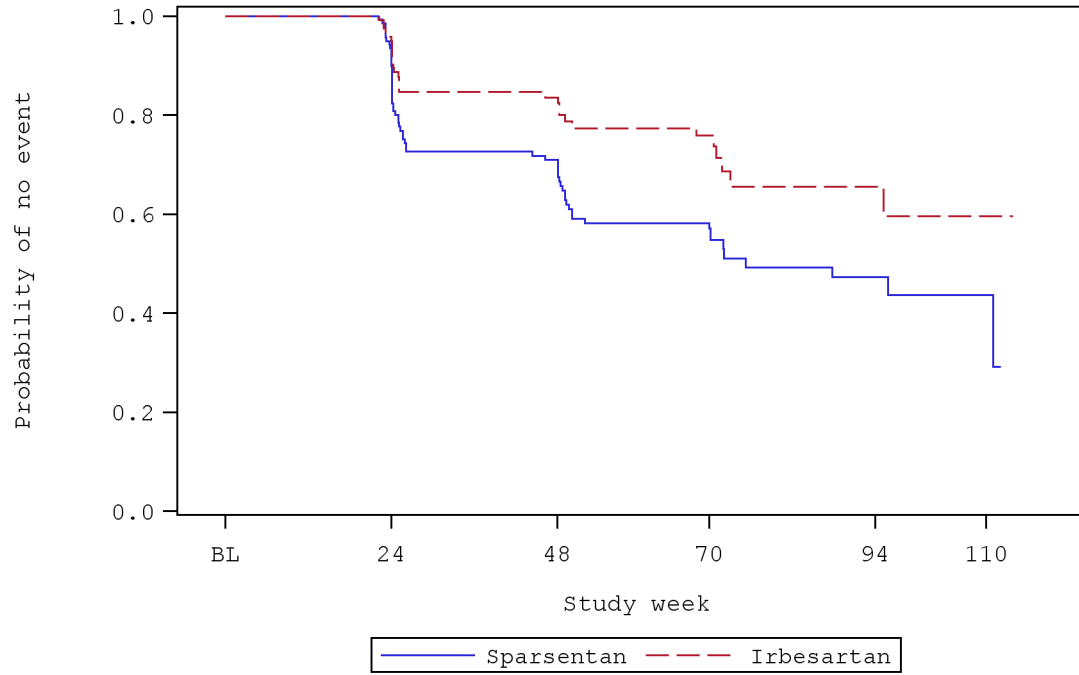
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

An increase reflects an improvement of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KBUIT\_FMK0: KDQOL: Time to increase in burden of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	127	81	56	21	10
Irbesartan	202	111	75	43	15	5

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KBUIT\_FMT0

Table PT1KBUDT\_FMT0: KDQOL: Time to decrease in burden of kidney disease by at least 15 points  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	25 (12.4)	112.1	(112.1, NE)	0.651	(0.382, 1.109)	0.115
Irbesartan	202	33 (16.3)	96.1	(94.6, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

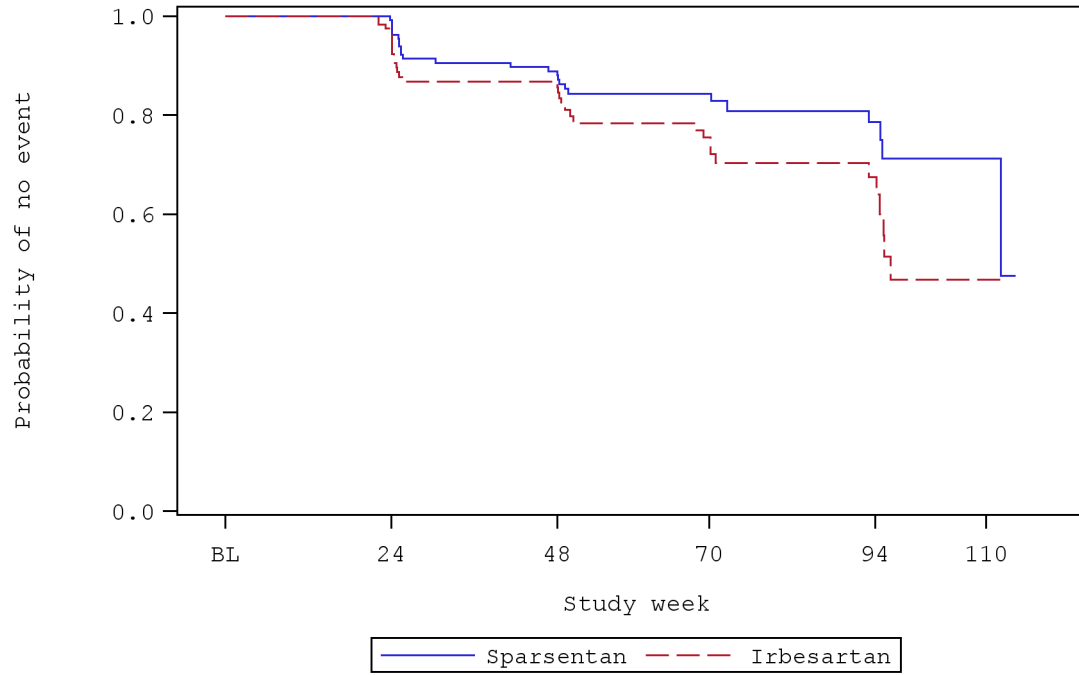
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KBUDT\_FMK0: KDQOL: Time to decrease in burden of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	135	104	73	32	11
Irbesartan	202	114	82	47	20	7

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KBUDT\_FMT0

Table PT1KEFC\_FMHO: KDQOL: Course of effect of kidney disease  
 Full Analysis Set

Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
KDQOL: effect of kidney disease	Baseline	Sparsentan	202	186 (92.1)	88.21 (13.70)	28.1	84.38	93.75	96.88	100.0		
		Irbesartan	202	179 (88.6)	88.88 (13.03)	37.5	84.38	93.75	100.00	100.0		
	Week 24	Sparsentan	202	125 (61.9)	89.68 (13.42)	9.4	87.50	93.75	100.00	100.0		
		Irbesartan	202	104 (51.5)	88.67 (16.94)	0.0	84.38	93.75	100.00	100.0		
	Week 48	Sparsentan	202	110 (54.5)	90.23 (13.12)	0.0	87.50	93.75	100.00	100.0		
		Irbesartan	202	77 (38.1)	92.78 (10.80)	40.6	90.63	96.88	100.00	100.0		
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	202	125 (61.9)	1.58 (13.88)	-75.0	-3.13	0.00	6.25	34.4	0.23 [-0.03, 0.50]
			Irbesartan	202	104 (51.5)	-1.77 (14.75)	-100.0	-3.13	0.00	4.69	31.3	
Week 48		Sparsentan	202	110 (54.5)	2.07 (15.78)	-100.0	-3.13	3.13	6.25	56.3	0.05 [-0.24, 0.35]	
		Irbesartan	202	77 (38.1)	1.34 (9.17)	-25.0	0.00	0.00	3.13	25.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FMC0: KDQOL: Change from baseline in effect of kidney disease  
 Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease				Repeated measures analysis				
				Change from Baseline		Treatment Difference		
Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Week 24	Sparsentan	202	125 (61.9)	1.15 (1.13)	(-1.08, 3.37)	2.29 (1.68)	(-1.02, 5.61)	0.175
	Irbesartan	202	104 (51.5)	-1.15 (1.25)	(-3.59, 1.30)			
Week 48	Sparsentan	202	110 (54.5)	0.99 (1.20)	(-1.36, 3.34)	0.02 (1.86)	(-3.64, 3.68)	0.991
	Irbesartan	202	77 (38.1)	0.97 (1.42)	(-1.83, 3.77)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

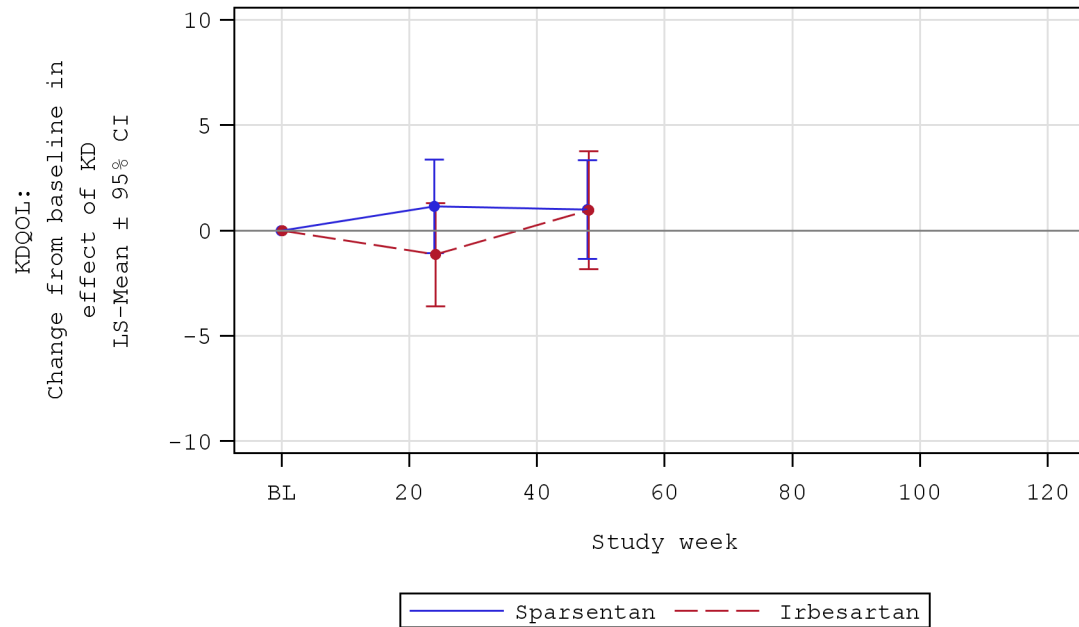
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 04APR2024

Figure PF1KEFC\_FMG0: KDQOL: Change from baseline in effect of kidney disease  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	125	110
Irbesartan	104	77

Number of available records are presented per time-point. LS-Mean = Least square mean. 95% CI = 95% confidence interval.  
 A decrease reflects a worsening of the status of the patient.  
 KD = kidney disease. Reference table: PT1KEFC\_FMG0

Table PT1KEFIT\_FMT0: KDQOL: Time to increase in effect of kidney disease by at least 15 points  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	23 (11.4)	NE		1.192	(0.589, 2.413)	0.625
Irbesartan	202	14 (6.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

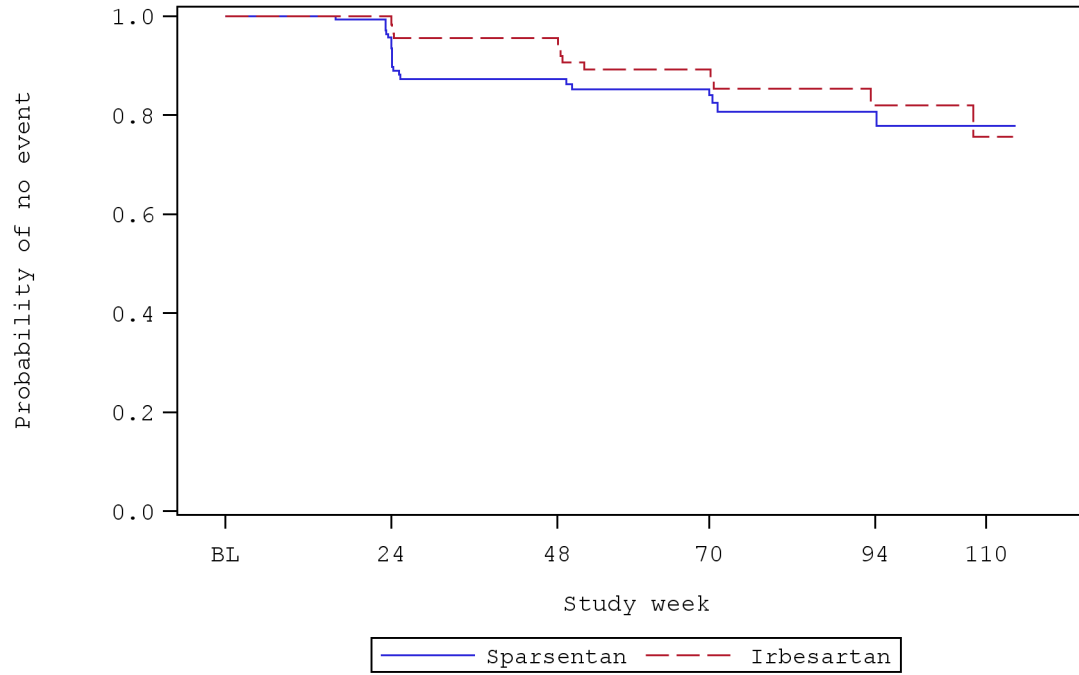
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

An increase reflects an improvement of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024



Figure PF1KEFIT\_FMK0: KDQOL: Time to increase in effect of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	131	99	70	29	12
Irbesartan	202	115	85	51	21	9

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FMT0

Table PT1KEFDT\_FMT0: KDQOL: Time to decrease in effect of kidney disease by at least 15 points  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	18 (8.9)	NE		0.628	(0.329, 1.196)	0.157
Irbesartan	202	20 (9.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

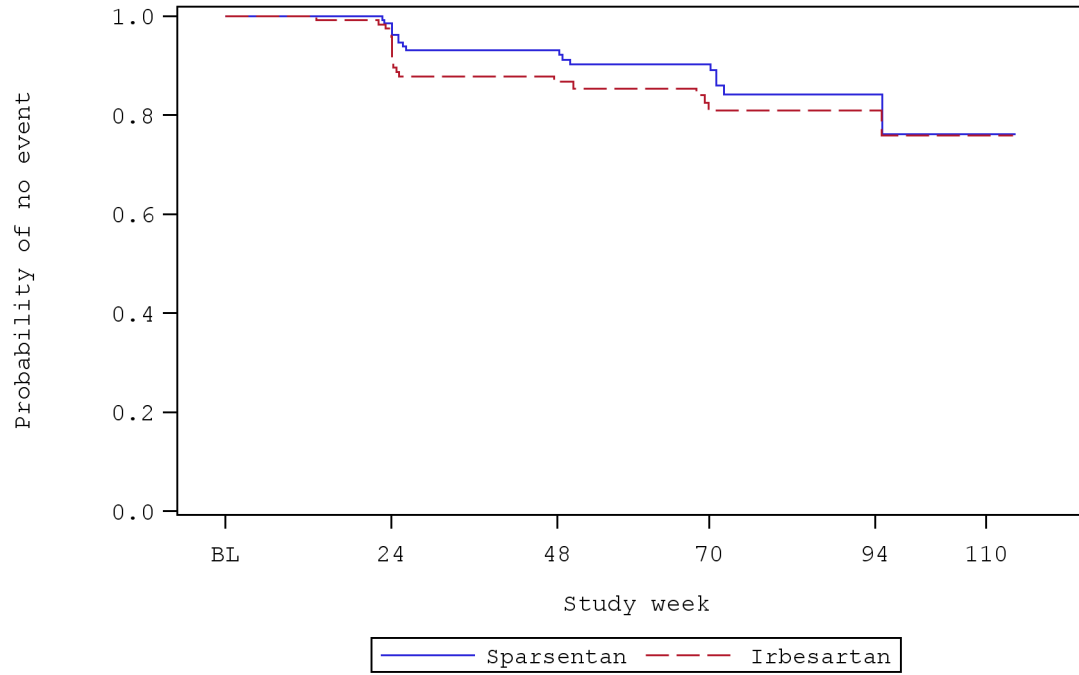
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KEFDT\_FMK0: KDQOL: Time to decrease in effect of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	133	109	81	34	11
Irbesartan	202	114	82	51	21	8

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KEFDT\_FMT0

Table PT1KSYC\_FMH0: KDQOL: Course of symptom/problems of kidney disease  
 Full Analysis Set

Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	202	186 (92.1)	88.84 (11.24)	31.8	84.09	90.91	97.73	100.0		
		Irbesartan	202	179 (88.6)	89.36 (11.04)	38.6	86.36	93.18	97.73	100.0		
	Week 24	Sparsentan	202	125 (61.9)	88.53 (12.87)	4.5	84.09	93.18	97.73	100.0		
		Irbesartan	202	104 (51.5)	87.22 (17.74)	4.5	84.09	93.18	97.73	100.0		
	Week 48	Sparsentan	202	110 (54.5)	88.02 (15.92)	0.0	81.82	93.18	97.73	100.0		
		Irbesartan	202	77 (38.1)	89.64 (12.52)	47.7	86.36	93.18	100.00	100.0		
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	202	125 (61.9)	0.16 (12.59)	-72.7	-4.55	0.00	4.55	63.6	0.17 [-0.09, 0.44]
			Irbesartan	202	104 (51.5)	-2.43 (17.14)	-90.9	-4.55	0.00	2.27	52.3	
Week 48		Sparsentan	202	110 (54.5)	0.04 (17.54)	-100.0	-4.55	0.00	6.82	59.1	0.04 [-0.25, 0.33]	
		Irbesartan	202	77 (38.1)	-0.53 (9.48)	-27.3	-4.55	0.00	2.27	34.1		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FMC0: KDQOL: Change from baseline in symptom/problems of kidney disease  
 Full Analysis Set

				Repeated measures analysis					
KDQOL: Change from baseline in symptom/problems of kidney disease				Change from Baseline		Treatment Difference			
Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Week 24	Sparsentan	202	125 (61.9)	-0.25 (1.23)	(-2.67, 2.17)	1.80 (1.83)	(-1.80, 5.39)	0.326	
	Irbesartan	202	104 (51.5)	-2.05 (1.35)	(-4.71, 0.61)				
Week 48	Sparsentan	202	110 (54.5)	-0.50 (1.31)	(-3.08, 2.07)	0.02 (2.05)	(-4.00, 4.04)	0.993	
	Irbesartan	202	77 (38.1)	-0.52 (1.57)	(-3.60, 2.56)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

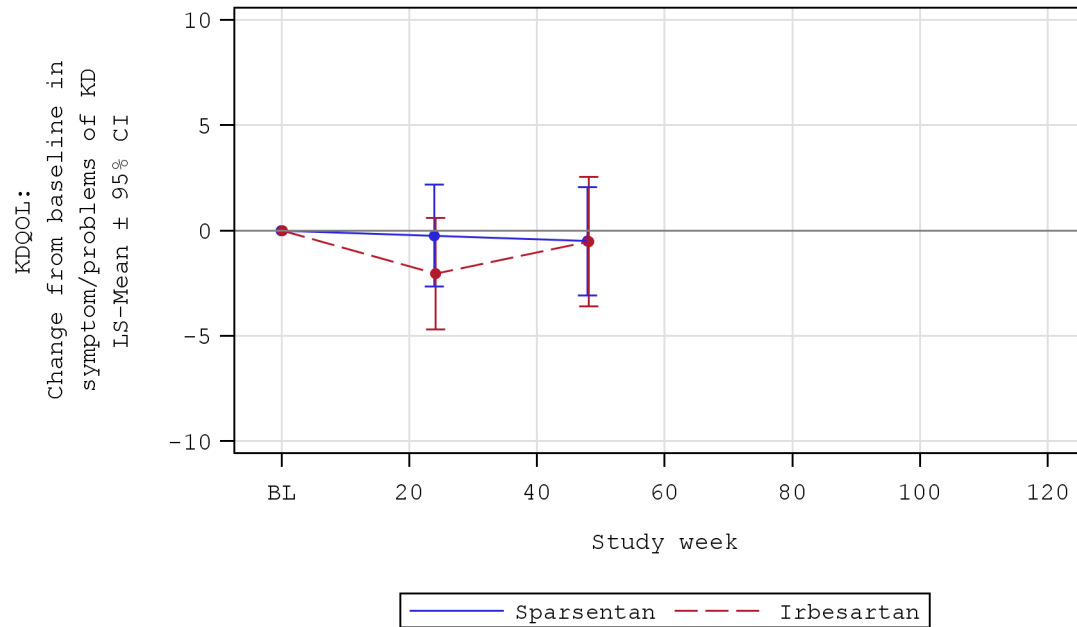
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 04APR2024

Figure PF1KSYC\_FMG0: KDQOL: Change from baseline in symptom/problems of kidney disease  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	125	110
Irbesartan	104	77

Number of available records are presented per time-point. LS-Mean = Least square mean. 95% CI = 95% confidence interval.  
 A decrease reflects a worsening of the status of the patient.  
 KD = kidney disease. Reference table: PT1KSYC\_FMC0

Table PT1KSYIT\_FMT0: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	15 (7.4)	NE		1.237	(0.504, 3.037)	0.643
Irbesartan	202	9 (4.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

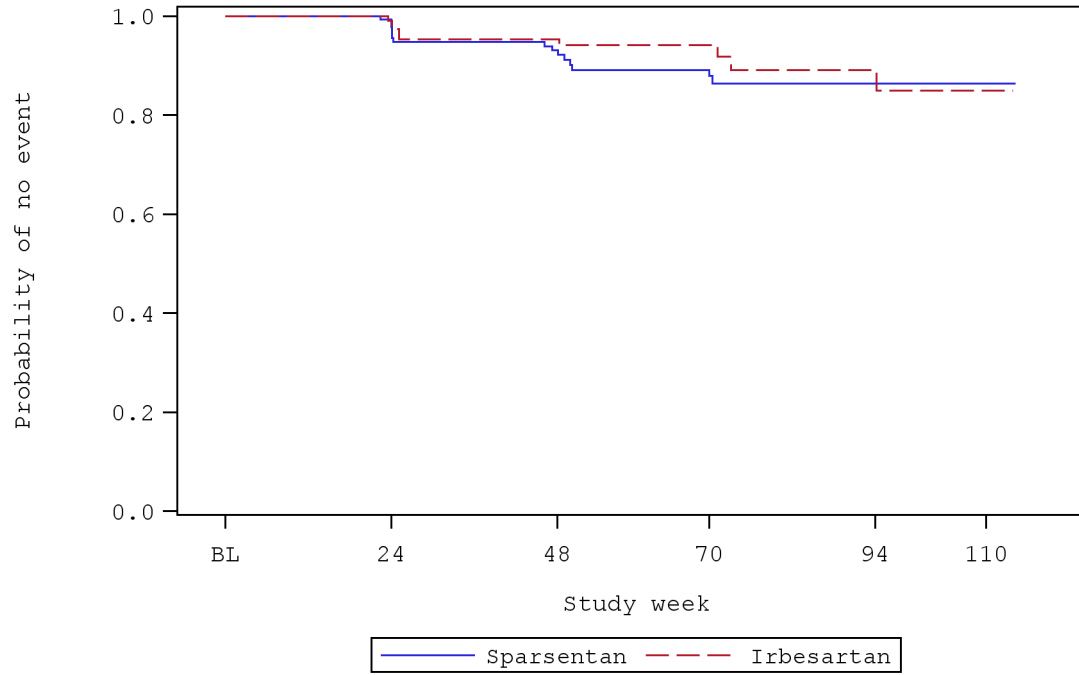
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

An increase reflects an improvement of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KSYIT\_FMK0: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	135	106	74	31	12
Irbesartan	202	114	85	54	23	8

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KSYIT\_FMT0



Table PT1KSYDT\_FMT0: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	23 (11.4)	NE		0.967	(0.519, 1.804)	0.916
Irbesartan	202	18 (8.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

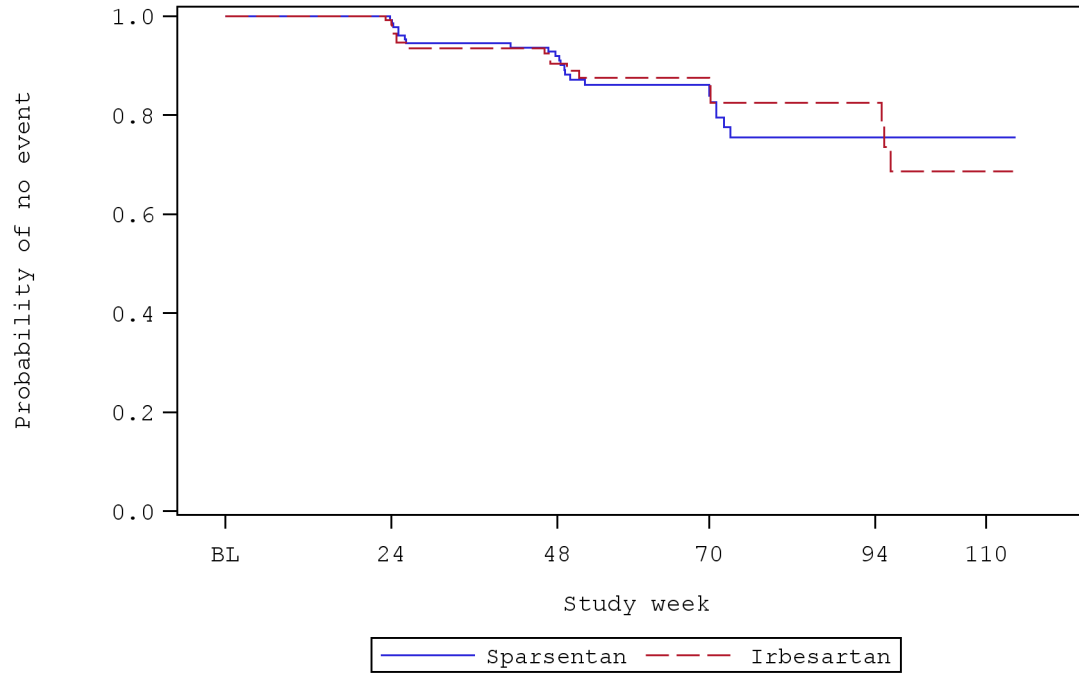
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KSYDT\_FMK0: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	135	107	75	29	10
Irbesartan	202	115	83	53	24	10

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KSYDT\_FMT0

Table PT1KPSC\_FMH0: KDQOL-SF12: Course of PCS  
 Full Analysis Set

Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
KDQOL-SF12: PCS	Baseline	Sparsentan	202	187 (92.6)	51.05 (8.10)	12.4	46.73	53.18	56.71	63.5	
		Irbesartan	202	184 (91.1)	51.70 (7.02)	21.6	49.37	53.46	56.15	64.3	
	Week 24	Sparsentan	202	128 (63.4)	51.04 (7.48)	16.3	47.08	52.67	56.36	62.9	
		Irbesartan	202	112 (55.4)	51.32 (7.20)	20.8	46.64	53.46	56.15	64.3	
	Week 48	Sparsentan	202	112 (55.4)	51.01 (6.96)	29.8	46.60	53.17	55.96	66.1	
		Irbesartan	202	80 (39.6)	51.58 (7.19)	32.8	46.78	54.53	56.59	64.5	
KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	202	128 (63.4)	0.38 (6.78)	-24.8	-3.59	0.07	3.92	20.4	0.09 [-0.16, 0.34]
		Irbesartan	202	112 (55.4)	-0.20 (5.70)	-20.0	-2.27	0.00	2.37	16.9	
	Week 48	Sparsentan	202	112 (55.4)	0.53 (7.31)	-22.3	-3.19	0.00	4.34	24.7	0.09 [-0.19, 0.38]
		Irbesartan	202	80 (39.6)	-0.17 (7.71)	-24.7	-4.06	0.14	3.01	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FMC0: KDQOL-SF12: Change from baseline in PCS  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS				Repeated measures analysis				
				Change from Baseline		Treatment Difference		
				Time	Treatment	N	n (%)	LS-Mean (STE)
Week 24	Sparsentan	202	128 (63.4)	0.35 (0.52)	(-0.68, 1.37)	0.29 (0.77)	(-1.22, 1.80)	0.704
	Irbesartan	202	112 (55.4)	0.06 (0.56)	(-1.05, 1.16)			
Week 48	Sparsentan	202	112 (55.4)	0.16 (0.55)	(-0.92, 1.24)	0.24 (0.85)	(-1.43, 1.91)	0.781
	Irbesartan	202	80 (39.6)	-0.08 (0.65)	(-1.35, 1.19)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

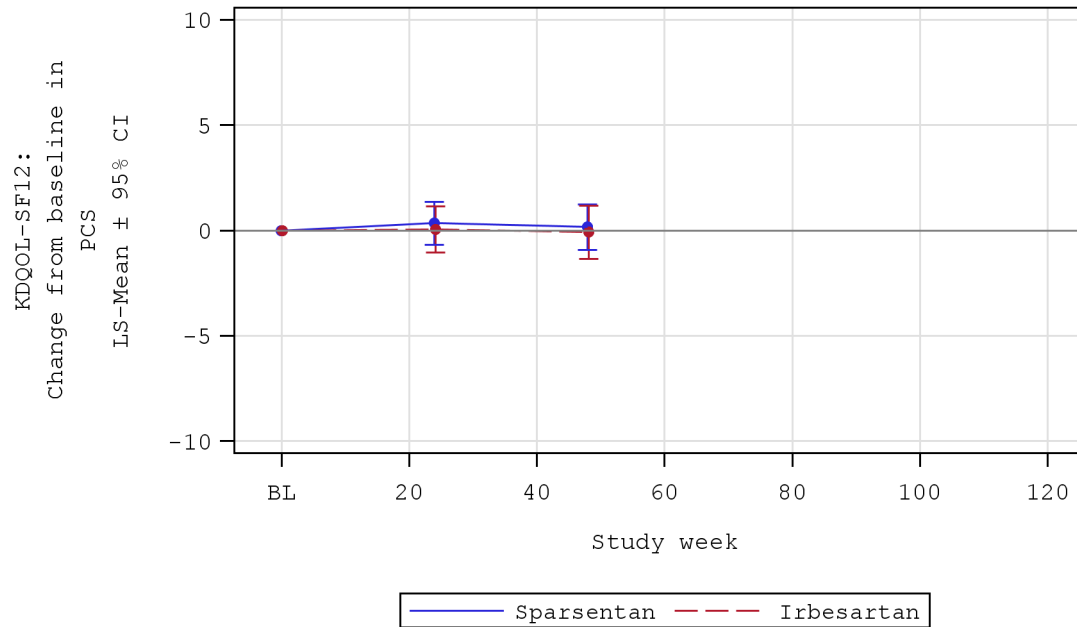
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 04APR2024

Figure PF1KPSC\_FMG0: KDQOL-SF12: Change from baseline in PCS  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	128	112
Irbesartan	112	80

Number of available records are presented per time-point. LS-Mean = Least square mean. 95% CI = 95% confidence interval.  
 A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KPSC\_FMC0

Table PT1KPSIT\_FMT0: KDQOL-SF12: Time to increase in PCS by at least 8.4 points  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	22 (10.9)	NE		1.067	(0.537, 2.119)	0.853
Irbesartan	202	15 (7.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

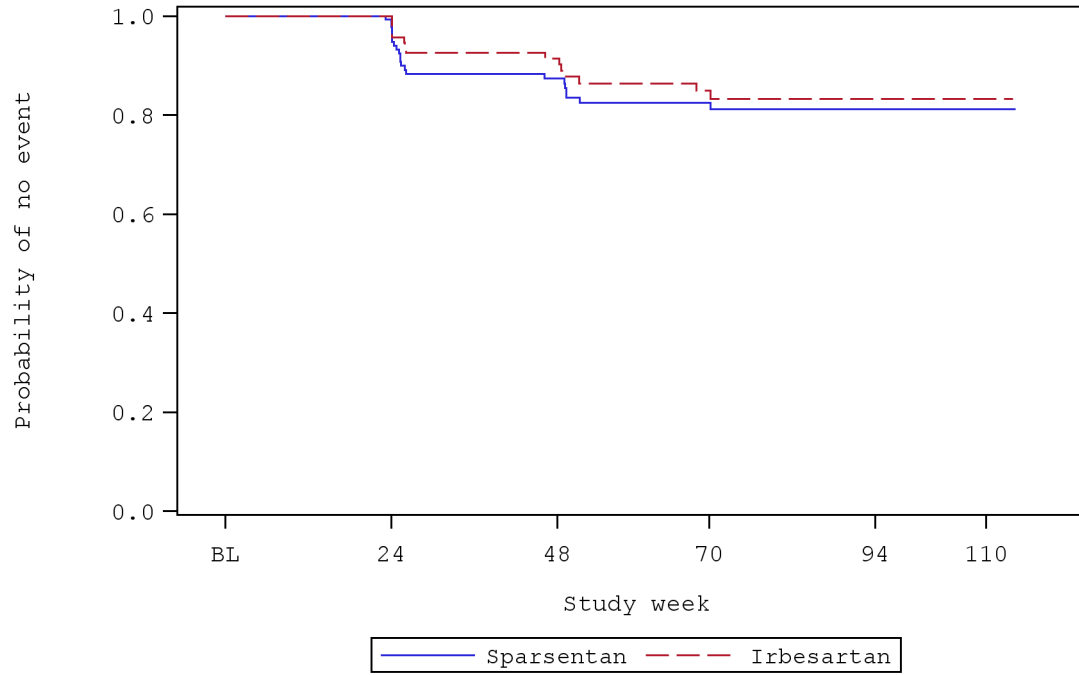
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

An increase reflects an improvement of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KPSIT\_FMK0: KDQOL-SF12: Time to increase in PCS by at least 8.4 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	136	101	72	32	15
Irbesartan	202	119	83	51	17	5

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KPSIT\_FMT0

Table PT1KPSDT\_FMT0: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	25 (12.4)	NE		1.068	(0.578, 1.973)	0.833
Irbesartan	202	18 (8.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

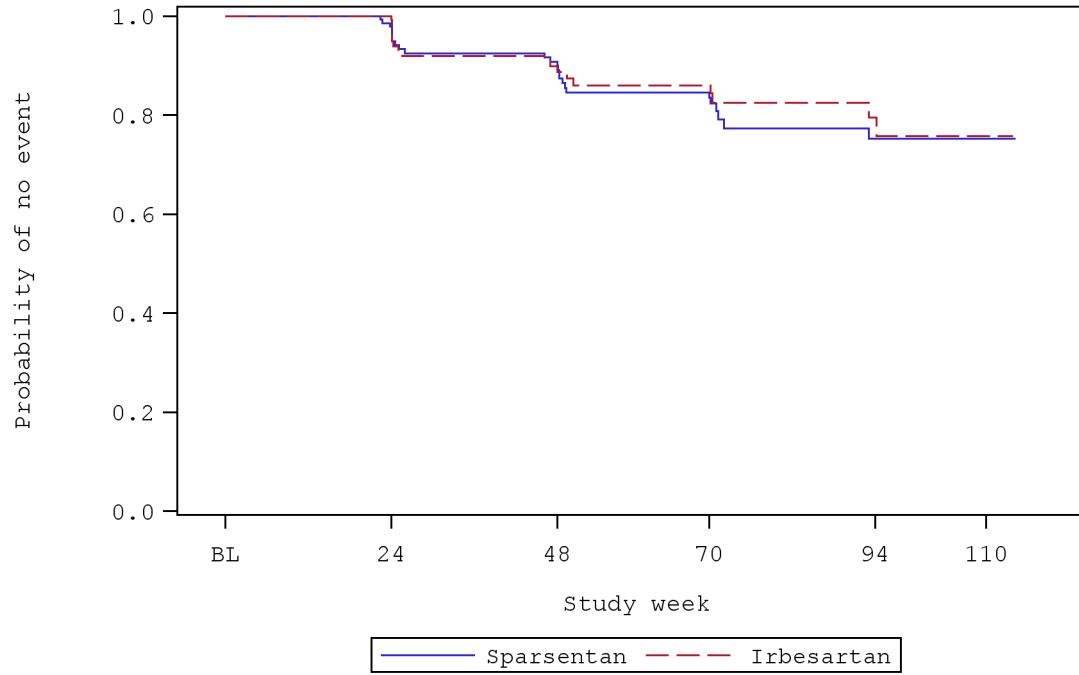
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024



Figure PF1KPSDT\_FMK0: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	135	107	76	32	10
Irbesartan	202	119	82	54	22	9

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KPSDT\_FMT0

Table PT1KMSC\_FMH0: KDQOL-SF12: Course of MCS  
 Full Analysis Set

Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
KDQOL-SF12: MCS	Baseline	Sparsentan	202	187 (92.6)	50.39 (8.31)	16.2	44.98	51.55	56.68	66.5	
		Irbesartan	202	184 (91.1)	51.37 (8.63)	24.3	47.20	52.88	57.16	66.6	
	Week 24	Sparsentan	202	128 (63.4)	51.58 (7.68)	20.8	47.69	52.83	57.27	66.8	
		Irbesartan	202	112 (55.4)	52.71 (8.31)	24.2	48.87	55.37	57.95	64.2	
	Week 48	Sparsentan	202	112 (55.4)	50.72 (8.35)	27.5	44.95	51.89	57.25	63.4	
		Irbesartan	202	80 (39.6)	51.28 (9.92)	19.6	47.43	54.20	57.66	64.6	
KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	202	128 (63.4)	0.73 (8.28)	-23.6	-3.61	0.49	4.82	26.4	0.03 [-0.23, 0.28]
		Irbesartan	202	112 (55.4)	0.54 (6.77)	-15.4	-4.23	0.00	3.83	28.7	
	Week 48	Sparsentan	202	112 (55.4)	0.64 (9.02)	-23.7	-4.53	0.21	6.18	28.1	0.21 [-0.08, 0.50]
		Irbesartan	202	80 (39.6)	-1.20 (8.39)	-33.9	-3.72	0.00	4.07	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FMC0: KDQOL-SF12: Change from baseline in MCS  
 Full Analysis Set

KDQOL-SF12: change from baseline in MCS				Repeated measures analysis					
Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Week 24	Sparsentan	202	128 (63.4)	0.54 (0.63)	(-0.71, 1.78)	-0.42 (0.93)	(-2.25, 1.41)	0.654	
	Irbesartan	202	112 (55.4)	0.95 (0.68)	(-0.38, 2.29)				
Week 48	Sparsentan	202	112 (55.4)	0.05 (0.68)	(-1.28, 1.38)	0.69 (1.05)	(-1.37, 2.75)	0.510	
	Irbesartan	202	80 (39.6)	-0.64 (0.80)	(-2.21, 0.93)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

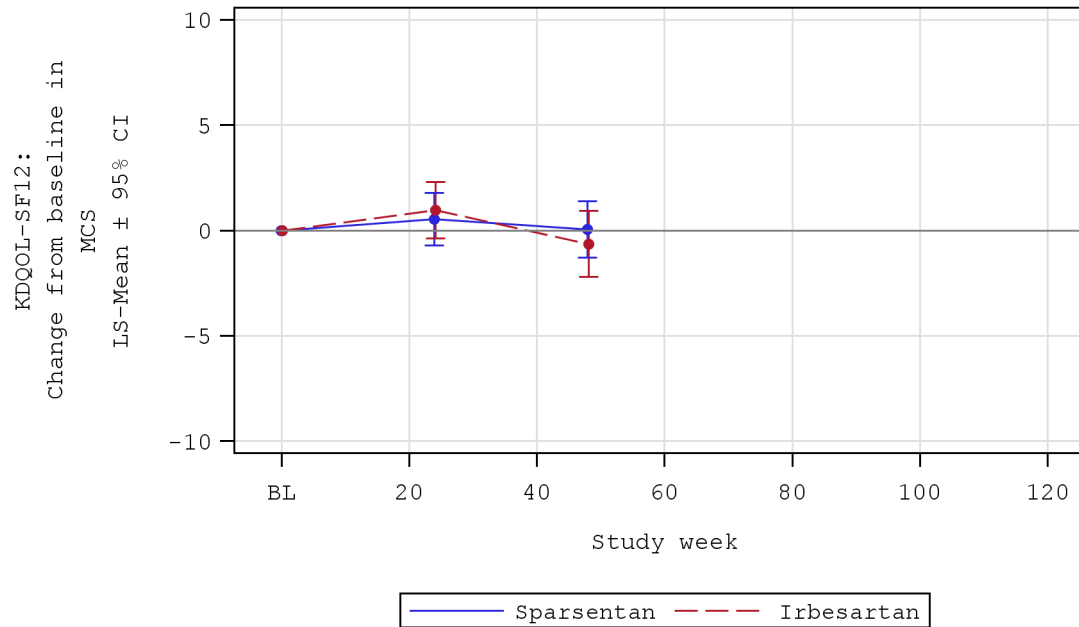
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 04APR2024

Figure PF1KMSC\_FMG0: KDQOL-SF12: Change from baseline in MCS  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	128	112
Irbesartan	112	80

Number of available records are presented per time-point. LS-Mean = Least square mean. 95% CI = 95% confidence interval.  
 A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KMSC\_FMG0

Table PT1KMSIT\_FMT0: KDQOL-SF12: Time to increase in MCS by at least 9.0 points  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	33 (16.3)	NE		1.821	(0.968, 3.426)	0.063
Irbesartan	202	15 (7.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

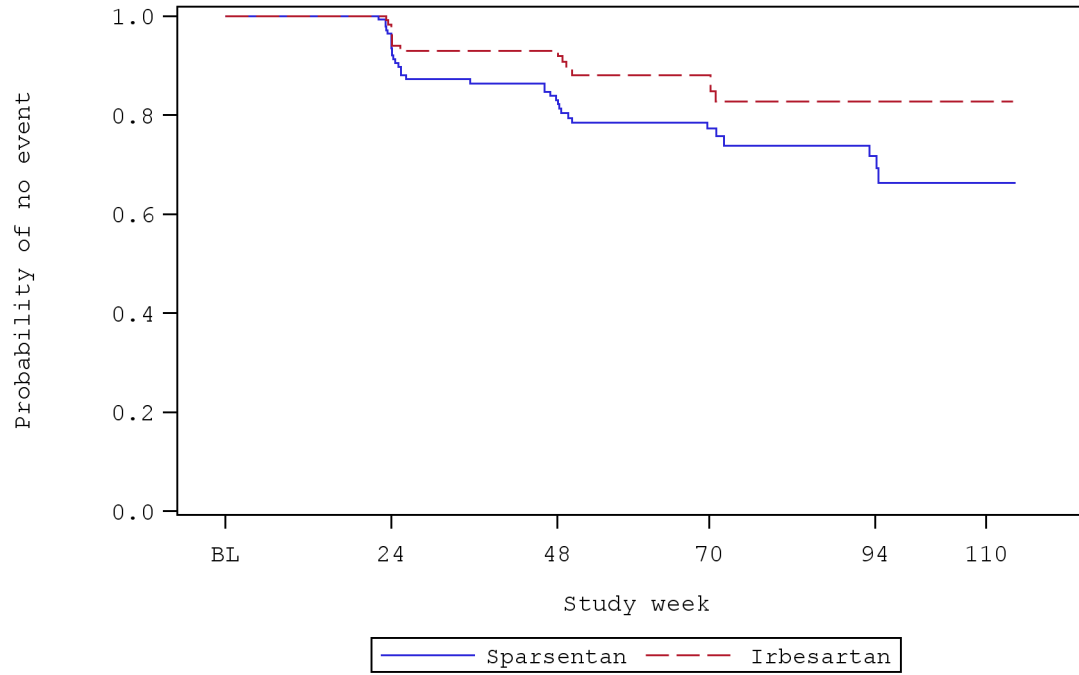
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

An increase reflects an improvement of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KMSIT\_FMK0: KDQOL-SF12: Time to increase in MCS by at least 9.0 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	133	97	67	32	13
Irbesartan	202	117	85	55	19	7

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KMSIT\_FMT0

Table PT1KMSDT\_FMT0: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0			Kaplan-Meier analysis		Cox regression		
Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sparsentan	202	34 (16.8)	111.1	(96.1, 114.3)	1.409	(0.808, 2.455)	0.227
Irbesartan	202	22 (10.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable. \* = significant treatment effect.

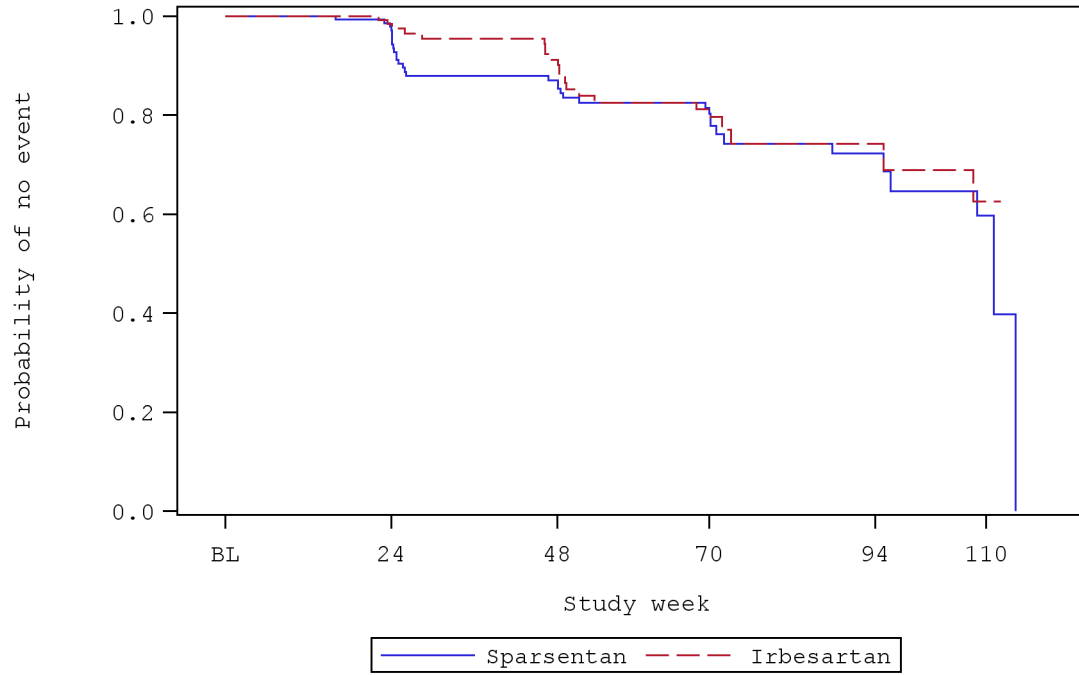
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient. Time is presented in weeks.

Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KMSDT\_FMK0: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	136	102	71	31	11
Irbesartan	202	118	84	52	19	7

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KMSDT\_FMT0



Table PT1A\_SMI0: Incidence of TEAEs during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
TEAEs	Double-blind period	Sparsentan	202	166 (82.2)	1.122 [1.010, 1.246]	1.682 [1.045, 2.709]	8.9 [0.3, 17.5]	0.042 *
		Irbesartan	202	148 (73.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AN\_SMI0: Incidence of non-severe TEAEs during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Non-severe TEAEs	Double-blind period	Sparsentan	202	49 (24.3)	1.195 [0.829, 1.724]	1.258 [0.786, 2.013]	4.0 [-4.6, 12.6]	0.403
		Irbesartan	202	41 (20.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AC\_SMI0: Incidence of severe TEAEs during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Severe TEAEs	Double-blind period	Sparsentan	202	14 (6.9)	0.933 [0.463, 1.883]	0.928 [0.436, 1.977]	-0.5 [-6.0, 5.0]	1.000
		Irbesartan	202	15 (7.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AS\_SMI0: Incidence of serious TEAEs during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Serious TEAEs	Double-blind period	Sparsentan	202	28 (13.9)	0.903 [0.563, 1.449]	0.888 [0.511, 1.543]	-1.5 [-8.9, 5.9]	0.778
		Irbesartan	202	31 (15.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AT\_SMI0: Incidence of TEAEs leading to study drug discontinuation during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
TEAEs leading to study drug discontinuation	Double-blind period	Sparsentan	202	16 (7.9)	1.778 [0.804, 3.929]	1.845 [0.795, 4.278]	3.5 [-1.7, 8.6]	0.215
		Irbesartan	202	9 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AD\_SMI0: Incidence of fatal TEAEs during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Fatal TEAEs	Double-blind period	Sparsentan	202	0 (0.0)				NE
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AA\_SMI0: Incidence of non-disease related TEAEs during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Non-disease related TEAEs	Double-blind period	Sparsentan	202	162 (80.2)	1.117 [1.001, 1.248]	1.592 [1.003, 2.528]	8.4 [-0.4, 17.2]	0.062
		Irbesartan	202	145 (71.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SMI0: Incidence of non-disease related non-severe TEAEs during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Non-disease related non-severe TEAEs	Double-blind period	Sparsentan	202	162 (80.2)	1.125 [1.007, 1.257]	1.631 [1.029, 2.587]	8.9 [0.1, 17.7]	0.048 *
		Irbesartan	202	144 (71.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024



Table PT1AAC\_SMI0: Incidence of non-disease related severe TEAEs during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Non-disease related severe TEAEs	Double-blind period	Sparsentan	202	11 (5.4)	1.000 [0.444, 2.254]	1.000 [0.423, 2.362]	0.0 [-4.9, 4.9]	1.000
		Irbesartan	202	11 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SMI0: Incidence of non-disease related serious TEAEs during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Non-disease related serious TEAEs	Double-blind period	Sparsentan	202	25 (12.4)	0.893 [0.540, 1.476]	0.878 [0.492, 1.565]	-1.5 [-8.6, 5.6]	0.768
		Irbesartan	202	28 (13.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SMS0: Incidence of TEAEs during double-blind period by SOC/PT Safety Set

TEAEs						RR	OR	RD	
SOC/PT	Time	Treatment	N	n (%)		[95 % CI]	[95 % CI]	[95 % CI]	p-value
SOC: Blood and lymphatic system disorders	Double-blind period	Sparsentan	202	13 (6.4)	1.625	[0.688, 3.836]	1.668	2.5	0.370
		Irbesartan	202	8 (4.0)				[-2.3, 7.3]	
SOC: Cardiac disorders	Double-blind period	Sparsentan	202	6 (3.0)	0.462	[0.179, 1.190]	0.445	-3.5	0.157
		Irbesartan	202	13 (6.4)				[-8.1, 1.1]	
SOC: Eye disorders	Double-blind period	Sparsentan	202	10 (5.0)	2.000	[0.696, 5.748]	2.052	2.5	0.293
		Irbesartan	202	5 (2.5)				[-1.7, 6.6]	
SOC: Gastrointestinal disorders	Double-blind period	Sparsentan	202	47 (23.3)	1.175	[0.809, 1.708]	1.228	3.5	0.468
		Irbesartan	202	40 (19.8)				[-5.0, 12.0]	\$
PT: Diarrhoea	Double-blind period	Sparsentan	202	7 (3.5)	0.500	[0.206, 1.213]	0.482	-3.5	0.178
		Irbesartan	202	14 (6.9)				[-8.3, 1.3]	
SOC: General disorders and administration site conditions	Double-blind period	Sparsentan	202	50 (24.8)	1.316	[0.905, 1.913]	1.420	5.9	0.185
		Irbesartan	202	38 (18.8)				[-2.6, 14.5]	\$
PT: Fatigue	Double-blind period	Sparsentan	202	16 (7.9)	2.667	[1.065, 6.677]	2.810	5.0	0.046 *
		Irbesartan	202	6 (3.0)				[0.1, 9.8]	
PT: Oedema peripheral	Double-blind period	Sparsentan	202	25 (12.4)	1.563	[0.861, 2.837]	1.642	4.5	0.187
		Irbesartan	202	16 (7.9)				[-1.9, 10.8]	\$

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher was used. \* = significant treatment effect.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Only events are reported with at least 1% of affected patients and 10 affected patients in at least 1 treatment arm. \$ = events with at least 10% affected patients in at least 1 treatment arm.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SMS0: Incidence of TEAEs during double-blind period by SOC/PT Safety Set

TEAEs						RR	OR	RD		
SOC/PT	Time	Treatment	N	n (%)		[95 % CI]	[95 % CI]	[95 % CI]	p-value	
SOC: Infections and infestations	Double-blind period	Sparsentan	202	56 (27.7)	0.918	0.887	-2.5	0.661	\$	
		Irbesartan	202	61 (30.2)	[0.676, 1.246]	[0.577, 1.363]	[-11.8, 6.9]			
PT: Upper respiratory tract infection	Double-blind period	Sparsentan	202	12 (5.9)	1.000	1.000	0.0	1.000		
		Irbesartan	202	12 (5.9)	[0.460, 2.173]	[0.438, 2.282]	[-5.1, 5.1]			
SOC: Injury, poisoning and procedural complications	Double-blind period	Sparsentan	202	11 (5.4)	0.611	0.589	-3.5	0.247		
		Irbesartan	202	18 (8.9)	[0.296, 1.261]	[0.271, 1.280]	[-9.0, 2.1]			
SOC: Investigations	Double-blind period	Sparsentan	202	47 (23.3)	1.516	1.673	7.9	0.058	\$	
		Irbesartan	202	31 (15.3)	[1.007, 2.283]	[1.012, 2.766]	[-0.2, 16.1]			
PT: Lipase increased	Double-blind period	Sparsentan	202	10 (5.0)	2.000	2.052	2.5	0.293		
		Irbesartan	202	5 (2.5)	[0.696, 5.748]	[0.689, 6.114]	[-1.7, 6.6]			
SOC: Metabolism and nutrition disorders	Double-blind period	Sparsentan	202	44 (21.8)	0.880	0.847	-3.0	0.556	\$	
		Irbesartan	202	50 (24.8)	[0.617, 1.255]	[0.533, 1.344]	[-11.7, 5.8]			
PT: Hyperkalaemia	Double-blind period	Sparsentan	202	21 (10.4)	1.167	1.186	1.5	0.737	\$	
		Irbesartan	202	18 (8.9)	[0.641, 2.123]	[0.612, 2.300]	[-4.8, 7.7]			
SOC: Musculoskeletal and connective tissue disorders	Double-blind period	Sparsentan	202	46 (22.8)	1.000	1.000	0.0	1.000	\$	
		Irbesartan	202	46 (22.8)	[0.698, 1.432]	[0.628, 1.592]	[-8.7, 8.7]			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher was used. \* = significant treatment effect.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Only events are reported with at least 1% of affected patients and 10 affected patients in at least 1 treatment arm. § = events with at least 10% affected patients in at least 1 treatment arm.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SMS0: Incidence of TEAEs during double-blind period by SOC/PT Safety Set

TEAEs						RR	OR	RD		
SOC/PT	Time	Treatment	N	n (%)		[95 % CI]	[95 % CI]	[95 % CI]	p-value	
PT: Back pain	Double-blind period	Sparsentan	202	8 (4.0)	0.615	[0.261, 1.453]	0.600	-2.5	0.370	
		Irbesartan	202	13 (6.4)			[0.243, 1.479]	[-7.3, 2.3]		
PT: Muscle spasms	Double-blind period	Sparsentan	202	8 (4.0)	0.727	[0.299, 1.770]	0.716	-1.5	0.639	
		Irbesartan	202	11 (5.4)			[0.282, 1.819]	[-6.1, 3.1]		
SOC: Nervous system disorders	Double-blind period	Sparsentan	202	53 (26.2)	1.262	[0.885, 1.799]	1.355	5.4	0.241	\$
		Irbesartan	202	42 (20.8)			[0.853, 2.151]	[-3.3, 14.2]		\$
PT: Dizziness	Double-blind period	Sparsentan	202	26 (12.9)	2.889	[1.389, 6.010]	3.168	8.4	0.004	* \$
		Irbesartan	202	9 (4.5)			[1.445, 6.946]	[2.5, 14.3]		\$
PT: Headache	Double-blind period	Sparsentan	202	20 (9.9)	0.952	[0.533, 1.702]	0.947	-0.5	1.000	\$
		Irbesartan	202	21 (10.4)			[0.496, 1.807]	[-6.9, 5.9]		\$
SOC: Psychiatric disorders	Double-blind period	Sparsentan	202	12 (5.9)	1.000	[0.460, 2.173]	1.000	0.0	1.000	
		Irbesartan	202	12 (5.9)			[0.438, 2.282]	[-5.1, 5.1]		
SOC: Renal and urinary disorders	Double-blind period	Sparsentan	202	29 (14.4)	1.115	[0.682, 1.825]	1.135	1.5	0.772	\$
		Irbesartan	202	26 (12.9)			[0.642, 2.005]	[-5.7, 8.7]		\$
SOC: Respiratory, thoracic and mediastinal disorders	Double-blind period	Sparsentan	202	25 (12.4)	1.471	[0.820, 2.638]	1.537	4.0	0.254	\$
		Irbesartan	202	17 (8.4)			[0.803, 2.943]	[-2.5, 10.4]		\$

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Only events are reported with at least 1% of affected patients and 10 affected patients in at least 1 treatment arm. \$ = events with at least 10% affected patients in at least 1 treatment arm.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SMS0: Incidence of TEAEs during double-blind period by SOC/PT Safety Set

TEAEs									
SOC/PT	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
SOC: Skin and subcutaneous tissue disorders	Double-blind period	Sparsentan	202	22 (10.9)	0.917 [0.532, 1.580]	0.906 [0.490, 1.676]	-1.0 [-7.7, 5.7]	0.876	\$
		Irbesartan	202	24 (11.9)					\$
SOC: Vascular disorders	Double-blind period	Sparsentan	202	47 (23.3)	1.679 [1.097, 2.568]	1.884 [1.125, 3.155]	9.4 [1.4, 17.4]	0.021	* \$
		Irbesartan	202	28 (13.9)					\$
PT: Hypertension	Double-blind period	Sparsentan	202	16 (7.9)	0.941 [0.489, 1.811]	0.936 [0.459, 1.909]	-0.5 [-6.3, 5.3]	1.000	
		Irbesartan	202	17 (8.4)					
PT: Hypotension	Double-blind period	Sparsentan	202	20 (9.9)	3.333 [1.367, 8.127]	3.590 [1.410, 9.138]	6.9 [1.7, 12.2]	0.007	* \$
		Irbesartan	202	6 (3.0)					

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher was used. \* = significant treatment effect.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Only events are reported with at least 1% of affected patients and 10 affected patients in at least 1 treatment arm. § = events with at least 10% affected patients in at least 1 treatment arm.  
Source Data: aae, created on: 30JAN2024

Table PT1AC\_SMS0: Incidence of severe TEAEs during double-blind period by SOC/PT  
Safety Set

Not done. No valid soc/pt combination with sufficient high incidence found.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher was used. \* = significant treatment effect.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Only events are reported with at least 5% of affected patients in at least 1 treatment arm.  
Source Data: aae, created on: 30JAN2024

Table PT1AS\_SMS0: Incidence of serious TEAEs during double-blind period by SOC/PT Safety Set

Serious TEAEs								
SOC/PT	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Infections and infestations	Double-blind period	Sparsentan	202	7 (3.5)	0.636 [0.252, 1.609]	0.623 [0.237, 1.642]	-2.0 [-6.5, 2.5]	0.470
		Irbesartan	202	11 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Only events are reported with at least 5% of affected patients in at least 1 treatment arm.  
 Source Data: aae, created on: 30JAN2024



Table PT1AT\_SMSD: Incidence of TEAEs leading to study drug discontinuation during double-blind period by SOC/PT  
Safety Set

TEAEs leading to study drug discontinuation				
SOC/PT	Time	Treatment	N	n (%)
<b>SOC:</b> Gastrointestinal disorders	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	1 (0.5)
PT: Abdominal pain	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
PT: Frequent bowel movements	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)
PT: Nausea	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)
PT: Rectal haemorrhage	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
PT: Vomiting	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
<b>SOC:</b> Infections and infestations	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)
PT: COVID-19	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Source Data: aae, created on: 30JAN2024

Table PT1AT\_SMSD: Incidence of TEAEs leading to study drug discontinuation during double-blind period by SOC/PT Safety Set

TEAEs leading to study drug discontinuation				
SOC/PT	Time	Treatment	N	n (%)
<b>SOC:</b> Injury, poisoning and procedural complications	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)
PT: Foetal exposure during pregnancy	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)
<b>SOC:</b> Investigations	Double-blind period	Sparsentan	202	4 (2.0)
		Irbesartan	202	1 (0.5)
PT: Alanine aminotransferase increased	Double-blind period	Sparsentan	202	2 (1.0)
		Irbesartan	202	0 (0.0)
PT: Aspartate aminotransferase increased	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
PT: Blood creatinine increased	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
PT: Lipase increased	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
PT: Pregnancy test urine positive	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Source Data: aae, created on: 30JAN2024

Table PT1AT\_SMSD: Incidence of TEAEs leading to study drug discontinuation during double-blind period by SOC/PT  
 Safety Set

TEAEs leading to study drug discontinuation				
SOC/PT	Time	Treatment	N	n (%)
<b>SOC:</b> Neoplasms benign, malignant and unspecified (incl cysts and polyps)	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
PT: Diffuse large B-cell lymphoma	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
<b>SOC:</b> Nervous system disorders	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
PT: Dizziness	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
<b>SOC:</b> Pregnancy, puerperium and perinatal conditions	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	1 (0.5)
PT: Pregnancy	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	1 (0.5)
<b>SOC:</b> Psychiatric disorders	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
PT: Depressed mood	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Source Data: aae, created on: 30JAN2024

Table PT1AT\_SMSD: Incidence of TEAEs leading to study drug discontinuation during double-blind period by SOC/PT Safety Set

TEAEs leading to study drug discontinuation				
SOC/PT	Time	Treatment	N	n (%)
<b>SOC:</b> Renal and urinary disorders	Double-blind period	Sparsentan	202	6 (3.0)
		Irbesartan	202	4 (2.0)
PT: Acute kidney injury	Double-blind period	Sparsentan	202	3 (1.5)
		Irbesartan	202	0 (0.0)
PT: Chronic kidney disease	Double-blind period	Sparsentan	202	2 (1.0)
		Irbesartan	202	0 (0.0)
PT: Glomerulonephritis rapidly progressive	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)
PT: IgA nephropathy	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)
PT: Nephrotic syndrome	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)
PT: Renal impairment	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	2 (1.0)
<b>SOC:</b> Skin and subcutaneous tissue disorders	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	1 (0.5)

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Source Data: aae, created on: 30JAN2024

Table PT1AT\_SMSD: Incidence of TEAEs leading to study drug discontinuation during double-blind period by SOC/PT  
 Safety Set

TEAEs leading to study drug discontinuation				
SOC/PT	Time	Treatment	N	n (%)
PT: Rash	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	1 (0.5)
<b>SOC:</b> Vascular disorders	Double-blind period	Sparsentan	202	2 (1.0)
		Irbesartan	202	0 (0.0)
PT: Hypotension	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)
PT: Orthostatic hypotension	Double-blind period	Sparsentan	202	1 (0.5)
		Irbesartan	202	0 (0.0)

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Source Data: aae, created on: 30JAN2024

Table PT1AD\_SMSD: Incidence of fatal TEAEs during double-blind period by SOC/PT  
Safety Set

No deaths observed.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Source Data: aae, created on: 30JAN2024

Table PT1AEF\_SMI0: Incidence of AESI abnormal liver function during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI abnormal liver function	Double-blind period	Sparsentan	202	4 (2.0)	1.000 [0.254, 3.944]	1.000 [0.247, 4.055]	0.0 [-3.2, 3.2]	1.000
		Irbesartan	202	4 (2.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEFN\_SMI0: Incidence of AESI abnormal liver function during double-blind period - non-severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI abnormal liver function - non-severe	Double-blind period	Sparsentan	202	4 (2.0)	1.000 [0.254, 3.944]	1.000 [0.247, 4.055]	0.0 [-3.2, 3.2]	1.000
		Irbesartan	202	4 (2.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEFC\_SMI0: Incidence of AESI abnormal liver function during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI abnormal liver function - severe	Double-blind period	Sparsentan	202	1 (0.5)	3.000 + [0.123, 73.208]	3.015 + [0.122, 74.448]	0.5 [-1.0, 2.0]	1.000
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEFS\_SMI0: Incidence of AESI abnormal liver function during double-blind period - serious  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI abnormal liver function - serious	Double-blind period	Sparsentan	202	0 (0.0)	0.333 + [0.014, 8.134]	0.332 + [0.013, 8.191]	-0.5 [-2.0, 1.0]	1.000
		Irbesartan	202	1 (0.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SMI0: Incidence of AESI COVID-19 during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI COVID-19	Double-blind period	Sparsentan	202	5 (2.5)	0.625 [0.208, 1.878]	0.615 [0.198, 1.914]	-1.5 [-5.4, 2.4]	0.575
		Irbesartan	202	8 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEVN\_SMI0: Incidence of AESI COVID-19 during double-blind period - non-severe Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI COVID-19 - non-severe	Double-blind period	Sparsentan	202	3 (1.5)	0.429 [0.112, 1.634]	0.420 [0.107, 1.647]	-2.0 [-5.5, 1.5]	0.338
		Irbesartan	202	7 (3.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEVC\_SMI0: Incidence of AESI COVID-19 during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI COVID-19 - severe	Double-blind period	Sparsentan	202	2 (1.0)	2.000 [0.183, 21.882]	2.010 [0.181, 22.344]	0.5 [-1.7, 2.7]	1.000
		Irbesartan	202	1 (0.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEVS\_SMI0: Incidence of AESI COVID-19 during double-blind period - serious  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI COVID-19 - serious	Double-blind period	Sparsentan	202	3 (1.5)	0.500	0.492	-1.5	0.503
		Irbesartan	202	6 (3.0)	[0.127, 1.972]	[0.121, 1.997]	[-4.9, 1.9]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SMI0: Incidence of AESI cardiovascular system during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI cardiovascular system	Double-blind period	Sparsentan	202	51 (25.2)	1.244 [0.866, 1.786]	1.326 [0.831, 2.116]	5.0 [-3.7, 13.6]	0.286
		Irbesartan	202	41 (20.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SMI0: Incidence of AESI cardiovascular system during double-blind period - non-severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI cardiovascular system - non-severe	Double-blind period	Sparsentan	202	50 (24.8)	1.250 [0.866, 1.805]	1.332 [0.832, 2.134]	5.0 [-3.6, 13.5]	0.282
		Irbesartan	202	40 (19.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024



Table PT1AECC\_SMI0: Incidence of AESI cardiovascular system during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI cardiovascular system - severe	Double-blind period	Sparsentan	202	1 (0.5)	1.000 [0.063, 15.878]	1.000 [0.062, 16.098]	0.0 [-1.9, 1.9]	1.000
		Irbesartan	202	1 (0.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AECS\_SMI0: Incidence of AESI cardiovascular system during double-blind period - serious  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI cardiovascular system - serious	Double-blind period	Sparsentan	202	1 (0.5)	0.500 [0.046, 5.471]	0.498 [0.045, 5.531]	-0.5 [-2.7, 1.7]	1.000
		Irbesartan	202	2 (1.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SMI0: Incidence of AESI hypotension during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hypotension	Double-blind period	Sparsentan	202	50 (24.8)	2.381 [1.487, 3.813]	2.835 [1.630, 4.931]	14.4 [6.6, 22.1]	<0.001 *
		Irbesartan	202	21 (10.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SMI0: Incidence of AESI hypotension during double-blind period - non-severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hypotension - non-severe	Double-blind period	Sparsentan	202	49 (24.3)	2.333 [1.455, 3.743]	2.760 [1.585, 4.807]	13.9 [6.1, 21.6]	<0.001 *
		Irbesartan	202	21 (10.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEHC\_SMI0: Incidence of AESI hypotension during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hypotension - severe	Double-blind period	Sparsentan	202	1 (0.5)	3.000 + [0.123, 73.208]	3.015 + [0.122, 74.448]	0.5 [-1.0, 2.0]	1.000
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEHS\_SMI0: Incidence of AESI hypotension during double-blind period - serious Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hypotension - serious	Double-blind period	Sparsentan	202	3 (1.5)	3.000 [0.315, 28.599]	3.030 [0.313, 29.378]	1.0 [-1.4, 3.4]	0.623
		Irbesartan	202	1 (0.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SMI0: Incidence of AESI hepatic disorders during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hepatic disorders	Double-blind period	Sparsentan	202	13 (6.4)	1.857 [0.757, 4.558]	1.916 [0.748, 4.907]	3.0 [-1.7, 7.7]	0.251
		Irbesartan	202	7 (3.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SMI0: Incidence of AESI hepatic disorders during double-blind period - non-severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hepatic disorders - non-severe	Double-blind period	Sparsentan	202	13 (6.4)	1.857 [0.757, 4.558]	1.916 [0.748, 4.907]	3.0 [-1.7, 7.7]	0.251
		Irbesartan	202	7 (3.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024



Table PT1AELC\_SMI0: Incidence of AESI hepatic disorders during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hepatic disorders - severe	Double-blind period	Sparsentan	202	1 (0.5)	3.000 + [0.123, 73.208]	3.015 + [0.122, 74.448]	0.5 [-1.0, 2.0]	1.000
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AELS\_SMI0: Incidence of AESI hepatic disorders during double-blind period - serious  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hepatic disorders - serious	Double-blind period	Sparsentan	202	0 (0.0)	0.333 + [0.014, 8.134]	0.332 + [0.013, 8.191]	-0.5 [-2.0, 1.0]	1.000
		Irbesartan	202	1 (0.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SMI0: Incidence of AESI acute pancreatitis during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI acute pancreatitis	Double-blind period	Sparsentan	202	13 (6.4)	1.625 [0.688, 3.836]	1.668 [0.676, 4.116]	2.5 [-2.3, 7.3]	0.370
		Irbesartan	202	8 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SMI0: Incidence of AESI acute pancreatitis during double-blind period - non-severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI acute pancreatitis - non-severe	Double-blind period	Sparsentan	202	13 (6.4)	1.625 [0.688, 3.836]	1.668 [0.676, 4.116]	2.5 [-2.3, 7.3]	0.370
		Irbesartan	202	8 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEPC\_SMI0: Incidence of AESI acute pancreatitis during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR	OR	RD	p-value
					[95 % CI]	[95 % CI]	[95 % CI]	
AESI acute pancreatitis - severe	Double-blind period	Sparsentan	202	0 (0.0)				NE
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEPS\_SMI0: Incidence of AESI acute pancreatitis during double-blind period - serious Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI acute pancreatitis - serious	Double-blind period	Sparsentan	202	0 (0.0)				NE
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SMI0: Incidence of AESI fluid retention during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI fluid retention	Double-blind period	Sparsentan	202	28 (13.9)	1.556 [0.889, 2.721]	1.645 [0.878, 3.080]	5.0 [-1.7, 11.6]	0.158
		Irbesartan	202	18 (8.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SMI0: Incidence of AESI fluid retention during double-blind period - non-severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI fluid retention - non-severe	Double-blind period	Sparsentan	202	28 (13.9)	1.556 [0.889, 2.721]	1.645 [0.878, 3.080]	5.0 [-1.7, 11.6]	0.158
		Irbesartan	202	18 (8.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEOC\_SMI0: Incidence of AESI fluid retention during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI fluid retention - severe	Double-blind period	Sparsentan	202	0 (0.0)				NE
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEOS\_SMI0: Incidence of AESI fluid retention during double-blind period - serious  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI fluid retention - serious	Double-blind period	Sparsentan	202	1 (0.5)	3.000 + [0.123, 73.208]	3.015 + [0.122, 74.448]	0.5 [-1.0, 2.0]	1.000
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SMI0: Incidence of AESI anemia during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI anemia	Double-blind period	Sparsentan	202	13 (6.4)	1.857 [0.757, 4.558]	1.916 [0.748, 4.907]	3.0 [-1.7, 7.7]	0.251
		Irbesartan	202	7 (3.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SMI0: Incidence of AESI anemia during double-blind period - non-severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI anemia - non-severe	Double-blind period	Sparsentan	202	13 (6.4)	1.857 [0.757, 4.558]	1.916 [0.748, 4.907]	3.0 [-1.7, 7.7]	0.251
		Irbesartan	202	7 (3.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEAC\_SMI0: Incidence of AESI anemia during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI anemia - severe	Double-blind period	Sparsentan	202	0 (0.0)				NE
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEAS\_SMI0: Incidence of AESI anemia during double-blind period - serious  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI anemia - serious	Double-blind period	Sparsentan	202	1 (0.5)	3.000 + [0.123, 73.208]	3.015 + [0.122, 74.448]	0.5 [-1.0, 2.0]	1.000
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SMI0: Incidence of AESI hyperkalemia during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hyperkalemia	Double-blind period	Sparsentan	202	23 (11.4)	1.211 [0.681, 2.152]	1.238 [0.652, 2.351]	2.0 [-4.5, 8.4]	0.625
		Irbesartan	202	19 (9.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SMI0: Incidence of AESI hyperkalemia during double-blind period - non-severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hyperkalemia - non-severe	Double-blind period	Sparsentan	202	23 (11.4)	1.211 [0.681, 2.152]	1.238 [0.652, 2.351]	2.0 [-4.5, 8.4]	0.625
		Irbesartan	202	19 (9.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEUC\_SMI0: Incidence of AESI hyperkalemia during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hyperkalemia - severe	Double-blind period	Sparsentan	202	0 (0.0)				NE
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEUS\_SMI0: Incidence of AESI hyperkalemia during double-blind period - serious  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI hyperkalemia - serious	Double-blind period	Sparsentan	202	0 (0.0)				NE
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SMI0: Incidence of AESI cardiac arrhythmias during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI cardiac arrhythmias	Double-blind period	Sparsentan	202	9 (4.5)	0.600 [0.269, 1.339]	0.581 [0.248, 1.361]	-3.0 [-8.1, 2.1]	0.293
		Irbesartan	202	15 (7.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SMI0: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI cardiac arrhythmias - non-severe	Double-blind period	Sparsentan	202	9 (4.5)	0.600 [0.269, 1.339]	0.581 [0.248, 1.361]	-3.0 [-8.1, 2.1]	0.293
		Irbesartan	202	15 (7.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SMI0: Incidence of AESI cardiac arrhythmias during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI cardiac arrhythmias - severe	Double-blind period	Sparsentan	202	0 (0.0)				NE
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEYS\_SMI0: Incidence of AESI cardiac arrhythmias during double-blind period - serious Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI cardiac arrhythmias - serious	Double-blind period	Sparsentan	202	0 (0.0)				NE
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEI\_SMI0: Incidence of AESI acute kidney injury during double-blind period  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI acute kidney injury	Double-blind period	Sparsentan	202	8 (4.0)	2.667 [0.718, 9.908]	2.735 [0.715, 10.463]	2.5 [-1.2, 6.1]	0.220
		Irbesartan	202	3 (1.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEIN\_SMI0: Incidence of AESI acute kidney injury during double-blind period - non-severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI acute kidney injury - non-severe	Double-blind period	Sparsentan	202	5 (2.5)	1.667 [0.404, 6.881]	1.684 [0.397, 7.140]	1.0 [-2.2, 4.2]	0.724
		Irbesartan	202	3 (1.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEIC\_SMI0: Incidence of AESI acute kidney injury during double-blind period - severe  
 Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI acute kidney injury - severe	Double-blind period	Sparsentan	202	3 (1.5)	7.000 + [0.364, 134.653]	7.105 + [0.365, 138.444]	1.5 [-0.7, 3.6]	0.248
		Irbesartan	202	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEIS\_SMI0: Incidence of AESI acute kidney injury during double-blind period - serious Safety Set

Variable	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
AESI acute kidney injury - serious	Double-blind period	Sparsentan	202	4 (2.0)	4.000 [0.451, 35.478]	4.061 [0.450, 36.650]	1.5 [-1.2, 4.1]	0.372
		Irbesartan	202	1 (0.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher was used. \* = significant treatment effect.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Source Data: aae, created on: 30JAN2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Sex	Overall	Sparsentan						Interaction:	0.007	#
Male	Week 6	Sparsentan	139	132 (95.0)	-41.52	(-47.37, -35.02)	0.596	(0.514, 0.691)	<0.001	*
		Irbesartan	143	137 (95.8)	-1.91	(-11.56, 8.79)				
	Week 36	Sparsentan	139	92 (66.2)	-47.16	(-53.12, -40.44)	0.584	(0.494, 0.691)	<0.001	*
		Irbesartan	143	97 (67.8)	-9.58	(-19.51, 1.57)				
	Week 58	Sparsentan	139	69 (49.6)	-46.10	(-52.99, -38.20)	0.583	(0.482, 0.706)	<0.001	*
		Irbesartan	143	73 (51.0)	-7.58	(-19.06, 5.52)				
Female	Week 6	Sparsentan	63	59 (93.7)	-34.05	(-43.52, -22.99)	0.813	(0.650, 1.017)	0.070	
		Irbesartan	59	54 (91.5)	-18.86	(-30.90, -4.71)				
	Week 36	Sparsentan	63	47 (74.6)	-49.63	(-57.44, -40.38)	0.709	(0.551, 0.913)	0.008	*
		Irbesartan	59	35 (59.3)	-28.97	(-41.10, -14.34)				
	Week 58	Sparsentan	63	44 (69.8)	-46.91	(-55.50, -36.66)	0.788	(0.596, 1.042)	0.094	
		Irbesartan	59	26 (44.1)	-32.61	(-45.67, -16.41)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Age	Overall	Sparsentan						Interaction:	0.591	
<= 45 years	Week 6	Sparsentan	96	91 (94.8)	-44.39	(-51.02, -36.86)	0.613	(0.513, 0.732)	<0.001	*
		Irbesartan	99	95 (96.0)	-9.23	(-19.88, 2.84)				
	Week 36	Sparsentan	96	66 (68.8)	-50.03	(-56.62, -42.44)	0.658	(0.539, 0.804)	<0.001	*
		Irbesartan	99	64 (64.6)	-24.09	(-34.09, -12.57)				
	Week 58	Sparsentan	96	51 (53.1)	-47.04	(-54.84, -37.90)	0.672	(0.534, 0.845)	<0.001	*
		Irbesartan	99	45 (45.5)	-21.20	(-33.16, -7.11)				
> 45 years	Week 6	Sparsentan	106	100 (94.3)	-34.61	(-41.96, -26.32)	0.691	(0.583, 0.819)	<0.001	*
		Irbesartan	103	96 (93.2)	-5.36	(-16.16, 6.84)				
	Week 36	Sparsentan	106	73 (68.9)	-46.28	(-53.04, -38.54)	0.576	(0.475, 0.699)	<0.001	*
		Irbesartan	103	68 (66.0)	-6.79	(-18.79, 6.99)				
	Week 58	Sparsentan	106	62 (58.5)	-45.60	(-53.02, -37.00)	0.599	(0.484, 0.742)	<0.001	*
		Irbesartan	103	54 (52.4)	-9.21	(-22.22, 5.97)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.490	
<= 18 years	Week 6	Sparsentan	9	8 (88.9)	-48.45	(-70.11, -11.09)	0.554	(0.210, 1.461)	0.212	
		Irbesartan	5	5 (100.0)	-6.98	(-55.78, 95.70)				
	Week 36	Sparsentan	9	6 (66.7)	-35.09	(-64.13, 17.45)	0.921	(0.296, 2.868)	0.882	
		Irbesartan	5	2 (40.0)	-29.55	(-72.79, 82.41)				
	Week 58	Sparsentan	9	4 (44.4)	-52.57	(-75.56, -7.94)	1.441	(0.369, 5.630)	0.587	
		Irbesartan	5	1 (20.0)	-67.08	(-89.97, 8.07)				
> 18 to 40 years	Week 6	Sparsentan	102	98 (96.1)	-43.71	(-49.93, -36.71)	0.628	(0.534, 0.740)	<0.001	*
		Irbesartan	109	104 (95.4)	-10.42	(-20.04, 0.36)				
	Week 36	Sparsentan	102	69 (67.6)	-50.14	(-56.34, -43.07)	0.627	(0.521, 0.754)	<0.001	*
		Irbesartan	109	72 (66.1)	-20.47	(-30.05, -9.57)				
	Week 58	Sparsentan	102	58 (56.9)	-48.71	(-55.67, -40.65)	0.623	(0.506, 0.768)	<0.001	*
		Irbesartan	109	52 (47.7)	-17.73	(-29.08, -4.55)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
> 40 years	Week 6	Sparsentan	91	85 (93.4)	-32.26	(-40.74, -22.56)	0.711	(0.587, 0.860)	<0.001	*
		Irbesartan	88	82 (93.2)	-4.67	(-16.75, 9.18)				
	Week 36	Sparsentan	91	64 (70.3)	-46.19	(-53.61, -37.59)	0.599	(0.484, 0.741)	<0.001	*
		Irbesartan	88	58 (65.9)	-10.13	(-22.92, 4.79)				
	Week 58	Sparsentan	91	51 (56.0)	-42.02	(-50.84, -31.62)	0.655	(0.516, 0.832)	<0.001	*
		Irbesartan	88	46 (52.3)	-11.53	(-25.55, 5.14)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Geographic region	Overall	Sparsentan						Interaction:	0.597	
North America	Week 6	Sparsentan	35	32 (91.4)	-29.17	(-41.34, -14.47)	0.688	(0.536, 0.884)	0.004	*
		Irbesartan	46	43 (93.5)	2.89	(-12.62, 21.15)				
	Week 36	Sparsentan	35	18 (51.4)	-47.67	(-58.60, -33.87)	0.465	(0.341, 0.633)	<0.001	*
		Irbesartan	46	25 (54.3)	12.60	(-7.81, 37.52)				
	Week 58	Sparsentan	35	12 (34.3)	-37.97	(-53.27, -17.67)	0.588	(0.407, 0.850)	0.005	*
		Irbesartan	46	18 (39.1)	5.52	(-16.53, 33.39)				
Europe	Week 6	Sparsentan	98	91 (92.9)	-43.23	(-49.90, -35.68)	0.626	(0.528, 0.742)	<0.001	*
		Irbesartan	115	108 (93.9)	-9.26	(-19.15, 1.83)				
	Week 36	Sparsentan	98	60 (61.2)	-47.91	(-54.96, -39.77)	0.642	(0.529, 0.780)	<0.001	*
		Irbesartan	115	78 (67.8)	-18.92	(-28.75, -7.73)				
	Week 58	Sparsentan	98	48 (49.0)	-44.23	(-52.68, -34.27)	0.657	(0.527, 0.820)	<0.001	*
		Irbesartan	115	57 (49.6)	-15.11	(-26.78, -1.59)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Asia Pacific	Week 6	Sparsentan	69	68 (98.6)	-39.98	(-48.57, -29.96)	0.656	(0.509, 0.845)	0.001	*
		Irbesartan	41	40 (97.6)	-8.54	(-25.13, 11.73)				
	Week 36	Sparsentan	69	61 (88.4)	-48.71	(-56.29, -39.82)	0.684	(0.521, 0.897)	0.006	*
		Irbesartan	41	29 (70.7)	-24.97	(-39.74, -6.58)				
	Week 58	Sparsentan	69	53 (76.8)	-50.99	(-58.60, -41.97)	0.676	(0.502, 0.910)	0.010	*
		Irbesartan	41	24 (58.5)	-27.50	(-43.21, -7.43)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024



Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Baseline BMI	Overall	Sparsentan						Interaction:	0.714	
< 27 kg/m**2	Week 6	Sparsentan	84	77 (91.7)	-43.64	(-50.72, -35.54)	0.644	(0.536, 0.775)	<0.001	*
		Irbesartan	94	90 (95.7)	-12.52	(-22.83, -0.83)				
	Week 36	Sparsentan	84	59 (70.2)	-52.39	(-58.95, -44.78)	0.672	(0.547, 0.827)	<0.001	*
		Irbesartan	94	59 (62.8)	-29.19	(-38.65, -18.28)				
	Week 58	Sparsentan	84	46 (54.8)	-57.42	(-63.86, -49.84)	0.595	(0.470, 0.753)	<0.001	*
		Irbesartan	94	41 (43.6)	-28.44	(-39.51, -15.34)				
≥ 27 kg/m**2	Week 6	Sparsentan	118	114 (96.6)	-35.51	(-42.38, -27.81)	0.661	(0.561, 0.779)	<0.001	*
		Irbesartan	107	100 (93.5)	-2.46	(-13.42, 9.89)				
	Week 36	Sparsentan	118	80 (67.8)	-44.50	(-51.16, -36.94)	0.568	(0.472, 0.684)	<0.001	*
		Irbesartan	107	72 (67.3)	-2.31	(-14.60, 11.75)				
	Week 58	Sparsentan	118	67 (56.8)	-36.42	(-44.80, -26.76)	0.661	(0.538, 0.813)	<0.001	*
		Irbesartan	107	58 (54.2)	-3.84	(-17.23, 11.71)				

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Randomization strata	Overall	Sparsentan						Interaction:	0.549	
eGFR Low and UP High	Week 6	Sparsentan	71	67 (94.4)	-33.04	(-41.81, -22.94)	0.673	(0.553, 0.819)	<0.001	*
		Irbesartan	74	70 (94.6)	-0.46	(-13.26, 14.24)				
	Week 36	Sparsentan	71	49 (69.0)	-41.33	(-49.90, -31.30)	0.640	(0.512, 0.799)	<0.001	*
		Irbesartan	74	48 (64.9)	-8.27	(-21.63, 7.36)				
	Week 58	Sparsentan	71	42 (59.2)	-44.06	(-52.90, -33.56)	0.591	(0.461, 0.757)	<0.001	*
		Irbesartan	74	37 (50.0)	-5.37	(-20.82, 13.09)				
eGFR Low and UP Low	Week 6	Sparsentan	55	51 (92.7)	-37.53	(-47.44, -25.76)	0.713	(0.559, 0.910)	0.007	*
		Irbesartan	55	53 (96.4)	-12.42	(-26.22, 3.96)				
	Week 36	Sparsentan	55	37 (67.3)	-42.23	(-52.27, -30.08)	0.654	(0.500, 0.854)	0.002	*
		Irbesartan	55	39 (70.9)	-11.61	(-26.72, 6.61)				
	Week 58	Sparsentan	55	29 (52.7)	-41.89	(-53.12, -27.98)	0.766	(0.567, 1.036)	0.083	
		Irbesartan	55	30 (54.5)	-24.18	(-38.60, -6.38)				

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
eGFR High and UP High	Week 6	Sparsentan	37	34 (91.9)	-46.21	(-56.95, -32.78)	0.574	(0.417, 0.789)	<0.001	*
		Irbesartan	36	32 (88.9)	-6.25	(-25.31, 17.68)				
	Week 36	Sparsentan	37	26 (70.3)	-57.31	(-66.64, -45.37)	0.544	(0.379, 0.782)	0.001	*
		Irbesartan	36	20 (55.6)	-21.57	(-39.86, 2.28)				
	Week 58	Sparsentan	37	19 (51.4)	-56.06	(-66.78, -41.87)	0.530	(0.349, 0.804)	0.003	*
		Irbesartan	36	15 (41.7)	-17.06	(-39.10, 12.95)				
eGFR High and UP Low	Week 6	Sparsentan	39	39 (100.0)	-46.90	(-56.32, -35.46)	0.604	(0.457, 0.799)	<0.001	*
		Irbesartan	37	36 (97.3)	-12.08	(-28.03, 7.41)				
	Week 36	Sparsentan	39	27 (69.2)	-58.69	(-66.98, -48.31)	0.581	(0.421, 0.801)	0.001	*
		Irbesartan	37	25 (67.6)	-28.87	(-43.53, -10.42)				
	Week 58	Sparsentan	39	23 (59.0)	-48.85	(-59.91, -34.75)	0.611	(0.425, 0.880)	0.008	*
		Irbesartan	37	17 (45.9)	-16.32	(-36.14, 9.67)				

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.020	#
< 60 mL/min/1.73 m**2	Week 6	Sparsentan	127	119 (93.7)	-35.57	(-42.18, -28.21)	0.679	(0.583, 0.791)	<0.001	*
		Irbesartan	129	124 (96.1)	-5.13	(-14.76, 5.58)				
	Week 36	Sparsentan	127	88 (69.3)	-40.53	(-47.26, -32.93)	0.666	(0.562, 0.789)	<0.001	*
		Irbesartan	129	88 (68.2)	-10.70	(-20.75, 0.64)				
	Week 58	Sparsentan	127	71 (55.9)	-42.52	(-49.71, -34.29)	0.674	(0.558, 0.815)	<0.001	*
		Irbesartan	129	70 (54.3)	-14.72	(-25.39, -2.52)				
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	48 (98.0)	-46.47	(-55.50, -35.61)	0.520	(0.399, 0.679)	<0.001	*
		Irbesartan	48	44 (91.7)	2.89	(-14.85, 24.33)				
	Week 36	Sparsentan	49	31 (63.3)	-58.39	(-66.51, -48.30)	0.439	(0.323, 0.596)	<0.001	*
		Irbesartan	48	32 (66.7)	-5.14	(-23.44, 17.54)				
	Week 58	Sparsentan	49	24 (49.0)	-51.34	(-61.92, -37.82)	0.478	(0.334, 0.685)	<0.001	*
		Irbesartan	48	19 (39.6)	1.73	(-21.80, 32.33)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)		
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	24 (92.3)	-44.00	(-55.69, -29.24)	0.785	(0.561, 1.099)	0.157
		Irbesartan	25	23 (92.0)	-28.68	(-43.87, -9.36)			
	Week 36	Sparsentan	26	20 (76.9)	-61.41	(-70.12, -50.18)	0.802	(0.540, 1.192)	0.273
		Irbesartan	25	12 (48.0)	-51.89	(-64.37, -35.02)			
	Week 58	Sparsentan	26	18 (69.2)	-56.32	(-66.65, -42.79)	0.723	(0.467, 1.121)	0.146
		Irbesartan	25	10 (40.0)	-39.62	(-57.15, -14.92)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

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Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.008	#
< 45 mL/min/1.73 m**2	Week 6	Sparsentan	82	77 (93.9)	-30.54	(-39.74, -19.94)	0.756	(0.618, 0.924)	0.007	*
		Irbesartan	80	78 (97.5)	-8.07	(-20.30, 6.04)				
	Week 36	Sparsentan	82	55 (67.1)	-36.00	(-45.40, -24.99)	0.729	(0.584, 0.911)	0.006	*
		Irbesartan	80	57 (71.3)	-12.24	(-24.94, 2.60)				
	Week 58	Sparsentan	82	45 (54.9)	-41.80	(-51.19, -30.61)	0.728	(0.568, 0.932)	0.012	*
		Irbesartan	80	45 (56.3)	-20.07	(-32.82, -4.89)				
45 to < 60 mL/min/1.73 m**2	Week 6	Sparsentan	45	42 (93.3)	-43.85	(-52.20, -34.05)	0.562	(0.449, 0.703)	<0.001	*
		Irbesartan	49	46 (93.9)	-0.04	(-14.38, 16.69)				
	Week 36	Sparsentan	45	33 (73.3)	-47.50	(-56.01, -37.34)	0.567	(0.439, 0.731)	<0.001	*
		Irbesartan	49	31 (63.3)	-7.34	(-22.66, 11.02)				
	Week 58	Sparsentan	45	26 (57.8)	-43.16	(-53.42, -30.65)	0.583	(0.438, 0.777)	<0.001	*
		Irbesartan	49	25 (51.0)	-2.53	(-20.53, 19.55)				

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

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Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	48 (98.0)	-46.47	(-55.50, -35.61)	0.520	(0.399, 0.679)	<0.001	*
		Irbesartan	48	44 (91.7)	2.89	(-14.85, 24.33)				
	Week 36	Sparsentan	49	31 (63.3)	-58.39	(-66.51, -48.30)	0.439	(0.323, 0.596)	<0.001	*
		Irbesartan	48	32 (66.7)	-5.14	(-23.44, 17.54)				
	Week 58	Sparsentan	49	24 (49.0)	-51.34	(-61.92, -37.82)	0.478	(0.334, 0.685)	<0.001	*
		Irbesartan	48	19 (39.6)	1.73	(-21.80, 32.33)				
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	24 (92.3)	-44.00	(-55.69, -29.24)	0.785	(0.561, 1.099)	0.157	
		Irbesartan	25	23 (92.0)	-28.68	(-43.87, -9.36)				
	Week 36	Sparsentan	26	20 (76.9)	-61.41	(-70.12, -50.18)	0.802	(0.540, 1.192)	0.273	
		Irbesartan	25	12 (48.0)	-51.89	(-64.37, -35.02)				
	Week 58	Sparsentan	26	18 (69.2)	-56.32	(-66.65, -42.79)	0.723	(0.467, 1.121)	0.146	
		Irbesartan	25	10 (40.0)	-39.62	(-57.15, -14.92)				

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.252	
<= 1.75 g/day	Week 6	Sparsentan	98	93 (94.9)	-34.02	(-42.13, -24.78)	0.694	(0.575, 0.837)	<0.001	*
		Irbesartan	94	89 (94.7)	-4.91	(-16.79, 8.66)				
	Week 36	Sparsentan	98	73 (74.5)	-43.10	(-50.70, -34.33)	0.634	(0.514, 0.783)	<0.001	*
		Irbesartan	94	61 (64.9)	-10.31	(-23.04, 4.51)				
	Week 58	Sparsentan	98	61 (62.2)	-39.64	(-48.34, -29.48)	0.711	(0.558, 0.905)	0.006	*
		Irbesartan	94	38 (40.4)	-15.08	(-29.45, 2.22)				
> 1.75 g/day	Week 6	Sparsentan	104	98 (94.2)	-44.08	(-50.18, -37.23)	0.618	(0.526, 0.727)	<0.001	*
		Irbesartan	108	102 (94.4)	-9.50	(-19.20, 1.36)				
	Week 36	Sparsentan	104	66 (63.5)	-53.13	(-58.95, -46.48)	0.585	(0.486, 0.703)	<0.001	*
		Irbesartan	108	71 (65.7)	-19.84	(-29.44, -8.94)				
	Week 58	Sparsentan	104	52 (50.0)	-53.14	(-59.68, -45.54)	0.559	(0.455, 0.686)	<0.001	*
		Irbesartan	108	61 (56.5)	-16.12	(-27.07, -3.52)				

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024



Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.576	
Yes	Week 6	Sparsentan	88	82 (93.2)	-40.00	(-47.34, -31.64)	0.648	(0.537, 0.782)	<0.001	*
		Irbesartan	83	76 (91.6)	-7.36	(-19.08, 6.06)				
	Week 36	Sparsentan	88	57 (64.8)	-46.36	(-53.89, -37.61)	0.602	(0.486, 0.745)	<0.001	*
		Irbesartan	83	58 (69.9)	-10.91	(-23.33, 3.53)				
	Week 58	Sparsentan	88	46 (52.3)	-40.65	(-49.91, -29.68)	0.748	(0.589, 0.950)	0.017	*
		Irbesartan	83	46 (55.4)	-20.64	(-32.97, -6.05)				
No	Week 6	Sparsentan	114	109 (95.6)	-38.54	(-45.32, -30.92)	0.666	(0.565, 0.784)	<0.001	*
		Irbesartan	119	115 (96.6)	-7.65	(-17.57, 3.46)				
	Week 36	Sparsentan	114	82 (71.9)	-49.10	(-55.21, -42.15)	0.635	(0.529, 0.762)	<0.001	*
		Irbesartan	119	74 (62.2)	-19.80	(-29.60, -8.63)				
	Week 58	Sparsentan	114	67 (58.8)	-49.88	(-56.43, -42.36)	0.565	(0.460, 0.694)	<0.001	*
		Irbesartan	119	53 (44.5)	-11.33	(-23.72, 3.08)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Time since renal biopsy Overall		Sparsentan						Interaction:	0.210	
<= 5 years	Week 6	Sparsentan	113	104 (92.0)	-40.96	(-47.69, -33.37)	0.659	(0.558, 0.777)	<0.001	*
		Irbesartan	127	121 (95.3)	-10.38	(-19.95, 0.35)				
	Week 36	Sparsentan	113	73 (64.6)	-50.87	(-57.13, -43.69)	0.642	(0.531, 0.775)	<0.001	*
		Irbesartan	127	79 (62.2)	-23.43	(-32.78, -12.78)				
	Week 58	Sparsentan	113	58 (51.3)	-43.60	(-51.64, -34.22)	0.755	(0.610, 0.936)	0.010	*
		Irbesartan	127	60 (47.2)	-25.33	(-35.69, -13.31)				
> 5 years	Week 6	Sparsentan	89	87 (97.8)	-37.40	(-44.70, -29.12)	0.637	(0.529, 0.766)	<0.001	*
		Irbesartan	75	70 (93.3)	-1.71	(-14.28, 12.71)				
	Week 36	Sparsentan	89	66 (74.2)	-44.36	(-51.53, -36.12)	0.566	(0.461, 0.694)	<0.001	*
		Irbesartan	75	53 (70.7)	-1.63	(-15.44, 14.43)				
	Week 58	Sparsentan	89	55 (61.8)	-48.75	(-55.89, -40.46)	0.496	(0.395, 0.623)	<0.001	*
		Irbesartan	75	39 (52.0)	3.35	(-12.99, 22.76)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT1RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
History of hypertension Overall		Sparsentan						Interaction:	0.196	
Yes	Week 6	Sparsentan	153	143 (93.5)	-37.84	(-43.71, -31.35)	0.679	(0.591, 0.781)	<0.001	*
		Irbesartan	157	146 (93.0)	-8.46	(-17.00, 0.96)				
	Week 36	Sparsentan	153	104 (68.0)	-45.14	(-50.92, -38.68)	0.644	(0.551, 0.753)	<0.001	*
		Irbesartan	157	105 (66.9)	-14.80	(-23.68, -4.88)				
	Week 58	Sparsentan	153	83 (54.2)	-42.85	(-49.52, -35.30)	0.679	(0.569, 0.809)	<0.001	*
		Irbesartan	157	80 (51.0)	-15.78	(-25.68, -4.57)				
No	Week 6	Sparsentan	49	48 (98.0)	-43.42	(-52.90, -32.03)	0.595	(0.456, 0.777)	<0.001	*
		Irbesartan	45	45 (100.0)	-4.89	(-21.43, 15.12)				
	Week 36	Sparsentan	49	35 (71.4)	-55.69	(-63.88, -45.64)	0.556	(0.410, 0.755)	<0.001	*
		Irbesartan	45	27 (60.0)	-20.36	(-36.33, -0.38)				
	Week 58	Sparsentan	49	30 (61.2)	-54.60	(-63.63, -43.31)	0.538	(0.381, 0.760)	<0.001	*
		Irbesartan	45	19 (42.2)	-15.67	(-35.12, 9.60)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 56 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 58 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

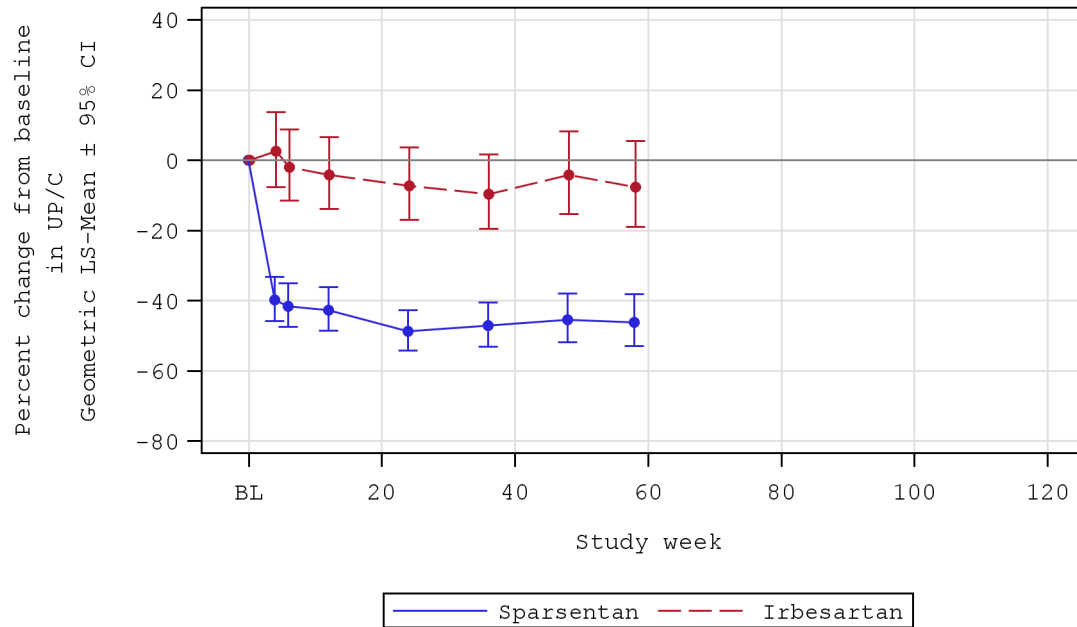
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

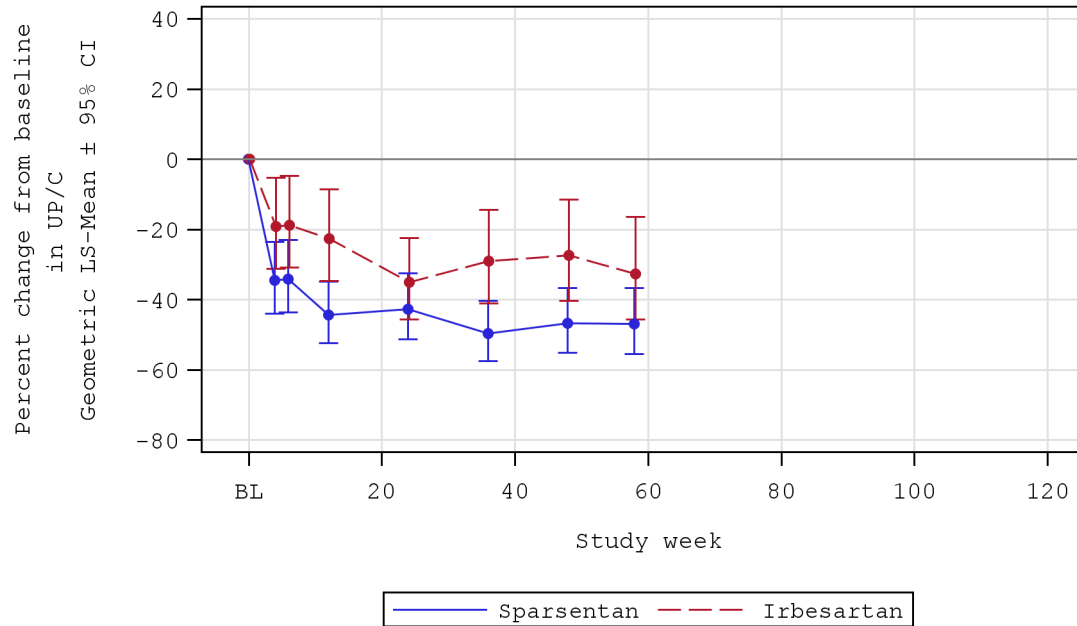
Figure PF1RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Sex: Male



Sparsentan	132	92	69
Irbesartan	137	97	73

An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data. Number of available records are presented for key visits up to Week 58 only.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 UP/C = urine protein/creatinine ratio. Reference table: PT1RPR\_FSLM

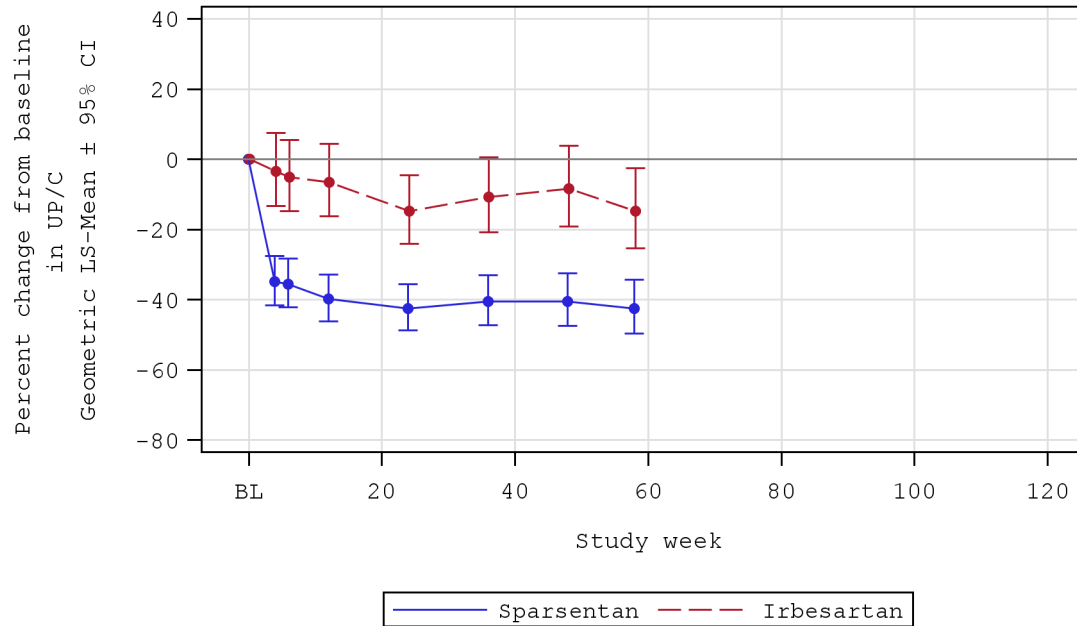
Figure PF1RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Sex: Female



Sparsentan	59	47	44
Irbesartan	54	35	26

An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data. Number of available records are presented for key visits up to Week 58 only.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 UP/C = urine protein/creatinine ratio. Reference table: PT1RPR\_FSLM

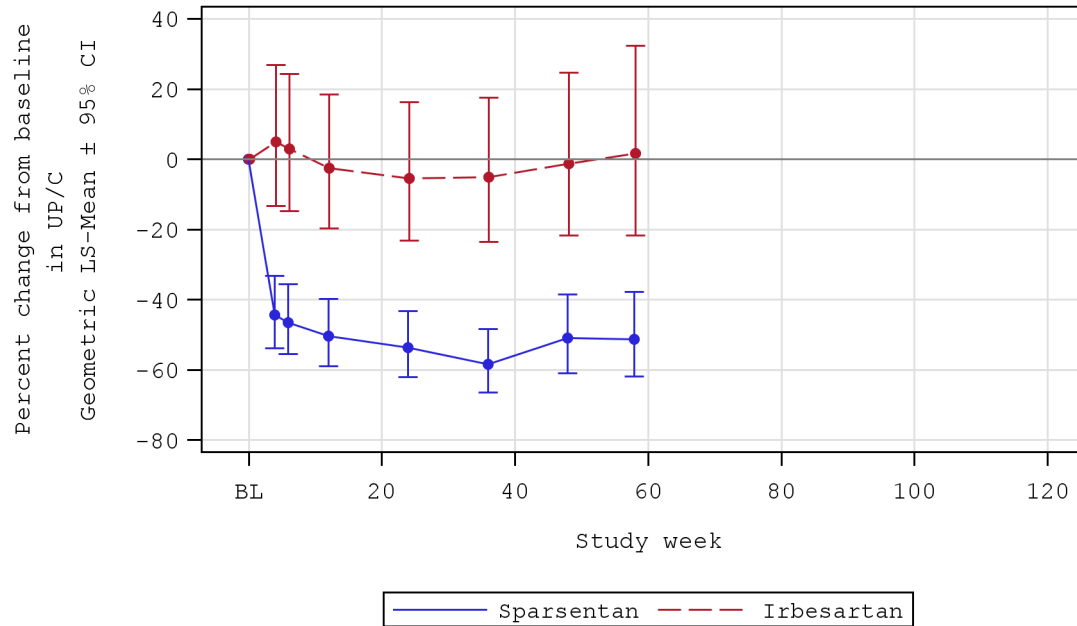
Figure PF1RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1: < 60 mL/min/1.73 m\*\*2



Sparsentan	119	88	71
Irbesartan	124	88	70

An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data. Number of available records are presented for key visits up to Week 58 only.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 UP/C = urine protein/creatinine ratio. Reference table: PT1RPR\_FSLM

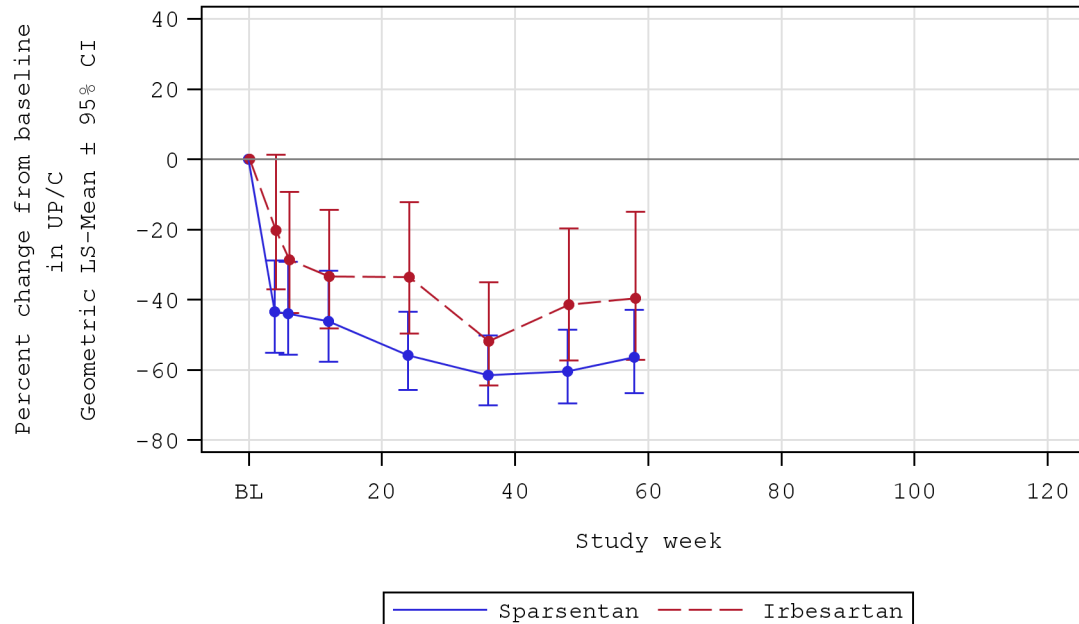
Figure PF1RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1: 60 to < 90 mL/min/1.73 m\*\*2



Sparsentan	48	31	24
Irbesartan	44	32	19

An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data. Number of available records are presented for key visits up to Week 58 only.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 UP/C = urine protein/creatinine ratio. Reference table: PT1RPR\_FSLM

Figure PF1RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1:  $\geq 90$  mL/min/1.73 m<sup>2</sup>

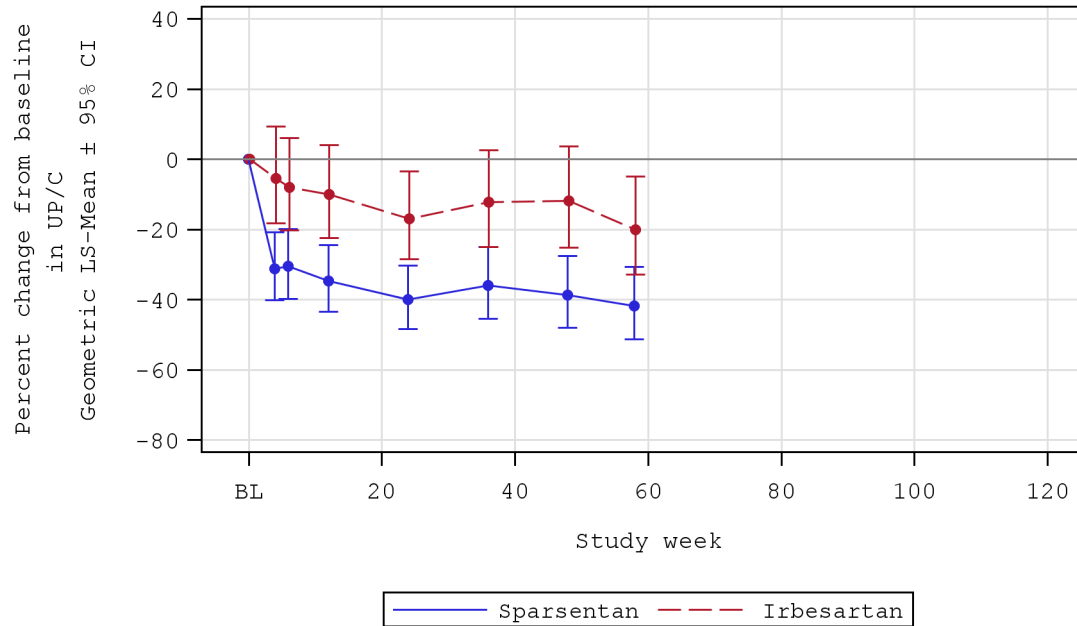


Sparsentan	24	20	18
Irbesartan	23	12	10

An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data. Number of available records are presented for key visits up to Week 58 only.  
 eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High  $\geq 60$  ml/min/1.73m<sup>2</sup>, UP Low =  $\leq 1.75$  g/day, UP High =  $> 1.75$  g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 UP/C = urine protein/creatinine ratio. Reference table: PT1RPR\_FSLM



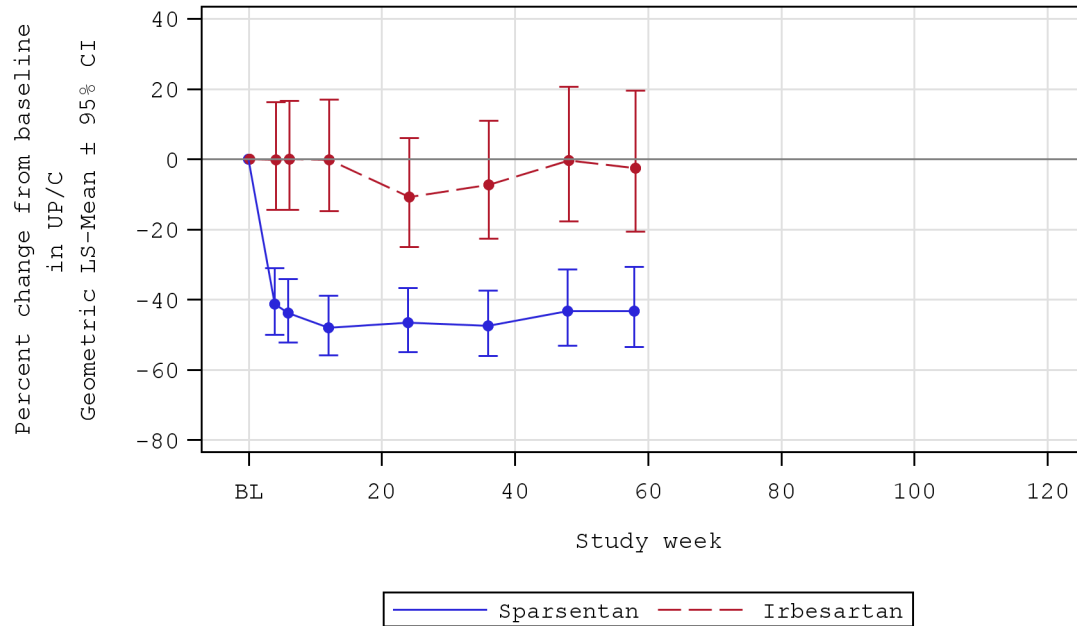
Figure PF1RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: < 45 mL/min/1.73 m\*\*2



Sparsentan	77	55	45
Irbesartan	78	57	45

An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data. Number of available records are presented for key visits up to Week 58 only.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 UP/C = urine protein/creatinine ratio. Reference table: PT1RPR\_FSLM

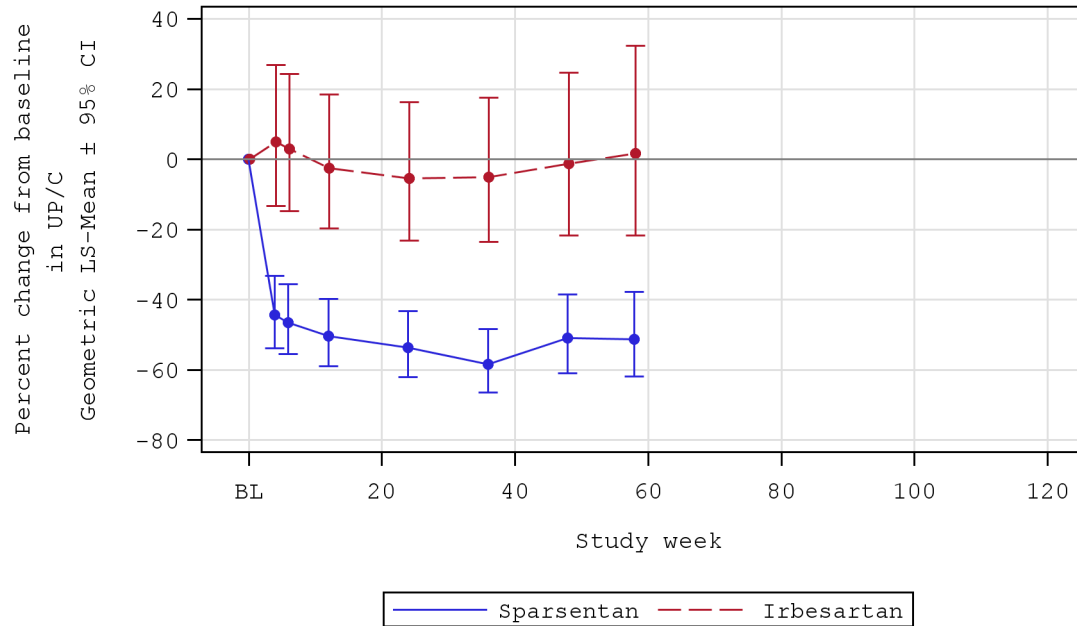
Figure PF1RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: 45 to < 60 mL/min/1.73 m\*\*2



Sparsentan	42	33	26
Irbesartan	46	31	25

An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data. Number of available records are presented for key visits up to Week 58 only.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 UP/C = urine protein/creatinine ratio. Reference table: PT1RPR\_FSLM

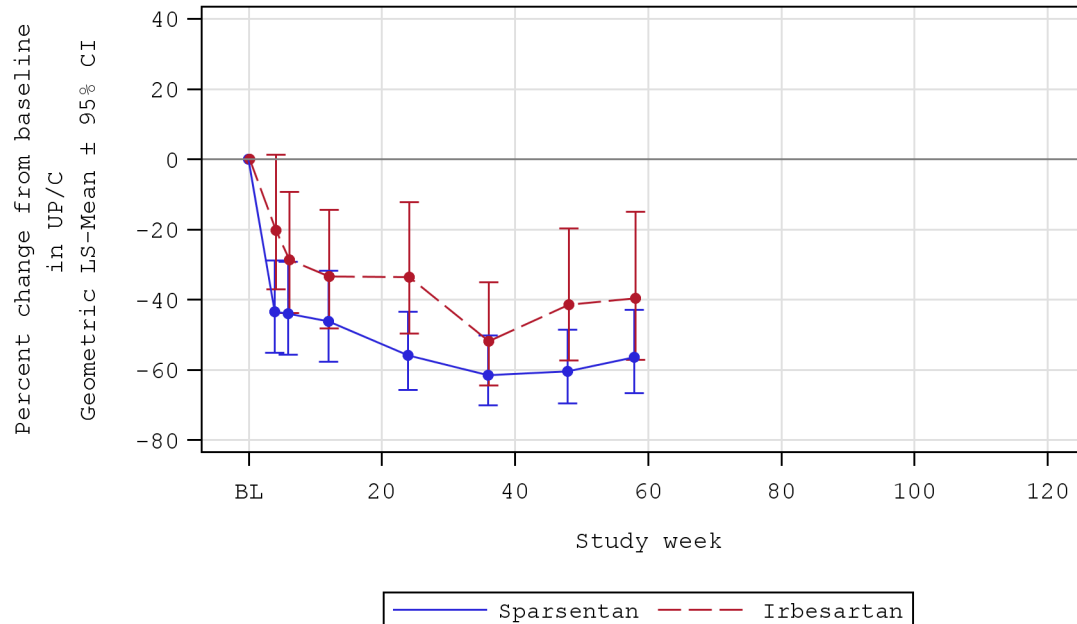
Figure PF1RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: 60 to < 90 mL/min/1.73 m\*\*2



Sparsentan	48	31	24
Irbesartan	44	32	19

An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data. Number of available records are presented for key visits up to Week 58 only.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 UP/C = urine protein/creatinine ratio. Reference table: PT1RPR\_FSLM

Figure PF1RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2:  $\geq 90$  mL/min/1.73 m<sup>2</sup>



Sparsentan	24	20	18
Irbesartan	23	12	10

An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data. Number of available records are presented for key visits up to Week 58 only.  
 eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High  $\geq 60$  ml/min/1.73m<sup>2</sup>, UP Low =  $\leq 1.75$  g/day, UP High =  $> 1.75$  g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 UP/C = urine protein/creatinine ratio. Reference table: PT1RPR\_FSLM

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Sex													
Male	Urine protein excretion	Baseline	Sparsentan	139	139 (100.0)	2.28 (1.37)	0.1	1.21	1.96	3.12	7.2		
			Irbesartan	143	143 (100.0)	2.25 (1.33)	0.5	1.34	1.85	2.85	7.5		
		Week 6	Sparsentan	139	132 (95.0)	1.59 (1.34)	0.2	0.63	1.17	2.08	6.5		
			Irbesartan	143	137 (95.8)	2.38 (1.69)	0.4	1.28	1.89	3.07	10.3		
		Week 36	Sparsentan	139	92 (66.2)	1.71 (1.78)	0.1	0.57	0.99	2.30	9.4		
			Irbesartan	143	98 (68.5)	2.26 (1.73)	0.2	1.02	1.87	2.84	9.2		
		Week 58	Sparsentan	139	69 (49.6)	1.70 (1.70)	0.1	0.49	1.21	2.11	7.5		
			Irbesartan	143	73 (51.0)	2.39 (1.77)	0.1	1.08	2.06	3.06	8.6		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	139	132 (95.0)	-0.75 (1.19)	-4.2	-1.41	-0.72	-0.11	5.2	-0.66 [-0.91, -0.41]
			Irbesartan	143	137 (95.8)	0.11 (1.39)	-3.7	-0.54	-0.11	0.72	8.4		
			Week 36	Sparsentan	139	92 (66.2)	-0.61 (1.36)	-4.1	-1.36	-0.67	-0.07	4.6	-0.40 [-0.69, -0.11]
				Irbesartan	143	98 (68.5)	-0.03 (1.55)	-3.3	-0.92	-0.21	0.65	6.5	
			Week 58	Sparsentan	139	69 (49.6)	-0.60 (1.43)	-4.4	-1.30	-0.62	0.07	3.3	-0.36 [-0.70, -0.03]
				Irbesartan	143	73 (51.0)	-0.06 (1.53)	-2.7	-0.89	-0.34	0.46	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Female	Urine protein excretion	Baseline	Sparsentan	63	63 (100.0)	2.09 (2.14)	0.5	1.15	1.45	2.32	14.7		
			Irbesartan	59	59 (100.0)	1.94 (0.97)	0.6	1.33	1.77	2.33	5.7		
		Week 6	Sparsentan	63	60 (95.2)	1.28 (0.83)	0.2	0.69	1.03	1.72	3.9		
			Irbesartan	59	54 (91.5)	1.67 (1.15)	0.4	0.93	1.36	2.24	7.6		
		Week 36	Sparsentan	63	47 (74.6)	1.02 (0.74)	0.1	0.56	0.80	1.32	3.5		
			Irbesartan	59	35 (59.3)	1.73 (1.43)	0.2	0.80	1.20	2.44	7.1		
		Week 58	Sparsentan	63	44 (69.8)	1.05 (0.70)	0.1	0.63	0.92	1.35	3.4		
			Irbesartan	59	25 (42.4)	1.42 (1.34)	0.2	0.65	1.03	1.94	6.8		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	63	60 (95.2)	-0.67 (1.84)	-13.0	-0.89	-0.45	-0.01	1.8	-0.29 [-0.66, 0.08]
				Irbesartan	59	54 (91.5)	-0.24 (0.86)	-1.8	-0.81	-0.19	0.12	3.0	
	Week 36		Sparsentan	63	47 (74.6)	-0.71 (1.04)	-4.2	-1.06	-0.66	-0.20	1.7	-0.50 [-0.95, -0.06]	
			Irbesartan	59	35 (59.3)	-0.17 (1.12)	-2.0	-0.91	-0.29	0.18	2.6		
	Week 58	Sparsentan	63	44 (69.8)	-0.66 (1.04)	-3.7	-1.04	-0.59	-0.02	1.2	-0.16 [-0.65, 0.33]		
		Irbesartan	59	25 (42.4)	-0.50 (0.89)	-2.1	-0.99	-0.55	-0.18	2.3			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Age													
<= 45 years	Urine protein excretion	Baseline	Sparsentan	96	96 (100.0)	2.39 (2.00)	0.4	1.23	1.81	2.91	14.7		
			Irbesartan	99	99 (100.0)	2.18 (1.06)	0.5	1.41	1.92	2.70	5.9		
		Week 6	Sparsentan	96	91 (94.8)	1.45 (1.18)	0.2	0.60	1.07	1.95	6.2		
			Irbesartan	99	95 (96.0)	2.19 (1.63)	0.4	1.08	1.84	2.67	10.3		
		Week 36	Sparsentan	96	66 (68.8)	1.52 (1.49)	0.1	0.55	0.92	2.26	8.3		
			Irbesartan	99	65 (65.7)	2.08 (1.81)	0.2	0.81	1.45	2.74	9.2		
		Week 58	Sparsentan	96	51 (53.1)	1.56 (1.61)	0.1	0.57	0.96	2.11	7.5		
			Irbesartan	99	44 (44.4)	2.06 (1.94)	0.1	0.66	1.42	2.47	8.6		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	96	91 (94.8)	-0.90 (1.69)	-13.0	-1.29	-0.71	-0.03	2.1	-0.57 [-0.86, -0.27]
			Irbesartan	99	95 (96.0)	-0.02 (1.40)	-3.2	-0.70	-0.25	0.46	8.4		
			Week 36	Sparsentan	96	66 (68.8)	-0.62 (1.19)	-4.1	-1.09	-0.66	0.15	2.6	-0.33 [-0.67, 0.02]
				Irbesartan	99	65 (65.7)	-0.16 (1.61)	-3.3	-1.10	-0.32	0.49	6.5	
			Week 58	Sparsentan	96	51 (53.1)	-0.59 (1.42)	-4.2	-1.29	-0.46	0.17	3.3	-0.21 [-0.61, 0.20]
				Irbesartan	99	44 (44.4)	-0.26 (1.69)	-2.7	-1.07	-0.58	0.12	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 45 years	Urine protein excretion	Baseline	Sparsentan	106	106 (100.0)	2.08 (1.24)	0.1	1.12	1.76	2.81	6.3	
			Irbesartan	103	103 (100.0)	2.15 (1.40)	0.6	1.25	1.77	2.43	7.5	
		Week 6	Sparsentan	106	101 (95.3)	1.53 (1.24)	0.2	0.73	1.09	2.01	6.5	
			Irbesartan	103	96 (93.2)	2.17 (1.56)	0.4	1.09	1.65	2.74	9.5	
		Week 36	Sparsentan	106	73 (68.9)	1.43 (1.59)	0.1	0.60	0.91	1.59	9.4	
			Irbesartan	103	68 (66.0)	2.16 (1.52)	0.2	1.12	1.78	2.64	7.7	
	Week 58	Sparsentan	106	62 (58.5)	1.35 (1.26)	0.1	0.57	1.14	1.62	6.8		
		Irbesartan	103	54 (52.4)	2.20 (1.52)	0.1	1.05	1.99	2.94	7.2		
	Change from baseline in urine protein excretion	Week 6	Sparsentan	106	101 (95.3)	-0.56 (1.12)	-3.8	-1.05	-0.57	-0.10	5.2	-0.54 [-0.82, -0.25]
			Irbesartan	103	96 (93.2)	0.04 (1.14)	-3.7	-0.54	-0.05	0.62	4.2	
		Week 36	Sparsentan	106	73 (68.9)	-0.67 (1.32)	-4.2	-1.27	-0.68	-0.27	4.6	-0.53 [-0.87, -0.19]
			Irbesartan	103	68 (66.0)	0.02 (1.28)	-2.5	-0.77	-0.13	0.59	4.2	
		Week 58	Sparsentan	106	62 (58.5)	-0.66 (1.17)	-4.4	-1.27	-0.65	-0.02	2.4	-0.48 [-0.85, -0.11]
			Irbesartan	103	54 (52.4)	-0.10 (1.14)	-2.1	-0.83	-0.27	0.39	3.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age at IgAN diagnosis												
<= 18 years	Urine protein excretion	Baseline	Sparsentan	9	9 (100.0)	3.85 (2.41)	1.2	2.78	3.07	4.00	9.7	
			Irbesartan	5	5 (100.0)	1.91 (1.00)	0.6	1.22	2.14	2.56	3.1	
		Week 6	Sparsentan	9	8 (88.9)	2.24 (1.77)	0.4	0.81	1.66	3.84	4.9	
			Irbesartan	5	5 (100.0)	2.01 (1.42)	0.6	1.18	1.73	2.27	4.3	
		Week 36	Sparsentan	9	6 (66.7)	2.63 (1.60)	0.6	1.50	2.62	3.82	4.6	
			Irbesartan	5	2 (40.0)	2.02 (1.30)	1.1	1.11	2.02	2.94	2.9	
	Week 58	Sparsentan	9	4 (44.4)	2.17 (2.17)	0.1	0.49	1.78	3.84	5.0		
		Irbesartan	5	1 (20.0)	0.57	0.6	0.57	0.57	0.57	0.6		
	Change from baseline in urine protein excretion	Week 6	Sparsentan	9	8 (88.9)	-0.87 (1.57)	-2.5	-2.18	-1.03	-0.11	2.1	-0.66 [-1.80, 0.49]
			Irbesartan	5	5 (100.0)	0.09 (1.30)	-1.9	-0.02	0.13	0.51	1.7	
		Week 36	Sparsentan	9	6 (66.7)	-0.61 (1.48)	-3.1	-1.42	-0.32	0.60	0.9	-0.03 [-1.63, 1.57]
			Irbesartan	5	2 (40.0)	-0.57 (1.94)	-1.9	-1.95	-0.57	0.80	0.8	
Week 58		Sparsentan	9	4 (44.4)	-1.04 (2.00)	-3.7	-2.42	-0.70	0.35	1.0	NE	
		Irbesartan	5	1 (20.0)	-2.49	-2.5	-2.49	-2.49	-2.49	-2.5		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 18 to 40 years	Urine protein excretion	Baseline	Sparsentan	102	102 (100.0)	2.26 (1.79)	0.4	1.21	1.81	2.81	14.7			
			Irbesartan	109	109 (100.0)	2.21 (1.08)	0.5	1.42	1.96	2.70	5.9			
		Week 6	Sparsentan	102	98 (96.1)	1.40 (1.13)	0.2	0.60	1.06	1.80	6.2			
			Irbesartan	109	105 (96.3)	2.16 (1.48)	0.4	1.18	1.82	2.61	7.7			
		Week 36	Sparsentan	102	69 (67.6)	1.52 (1.68)	0.1	0.55	0.90	2.14	9.4			
			Irbesartan	109	73 (67.0)	2.00 (1.58)	0.2	1.01	1.68	2.52	9.2			
		Week 58	Sparsentan	102	58 (56.9)	1.48 (1.62)	0.1	0.57	0.92	1.86	7.5			
			Irbesartan	109	51 (46.8)	1.99 (1.54)	0.1	0.80	1.69	2.50	6.8			
		Change from baseline in urine protein excretion	Week 6	Sparsentan	102	98 (96.1)	-0.90 (1.60)	-13.0	-1.22	-0.69	-0.22	1.8	-0.59 [-0.87, -0.31]	
				Irbesartan	109	105 (96.3)	-0.07 (1.16)	-3.2	-0.68	-0.25	0.44	4.2		
				Week 36	Sparsentan	102	69 (67.6)	-0.61 (1.31)	-4.1	-1.28	-0.72	-0.03	4.6	-0.22 [-0.55, 0.11]
					Irbesartan	109	73 (67.0)	-0.30 (1.48)	-3.3	-1.17	-0.36	0.31	6.5	
				Week 58	Sparsentan	102	58 (56.9)	-0.66 (1.35)	-4.2	-1.34	-0.73	-0.03	3.3	-0.16 [-0.54, 0.21]
					Irbesartan	109	51 (46.8)	-0.45 (1.25)	-2.7	-1.44	-0.58	0.02	3.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	Urine protein excretion	Baseline	Sparsentan	91	91 (100.0)	2.02 (1.29)	0.1	1.06	1.54	2.70	6.3	
			Irbesartan	88	88 (100.0)	2.11 (1.43)	0.6	1.27	1.66	2.37	7.5	
		Week 6	Sparsentan	91	86 (94.5)	1.53 (1.23)	0.2	0.72	1.13	2.12	6.5	
			Irbesartan	88	81 (92.0)	2.22 (1.75)	0.4	1.03	1.66	2.78	10.3	
		Week 36	Sparsentan	91	64 (70.3)	1.32 (1.34)	0.1	0.59	0.87	1.46	6.4	
			Irbesartan	88	58 (65.9)	2.27 (1.78)	0.2	0.88	1.63	3.30	7.7	
	Week 58	Sparsentan	91	51 (56.0)	1.35 (1.10)	0.1	0.57	1.23	1.67	4.8		
		Irbesartan	88	46 (52.3)	2.34 (1.89)	0.1	0.98	2.02	3.08	8.6		
	Change from baseline in urine protein excretion	Week 6	Sparsentan	91	86 (94.5)	-0.51 (1.15)	-3.8	-1.03	-0.51	-0.03	5.2	-0.49 [-0.80, -0.18]
			Irbesartan	88	81 (92.0)	0.12 (1.40)	-3.7	-0.54	0.02	0.62	8.4	
		Week 36	Sparsentan	91	64 (70.3)	-0.69 (1.20)	-4.2	-1.17	-0.66	-0.27	2.6	-0.72 [-1.09, -0.36]
			Irbesartan	88	58 (65.9)	0.23 (1.37)	-2.1	-0.62	-0.11	0.98	5.1	
		Week 58	Sparsentan	91	51 (56.0)	-0.55 (1.17)	-4.4	-1.23	-0.58	0.12	2.4	-0.55 [-0.96, -0.15]
			Irbesartan	88	46 (52.3)	0.18 (1.49)	-2.1	-0.58	-0.11	0.58	6.7	

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95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Geographic region													
North America	Urine protein excretion	Baseline	Sparsentan	35	35 (100.0)	2.73 (2.53)	0.1	1.18	2.19	3.14	14.7		
			Irbesartan	46	46 (100.0)	2.53 (1.47)	0.6	1.67	2.14	3.00	7.5		
		Week 6	Sparsentan	35	33 (94.3)	1.99 (1.37)	0.3	1.09	1.71	2.64	6.5		
			Irbesartan	46	43 (93.5)	2.69 (1.73)	0.5	1.41	2.45	3.78	7.7		
		Week 36	Sparsentan	35	18 (51.4)	1.55 (1.23)	0.3	0.69	1.00	2.50	4.0		
			Irbesartan	46	25 (54.3)	2.87 (1.71)	0.5	1.84	2.68	4.02	7.1		
		Week 58	Sparsentan	35	12 (34.3)	1.57 (1.28)	0.5	0.66	1.14	1.90	4.7		
			Irbesartan	46	18 (39.1)	3.18 (2.02)	0.5	1.46	2.87	4.61	7.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	35	33 (94.3)	-0.88 (2.75)	-13.0	-1.65	-0.47	0.20	5.2	-0.46 [-0.92, -0.01]
			Irbesartan	46	43 (93.5)	0.13 (1.57)	-3.7	-0.54	-0.13	0.96	4.2		
	Week 36		Sparsentan	35	18 (51.4)	-0.80 (1.51)	-4.2	-1.54	-0.53	-0.11	2.6	-0.62 [-1.24, -0.00]	
			Irbesartan	46	25 (54.3)	0.20 (1.66)	-2.5	-0.50	-0.06	0.93	4.2		
	Week 58		Sparsentan	35	12 (34.3)	-0.42 (1.76)	-3.7	-1.21	-0.11	0.39	3.3	-0.35 [-1.08, 0.39]	
			Irbesartan	46	18 (39.1)	0.21 (1.83)	-2.7	-0.88	-0.29	1.24	3.7		

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Europe	Urine protein excretion	Baseline	Sparsentan	98	98 (100.0)	2.17 (1.40)	0.4	1.21	1.86	2.86	9.7			
			Irbesartan	115	115 (100.0)	2.12 (1.17)	0.5	1.32	1.82	2.60	5.9			
		Week 6	Sparsentan	98	91 (92.9)	1.36 (1.03)	0.2	0.66	1.08	1.95	5.5			
			Irbesartan	115	109 (94.8)	2.06 (1.51)	0.5	1.09	1.65	2.56	10.3			
		Week 36	Sparsentan	98	60 (61.2)	1.53 (1.47)	0.1	0.60	1.01	2.07	6.4			
			Irbesartan	115	78 (67.8)	1.93 (1.42)	0.2	0.81	1.55	2.52	7.0			
		Week 58	Sparsentan	98	48 (49.0)	1.50 (1.43)	0.1	0.49	1.10	1.96	7.5			
			Irbesartan	115	56 (48.7)	1.99 (1.54)	0.1	0.88	1.72	2.75	8.6			
		Change from baseline in urine protein excretion	Week 6	Sparsentan	98	91 (92.9)	-0.74 (1.07)	-4.2	-1.21	-0.70	-0.12	2.7	-0.60 [-0.88, -0.31]	
					Irbesartan	115	109 (94.8)	-0.03 (1.26)	-2.4	-0.77	-0.12	0.44	8.4	
				Week 36	Sparsentan	98	60 (61.2)	-0.56 (1.25)	-4.1	-1.18	-0.70	0.10	2.6	-0.28 [-0.62, 0.05]
					Irbesartan	115	78 (67.8)	-0.19 (1.36)	-3.3	-1.10	-0.29	0.63	5.1	
				Week 58	Sparsentan	98	48 (49.0)	-0.49 (1.27)	-4.2	-1.14	-0.75	0.14	2.5	-0.17 [-0.55, 0.22]
					Irbesartan	115	56 (48.7)	-0.26 (1.40)	-2.5	-0.99	-0.39	0.14	6.7	

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Asia Pacific	Urine protein excretion	Baseline	Sparsentan	69	69 (100.0)	2.04 (1.36)	0.4	1.15	1.49	2.51	7.2			
			Irbesartan	41	41 (100.0)	1.87 (1.08)	0.6	1.25	1.59	2.29	6.8			
		Week 6	Sparsentan	69	68 (98.6)	1.43 (1.32)	0.2	0.61	0.95	1.79	6.2			
			Irbesartan	41	39 (95.1)	1.93 (1.56)	0.4	0.93	1.55	2.64	9.5			
		Week 36	Sparsentan	69	61 (88.4)	1.39 (1.70)	0.1	0.50	0.83	1.74	9.4			
			Irbesartan	41	30 (73.2)	1.99 (2.06)	0.2	0.73	1.23	2.48	9.2			
		Week 58	Sparsentan	69	53 (76.8)	1.37 (1.48)	0.1	0.57	0.88	1.53	6.8			
			Irbesartan	41	24 (58.5)	1.73 (1.62)	0.1	0.68	1.08	2.27	6.6			
		Change from baseline in urine protein excretion	Week 6	Sparsentan	69	68 (98.6)	-0.62 (0.78)	-2.4	-1.01	-0.61	-0.10	0.9	-0.78 [-1.18, -0.37]	
				Irbesartan	41	39 (95.1)	0.02 (0.88)	-1.7	-0.50	-0.20	0.42	2.8		
				Week 36	Sparsentan	69	61 (88.4)	-0.69 (1.19)	-3.1	-1.23	-0.71	-0.27	4.6	-0.54 [-0.99, -0.10]
					Irbesartan	41	30 (73.2)	0.02 (1.50)	-2.0	-0.75	-0.28	0.02	6.5	
				Week 58	Sparsentan	69	53 (76.8)	-0.79 (1.19)	-4.4	-1.44	-0.59	-0.04	2.0	-0.48 [-0.97, 0.01]
					Irbesartan	41	24 (58.5)	-0.25 (1.01)	-1.7	-0.91	-0.42	0.06	2.2	

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline BMI												
< 27 kg/m**2	Urine protein excretion	Baseline	Sparsentan	84	84 (100.0)	1.88 (1.02)	0.4	1.15	1.58	2.40	5.0	
			Irbesartan	94	94 (100.0)	2.13 (1.13)	0.6	1.37	1.81	2.56	6.8	
		Week 6	Sparsentan	84	77 (91.7)	1.33 (1.19)	0.2	0.64	0.95	1.60	6.5	
			Irbesartan	94	90 (95.7)	2.07 (1.57)	0.5	1.06	1.56	2.43	9.5	
		Week 36	Sparsentan	84	59 (70.2)	1.20 (1.20)	0.1	0.50	0.79	1.47	6.3	
			Irbesartan	94	59 (62.8)	1.79 (1.39)	0.2	0.81	1.43	2.52	7.7	
	Week 58	Sparsentan	84	46 (54.8)	0.96 (0.90)	0.1	0.30	0.74	1.30	5.0		
		Irbesartan	94	40 (42.6)	1.70 (1.40)	0.1	0.78	1.22	2.37	6.6		
	Change from baseline in urine protein excretion	Week 6	Sparsentan	84	77 (91.7)	-0.56 (1.12)	-2.4	-1.19	-0.65	-0.10	5.2	-0.43 [-0.74, -0.12]
			Irbesartan	94	90 (95.7)	-0.10 (1.04)	-2.4	-0.73	-0.29	0.42	2.9	
		Week 36	Sparsentan	84	59 (70.2)	-0.65 (1.06)	-3.1	-1.23	-0.65	-0.07	2.2	-0.18 [-0.54, 0.18]
			Irbesartan	94	59 (62.8)	-0.47 (0.99)	-3.3	-1.07	-0.44	0.26	1.2	
Week 58		Sparsentan	84	46 (54.8)	-0.93 (1.01)	-3.7	-1.36	-0.80	-0.30	1.0	-0.30 [-0.73, 0.13]	
		Irbesartan	94	40 (42.6)	-0.64 (0.87)	-2.5	-1.29	-0.53	-0.17	1.5		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 27 kg/m**2	Urine protein excretion	Baseline	Sparsentan	118	118 (100.0)	2.47 (1.95)	0.1	1.18	1.92	3.16	14.7		
			Irbesartan	107	107 (100.0)	2.20 (1.33)	0.6	1.33	1.82	2.78	7.5		
		Week 6	Sparsentan	118	115 (97.5)	1.60 (1.22)	0.2	0.69	1.20	2.32	6.2		
			Irbesartan	107	100 (93.5)	2.29 (1.61)	0.4	1.18	1.92	2.80	10.3		
		Week 36	Sparsentan	118	80 (67.8)	1.67 (1.73)	0.1	0.60	1.04	2.21	9.4		
			Irbesartan	107	73 (68.2)	2.41 (1.82)	0.2	1.12	1.96	3.35	9.2		
		Week 58	Sparsentan	118	67 (56.8)	1.78 (1.62)	0.1	0.65	1.21	2.18	7.5		
			Irbesartan	107	58 (54.2)	2.44 (1.85)	0.1	0.95	2.03	3.13	8.6		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	118	115 (97.5)	-0.83 (1.59)	-13.0	-1.28	-0.67	-0.09	2.7	-0.61 [-0.89, -0.34]
				Irbesartan	107	100 (93.5)	0.11 (1.45)	-3.7	-0.54	-0.05	0.62	8.4	
			Week 36	Sparsentan	118	80 (67.8)	-0.64 (1.39)	-4.2	-1.22	-0.70	-0.08	4.6	-0.58 [-0.91, -0.26]
				Irbesartan	107	73 (68.2)	0.25 (1.68)	-2.5	-0.78	-0.12	1.00	6.5	
			Week 58	Sparsentan	118	67 (56.8)	-0.42 (1.42)	-4.4	-1.19	-0.45	0.26	3.3	-0.38 [-0.73, -0.02]
				Irbesartan	107	58 (54.2)	0.15 (1.61)	-2.7	-0.74	-0.22	1.01	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Randomization strata													
eGFR Low and UP High	Urine protein excretion	Baseline	Sparsentan	71	71 (100.0)	3.07 (1.58)	0.1	1.96	2.81	3.74	9.7		
			Irbesartan	74	74 (100.0)	2.96 (1.43)	0.5	1.86	2.59	3.77	7.5		
		Week 6	Sparsentan	71	67 (94.4)	2.10 (1.20)	0.5	1.24	1.88	2.51	5.7		
			Irbesartan	74	70 (94.6)	2.98 (1.63)	0.8	1.89	2.66	3.75	7.7		
		Week 36	Sparsentan	71	49 (69.0)	2.20 (1.87)	0.3	0.87	1.49	2.89	9.4		
			Irbesartan	74	48 (64.9)	2.84 (1.63)	0.2	1.81	2.56	3.56	7.1		
		Week 58	Sparsentan	71	42 (59.2)	1.98 (1.67)	0.2	0.81	1.49	2.47	7.5		
			Irbesartan	74	36 (48.6)	2.93 (1.68)	0.1	1.97	2.73	3.43	7.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	71	67 (94.4)	-0.92 (1.29)	-4.2	-1.65	-0.85	0.09	1.9	-0.68 [-1.03, -0.34]
				Irbesartan	74	70 (94.6)	0.03 (1.47)	-3.7	-0.84	-0.08	0.73	4.2	
			Week 36	Sparsentan	71	49 (69.0)	-0.80 (1.68)	-4.2	-1.64	-0.96	0.21	4.6	-0.41 [-0.81, -0.01]
				Irbesartan	74	48 (64.9)	-0.13 (1.60)	-3.3	-1.32	-0.09	1.00	4.2	
			Week 58	Sparsentan	71	42 (59.2)	-0.82 (1.61)	-4.4	-1.80	-0.83	-0.01	2.5	-0.43 [-0.88, 0.02]
				Irbesartan	74	36 (48.6)	-0.16 (1.41)	-2.7	-1.30	-0.09	0.73	3.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR Low and UP Low	Urine protein excretion	Baseline	Sparsentan	55	55 (100.0)	1.56 (1.91)	0.4	0.97	1.19	1.51	14.7		
			Irbesartan	55	55 (100.0)	1.42 (0.44)	0.6	1.10	1.33	1.68	3.1		
		Week 6	Sparsentan	55	51 (92.7)	1.06 (0.97)	0.2	0.60	0.86	1.36	6.5		
			Irbesartan	55	53 (96.4)	1.32 (0.61)	0.6	0.89	1.29	1.64	4.0		
		Week 36	Sparsentan	55	37 (67.3)	1.02 (0.92)	0.1	0.52	0.62	1.31	4.0		
			Irbesartan	55	39 (70.9)	1.38 (1.02)	0.3	0.73	1.03	1.54	4.4		
		Week 58	Sparsentan	55	29 (52.7)	0.95 (0.88)	0.1	0.46	0.78	1.03	4.7		
			Irbesartan	55	30 (54.5)	1.22 (1.11)	0.1	0.65	0.97	1.35	5.0		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	55	51 (92.7)	-0.55 (2.08)	-13.0	-0.80	-0.41	0.08	5.2	-0.29 [-0.68, 0.10]
				Irbesartan	55	53 (96.4)	-0.11 (0.70)	-1.5	-0.48	-0.19	0.31	2.9	
			Week 36	Sparsentan	55	37 (67.3)	-0.17 (0.96)	-2.1	-0.79	-0.28	0.18	2.6	-0.08 [-0.53, 0.37]
				Irbesartan	55	39 (70.9)	-0.09 (1.04)	-1.3	-0.75	-0.35	-0.08	3.6	
			Week 58	Sparsentan	55	29 (52.7)	-0.22 (0.90)	-1.5	-0.82	-0.36	0.12	3.3	0.10 [-0.41, 0.61]
	Irbesartan			55	30 (54.5)	-0.32 (1.02)	-1.5	-0.74	-0.46	-0.18	3.7		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP High	Urine protein excretion	Baseline	Sparsentan	37	37 (100.0)	2.60 (1.20)	1.0	1.96	2.42	2.99	7.2			
			Irbesartan	36	36 (100.0)	2.47 (1.05)	0.6	1.95	2.32	2.84	6.8			
		Week 6	Sparsentan	37	35 (94.6)	1.82 (1.45)	0.2	0.76	1.20	2.47	6.2			
			Irbesartan	36	33 (91.7)	2.69 (2.11)	0.6	1.47	2.27	3.11	10.3			
		Week 36	Sparsentan	37	26 (70.3)	1.62 (1.69)	0.1	0.65	1.16	1.74	8.3			
			Irbesartan	36	21 (58.3)	2.86 (2.33)	0.6	1.52	1.96	2.83	9.2			
		Week 58	Sparsentan	37	19 (51.4)	1.66 (1.70)	0.1	0.23	1.21	1.90	5.4			
			Irbesartan	36	15 (41.7)	2.70 (2.32)	0.7	0.90	1.98	2.98	8.6			
		Change from baseline in urine protein excretion	Week 6	Sparsentan	37	35 (94.6)	-0.79 (1.16)	-2.4	-1.47	-0.97	-0.31	2.7	-0.62 [-1.11, -0.13]	
				Irbesartan	36	33 (91.7)	0.18 (1.89)	-3.2	-0.78	-0.04	0.72	8.4		
				Week 36	Sparsentan	37	26 (70.3)	-0.96 (1.05)	-3.1	-1.42	-0.66	-0.48	1.0	-0.70 [-1.29, -0.11]
					Irbesartan	36	21 (58.3)	0.23 (2.25)	-2.5	-1.34	-0.15	0.80	6.5	
				Week 58	Sparsentan	37	19 (51.4)	-1.01 (1.27)	-3.7	-1.83	-1.02	-0.16	1.7	-0.50 [-1.19, 0.19]
					Irbesartan	36	15 (41.7)	-0.11 (2.28)	-2.7	-1.62	-0.79	1.01	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP Low	Urine protein excretion	Baseline	Sparsentan	39	39 (100.0)	1.26 (0.48)	0.4	0.96	1.18	1.51	3.3			
			Irbesartan	37	37 (100.0)	1.37 (0.44)	0.6	1.11	1.41	1.60	2.9			
		Week 6	Sparsentan	39	39 (100.0)	0.72 (0.38)	0.2	0.46	0.64	0.90	1.8			
			Irbesartan	37	35 (94.6)	1.39 (0.80)	0.4	0.87	1.19	1.91	3.6			
		Week 36	Sparsentan	39	27 (69.2)	0.64 (0.47)	0.1	0.30	0.56	0.88	2.3			
			Irbesartan	37	25 (67.6)	1.28 (0.84)	0.2	0.78	1.13	1.58	3.6			
		Week 58	Sparsentan	39	23 (59.0)	0.91 (0.78)	0.1	0.30	0.59	1.21	3.1			
			Irbesartan	37	17 (45.9)	1.59 (1.07)	0.2	0.90	1.21	2.41	3.6			
		Change from baseline in urine protein excretion		Week 6	Sparsentan	39	39 (100.0)	-0.54 (0.53)	-2.6	-0.79	-0.51	-0.19	0.6	-0.91 [-1.39, -0.43]
					Irbesartan	37	35 (94.6)	0.01 (0.67)	-1.1	-0.52	-0.25	0.72	1.4	
				Week 36	Sparsentan	39	27 (69.2)	-0.72 (0.60)	-2.4	-1.06	-0.73	-0.51	0.7	-0.83 [-1.40, -0.26]
					Irbesartan	37	25 (67.6)	-0.17 (0.73)	-1.5	-0.67	-0.24	0.36	1.0	
				Week 58	Sparsentan	39	23 (59.0)	-0.45 (0.88)	-2.7	-1.05	-0.51	0.11	1.4	-0.48 [-1.12, 0.15]
					Irbesartan	37	17 (45.9)	0.01 (1.06)	-1.5	-0.52	-0.34	0.25	2.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 1													
< 60 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	127	127 (100.0)	2.27 (1.52)	0.1	1.19	1.85	3.10	9.7		
		Week 6	Irbesartan	129	129 (100.0)	2.28 (1.34)	0.5	1.33	1.82	2.80	7.5		
			Sparsentan	127	119 (93.7)	1.60 (1.22)	0.2	0.72	1.25	2.12	6.5		
		Week 36	Irbesartan	129	124 (96.1)	2.20 (1.46)	0.5	1.15	1.83	2.71	7.7		
			Sparsentan	127	88 (69.3)	1.66 (1.63)	0.1	0.60	1.04	2.21	9.4		
		Week 58	Irbesartan	129	88 (68.2)	2.10 (1.52)	0.2	0.98	1.74	2.79	7.1		
			Sparsentan	127	71 (55.9)	1.53 (1.48)	0.1	0.59	1.13	1.92	7.5		
		Week 58	Irbesartan	129	69 (53.5)	2.09 (1.63)	0.1	0.87	1.69	2.80	7.2		
			Change from baseline in urine protein excretion	Week 6	Sparsentan	127	119 (93.7)	-0.67 (1.23)	-4.2	-1.36	-0.67	0.03	5.2
		Week 36	Irbesartan	129	124 (96.1)	-0.06 (1.18)	-3.7	-0.68	-0.19	0.43	4.2		
			Sparsentan	127	88 (69.3)	-0.51 (1.42)	-4.2	-1.21	-0.64	0.19	4.6	-0.25 [-0.55, 0.05]	
		Week 58	Irbesartan	129	88 (68.2)	-0.17 (1.32)	-3.3	-1.01	-0.27	0.39	4.2		
			Sparsentan	127	71 (55.9)	-0.56 (1.39)	-4.4	-1.29	-0.48	0.12	3.3	-0.22 [-0.56, 0.11]	
		Week 58	Irbesartan	129	69 (53.5)	-0.27 (1.19)	-2.7	-0.88	-0.40	0.21	3.7		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	49	49 (100.0)	2.14 (2.03)	0.6	1.12	1.62	2.62	14.7		
			Irbesartan	48	48 (100.0)	1.97 (0.91)	0.6	1.36	1.84	2.45	4.6		
		Week 6	Sparsentan	49	48 (98.0)	1.33 (1.16)	0.2	0.56	0.86	1.84	5.5		
			Irbesartan	48	43 (89.6)	2.33 (1.76)	0.4	1.28	1.93	3.01	10.3		
		Week 36	Sparsentan	49	31 (63.3)	1.07 (0.90)	0.1	0.60	0.86	1.32	3.8		
			Irbesartan	48	33 (68.8)	2.37 (1.87)	0.2	1.24	1.77	2.60	9.2		
		Week 58	Sparsentan	49	24 (49.0)	1.16 (1.06)	0.1	0.25	1.17	1.56	4.6		
			Irbesartan	48	19 (39.6)	2.58 (1.92)	0.2	1.21	2.27	3.06	8.6		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	49	48 (98.0)	-0.80 (2.04)	-13.0	-1.09	-0.67	-0.15	2.7	-0.59 [-1.01, -0.17]
				Irbesartan	48	43 (89.6)	0.29 (1.61)	-3.2	-0.50	0.05	0.80	8.4	
			Week 36	Sparsentan	49	31 (63.3)	-0.79 (0.69)	-2.5	-1.32	-0.67	-0.52	0.9	-0.91 [-1.42, -0.39]
				Irbesartan	48	33 (68.8)	0.44 (1.76)	-2.2	-0.52	0.15	0.90	6.5	
			Week 58	Sparsentan	49	24 (49.0)	-0.62 (0.91)	-2.7	-1.07	-0.75	-0.16	1.7	-0.68 [-1.30, -0.06]
		Irbesartan		48	19 (39.6)	0.41 (2.06)	-2.7	-1.05	-0.08	1.46	6.7		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	26	26 (100.0)	2.15 (1.51)	0.7	1.18	1.64	2.48	7.2		
		Week 6	Irbesartan	25	25 (100.0)	1.94 (1.21)	0.6	1.30	1.63	2.10	6.8		
			Sparsentan	26	25 (96.2)	1.31 (1.26)	0.4	0.57	0.90	1.51	6.2		
		Week 36	Irbesartan	25	24 (96.0)	1.80 (1.91)	0.4	0.84	1.22	1.95	9.5		
			Sparsentan	26	20 (76.9)	1.30 (1.84)	0.1	0.35	0.63	1.61	8.3		
		Week 58	Irbesartan	25	12 (48.0)	1.57 (2.09)	0.2	0.43	0.78	1.96	7.7		
			Sparsentan	26	18 (69.2)	1.49 (1.64)	0.2	0.34	0.85	1.81	5.4		
		Irbesartan	25	10 (40.0)	1.66 (1.90)	0.3	0.55	0.99	2.06	6.6			
		Change from baseline in urine protein excretion	Week 6	Sparsentan	26	25 (96.2)	-0.83 (0.73)	-2.4	-1.20	-0.65	-0.44	0.4	-0.83 [-1.41, -0.25]
		Week 36	Irbesartan	25	24 (96.0)	-0.13 (0.95)	-1.9	-0.63	-0.33	0.35	2.8		
			Sparsentan	26	20 (76.9)	-1.02 (1.08)	-3.1	-1.50	-0.89	-0.45	1.0	-0.28 [-1.00, 0.43]	
		Week 58	Irbesartan	25	12 (48.0)	-0.72 (1.01)	-2.5	-1.46	-0.58	-0.13	1.0		
			Sparsentan	26	18 (69.2)	-0.89 (1.30)	-3.7	-1.36	-0.83	0.07	1.4	-0.21 [-0.99, 0.56]	
	Irbesartan	25	10 (40.0)	-0.63 (1.04)	-2.3	-1.55	-0.43	-0.12	1.4				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 2													
< 45 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	82	82 (100.0)	2.37 (1.59)	0.1	1.28	1.94	3.14	9.7		
		Week 6	Irbesartan	80	80 (100.0)	2.32 (1.40)	0.5	1.33	1.95	2.94	7.5		
			Sparsentan	82	77 (93.9)	1.75 (1.35)	0.2	0.92	1.28	2.15	6.5		
		Week 36	Irbesartan	80	78 (97.5)	2.18 (1.37)	0.6	1.04	1.84	2.70	7.0		
			Sparsentan	82	55 (67.1)	1.87 (1.78)	0.1	0.61	1.27	2.66	9.4		
		Week 58	Irbesartan	80	57 (71.3)	2.21 (1.52)	0.2	1.00	2.05	2.85	6.7		
			Sparsentan	82	45 (54.9)	1.67 (1.73)	0.1	0.59	1.01	1.99	7.5		
		Week 58	Irbesartan	80	44 (55.0)	1.99 (1.56)	0.1	0.70	1.74	2.79	7.2		
			Change from baseline in urine protein excretion	Week 6	Sparsentan	82	77 (93.9)	-0.59 (1.32)	-4.2	-1.36	-0.67	0.10	5.2
		Week 36	Change from baseline in urine protein excretion	Irbesartan	80	78 (97.5)	-0.14 (1.22)	-3.7	-0.78	-0.21	0.38	3.2	
				Sparsentan	82	55 (67.1)	-0.44 (1.57)	-4.2	-1.27	-0.60	0.51	4.6	-0.22 [-0.59, 0.15]
				Irbesartan	80	57 (71.3)	-0.12 (1.38)	-2.5	-0.99	-0.29	0.63	4.2	
				Sparsentan	82	45 (54.9)	-0.53 (1.51)	-4.4	-1.45	-0.55	0.42	3.3	-0.09 [-0.50, 0.33]
		Week 58	Change from baseline in urine protein excretion	Sparsentan	82	45 (54.9)	-0.53 (1.51)	-4.4	-1.45	-0.55	0.42	3.3	-0.09 [-0.50, 0.33]
Irbesartan	80			44 (55.0)	-0.41 (1.11)	-2.7	-1.15	-0.38	0.18	3.4			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
45 to < 60 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	45	45 (100.0)	2.08 (1.39)	0.4	1.18	1.76	2.75	6.4		
			Irbesartan	49	49 (100.0)	2.20 (1.26)	0.6	1.34	1.76	2.64	5.7		
		Week 6	Sparsentan	45	42 (93.3)	1.32 (0.90)	0.2	0.60	1.09	2.01	3.5		
			Irbesartan	49	46 (93.9)	2.23 (1.62)	0.5	1.19	1.83	2.74	7.7		
		Week 36	Sparsentan	45	33 (73.3)	1.29 (1.28)	0.1	0.59	0.83	1.47	6.4		
			Irbesartan	49	31 (63.3)	1.90 (1.52)	0.3	0.88	1.22	2.34	7.1		
		Week 58	Sparsentan	45	26 (57.8)	1.30 (0.90)	0.1	0.69	1.19	1.60	4.5		
			Irbesartan	49	25 (51.0)	2.26 (1.76)	0.3	0.98	1.69	2.91	6.8		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	45	42 (93.3)	-0.80 (1.04)	-4.0	-1.28	-0.70	-0.10	0.9	-0.82 [-1.26, -0.39]
				Irbesartan	49	46 (93.9)	0.08 (1.10)	-1.9	-0.56	0.03	0.44	4.2	
			Week 36	Sparsentan	45	33 (73.3)	-0.62 (1.16)	-4.1	-1.18	-0.76	-0.07	2.6	-0.31 [-0.80, 0.19]
				Irbesartan	49	31 (63.3)	-0.26 (1.24)	-3.3	-1.10	-0.15	0.25	2.8	
			Week 58	Sparsentan	45	26 (57.8)	-0.61 (1.20)	-4.2	-1.25	-0.44	-0.02	2.4	-0.47 [-1.03, 0.08]
		Irbesartan		49	25 (51.0)	-0.02 (1.31)	-1.9	-0.74	-0.42	0.31	3.7		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	49	49 (100.0)	2.14 (2.03)	0.6	1.12	1.62	2.62	14.7		
			Irbesartan	48	48 (100.0)	1.97 (0.91)	0.6	1.36	1.84	2.45	4.6		
		Week 6	Sparsentan	49	48 (98.0)	1.33 (1.16)	0.2	0.56	0.86	1.84	5.5		
			Irbesartan	48	43 (89.6)	2.33 (1.76)	0.4	1.28	1.93	3.01	10.3		
		Week 36	Sparsentan	49	31 (63.3)	1.07 (0.90)	0.1	0.60	0.86	1.32	3.8		
			Irbesartan	48	33 (68.8)	2.37 (1.87)	0.2	1.24	1.77	2.60	9.2		
		Week 58	Sparsentan	49	24 (49.0)	1.16 (1.06)	0.1	0.25	1.17	1.56	4.6		
			Irbesartan	48	19 (39.6)	2.58 (1.92)	0.2	1.21	2.27	3.06	8.6		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	49	48 (98.0)	-0.80 (2.04)	-13.0	-1.09	-0.67	-0.15	2.7	-0.59 [-1.01, -0.17]
		Week 36	Sparsentan	Irbesartan	48	43 (89.6)	0.29 (1.61)	-3.2	-0.50	0.05	0.80	8.4	
				Sparsentan	49	31 (63.3)	-0.79 (0.69)	-2.5	-1.32	-0.67	-0.52	0.9	-0.91 [-1.42, -0.39]
			Irbesartan	Sparsentan	48	33 (68.8)	0.44 (1.76)	-2.2	-0.52	0.15	0.90	6.5	
				Irbesartan	49	24 (49.0)	-0.62 (0.91)	-2.7	-1.07	-0.75	-0.16	1.7	-0.68 [-1.30, -0.06]
	Week 58		Sparsentan	48	19 (39.6)	0.41 (2.06)	-2.7	-1.05	-0.08	1.46	6.7		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	26	26 (100.0)	2.15 (1.51)	0.7	1.18	1.64	2.48	7.2		
		Week 6	Irbesartan	25	25 (100.0)	1.94 (1.21)	0.6	1.30	1.63	2.10	6.8		
			Sparsentan	26	25 (96.2)	1.31 (1.26)	0.4	0.57	0.90	1.51	6.2		
		Week 36	Irbesartan	25	24 (96.0)	1.80 (1.91)	0.4	0.84	1.22	1.95	9.5		
			Sparsentan	26	20 (76.9)	1.30 (1.84)	0.1	0.35	0.63	1.61	8.3		
		Week 58	Irbesartan	25	12 (48.0)	1.57 (2.09)	0.2	0.43	0.78	1.96	7.7		
			Sparsentan	26	18 (69.2)	1.49 (1.64)	0.2	0.34	0.85	1.81	5.4		
		Irbesartan	25	10 (40.0)	1.66 (1.90)	0.3	0.55	0.99	2.06	6.6			
		Change from baseline in urine protein excretion	Week 6	Sparsentan	26	25 (96.2)	-0.83 (0.73)	-2.4	-1.20	-0.65	-0.44	0.4	-0.83 [-1.41, -0.25]
		Week 36	Irbesartan	25	24 (96.0)	-0.13 (0.95)	-1.9	-0.63	-0.33	0.35	2.8		
			Sparsentan	26	20 (76.9)	-1.02 (1.08)	-3.1	-1.50	-0.89	-0.45	1.0	-0.28 [-1.00, 0.43]	
		Week 58	Irbesartan	25	12 (48.0)	-0.72 (1.01)	-2.5	-1.46	-0.58	-0.13	1.0		
			Sparsentan	26	18 (69.2)	-0.89 (1.30)	-3.7	-1.36	-0.83	0.07	1.4	-0.21 [-0.99, 0.56]	
	Irbesartan	25	10 (40.0)	-0.63 (1.04)	-2.3	-1.55	-0.43	-0.12	1.4				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline urine protein excretion													
<= 1.75 g/day	Urine protein excretion	Baseline	Sparsentan	98	98 (100.0)	1.14 (0.33)	0.1	0.97	1.16	1.40	1.7		
			Irbesartan	94	94 (100.0)	1.26 (0.31)	0.5	1.07	1.33	1.50	1.7		
		Week 6	Sparsentan	98	93 (94.9)	0.96 (0.80)	0.2	0.48	0.76	1.17	6.5		
			Irbesartan	94	88 (93.6)	1.46 (0.89)	0.4	0.89	1.27	1.83	4.8		
		Week 36	Sparsentan	98	73 (74.5)	0.97 (0.85)	0.1	0.40	0.70	1.27	4.0		
			Irbesartan	94	61 (64.9)	1.47 (1.11)	0.2	0.75	1.11	1.89	5.8		
		Week 58	Sparsentan	98	61 (62.2)	1.00 (0.85)	0.1	0.34	0.79	1.40	4.7		
			Irbesartan	94	38 (40.4)	1.30 (1.07)	0.1	0.65	1.04	1.56	5.0		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	98	93 (94.9)	-0.21 (0.80)	-1.2	-0.67	-0.40	0.02	5.2	-0.49 [-0.79, -0.20]
				Irbesartan	94	88 (93.6)	0.20 (0.84)	-1.1	-0.35	0.03	0.53	3.2	
			Week 36	Sparsentan	98	73 (74.5)	-0.22 (0.83)	-1.4	-0.79	-0.48	0.15	2.6	-0.42 [-0.76, -0.08]
				Irbesartan	94	61 (64.9)	0.18 (1.06)	-1.4	-0.44	-0.13	0.53	4.2	
			Week 58	Sparsentan	98	61 (62.2)	-0.20 (0.80)	-1.4	-0.76	-0.30	0.23	3.3	-0.18 [-0.59, 0.22]
				Irbesartan	94	38 (40.4)	-0.04 (1.03)	-1.5	-0.63	-0.35	0.10	3.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 1.75 g/day	Urine protein excretion	Baseline	Sparsentan	104	104 (100.0)	3.24 (1.75)	1.8	2.22	2.81	3.50	14.7			
			Irbesartan	108	108 (100.0)	2.95 (1.21)	1.8	2.05	2.54	3.39	7.5			
		Week 6	Sparsentan	104	99 (95.2)	1.99 (1.32)	0.3	1.03	1.62	2.51	6.2			
			Irbesartan	108	103 (95.4)	2.79 (1.79)	0.7	1.56	2.38	3.31	10.3			
		Week 36	Sparsentan	104	66 (63.5)	2.03 (1.90)	0.1	0.79	1.42	2.54	9.4			
			Irbesartan	108	72 (66.7)	2.67 (1.85)	0.2	1.42	2.43	3.40	9.2			
		Week 58	Sparsentan	104	52 (50.0)	1.97 (1.76)	0.2	0.73	1.33	2.58	7.5			
			Irbesartan	108	60 (55.6)	2.67 (1.84)	0.1	1.48	2.27	3.14	8.6			
		Change from baseline in urine protein excretion	Week 6	Sparsentan	104	99 (95.2)	-1.20 (1.69)	-13.0	-1.82	-1.20	-0.54	2.7	-0.66 [-0.94, -0.37]	
					Irbesartan	108	103 (95.4)	-0.14 (1.53)	-3.7	-1.00	-0.32	0.43	8.4	
				Week 36	Sparsentan	104	66 (63.5)	-1.12 (1.47)	-4.2	-2.05	-1.21	-0.53	4.6	-0.53 [-0.87, -0.19]
					Irbesartan	108	72 (66.7)	-0.28 (1.69)	-3.3	-1.34	-0.54	0.58	6.5	
				Week 58	Sparsentan	104	52 (50.0)	-1.12 (1.56)	-4.4	-2.12	-1.26	-0.54	2.5	-0.54 [-0.92, -0.17]
					Irbesartan	108	60 (55.6)	-0.26 (1.60)	-2.7	-1.45	-0.41	0.35	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline use of antihypertensives													
Yes	Urine protein excretion	Baseline	Sparsentan	88	88 (100.0)	2.57 (2.07)	0.4	1.30	2.13	3.15	14.7		
			Irbesartan	83	83 (100.0)	2.16 (1.42)	0.5	1.22	1.70	2.55	7.5		
		Week 6	Sparsentan	88	83 (94.3)	1.62 (1.31)	0.2	0.72	1.11	2.16	6.5		
			Irbesartan	83	77 (92.8)	2.22 (1.84)	0.4	1.05	1.64	2.70	10.3		
		Week 36	Sparsentan	88	57 (64.8)	1.56 (1.51)	0.3	0.67	0.97	1.72	8.3		
			Irbesartan	83	59 (71.1)	2.31 (1.93)	0.2	1.03	1.78	2.74	9.2		
		Week 58	Sparsentan	88	46 (52.3)	1.68 (1.58)	0.1	0.59	1.24	1.94	7.5		
			Irbesartan	83	46 (55.4)	2.14 (1.94)	0.1	0.86	1.62	2.80	8.6		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	88	83 (94.3)	-0.92 (1.86)	-13.0	-1.45	-0.77	-0.12	5.2	-0.60 [-0.92, -0.29]
				Irbesartan	83	77 (92.8)	0.09 (1.45)	-3.7	-0.56	-0.06	0.55	8.4	
			Week 36	Sparsentan	88	57 (64.8)	-0.80 (1.27)	-4.2	-1.36	-0.72	-0.29	2.6	-0.64 [-1.02, -0.27]
				Irbesartan	83	59 (71.1)	0.13 (1.58)	-3.3	-0.62	-0.11	0.63	6.5	
			Week 58	Sparsentan	88	46 (52.3)	-0.52 (1.42)	-4.4	-1.30	-0.53	0.11	3.3	-0.23 [-0.64, 0.18]
				Irbesartan	83	46 (55.4)	-0.18 (1.50)	-2.7	-1.08	-0.35	0.21	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
No	Urine protein excretion	Baseline	Sparsentan	114	114 (100.0)	1.95 (1.17)	0.1	1.10	1.56	2.58	6.4			
			Irbesartan	119	119 (100.0)	2.17 (1.11)	0.6	1.40	1.92	2.64	6.9			
		Week 6	Sparsentan	114	109 (95.6)	1.40 (1.13)	0.2	0.60	1.06	1.78	5.7			
			Irbesartan	119	114 (95.8)	2.15 (1.41)	0.4	1.09	1.84	2.69	7.7			
		Week 36	Sparsentan	114	82 (71.9)	1.41 (1.56)	0.1	0.50	0.83	2.00	9.4			
			Irbesartan	119	74 (62.2)	1.97 (1.42)	0.2	1.00	1.50	2.68	7.1			
		Week 58	Sparsentan	114	67 (58.8)	1.28 (1.30)	0.1	0.42	0.88	1.58	6.8			
			Irbesartan	119	52 (43.7)	2.14 (1.51)	0.3	0.90	1.84	2.90	6.8			
		Change from baseline in urine protein excretion		Week 6	Sparsentan	114	109 (95.6)	-0.57 (0.95)	-4.0	-1.00	-0.58	0.02	2.1	-0.50 [-0.77, -0.24]
					Irbesartan	119	114 (95.8)	-0.04 (1.14)	-3.2	-0.66	-0.19	0.44	4.2	
				Week 36	Sparsentan	114	82 (71.9)	-0.54 (1.24)	-4.1	-1.18	-0.65	0.15	4.6	-0.25 [-0.56, 0.07]
					Irbesartan	119	74 (62.2)	-0.22 (1.32)	-2.5	-1.07	-0.40	0.48	4.2	
				Week 58	Sparsentan	114	67 (58.8)	-0.69 (1.19)	-4.2	-1.25	-0.71	0.02	2.0	-0.42 [-0.79, -0.05]
					Irbesartan	119	52 (43.7)	-0.17 (1.33)	-2.7	-0.86	-0.40	0.39	3.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Time since renal biopsy													
<= 5 years	Urine protein excretion	Baseline	Sparsentan	113	113 (100.0)	2.28 (1.84)	0.1	1.17	1.67	2.95	14.7		
			Irbesartan	127	127 (100.0)	2.18 (1.33)	0.5	1.34	1.77	2.63	7.5		
		Week 6	Sparsentan	113	105 (92.9)	1.54 (1.32)	0.2	0.64	1.05	2.02	6.5		
			Irbesartan	127	120 (94.5)	2.10 (1.55)	0.4	1.05	1.56	2.68	9.5		
		Week 36	Sparsentan	113	73 (64.6)	1.45 (1.50)	0.1	0.56	0.88	1.77	8.3		
			Irbesartan	127	80 (63.0)	2.07 (1.84)	0.2	0.79	1.42	2.64	9.2		
		Week 58	Sparsentan	113	58 (51.3)	1.64 (1.59)	0.1	0.59	1.14	1.94	7.5		
			Irbesartan	127	59 (46.5)	2.01 (1.80)	0.1	0.69	1.35	2.75	7.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	113	105 (92.9)	-0.79 (1.70)	-13.0	-1.22	-0.71	-0.03	5.2	-0.51 [-0.78, -0.25]
			Irbesartan	127	120 (94.5)	-0.07 (1.09)	-3.7	-0.66	-0.18	0.48	3.2		
			Week 36	Sparsentan	113	73 (64.6)	-0.74 (1.21)	-4.2	-1.26	-0.68	-0.11	2.6	-0.45 [-0.77, -0.13]
				Irbesartan	127	80 (63.0)	-0.12 (1.50)	-3.3	-0.95	-0.30	0.58	6.5	
			Week 58	Sparsentan	113	58 (51.3)	-0.46 (1.36)	-4.4	-1.27	-0.55	0.39	3.3	-0.13 [-0.49, 0.23]
				Irbesartan	127	59 (46.5)	-0.29 (1.31)	-2.7	-1.05	-0.43	0.21	3.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 5 years	Urine protein excretion	Baseline	Sparsentan	89	89 (100.0)	2.15 (1.38)	0.4	1.18	1.77	2.78	9.7			
			Irbesartan	75	75 (100.0)	2.13 (1.09)	0.6	1.20	1.96	2.60	5.9			
		Week 6	Sparsentan	89	87 (97.8)	1.43 (1.07)	0.2	0.72	1.09	1.95	5.7			
			Irbesartan	75	71 (94.7)	2.30 (1.65)	0.5	1.20	1.89	2.74	10.3			
		Week 36	Sparsentan	89	66 (74.2)	1.50 (1.59)	0.1	0.60	0.96	1.92	9.4			
			Irbesartan	75	53 (70.7)	2.20 (1.37)	0.2	1.18	1.89	2.74	7.0			
		Week 58	Sparsentan	89	55 (61.8)	1.24 (1.22)	0.1	0.46	0.87	1.53	6.8			
			Irbesartan	75	39 (52.0)	2.33 (1.58)	0.3	1.12	2.06	2.98	8.6			
		Change from baseline in urine protein excretion	Week 6	Sparsentan	89	87 (97.8)	-0.63 (0.99)	-4.0	-1.00	-0.59	-0.12	2.1	-0.62 [-0.95, -0.30]	
				Irbesartan	75	71 (94.7)	0.15 (1.52)	-2.4	-0.54	-0.15	0.61	8.4		
				Week 36	Sparsentan	89	66 (74.2)	-0.54 (1.31)	-4.1	-1.16	-0.66	-0.07	4.6	-0.42 [-0.78, -0.05]
					Irbesartan	75	53 (70.7)	0.01 (1.38)	-2.5	-0.76	-0.17	0.53	5.1	
				Week 58	Sparsentan	89	55 (61.8)	-0.80 (1.19)	-4.2	-1.29	-0.76	-0.16	2.0	-0.59 [-1.01, -0.17]
					Irbesartan	75	39 (52.0)	-0.01 (1.54)	-2.5	-0.83	-0.30	0.46	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: History of hypertension													
Yes	Urine protein excretion	Baseline	Sparsentan	153	153 (100.0)	2.25 (1.67)	0.1	1.18	1.77	3.00	14.7		
			Irbesartan	157	157 (100.0)	2.20 (1.31)	0.5	1.34	1.78	2.64	7.5		
		Week 6	Sparsentan	153	144 (94.1)	1.54 (1.23)	0.2	0.70	1.09	2.07	6.5		
			Irbesartan	157	147 (93.6)	2.16 (1.59)	0.4	1.13	1.71	2.70	10.3		
		Week 36	Sparsentan	153	104 (68.0)	1.53 (1.48)	0.1	0.60	0.99	1.89	8.3		
			Irbesartan	157	106 (67.5)	2.21 (1.77)	0.2	1.01	1.72	2.74	9.2		
		Week 58	Sparsentan	153	83 (54.2)	1.56 (1.45)	0.1	0.59	1.13	1.92	7.5		
			Irbesartan	157	80 (51.0)	2.24 (1.82)	0.1	0.88	1.82	2.97	8.6		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	153	144 (94.1)	-0.74 (1.55)	-13.0	-1.29	-0.69	-0.03	5.2	-0.49 [-0.73, -0.26]
				Irbesartan	157	147 (93.6)	-0.04 (1.28)	-3.7	-0.63	-0.19	0.41	8.4	
			Week 36	Sparsentan	153	104 (68.0)	-0.63 (1.21)	-4.2	-1.25	-0.66	-0.05	2.6	-0.43 [-0.71, -0.16]
				Irbesartan	157	106 (67.5)	-0.04 (1.53)	-3.3	-0.92	-0.29	0.53	6.5	
			Week 58	Sparsentan	153	83 (54.2)	-0.53 (1.29)	-4.4	-1.18	-0.55	0.11	3.3	-0.29 [-0.60, 0.01]
				Irbesartan	157	80 (51.0)	-0.12 (1.48)	-2.7	-0.99	-0.34	0.39	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
No	Urine protein excretion	Baseline	Sparsentan	49	49 (100.0)	2.14 (1.58)	0.4	1.18	1.62	2.63	9.7			
			Irbesartan	45	45 (100.0)	2.04 (0.96)	0.6	1.33	1.96	2.54	4.6			
		Week 6	Sparsentan	49	48 (98.0)	1.33 (1.16)	0.2	0.59	0.92	1.57	5.7			
			Irbesartan	45	44 (97.8)	2.25 (1.59)	0.5	1.00	1.85	2.66	7.7			
		Week 36	Sparsentan	49	35 (71.4)	1.30 (1.71)	0.1	0.40	0.71	1.92	9.4			
			Irbesartan	45	27 (60.0)	1.75 (1.12)	0.2	0.96	1.48	2.52	4.4			
		Week 58	Sparsentan	49	30 (61.2)	1.14 (1.32)	0.1	0.34	0.81	1.31	6.8			
			Irbesartan	45	18 (40.0)	1.71 (1.08)	0.3	0.90	1.56	2.41	4.6			
		Change from baseline in urine protein excretion		Week 6	Sparsentan	49	48 (98.0)	-0.65 (0.96)	-4.0	-1.09	-0.57	-0.23	2.1	-0.77 [-1.20, -0.35]
					Irbesartan	45	44 (97.8)	0.19 (1.22)	-2.4	-0.57	0.12	0.73	4.2	
				Week 36	Sparsentan	49	35 (71.4)	-0.69 (1.39)	-4.1	-1.18	-0.77	-0.29	4.6	-0.39 [-0.89, 0.12]
					Irbesartan	45	27 (60.0)	-0.19 (1.09)	-2.5	-0.76	-0.14	0.79	1.6	
				Week 58	Sparsentan	49	30 (61.2)	-0.90 (1.27)	-4.2	-1.44	-0.80	-0.09	2.0	-0.40 [-0.99, 0.19]
					Irbesartan	45	18 (40.0)	-0.42 (0.99)	-2.1	-0.84	-0.42	-0.18	2.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.151
Male	Week 6	Sparsentan	139	132 (95.0)	-0.73 (0.11)	(-0.94, -0.52)	-0.81 (0.15)	(-1.11, -0.51)	<0.001 *
		Irbesartan	143	137 (95.8)	0.08 (0.11)	(-0.13, 0.29)			
	Week 36	Sparsentan	139	92 (66.2)	-0.71 (0.13)	(-0.95, -0.46)	-0.72 (0.17)	(-1.06, -0.37)	<0.001 *
		Irbesartan	143	98 (68.5)	0.01 (0.12)	(-0.23, 0.25)			
	Week 58	Sparsentan	139	69 (49.6)	-0.58 (0.14)	(-0.86, -0.30)	-0.59 (0.20)	(-0.98, -0.19)	0.003 *
		Irbesartan	143	73 (51.0)	0.01 (0.14)	(-0.26, 0.28)			
Female	Week 6	Sparsentan	63	60 (95.2)	-0.52 (0.14)	(-0.80, -0.23)	-0.25 (0.21)	(-0.66, 0.17)	0.240
		Irbesartan	59	54 (91.5)	-0.27 (0.15)	(-0.57, 0.03)			
	Week 36	Sparsentan	63	47 (74.6)	-0.83 (0.16)	(-1.14, -0.52)	-0.56 (0.24)	(-1.03, -0.10)	0.018 *
		Irbesartan	59	35 (59.3)	-0.27 (0.18)	(-0.62, 0.08)			
	Week 58	Sparsentan	63	44 (69.8)	-0.76 (0.17)	(-1.09, -0.43)	-0.35 (0.27)	(-0.87, 0.17)	0.189
		Irbesartan	59	25 (42.4)	-0.41 (0.21)	(-0.82, -0.00)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.972
<= 45 years	Week 6	Sparsentan	96	91 (94.8)	-0.81 (0.14)	(-1.09, -0.53)	-0.73 (0.20)	(-1.12, -0.34)	<0.001 *
		Irbesartan	99	95 (96.0)	-0.08 (0.14)	(-0.35, 0.19)			
	Week 36	Sparsentan	96	66 (68.8)	-0.78 (0.16)	(-1.09, -0.47)	-0.55 (0.22)	(-0.99, -0.11)	0.014 *
		Irbesartan	99	65 (65.7)	-0.23 (0.16)	(-0.54, 0.08)			
	Week 58	Sparsentan	96	51 (53.1)	-0.60 (0.18)	(-0.95, -0.25)	-0.42 (0.26)	(-0.93, 0.09)	0.107
		Irbesartan	99	44 (44.4)	-0.18 (0.19)	(-0.55, 0.19)			
> 45 years	Week 6	Sparsentan	106	101 (95.3)	-0.55 (0.11)	(-0.76, -0.35)	-0.57 (0.15)	(-0.87, -0.28)	<0.001 *
		Irbesartan	103	96 (93.2)	0.02 (0.11)	(-0.19, 0.23)			
	Week 36	Sparsentan	106	73 (68.9)	-0.72 (0.12)	(-0.96, -0.48)	-0.81 (0.17)	(-1.15, -0.46)	<0.001 *
		Irbesartan	103	68 (66.0)	0.09 (0.13)	(-0.16, 0.33)			
	Week 58	Sparsentan	106	62 (58.5)	-0.68 (0.13)	(-0.94, -0.42)	-0.67 (0.19)	(-1.05, -0.29)	<0.001 *
		Irbesartan	103	54 (52.4)	-0.01 (0.14)	(-0.28, 0.27)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.461
<= 18 years	Week 6	Sparsentan	9	8 (88.9)	-0.76 (0.57)	(-1.95, 0.43)	-0.44 (1.02)	(-2.60, 1.73)	0.676
		Irbesartan	5	5 (100.0)	-0.32 (0.77)	(-1.96, 1.31)			
	Week 36	Sparsentan	9	6 (66.7)	-0.79 (0.64)	(-2.10, 0.53)	-0.48 (1.24)	(-3.06, 2.09)	0.701
		Irbesartan	5	2 (40.0)	-0.30 (1.06)	(-2.47, 1.86)			
	Week 58	Sparsentan	9	4 (44.4)	-0.62 (0.72)	(-2.10, 0.87)	0.88 (1.53)	(-2.24, 4.00)	0.569
		Irbesartan	5	1 (20.0)	-1.50 (1.35)	(-4.24, 1.24)			
> 18 to 40 years	Week 6	Sparsentan	102	98 (96.1)	-0.86 (0.13)	(-1.11, -0.61)	-0.77 (0.18)	(-1.12, -0.42)	<0.001 *
		Irbesartan	109	105 (96.3)	-0.09 (0.12)	(-0.33, 0.15)			
	Week 36	Sparsentan	102	69 (67.6)	-0.76 (0.15)	(-1.05, -0.48)	-0.51 (0.20)	(-0.91, -0.11)	0.012 *
		Irbesartan	109	73 (67.0)	-0.26 (0.14)	(-0.53, 0.02)			
	Week 58	Sparsentan	102	58 (56.9)	-0.72 (0.16)	(-1.03, -0.41)	-0.46 (0.23)	(-0.92, -0.01)	0.045 *
		Irbesartan	109	51 (46.8)	-0.26 (0.17)	(-0.58, 0.07)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
> 40 years	Week 6	Sparsentan	91	86 (94.5)	-0.48 (0.12)	(-0.71, -0.24)	-0.56 (0.17)	(-0.90, -0.22)	0.001	*
		Irbesartan	88	81 (92.0)	0.08 (0.12)	(-0.16, 0.32)				
	Week 36	Sparsentan	91	64 (70.3)	-0.75 (0.14)	(-1.02, -0.48)	-0.95 (0.20)	(-1.34, -0.57)	<0.001	*
		Irbesartan	88	58 (65.9)	0.20 (0.14)	(-0.08, 0.48)				
	Week 58	Sparsentan	91	51 (56.0)	-0.58 (0.15)	(-0.88, -0.28)	-0.75 (0.22)	(-1.18, -0.31)	<0.001	*
		Irbesartan	88	46 (52.3)	0.17 (0.16)	(-0.15, 0.48)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.026 #
North America	Week 6	Sparsentan	35	33 (94.3)	-0.64 (0.29)	(-1.21, -0.08)	-0.65 (0.38)	(-1.40, 0.09)	0.086
		Irbesartan	46	43 (93.5)	0.01 (0.25)	(-0.48, 0.50)			
	Week 36	Sparsentan	35	18 (51.4)	-1.02 (0.37)	(-1.74, -0.30)	-1.22 (0.48)	(-2.17, -0.28)	0.012 *
		Irbesartan	46	25 (54.3)	0.21 (0.31)	(-0.41, 0.82)			
	Week 58	Sparsentan	35	12 (34.3)	-0.78 (0.44)	(-1.66, 0.09)	-0.99 (0.58)	(-2.13, 0.15)	0.087
		Irbesartan	46	18 (39.1)	0.21 (0.37)	(-0.51, 0.93)			
Europe	Week 6	Sparsentan	98	91 (92.9)	-0.75 (0.12)	(-0.97, -0.52)	-0.69 (0.16)	(-1.00, -0.38)	<0.001 *
		Irbesartan	115	109 (94.8)	-0.06 (0.11)	(-0.27, 0.15)			
	Week 36	Sparsentan	98	60 (61.2)	-0.69 (0.14)	(-0.96, -0.41)	-0.52 (0.19)	(-0.88, -0.16)	0.005 *
		Irbesartan	115	78 (67.8)	-0.17 (0.12)	(-0.41, 0.08)			
	Week 58	Sparsentan	98	48 (49.0)	-0.51 (0.16)	(-0.82, -0.20)	-0.39 (0.21)	(-0.80, 0.03)	0.069
		Irbesartan	115	56 (48.7)	-0.13 (0.14)	(-0.41, 0.15)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Asia Pacific	Week 6	Sparsentan	69	68 (98.6)	-0.62 (0.12)	(-0.86, -0.37)	-0.60 (0.20)	(-1.00, -0.20)	0.003	*
		Irbesartan	41	39 (95.1)	-0.01 (0.16)	(-0.33, 0.30)				
	Week 36	Sparsentan	69	61 (88.4)	-0.70 (0.13)	(-0.95, -0.45)	-0.65 (0.22)	(-1.08, -0.23)	0.003	*
		Irbesartan	41	30 (73.2)	-0.05 (0.17)	(-0.39, 0.30)				
	Week 58	Sparsentan	69	53 (76.8)	-0.73 (0.14)	(-0.99, -0.46)	-0.45 (0.24)	(-0.92, 0.02)	0.060	
		Irbesartan	41	24 (58.5)	-0.27 (0.20)	(-0.66, 0.11)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.174
< 27 kg/m**2	Week 6	Sparsentan	84	77 (91.7)	-0.57 (0.11)	(-0.79, -0.36)	-0.49 (0.15)	(-0.79, -0.20)	0.001 *
		Irbesartan	94	90 (95.7)	-0.08 (0.10)	(-0.28, 0.12)			
	Week 36	Sparsentan	84	59 (70.2)	-0.75 (0.12)	(-0.99, -0.51)	-0.36 (0.17)	(-0.70, -0.01)	0.041 *
		Irbesartan	94	59 (62.8)	-0.39 (0.12)	(-0.63, -0.15)			
	Week 58	Sparsentan	84	46 (54.8)	-0.85 (0.14)	(-1.12, -0.58)	-0.36 (0.20)	(-0.76, 0.04)	0.075
		Irbesartan	94	40 (42.6)	-0.49 (0.15)	(-0.77, -0.20)			
≥ 27 kg/m**2	Week 6	Sparsentan	118	115 (97.5)	-0.72 (0.13)	(-0.97, -0.48)	-0.72 (0.19)	(-1.09, -0.36)	<0.001 *
		Irbesartan	107	100 (93.5)	-0.00 (0.13)	(-0.27, 0.26)			
	Week 36	Sparsentan	118	80 (67.8)	-0.73 (0.14)	(-1.02, -0.45)	-0.91 (0.21)	(-1.32, -0.50)	<0.001 *
		Irbesartan	107	73 (68.2)	0.18 (0.15)	(-0.12, 0.48)			
	Week 58	Sparsentan	118	67 (56.8)	-0.51 (0.16)	(-0.83, -0.20)	-0.66 (0.23)	(-1.12, -0.20)	0.005 *
		Irbesartan	107	58 (54.2)	0.15 (0.17)	(-0.19, 0.48)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
					Change from Baseline		Treatment Difference			
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Randomization strata	Overall	Sparsentan								Interaction: 0.039 #
eGFR Low and UP High	Week 6	Sparsentan	71	67 (94.4)	-0.87 (0.18)	(-1.22, -0.52)	-0.85 (0.25)	(-1.34, -0.37)	<0.001	*
		Irbesartan	74	70 (94.6)	-0.02 (0.17)	(-0.36, 0.32)				
	Week 36	Sparsentan	71	49 (69.0)	-0.88 (0.20)	(-1.28, -0.48)	-0.77 (0.29)	(-1.33, -0.21)	0.007	*
		Irbesartan	74	48 (64.9)	-0.11 (0.20)	(-0.51, 0.29)				
	Week 58	Sparsentan	71	42 (59.2)	-0.90 (0.22)	(-1.34, -0.47)	-0.81 (0.32)	(-1.45, -0.18)	0.011	*
		Irbesartan	74	36 (48.6)	-0.09 (0.23)	(-0.55, 0.37)				
eGFR Low and UP Low	Week 6	Sparsentan	55	51 (92.7)	-0.40 (0.11)	(-0.62, -0.18)	-0.28 (0.15)	(-0.59, 0.02)	0.068	
		Irbesartan	55	53 (96.4)	-0.12 (0.11)	(-0.33, 0.10)				
	Week 36	Sparsentan	55	37 (67.3)	-0.41 (0.12)	(-0.66, -0.17)	-0.44 (0.17)	(-0.78, -0.09)	0.013	*
		Irbesartan	55	39 (70.9)	0.02 (0.12)	(-0.22, 0.26)				
	Week 58	Sparsentan	55	29 (52.7)	-0.41 (0.14)	(-0.69, -0.13)	-0.26 (0.20)	(-0.65, 0.13)	0.184	
		Irbesartan	55	30 (54.5)	-0.15 (0.14)	(-0.42, 0.12)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

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An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
eGFR High and UP High	Week 6	Sparsentan	37	35 (94.6)	-0.82 (0.26)	(-1.33, -0.32)	-0.90 (0.37)	(-1.62, -0.18)	0.015 *
		Irbesartan	36	33 (91.7)	0.08 (0.26)	(-0.44, 0.59)			
	Week 36	Sparsentan	37	26 (70.3)	-1.16 (0.29)	(-1.72, -0.59)	-1.17 (0.42)	(-2.00, -0.34)	0.006 *
		Irbesartan	36	21 (58.3)	0.01 (0.31)	(-0.59, 0.62)			
	Week 58	Sparsentan	37	19 (51.4)	-0.92 (0.33)	(-1.56, -0.28)	-0.72 (0.49)	(-1.68, 0.23)	0.138
		Irbesartan	36	15 (41.7)	-0.20 (0.36)	(-0.91, 0.51)			
eGFR High and UP Low	Week 6	Sparsentan	39	39 (100.0)	-0.59 (0.10)	(-0.80, -0.39)	-0.61 (0.15)	(-0.90, -0.32)	<0.001 *
		Irbesartan	37	35 (94.6)	0.02 (0.11)	(-0.19, 0.23)			
	Week 36	Sparsentan	39	27 (69.2)	-0.72 (0.12)	(-0.95, -0.49)	-0.53 (0.17)	(-0.87, -0.20)	0.002 *
		Irbesartan	37	25 (67.6)	-0.19 (0.12)	(-0.43, 0.05)			
	Week 58	Sparsentan	39	23 (59.0)	-0.42 (0.13)	(-0.67, -0.16)	-0.51 (0.19)	(-0.89, -0.13)	0.009 *
		Irbesartan	37	17 (45.9)	0.09 (0.14)	(-0.19, 0.38)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.189
< 60 mL/min/1.73 m**2	Week 6	Sparsentan	127	119 (93.7)	-0.63 (0.11)	(-0.85, -0.42)	-0.57 (0.15)	(-0.87, -0.27)	<0.001 *
		Irbesartan	129	124 (96.1)	-0.06 (0.11)	(-0.28, 0.15)			
	Week 36	Sparsentan	127	88 (69.3)	-0.60 (0.12)	(-0.85, -0.36)	-0.51 (0.18)	(-0.86, -0.17)	0.004 *
		Irbesartan	129	88 (68.2)	-0.09 (0.12)	(-0.33, 0.16)			
	Week 58	Sparsentan	127	71 (55.9)	-0.63 (0.14)	(-0.90, -0.35)	-0.49 (0.20)	(-0.88, -0.10)	0.013 *
		Irbesartan	129	69 (53.5)	-0.14 (0.14)	(-0.41, 0.14)			
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	48 (98.0)	-0.71 (0.19)	(-1.08, -0.33)	-0.91 (0.27)	(-1.45, -0.37)	<0.001 *
		Irbesartan	48	43 (89.6)	0.20 (0.19)	(-0.18, 0.58)			
	Week 36	Sparsentan	49	31 (63.3)	-1.10 (0.22)	(-1.53, -0.66)	-1.39 (0.31)	(-2.00, -0.78)	<0.001 *
		Irbesartan	48	33 (68.8)	0.30 (0.22)	(-0.13, 0.72)			
	Week 58	Sparsentan	49	24 (49.0)	-0.67 (0.25)	(-1.16, -0.18)	-0.97 (0.36)	(-1.68, -0.25)	0.009 *
		Irbesartan	48	19 (39.6)	0.30 (0.27)	(-0.23, 0.82)			

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	25 (96.2)	-0.84 (0.19)	(-1.22, -0.46)	-0.67 (0.27)	(-1.21, -0.13)	0.015 *
		Irbesartan	25	24 (96.0)	-0.17 (0.20)	(-0.55, 0.22)			
	Week 36	Sparsentan	26	20 (76.9)	-0.97 (0.21)	(-1.39, -0.55)	-0.23 (0.33)	(-0.89, 0.43)	0.484
		Irbesartan	25	12 (48.0)	-0.74 (0.26)	(-1.25, -0.23)			
	Week 58	Sparsentan	26	18 (69.2)	-0.86 (0.22)	(-1.30, -0.42)	-0.22 (0.37)	(-0.94, 0.51)	0.551
		Irbesartan	25	10 (40.0)	-0.64 (0.29)	(-1.22, -0.07)			

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.171
< 45 mL/min/1.73 m**2	Week 6	Sparsentan	82	77 (93.9)	-0.56 (0.14)	(-0.83, -0.29)	-0.41 (0.20)	(-0.79, -0.02)	0.040 *
		Irbesartan	80	78 (97.5)	-0.15 (0.14)	(-0.43, 0.12)			
	Week 36	Sparsentan	82	55 (67.1)	-0.52 (0.16)	(-0.84, -0.21)	-0.51 (0.22)	(-0.95, -0.07)	0.023 *
		Irbesartan	80	57 (71.3)	-0.02 (0.16)	(-0.32, 0.29)			
	Week 58	Sparsentan	82	45 (54.9)	-0.56 (0.18)	(-0.91, -0.21)	-0.34 (0.25)	(-0.83, 0.15)	0.179
		Irbesartan	80	44 (55.0)	-0.23 (0.18)	(-0.57, 0.12)			
45 to < 60 mL/min/1.73 m**2	Week 6	Sparsentan	45	42 (93.3)	-0.78 (0.18)	(-1.14, -0.43)	-0.89 (0.25)	(-1.38, -0.39)	<0.001 *
		Irbesartan	49	46 (93.9)	0.10 (0.17)	(-0.24, 0.45)			
	Week 36	Sparsentan	45	33 (73.3)	-0.75 (0.20)	(-1.15, -0.36)	-0.56 (0.29)	(-1.13, 0.01)	0.053
		Irbesartan	49	31 (63.3)	-0.19 (0.21)	(-0.60, 0.21)			
	Week 58	Sparsentan	45	26 (57.8)	-0.76 (0.23)	(-1.20, -0.31)	-0.82 (0.33)	(-1.47, -0.18)	0.013 *
		Irbesartan	49	25 (51.0)	0.06 (0.23)	(-0.39, 0.52)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	48 (98.0)	-0.71 (0.19)	(-1.08, -0.33)	-0.91 (0.27)	(-1.45, -0.37)	<0.001 *
		Irbesartan	48	43 (89.6)	0.20 (0.19)	(-0.18, 0.58)			
	Week 36	Sparsentan	49	31 (63.3)	-1.10 (0.22)	(-1.53, -0.66)	-1.39 (0.31)	(-2.00, -0.78)	<0.001 *
		Irbesartan	48	33 (68.8)	0.30 (0.22)	(-0.13, 0.72)			
	Week 58	Sparsentan	49	24 (49.0)	-0.67 (0.25)	(-1.16, -0.18)	-0.97 (0.36)	(-1.68, -0.25)	0.009 *
		Irbesartan	48	19 (39.6)	0.30 (0.27)	(-0.23, 0.82)			
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	25 (96.2)	-0.84 (0.19)	(-1.22, -0.46)	-0.67 (0.27)	(-1.21, -0.13)	0.015 *
		Irbesartan	25	24 (96.0)	-0.17 (0.20)	(-0.55, 0.22)			
	Week 36	Sparsentan	26	20 (76.9)	-0.97 (0.21)	(-1.39, -0.55)	-0.23 (0.33)	(-0.89, 0.43)	0.484
		Irbesartan	25	12 (48.0)	-0.74 (0.26)	(-1.25, -0.23)			
	Week 58	Sparsentan	26	18 (69.2)	-0.86 (0.22)	(-1.30, -0.42)	-0.22 (0.37)	(-0.94, 0.51)	0.551
		Irbesartan	25	10 (40.0)	-0.64 (0.29)	(-1.22, -0.07)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan							Interaction: <0.001 #
<= 1.75 g/day	Week 6	Sparsentan	98	93 (94.9)	-0.18 (0.08)	(-0.33, -0.02)	-0.35 (0.11)	(-0.58, -0.13)	0.002 *
		Irbesartan	94	88 (93.6)	0.18 (0.08)	(0.02, 0.33)			
	Week 36	Sparsentan	98	73 (74.5)	-0.23 (0.09)	(-0.41, -0.06)	-0.41 (0.13)	(-0.66, -0.16)	0.002 *
		Irbesartan	94	61 (64.9)	0.18 (0.09)	(-0.01, 0.36)			
	Week 58	Sparsentan	98	61 (62.2)	-0.14 (0.10)	(-0.33, 0.05)	-0.17 (0.15)	(-0.46, 0.12)	0.258
		Irbesartan	94	38 (40.4)	0.03 (0.12)	(-0.20, 0.26)			
> 1.75 g/day	Week 6	Sparsentan	104	99 (95.2)	-1.21 (0.16)	(-1.53, -0.89)	-1.04 (0.23)	(-1.49, -0.60)	<0.001 *
		Irbesartan	108	103 (95.4)	-0.17 (0.16)	(-0.48, 0.14)			
	Week 36	Sparsentan	104	66 (63.5)	-1.33 (0.19)	(-1.70, -0.96)	-1.07 (0.26)	(-1.58, -0.56)	<0.001 *
		Irbesartan	108	72 (66.7)	-0.26 (0.18)	(-0.61, 0.10)			
	Week 58	Sparsentan	104	52 (50.0)	-1.21 (0.21)	(-1.63, -0.79)	-0.99 (0.29)	(-1.57, -0.42)	<0.001 *
		Irbesartan	108	60 (55.6)	-0.22 (0.20)	(-0.61, 0.17)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.533
Yes	Week 6	Sparsentan	88	83 (94.3)	-0.79 (0.14)	(-1.07, -0.52)	-0.77 (0.21)	(-1.17, -0.36)	<0.001 *
		Irbesartan	83	77 (92.8)	-0.03 (0.15)	(-0.32, 0.26)			
	Week 36	Sparsentan	88	57 (64.8)	-0.91 (0.17)	(-1.23, -0.58)	-0.91 (0.23)	(-1.37, -0.46)	<0.001 *
		Irbesartan	83	59 (71.1)	0.01 (0.16)	(-0.32, 0.33)			
	Week 58	Sparsentan	88	46 (52.3)	-0.62 (0.19)	(-0.99, -0.26)	-0.43 (0.26)	(-0.95, 0.09)	0.103
		Irbesartan	83	46 (55.4)	-0.19 (0.19)	(-0.56, 0.17)			
No	Week 6	Sparsentan	114	109 (95.6)	-0.58 (0.11)	(-0.79, -0.36)	-0.54 (0.15)	(-0.84, -0.24)	<0.001 *
		Irbesartan	119	114 (95.8)	-0.04 (0.11)	(-0.24, 0.17)			
	Week 36	Sparsentan	114	82 (71.9)	-0.63 (0.12)	(-0.87, -0.39)	-0.48 (0.18)	(-0.83, -0.14)	0.006 *
		Irbesartan	119	74 (62.2)	-0.15 (0.13)	(-0.39, 0.10)			
	Week 58	Sparsentan	114	67 (58.8)	-0.67 (0.13)	(-0.93, -0.40)	-0.63 (0.20)	(-1.02, -0.24)	0.002 *
		Irbesartan	119	52 (43.7)	-0.04 (0.15)	(-0.33, 0.25)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.529
<= 5 years	Week 6	Sparsentan	113	105 (92.9)	-0.73 (0.12)	(-0.97, -0.48)	-0.61 (0.17)	(-0.95, -0.28)	<0.001 *
		Irbesartan	127	120 (94.5)	-0.11 (0.12)	(-0.34, 0.11)			
	Week 36	Sparsentan	113	73 (64.6)	-0.88 (0.14)	(-1.15, -0.60)	-0.71 (0.20)	(-1.10, -0.33)	<0.001 *
		Irbesartan	127	80 (63.0)	-0.16 (0.14)	(-0.43, 0.10)			
	Week 58	Sparsentan	113	58 (51.3)	-0.57 (0.16)	(-0.88, -0.25)	-0.35 (0.22)	(-0.79, 0.09)	0.115
		Irbesartan	127	59 (46.5)	-0.21 (0.16)	(-0.52, 0.09)			
> 5 years	Week 6	Sparsentan	89	87 (97.8)	-0.63 (0.12)	(-0.88, -0.39)	-0.76 (0.18)	(-1.13, -0.40)	<0.001 *
		Irbesartan	75	71 (94.7)	0.13 (0.14)	(-0.14, 0.40)			
	Week 36	Sparsentan	89	66 (74.2)	-0.60 (0.14)	(-0.88, -0.33)	-0.70 (0.21)	(-1.10, -0.29)	<0.001 *
		Irbesartan	75	53 (70.7)	0.10 (0.15)	(-0.20, 0.40)			
	Week 58	Sparsentan	89	55 (61.8)	-0.72 (0.15)	(-1.01, -0.42)	-0.83 (0.23)	(-1.28, -0.37)	<0.001 *
		Irbesartan	75	39 (52.0)	0.11 (0.17)	(-0.23, 0.45)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.861
Yes	Week 6	Sparsentan	153	144 (94.1)	-0.70 (0.11)	(-0.91, -0.49)	-0.61 (0.15)	(-0.90, -0.31)	<0.001 *
		Irbesartan	157	147 (93.6)	-0.09 (0.10)	(-0.30, 0.11)			
	Week 36	Sparsentan	153	104 (68.0)	-0.76 (0.12)	(-1.00, -0.53)	-0.73 (0.17)	(-1.06, -0.40)	<0.001 *
		Irbesartan	157	106 (67.5)	-0.03 (0.12)	(-0.27, 0.20)			
	Week 58	Sparsentan	153	83 (54.2)	-0.59 (0.13)	(-0.86, -0.33)	-0.52 (0.19)	(-0.89, -0.14)	0.007 *
		Irbesartan	157	80 (51.0)	-0.08 (0.14)	(-0.34, 0.19)			
No	Week 6	Sparsentan	49	48 (98.0)	-0.64 (0.15)	(-0.93, -0.34)	-0.82 (0.22)	(-1.25, -0.40)	<0.001 *
		Irbesartan	45	44 (97.8)	0.19 (0.16)	(-0.12, 0.49)			
	Week 36	Sparsentan	49	35 (71.4)	-0.73 (0.17)	(-1.06, -0.40)	-0.54 (0.25)	(-1.04, -0.05)	0.031 *
		Irbesartan	45	27 (60.0)	-0.19 (0.18)	(-0.55, 0.17)			
	Week 58	Sparsentan	49	30 (61.2)	-0.81 (0.18)	(-1.17, -0.45)	-0.64 (0.29)	(-1.21, -0.08)	0.026 *
		Irbesartan	45	18 (40.0)	-0.17 (0.22)	(-0.60, 0.27)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

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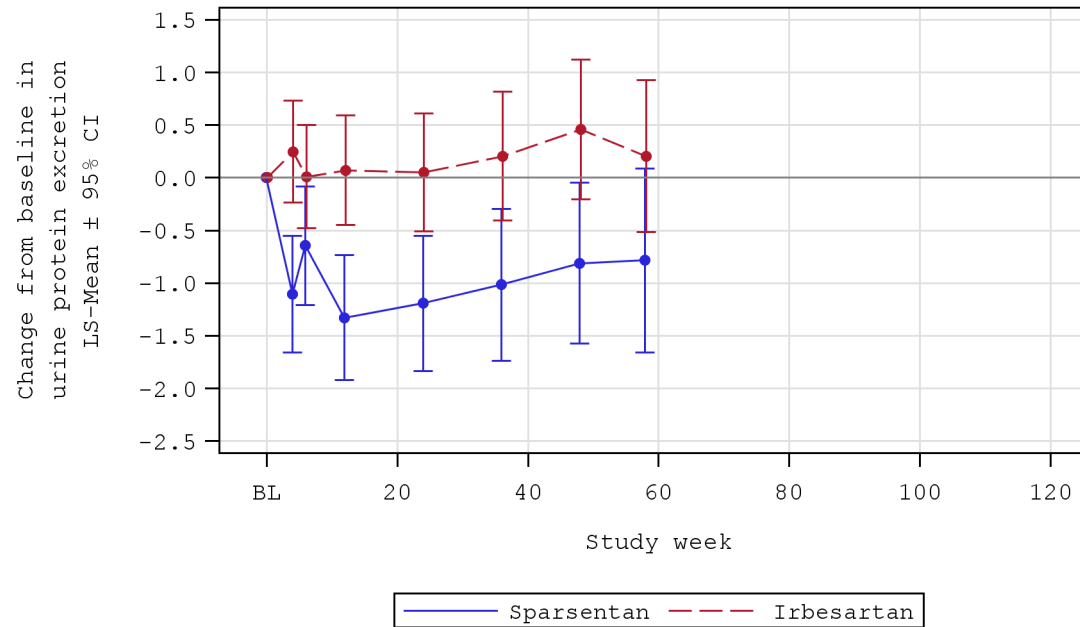
An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

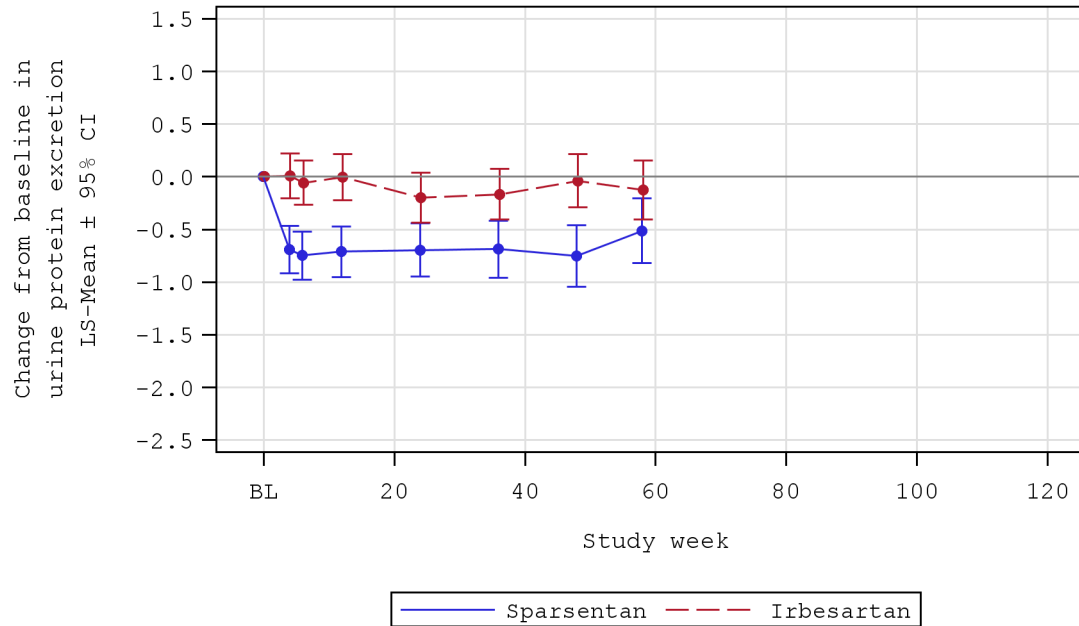
Figure PF1UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Geographic region: North America



Sparsentan	33	18	12
Irbesartan	43	25	18

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT1UEC\_FSCM.

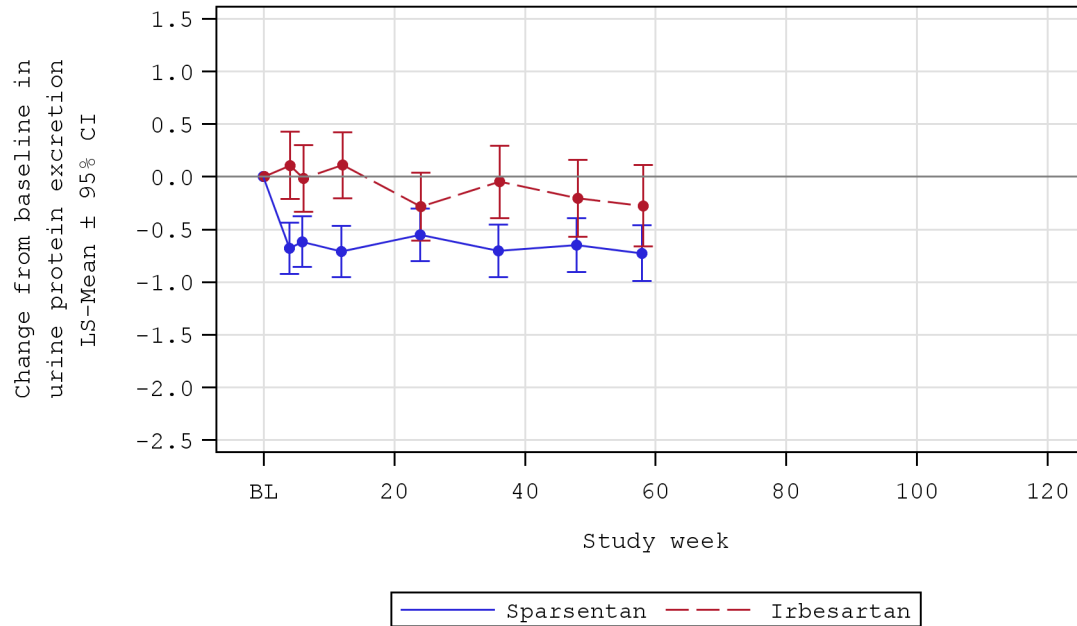
Figure PF1UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Geographic region: Europe



Sparsentan	91	60	48
Irbesartan	109	78	56

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT1UEC\_FSCM.

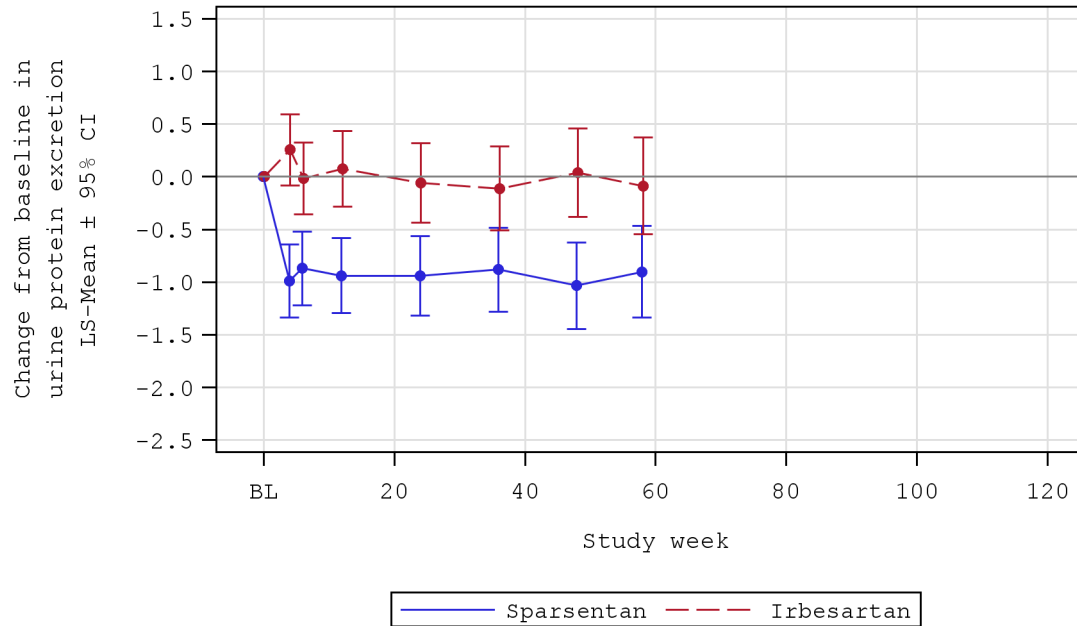
Figure PF1UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Geographic region: Asia Pacific



Sparsentan	68	61	53
Irbesartan	39	30	24

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT1UEC\_FSCM.

Figure PF1UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR Low and UP High

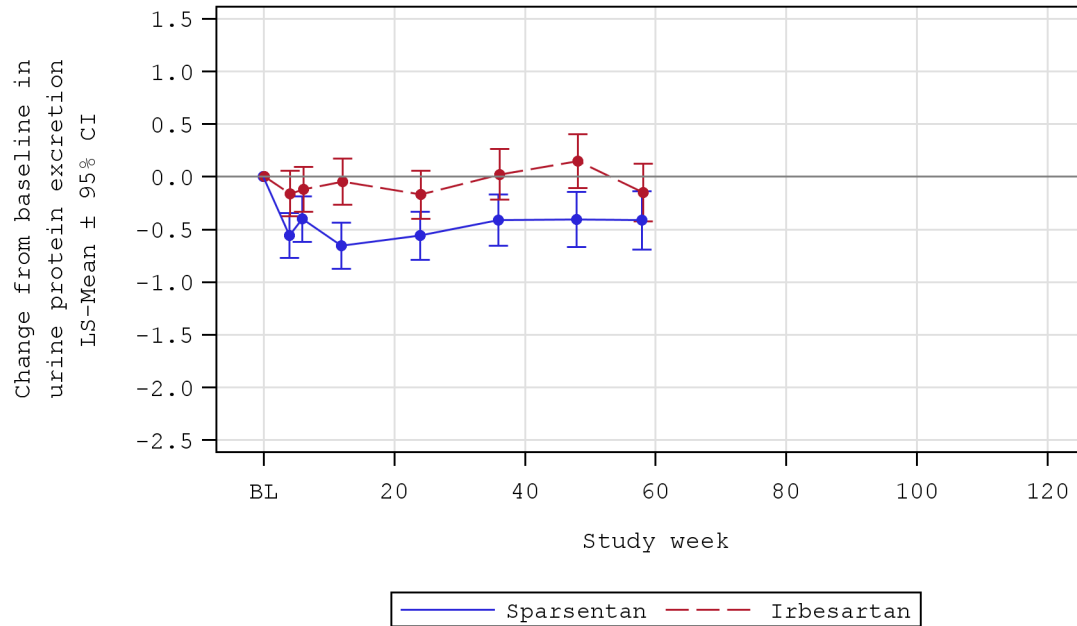


Sparsentan	67	49	42
Irbesartan	70	48	36

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT1UEC\_FSCM.



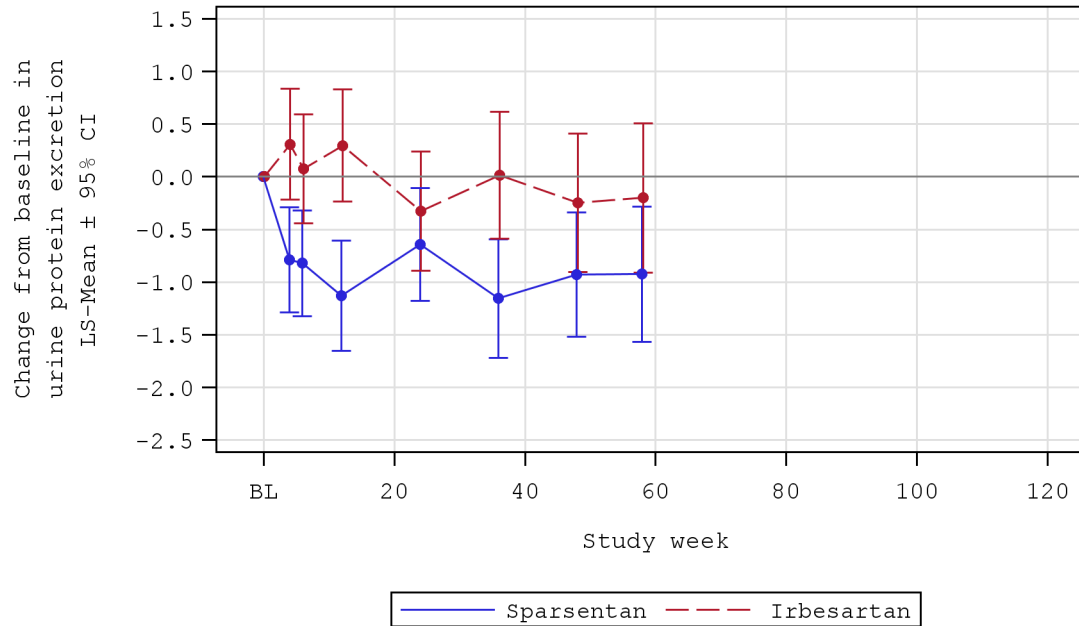
Figure PF1UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR Low and UP Low



Sparsentan	51	37	29
Irbesartan	53	39	30

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT1UEC\_FSCM.

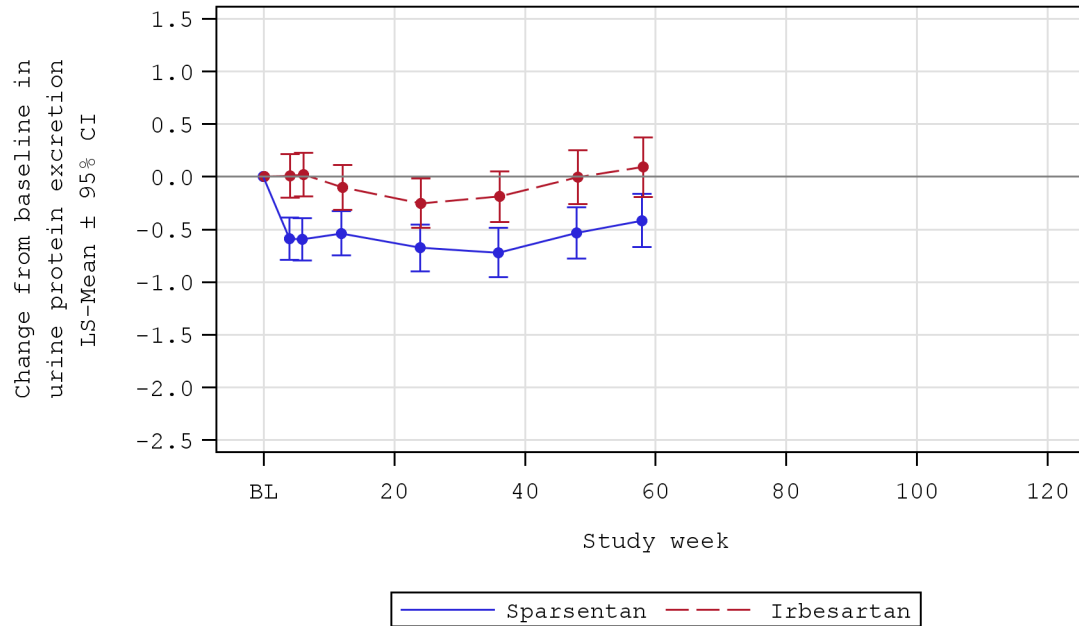
Figure PF1UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR High and UP High



Sparsentan	35	26	19
Irbesartan	33	21	15

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT1UEC\_FSCM.

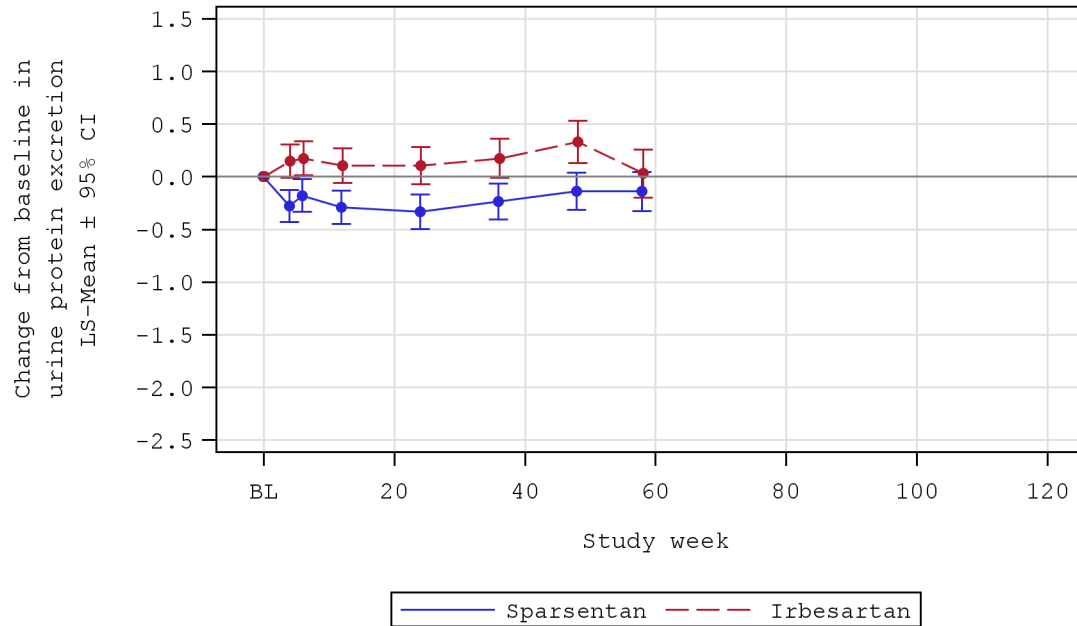
Figure PF1UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR High and UP Low



Sparsentan	39	27	23
Irbesartan	35	25	17

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT1UEC\_FSCM.

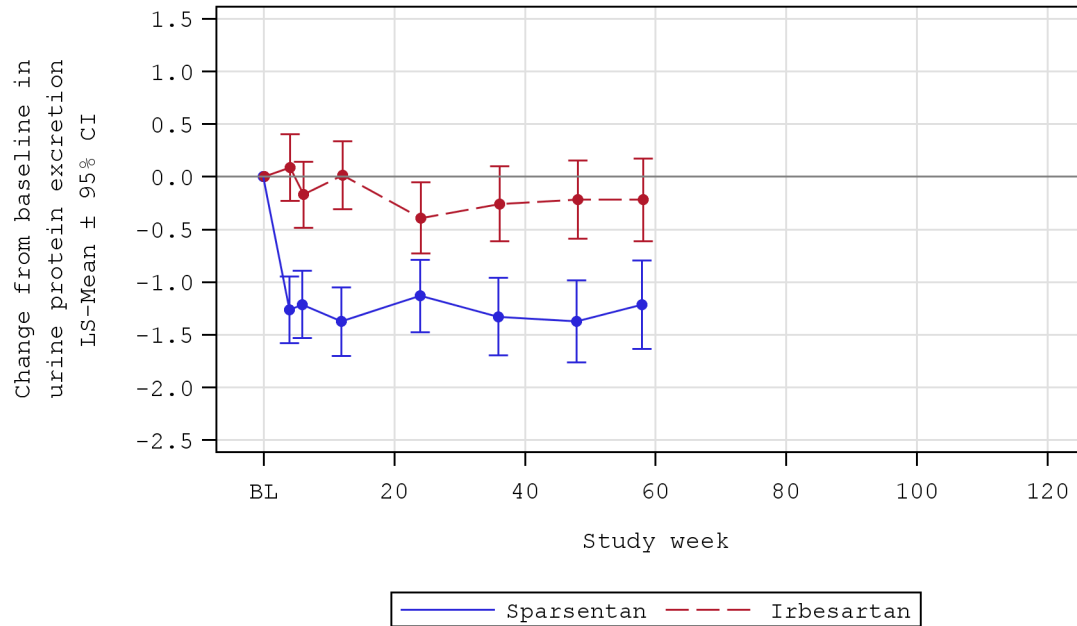
Figure PF1UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline urine protein excretion:  $\leq 1.75$  g/day



Sparsentan	93	73	61
Irbesartan	88	61	38

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low =  $30 < eGFR < 60$  ml/min/1.73m<sup>2</sup>, eGFR High  $\geq 60$  ml/min/1.73m<sup>2</sup>, UP Low =  $\leq 1.75$  g/day, UP High =  $> 1.75$  g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT1UEC\_FSCM.

Figure PF1UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline urine protein excretion: > 1.75 g/day



Sparsentan	99	66	52
Irbesartan	103	72	60

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT1UEC\_FSCM.

Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission										
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
Sex	Week 58	Sparsentan							Interaction test:	0.135
Male	Week 58	Sparsentan	139	69 (49.6)	31 (44.9)	1.929 [1.180, 3.154]	2.687 [1.307, 5.525]	21.6 [5.0, 38.3]	0.008	*
		Irbesartan	143	73 (51.0)	17 (23.3)					
Female	Week 58	Sparsentan	63	44 (69.8)	24 (54.5)	1.136 [0.697, 1.853]	1.300 [0.486, 3.477]	6.5 [-21.1, 34.2]	0.625	
		Irbesartan	59	25 (42.4)	12 (48.0)					

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Week 58	Sparsentan							Interaction test: 0.209
<= 45 years	Week 58	Sparsentan	96	51 (53.1)	26 (51.0)	1.319 [0.833, 2.089]	1.652 [0.729, 3.744]	12.3 [-9.7, 34.3]	0.302
		Irbesartan	99	44 (44.4)	17 (38.6)				
> 45 years	Week 58	Sparsentan	106	62 (58.5)	29 (46.8)	2.105 [1.196, 3.704]	3.076 [1.365, 6.933]	24.6 [6.2, 42.9]	0.007 *
		Irbesartan	103	54 (52.4)	12 (22.2)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Week 58	Sparsentan							Interaction test: 0.072
<= 18 years	Week 58	Sparsentan	9	4 (44.4)	2 (50.0)	0.500 [0.188, 1.332]	0.333 + [0.009, 12.815]	-50.0 [-100.0, 61.5]	1.000
		Irbesartan	5	1 (20.0)	1 (100.0)				
> 18 to 40 years	Week 58	Sparsentan	102	58 (56.9)	32 (55.2)	1.759 [1.102, 2.807]	2.692 [1.227, 5.908]	23.8 [3.9, 43.7]	0.020 *
		Irbesartan	109	51 (46.8)	16 (31.4)				
> 40 years	Week 58	Sparsentan	91	51 (56.0)	21 (41.2)	1.578 [0.878, 2.838]	1.983 [0.837, 4.699]	15.1 [-5.5, 35.7]	0.137
		Irbesartan	88	46 (52.3)	12 (26.1)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024



Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Week 58	Sparsentan							Interaction test: 0.329
North America	Week 58	Sparsentan	35	12 (34.3)	5 (41.7)	3.750 [0.864, 16.280]	5.714 [0.885, 36.888]	30.6 [-7.8, 68.9]	0.084
		Irbesartan	46	18 (39.1)	2 (11.1)				
Europe	Week 58	Sparsentan	98	48 (49.0)	21 (43.8)	1.531 [0.907, 2.585]	1.944 [0.862, 4.385]	15.2 [-5.1, 35.5]	0.150
		Irbesartan	115	56 (48.7)	16 (28.6)				
Asia Pacific	Week 58	Sparsentan	69	53 (76.8)	29 (54.7)	1.194 [0.725, 1.967]	1.428 [0.542, 3.760]	8.9 [-18.2, 35.9]	0.623
		Irbesartan	41	24 (58.5)	11 (45.8)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission										
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
Baseline BMI	Week 58	Sparsentan							Interaction test:	0.615
< 27 kg/m**2	Week 58	Sparsentan	84	46 (54.8)	29 (63.0)	1.801 [1.118, 2.902]	3.168 [1.309, 7.665]	28.0 [5.4, 50.7]	0.017	*
		Irbesartan	94	40 (42.6)	14 (35.0)					
≥ 27 kg/m**2	Week 58	Sparsentan	118	67 (56.8)	26 (38.8)	1.500 [0.884, 2.548]	1.818 [0.845, 3.911]	12.9 [-4.9, 30.8]	0.132	
		Irbesartan	107	58 (54.2)	15 (25.9)					

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Week 58	Sparsentan							Interaction test: 0.532
eGFR Low and UP High	Week 58	Sparsentan	71	42 (59.2)	11 (26.2)	2.357 [0.821, 6.764]	2.839 [0.816, 9.873]	15.1 [-4.3, 34.5]	0.149
		Irbesartan	74	36 (48.6)	4 (11.1)				
eGFR Low and UP Low	Week 58	Sparsentan	55	29 (52.7)	20 (69.0)	1.293 [0.854, 1.957]	1.944 [0.671, 5.638]	15.6 [-12.3, 43.6]	0.288
		Irbesartan	55	30 (54.5)	16 (53.3)				
eGFR High and UP High	Week 58	Sparsentan	37	19 (51.4)	9 (47.4)	1.776 [0.678, 4.657]	2.475 [0.577, 10.617]	20.7 [-17.0, 58.4]	0.296
		Irbesartan	36	15 (41.7)	4 (26.7)				
eGFR High and UP Low	Week 58	Sparsentan	39	23 (59.0)	15 (65.2)	2.217 [1.002, 4.908]	4.500 [1.166, 17.373]	35.8 [1.6, 70.0]	0.054
		Irbesartan	37	17 (45.9)	5 (29.4)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Week 58	Sparsentan							Interaction test: 0.494
< 60 mL/min/1.73 m**2	Week 58	Sparsentan	127	71 (55.9)	32 (45.1)	1.481 [0.954, 2.298]	1.875 [0.937, 3.754]	14.6 [-2.7, 31.9]	0.084
		Irbesartan	129	69 (53.5)	21 (30.4)				
60 to < 90 mL/min/1.73 m**2	Week 58	Sparsentan	49	24 (49.0)	11 (45.8)	2.903 [0.942, 8.949]	4.513 [1.036, 19.657]	30.0 [-0.5, 60.6]	0.052
		Irbesartan	48	19 (39.6)	3 (15.8)				
≥ 90 mL/min/1.73 m**2	Week 58	Sparsentan	26	18 (69.2)	12 (66.7)	1.333 [0.662, 2.687]	2.000 [0.412, 9.712]	16.7 [-29.0, 62.3]	0.444
		Irbesartan	25	10 (40.0)	5 (50.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Week 58	Sparsentan							Interaction test: 0.692
< 45 mL/min/1.73 m**2	Week 58	Sparsentan	82	45 (54.9)	22 (48.9)	1.537 [0.908, 2.599]	2.050 [0.865, 4.856]	17.1 [-5.2, 39.4]	0.132
		Irbesartan	80	44 (55.0)	14 (31.8)				
45 to < 60 mL/min/1.73 m**2	Week 58	Sparsentan	45	26 (57.8)	10 (38.5)	1.374 [0.621, 3.041]	1.607 [0.495, 5.217]	10.5 [-19.1, 40.1]	0.555
		Irbesartan	49	25 (51.0)	7 (28.0)				
60 to < 90 mL/min/1.73 m**2	Week 58	Sparsentan	49	24 (49.0)	11 (45.8)	2.903 [0.942, 8.949]	4.513 [1.036, 19.657]	30.0 [-0.5, 60.6]	0.052
		Irbesartan	48	19 (39.6)	3 (15.8)				
>= 90 mL/min/1.73 m**2	Week 58	Sparsentan	26	18 (69.2)	12 (66.7)	1.333 [0.662, 2.687]	2.000 [0.412, 9.712]	16.7 [-29.0, 62.3]	0.444
		Irbesartan	25	10 (40.0)	5 (50.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URP\_FSPM: Patients with partial remission by subgroup  
Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Week 58	Sparsentan							Interaction test: 0.317
<= 1.75 g/day	Week 58	Sparsentan	98	61 (62.2)	37 (60.7)	1.281 [0.866, 1.894]	1.713 [0.756, 3.882]	13.3 [-8.9, 35.5]	0.218
		Irbesartan	94	38 (40.4)	18 (47.4)				
> 1.75 g/day	Week 58	Sparsentan	104	52 (50.0)	18 (34.6)	1.888 [0.984, 3.623]	2.358 [0.990, 5.620]	16.3 [-1.7, 34.3]	0.055
		Irbesartan	108	60 (55.6)	11 (18.3)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Partial remission means an urinary protein excretion of <1.0 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024

Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Week 58	Sparsentan							Interaction test: 0.293
Yes	Week 58	Sparsentan	88	46 (52.3)	18 (39.1)	1.286 [0.730, 2.265]	1.469 [0.620, 3.483]	8.7 [-12.9, 30.3]	0.512
		Irbesartan	83	46 (55.4)	14 (30.4)				
No	Week 58	Sparsentan	114	67 (58.8)	37 (55.2)	1.914 [1.187, 3.088]	3.042 [1.410, 6.566]	26.4 [7.5, 45.2]	0.005 *
		Irbesartan	119	52 (43.7)	15 (28.8)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission										
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
Time since renal biopsy	Week 58	Sparsentan							Interaction test:	0.015 #
<= 5 years	Week 58	Sparsentan	113	58 (51.3)	24 (41.4)	1.110 [0.707, 1.742]	1.187 [0.565, 2.495]	4.1 [-15.3, 23.5]	0.707	
		Irbesartan	127	59 (46.5)	22 (37.3)					
> 5 years	Week 58	Sparsentan	89	55 (61.8)	31 (56.4)	3.140 [1.544, 6.388]	5.905 [2.225, 15.672]	38.4 [18.4, 58.4]	<0.001	*
		Irbesartan	75	39 (52.0)	7 (17.9)					

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024



Table PT1URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Week 58	Sparsentan							Interaction test: 0.725
Yes	Week 58	Sparsentan	153	83 (54.2)	37 (44.6)	1.551 [1.019, 2.360]	1.993 [1.041, 3.816]	15.8 [0.0, 31.6]	0.051
		Irbesartan	157	80 (51.0)	23 (28.8)				
No	Week 58	Sparsentan	49	30 (61.2)	18 (60.0)	1.800 [0.880, 3.682]	3.000 [0.884, 10.184]	26.7 [-5.7, 59.1]	0.135
		Irbesartan	45	18 (40.0)	6 (33.3)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.167
Male	Sparsentan	139	91 (65.5)	7.1	(6.0, 24.1)	2.811	(2.006, 3.940)	<0.001 *
	Irbesartan	143	59 (41.3)	105.3	(50.3, NE)			
Female	Sparsentan	63	54 (85.7)	6.1	(4.1, 12.7)	2.178	(1.395, 3.400)	<0.001 *
	Irbesartan	59	35 (59.3)	12.4	(6.7, 59.6)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URPT\_FSTM: Time to partial remission by subgroup  
Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan							Interaction test: 0.110
<= 45 years	Sparsentan	96	66 (68.8)	7.1	(4.7, 24.0)	2.121	(1.463, 3.077)	<0.001 *
	Irbesartan	99	51 (51.5)	46.1	(25.1, 94.0)			
> 45 years	Sparsentan	106	79 (74.5)	6.1	(4.9, 13.3)	3.026	(2.067, 4.431)	<0.001 *
	Irbesartan	103	43 (41.7)	105.3	(37.3, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb\_tte, created on: 31MAY2024

Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: 0.889
<= 18 years	Sparsentan	9	4 (44.4)	NE		2.263	(0.207, 24.756)	0.503
	Irbesartan	5	2 (40.0)	24.0	(6.0, NE)			
> 18 to 40 years	Sparsentan	102	72 (70.6)	6.3	(4.7, 13.1)	2.437	(1.681, 3.532)	<0.001 *
	Irbesartan	109	51 (46.8)	59.6	(36.0, 105.3)			
> 40 years	Sparsentan	91	69 (75.8)	6.1	(4.9, 13.3)	2.530	(1.700, 3.767)	<0.001 *
	Irbesartan	88	41 (46.6)	57.4	(24.7, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.645
North America	Sparsentan	35	23 (65.7)	13.3	(4.7, 48.6)	3.514	(1.718, 7.188)	<0.001 *
	Irbesartan	46	13 (28.3)	NE				
Europe	Sparsentan	98	66 (67.3)	7.6	(5.0, 24.1)	2.543	(1.757, 3.680)	<0.001 *
	Irbesartan	115	55 (47.8)	58.1	(36.0, NE)			
Asia Pacific	Sparsentan	69	56 (81.2)	4.9	(4.1, 6.9)	2.040	(1.268, 3.283)	0.003 *
	Irbesartan	41	26 (63.4)	24.7	(6.9, 70.6)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.599
< 27 kg/m**2	Sparsentan	84	67 (79.8)	4.6	(4.1, 9.1)	2.199	(1.494, 3.235)	<0.001 *
	Irbesartan	94	48 (51.1)	38.0	(24.0, 114.1)			
≥ 27 kg/m**2	Sparsentan	118	78 (66.1)	12.0	(6.1, 34.6)	2.859	(1.962, 4.166)	<0.001 *
	Irbesartan	107	45 (42.1)	82.9	(37.3, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan							Interaction test: 0.961
eGFR Low and UP High	Sparsentan	71	36 (50.7)	51.1	(25.1, 112.1)	2.635	(1.494, 4.647)	<0.001 *
	Irbesartan	74	18 (24.3)	114.1	(94.0, NE)			
eGFR Low and UP Low	Sparsentan	55	50 (90.9)	4.3	(4.1, 4.7)	2.245	(1.458, 3.457)	<0.001 *
	Irbesartan	55	37 (67.3)	12.1	(4.9, 25.1)			
eGFR High and UP High	Sparsentan	37	22 (59.5)	24.7	(6.9, 81.9)	2.712	(1.308, 5.624)	0.007 *
	Irbesartan	36	11 (30.6)	82.9	(51.9, NE)			
eGFR High and UP Low	Sparsentan	39	37 (94.9)	4.1	(4.1, 4.4)	2.540	(1.522, 4.240)	<0.001 *
	Irbesartan	37	28 (75.7)	12.1	(6.1, 26.1)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.368
< 60 mL/min/1.73 m**2	Sparsentan	127	88 (69.3)	12.0	(6.0, 23.7)	2.393	(1.695, 3.378)	<0.001 *
	Irbesartan	129	58 (45.0)	70.6	(36.0, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	37 (75.5)	6.1	(4.6, 24.1)	3.705	(2.032, 6.753)	<0.001 *
	Irbesartan	48	19 (39.6)	82.9	(36.0, NE)			
≥ 90 mL/min/1.73 m**2	Sparsentan	26	20 (76.9)	4.1	(4.1, 24.7)	1.652	(0.834, 3.271)	0.150
	Irbesartan	25	17 (68.0)	24.7	(6.1, 36.3)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024



Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: 0.400
< 45 mL/min/1.73 m**2	Sparsentan	82	54 (65.9)	12.4	(6.1, 38.0)	2.170	(1.413, 3.331)	<0.001 *
	Irbesartan	80	35 (43.8)	105.3	(25.6, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	34 (75.6)	6.0	(4.1, 12.7)	3.054	(1.660, 5.618)	<0.001 *
	Irbesartan	49	23 (46.9)	70.6	(24.1, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	37 (75.5)	6.1	(4.6, 24.1)	3.705	(2.032, 6.753)	<0.001 *
	Irbesartan	48	19 (39.6)	82.9	(36.0, NE)			
≥ 90 mL/min/1.73 m**2	Sparsentan	26	20 (76.9)	4.1	(4.1, 24.7)	1.652	(0.834, 3.271)	0.150
	Irbesartan	25	17 (68.0)	24.7	(6.1, 36.3)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.210
<= 1.75 g/day	Sparsentan	98	88 (89.8)	4.3	(4.1, 4.7)	2.068	(1.494, 2.864)	<0.001 *
	Irbesartan	94	65 (69.1)	12.1	(6.1, 25.1)			
> 1.75 g/day	Sparsentan	104	57 (54.8)	37.3	(23.7, 69.3)	3.479	(2.175, 5.565)	<0.001 *
	Irbesartan	108	29 (26.9)	114.1	(94.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.042 #
Yes	Sparsentan	88	64 (72.7)	7.1	(6.0, 24.1)	3.658	(2.370, 5.645)	<0.001 *
	Irbesartan	83	36 (43.4)	70.6	(37.3, NE)			
No	Sparsentan	114	81 (71.1)	6.1	(4.7, 12.7)	1.909	(1.353, 2.692)	<0.001 *
	Irbesartan	119	58 (48.7)	46.1	(25.3, 114.1)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.102
<= 5 years	Sparsentan	113	81 (71.7)	6.9	(4.9, 22.1)	2.161	(1.554, 3.004)	<0.001 *
	Irbesartan	127	66 (52.0)	36.1	(25.1, 82.9)			
> 5 years	Sparsentan	89	64 (71.9)	6.9	(4.7, 24.0)	3.446	(2.169, 5.475)	<0.001 *
	Irbesartan	75	28 (37.3)	105.3	(57.4, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

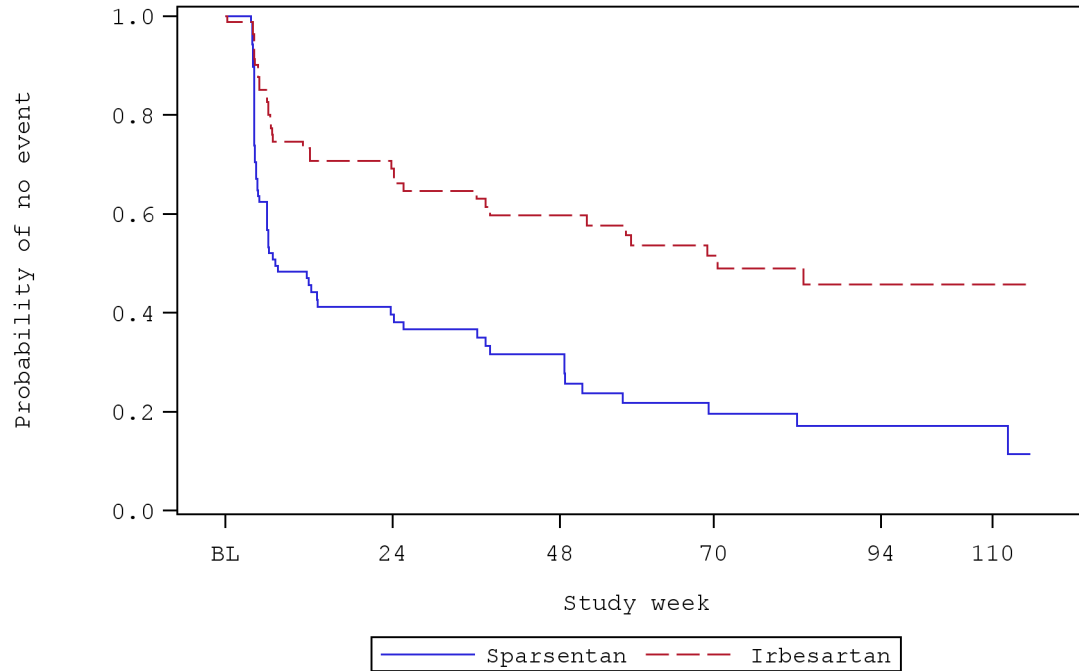
Table PT1URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension		Sparsentan				Interaction test: 0.647		
Yes	Sparsentan	153	110 (71.9)	7.6	(6.0, 22.1)	2.712	(1.998, 3.681)	<0.001 *
	Irbesartan	157	70 (44.6)	70.6	(36.3, NE)			
No	Sparsentan	49	35 (71.4)	4.4	(4.1, 12.7)	2.040	(1.186, 3.507)	0.010 *
	Irbesartan	45	24 (53.3)	26.1	(8.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Figure PF1URPT\_FSKM: Time to partial remission by subgroup  
 Full Analysis Set

Study: PROTECT  
 Baseline use of antihypertensives: Yes

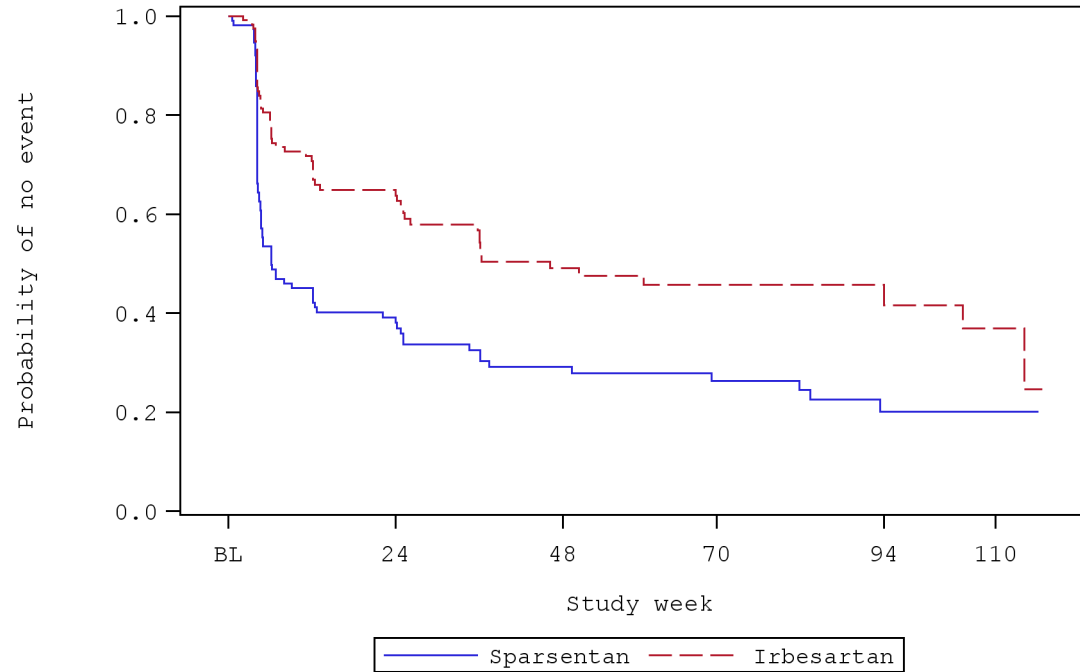


Sparsentan	88	26	17	9	3	3
Irbesartan	83	47	31	22	9	4

Partial remission means an urinary protein excretion of <1.0 g/day. No event means no partial remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT1URPT\_FSTM

Figure PF1URPT\_FSKM: Time to partial remission by subgroup  
 Full Analysis Set

Study: PROTECT  
 Baseline use of antihypertensives: No



Sparsentan	114	36	25	16	8	3
Irbesartan	119	59	38	23	11	4

Partial remission means an urinary protein excretion of <1.0 g/day. No event means no partial remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT1URPT\_FSTM

Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Week 58	Sparsentan							Interaction test: 0.792
Male	Week 58	Sparsentan	139	69 (49.6)	11 (15.9)	2.909 [0.972, 8.706]	3.272 [0.989, 10.823]	10.5 [-1.0, 22.0]	0.056
		Irbesartan	143	73 (51.0)	4 (5.5)				
Female	Week 58	Sparsentan	63	44 (69.8)	8 (18.2)	2.273 [0.523, 9.881]	2.556 [0.498, 13.114]	10.2 [-8.5, 28.9]	0.308
		Irbesartan	59	25 (42.4)	2 (8.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024



Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Week 58	Sparsentan							Interaction test: 0.539
<= 45 years	Week 58	Sparsentan	96	51 (53.1)	9 (17.6)	3.882 [0.885, 17.023]	4.500 [0.917, 22.083]	13.1 [-1.2, 27.4]	0.058
		Irbesartan	99	44 (44.4)	2 (4.5)				
> 45 years	Week 58	Sparsentan	106	62 (58.5)	10 (16.1)	2.177 [0.724, 6.546]	2.404 [0.708, 8.165]	8.7 [-4.5, 22.0]	0.168
		Irbesartan	103	54 (52.4)	4 (7.4)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Week 58	Sparsentan							Interaction test: 0.681
<= 18 years	Week 58	Sparsentan	9	4 (44.4)	1 (25.0)	1.200 + [0.077, 18.745]	1.286 + [0.031, 53.512]	25.0 [-79.9, 100.0]	1.000
		Irbesartan	5	1 (20.0)	0 (0.0)				
> 18 to 40 years	Week 58	Sparsentan	102	58 (56.9)	9 (15.5)	3.957 [0.896, 17.474]	4.500 [0.925, 21.902]	11.6 [-1.0, 24.2]	0.058
		Irbesartan	109	51 (46.8)	2 (3.9)				
> 40 years	Week 58	Sparsentan	91	51 (56.0)	9 (17.6)	2.029 [0.670, 6.148]	2.250 [0.643, 7.877]	9.0 [-6.4, 24.3]	0.242
		Irbesartan	88	46 (52.3)	4 (8.7)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Week 58	Sparsentan							Interaction test: 0.761
North America	Week 58	Sparsentan	35	12 (34.3)	0 (0.0)	NE	NE	0.0 [NE, NE]	NE
		Irbesartan	46	18 (39.1)	0 (0.0)				
Europe	Week 58	Sparsentan	98	48 (49.0)	8 (16.7)	3.111 [0.874, 11.074]	3.533 [0.881, 14.171]	11.3 [-2.7, 25.3]	0.107
		Irbesartan	115	56 (48.7)	3 (5.4)				
Asia Pacific	Week 58	Sparsentan	69	53 (76.8)	11 (20.8)	1.660 [0.509, 5.414]	1.833 [0.461, 7.287]	8.3 [-11.9, 28.4]	0.529
		Irbesartan	41	24 (58.5)	3 (12.5)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Week 58	Sparsentan							Interaction test: 0.064
< 27 kg/m**2	Week 58	Sparsentan	84	46 (54.8)	12 (26.1)	10.435 [1.419, 76.757]	13.765 [1.700, 111.419]	23.6 [7.7, 39.5]	0.002 *
		Irbesartan	94	40 (42.6)	1 (2.5)				
≥ 27 kg/m**2	Week 58	Sparsentan	118	67 (56.8)	7 (10.4)	1.212 [0.406, 3.614]	1.237 [0.370, 4.129]	1.8 [-10.1, 13.7]	0.771
		Irbesartan	107	58 (54.2)	5 (8.6)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Week 58	Sparsentan							Interaction test: NE
eGFR Low and UP High	Week 58	Sparsentan	71	42 (59.2)	2 (4.8)	all n<10			NE
		Irbesartan	74	36 (48.6)	1 (2.8)				
eGFR Low and UP Low	Week 58	Sparsentan	55	29 (52.7)	6 (20.7)	all n<10			NE
		Irbesartan	55	30 (54.5)	3 (10.0)				
eGFR High and UP High	Week 58	Sparsentan	37	19 (51.4)	5 (26.3)	all n<10			NE
		Irbesartan	36	15 (41.7)	0 (0.0)				
eGFR High and UP Low	Week 58	Sparsentan	39	23 (59.0)	6 (26.1)	all n<10			NE
		Irbesartan	37	17 (45.9)	2 (11.8)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Week 58	Sparsentan							Interaction test: 0.671
< 60 mL/min/1.73 m**2	Week 58	Sparsentan	127	71 (55.9)	8 (11.3)	1.944 [0.613, 6.161]	2.063 [0.592, 7.197]	5.5 [-5.2, 16.1]	0.367
		Irbesartan	129	69 (53.5)	4 (5.8)				
60 to < 90 mL/min/1.73 m**2	Week 58	Sparsentan	49	24 (49.0)	7 (29.2)	5.542 [0.745, 41.235]	7.412 [0.823, 66.734]	23.9 [-1.6, 49.4]	0.059
		Irbesartan	48	19 (39.6)	1 (5.3)				
≥ 90 mL/min/1.73 m**2	Week 58	Sparsentan	26	18 (69.2)	4 (22.2)	2.222 [0.286, 17.269]	2.571 [0.246, 26.851]	12.2 [-22.3, 46.7]	0.626
		Irbesartan	25	10 (40.0)	1 (10.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Week 58	Sparsentan							Interaction test: NE
< 45 mL/min/1.73 m**2	Week 58	Sparsentan	82	45 (54.9)	6 (13.3)	all n<10			NE
		Irbesartan	80	44 (55.0)	3 (6.8)				
45 to < 60 mL/min/1.73 m**2	Week 58	Sparsentan	45	26 (57.8)	2 (7.7)	all n<10			NE
		Irbesartan	49	25 (51.0)	1 (4.0)				
60 to < 90 mL/min/1.73 m**2	Week 58	Sparsentan	49	24 (49.0)	7 (29.2)	all n<10			NE
		Irbesartan	48	19 (39.6)	1 (5.3)				
>= 90 mL/min/1.73 m**2	Week 58	Sparsentan	26	18 (69.2)	4 (22.2)	all n<10			NE
		Irbesartan	25	10 (40.0)	1 (10.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URF\_FSPM: Patients with complete remission by subgroup  
Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Week 58	Sparsentan							Interaction test: 0.311
<= 1.75 g/day	Week 58	Sparsentan	98	61 (62.2)	14 (23.0)	1.744 [0.683, 4.453]	1.966 [0.645, 5.989]	9.8 [-7.4, 27.0]	0.298
		Irbesartan	94	38 (40.4)	5 (13.2)				
> 1.75 g/day	Week 58	Sparsentan	104	52 (50.0)	5 (9.6)	5.769 [0.696, 47.809]	6.277 [0.709, 55.578]	7.9 [-2.5, 18.4]	0.095
		Irbesartan	108	60 (55.6)	1 (1.7)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Complete remission means an urinary protein excretion of <0.3 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024



Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission										
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
Baseline use of antihypertensives	Week 58	Sparsentan								Interaction test: 0.026 #
Yes	Week 58	Sparsentan	88	46 (52.3)	4 (8.7)	0.800 [0.229, 2.791]	0.781 [0.196, 3.115]	-2.2 [-16.5, 12.1]	1.000	
		Irbesartan	83	46 (55.4)	5 (10.9)					
No	Week 58	Sparsentan	114	67 (58.8)	15 (22.4)	11.642 [1.589, 85.300]	14.712 [1.874, 115.515]	20.5 [8.1, 32.8]	<0.001	*
		Irbesartan	119	52 (43.7)	1 (1.9)					

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Week 58	Sparsentan							Interaction test: 0.322
<= 5 years	Week 58	Sparsentan	113	58 (51.3)	10 (17.2)	2.034 [0.741, 5.589]	2.250 [0.718, 7.048]	8.8 [-5.0, 22.5]	0.178
		Irbesartan	127	59 (46.5)	5 (8.5)				
> 5 years	Week 58	Sparsentan	89	55 (61.8)	9 (16.4)	6.382 [0.843, 48.341]	7.435 [0.901, 61.335]	13.8 [0.6, 27.0]	0.042 *
		Irbesartan	75	39 (52.0)	1 (2.6)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Week 58	Sparsentan							Interaction test: 0.300
Yes	Week 58	Sparsentan	153	83 (54.2)	12 (14.5)	1.928 [0.760, 4.889]	2.085 [0.742, 5.854]	7.0 [-3.8, 17.7]	0.212
		Irbesartan	157	80 (51.0)	6 (7.5)				
No	Week 58	Sparsentan	49	30 (61.2)	7 (23.3)	9.194 + [0.556, 151.969]	11.809 + [0.633, 220.441]	23.3 [3.8, 42.9]	0.036 *
		Irbesartan	45	18 (40.0)	0 (0.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.632
Male	Sparsentan	139	30 (21.6)	NE		2.906	(1.507, 5.602)	0.001 *
	Irbesartan	143	13 (9.1)	NE				
Female	Sparsentan	63	13 (20.6)	NE		2.086	(0.647, 6.724)	0.218
	Irbesartan	59	4 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.872
<= 45 years	Sparsentan	96	26 (27.1)	NE		2.750	(1.319, 5.730)	0.007 *
	Irbesartan	99	10 (10.1)	NE				
> 45 years	Sparsentan	106	17 (16.0)	NE		2.458	(1.011, 5.977)	0.047 *
	Irbesartan	103	7 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis		Sparsentan				Interaction test:		NE
<= 18 years	Sparsentan	9	1 (11.1) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	26 (25.5)	NE		2.731	(1.303, 5.726)	0.008 *
	Irbesartan	109	10 (9.2)	NE				
> 40 years	Sparsentan	91	16 (17.6)	NE		2.203	(0.899, 5.398)	0.084
	Irbesartan	88	7 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.118
North America	Sparsentan	35	4 (11.4)	NE		1.218	(0.244, 6.078)	0.810
	Irbesartan	46	3 (6.5)	NE				
Europe	Sparsentan	98	21 (21.4)	NE		4.370	(1.847, 10.336)	<0.001 *
	Irbesartan	115	7 (6.1)	NE				
Asia Pacific	Sparsentan	69	18 (26.1)	NE		1.430	(0.585, 3.496)	0.433
	Irbesartan	41	7 (17.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.528
< 27 kg/m**2	Sparsentan	84	22 (26.2)	NE		3.107	(1.362, 7.087)	0.007 *
	Irbesartan	94	8 (8.5)	NE				
≥ 27 kg/m**2	Sparsentan	118	21 (17.8)	NE		2.509	(1.102, 5.712)	0.028 *
	Irbesartan	107	8 (7.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024



Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.808
eGFR Low and UP High	Sparsentan	71	5 (7.0)	NE		1.591	(0.378, 6.684)	0.526
	Irbesartan	74	3 (4.1)	NE				
eGFR Low and UP Low	Sparsentan	55	18 (32.7)	94.3	(59.0, NE)	3.475	(1.376, 8.777)	0.008 *
	Irbesartan	55	6 (10.9)	NE				
eGFR High and UP High	Sparsentan	37	6 (16.2)	NE		2.831	(0.570, 14.070)	0.203
	Irbesartan	36	2 (5.6)	NE				
eGFR High and UP Low	Sparsentan	39	14 (35.9)	NE		2.115	(0.802, 5.573)	0.130
	Irbesartan	37	6 (16.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.058
< 60 mL/min/1.73 m**2	Sparsentan	127	23 (18.1)	NE		2.641	(1.207, 5.780)	0.015 *
	Irbesartan	129	9 (7.0)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	14 (28.6)	NE		4.870	(1.334, 17.774)	0.017 *
	Irbesartan	48	3 (6.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	6 (23.1)	NE		0.780	(0.232, 2.618)	0.688
	Irbesartan	25	5 (20.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: 0.011 #
< 45 mL/min/1.73 m**2	Sparsentan	82	12 (14.6)	NE		1.274	(0.507, 3.202)	0.606
	Irbesartan	80	8 (10.0)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	11 (24.4)	NE		13.639	(1.734, 107.307)	0.013 *
	Irbesartan	49	1 (2.0)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	14 (28.6)	NE		4.870	(1.334, 17.774)	0.017 *
	Irbesartan	48	3 (6.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	6 (23.1)	NE		0.780	(0.232, 2.618)	0.688
	Irbesartan	25	5 (20.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.426
<= 1.75 g/day	Sparsentan	98	31 (31.6)	NE		2.220	(1.160, 4.250)	0.016 *
	Irbesartan	94	13 (13.8)	NE				
> 1.75 g/day	Sparsentan	104	12 (11.5)	NE		4.243	(1.296, 13.891)	0.017 *
	Irbesartan	108	4 (3.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.034 #
Yes	Sparsentan	88	11 (12.5)	NE		1.294	(0.542, 3.090)	0.562
	Irbesartan	83	10 (12.0)	NE				
No	Sparsentan	114	32 (28.1)	NE		4.035	(1.761, 9.247)	<0.001 *
	Irbesartan	119	7 (5.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.210
<= 5 years	Sparsentan	113	25 (22.1)	NE		2.232	(1.156, 4.308)	0.017 *
	Irbesartan	127	14 (11.0)	NE				
> 5 years	Sparsentan	89	18 (20.2)	NE		5.140	(1.496, 17.657)	0.009 *
	Irbesartan	75	3 (4.0)	NE				

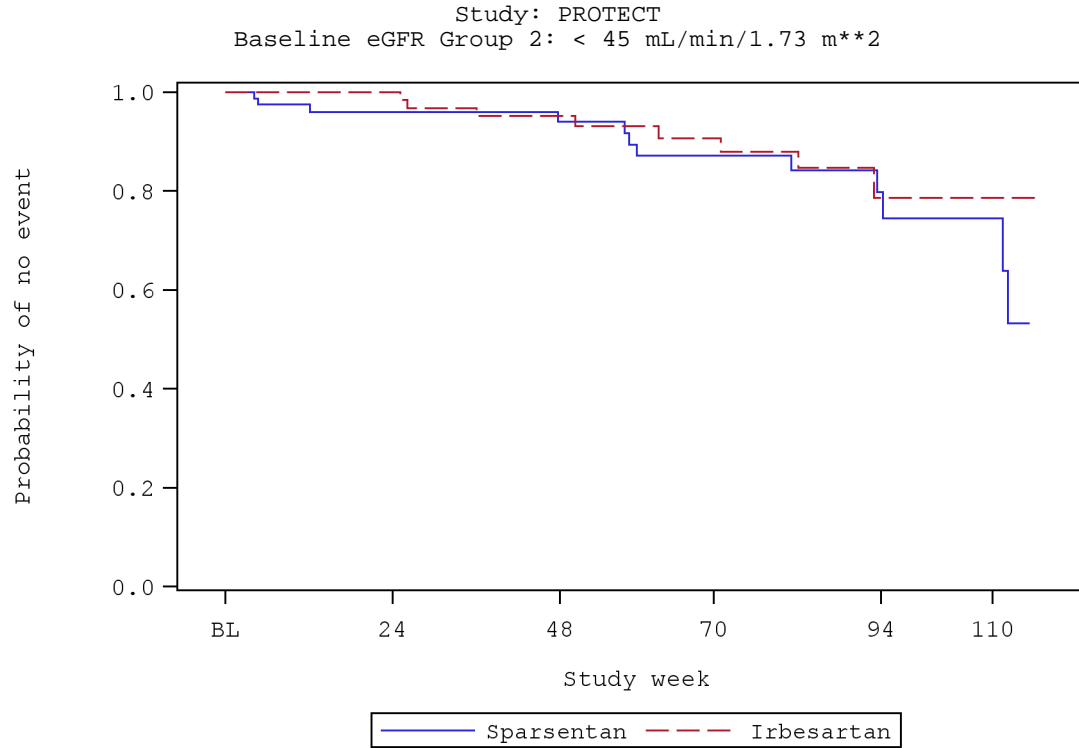
N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Table PT1URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension						Interaction test:		0.366
Yes	Sparsentan	153	31 (20.3)	NE		3.082	(1.547, 6.138)	0.001 *
	Irbesartan	157	11 (7.0)	NE				
No	Sparsentan	49	12 (24.5)	NE		1.401	(0.494, 3.970)	0.526
	Irbesartan	45	6 (13.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 31MAY2024

Figure PF1URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set

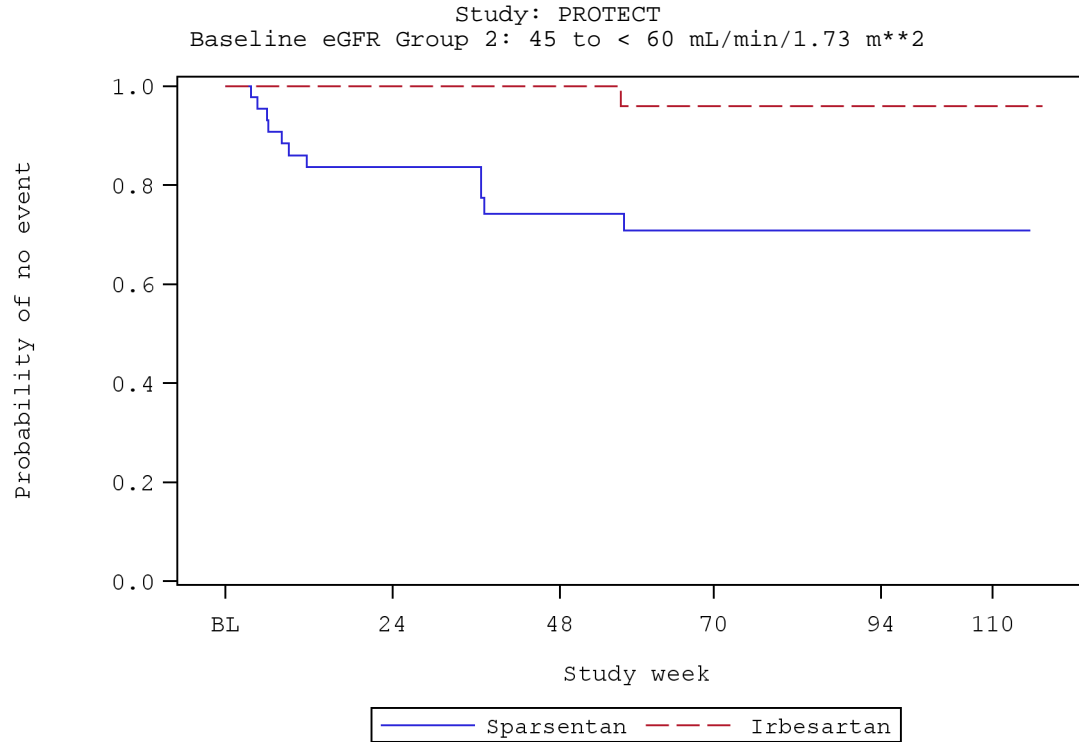


Sparsentan	82	59	47	35	18	8
Irbesartan	80	67	51	35	11	2

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT1URFT\_FSTM



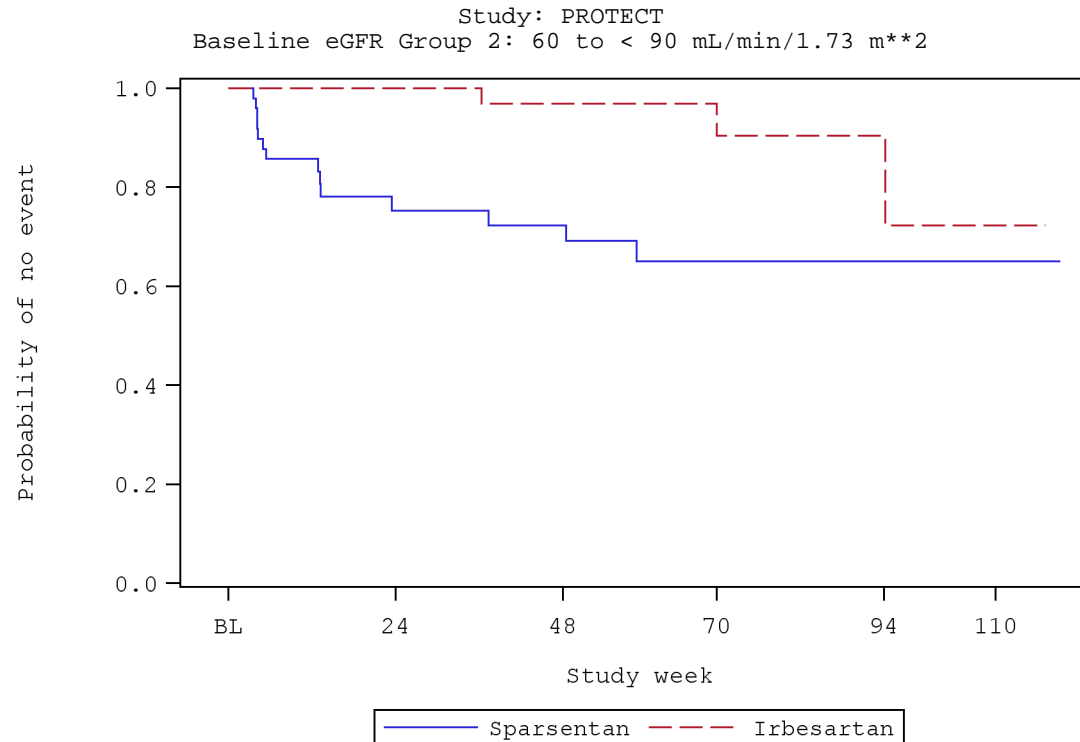
Figure PF1URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	45	30	23	18	8	4
Irbesartan	49	35	28	22	13	6

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT1URFT\_FSTM

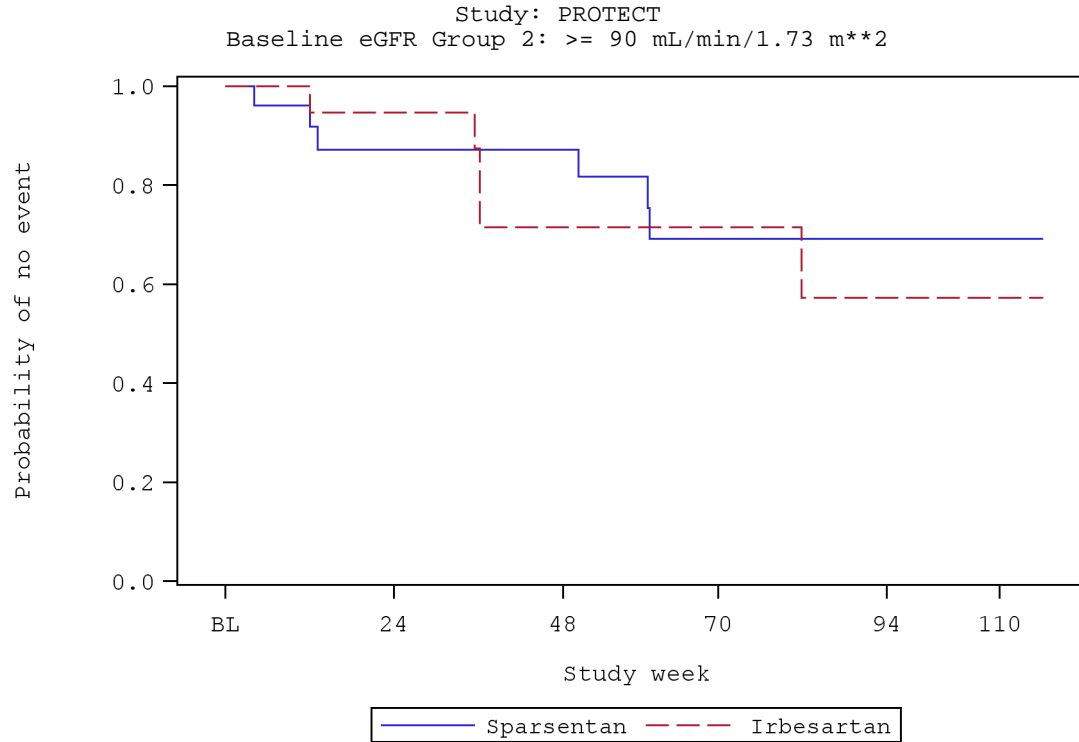
Figure PF1URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	49	27	23	15	6	2
Irbesartan	48	34	27	15	5	2

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT1URFT\_FSTM

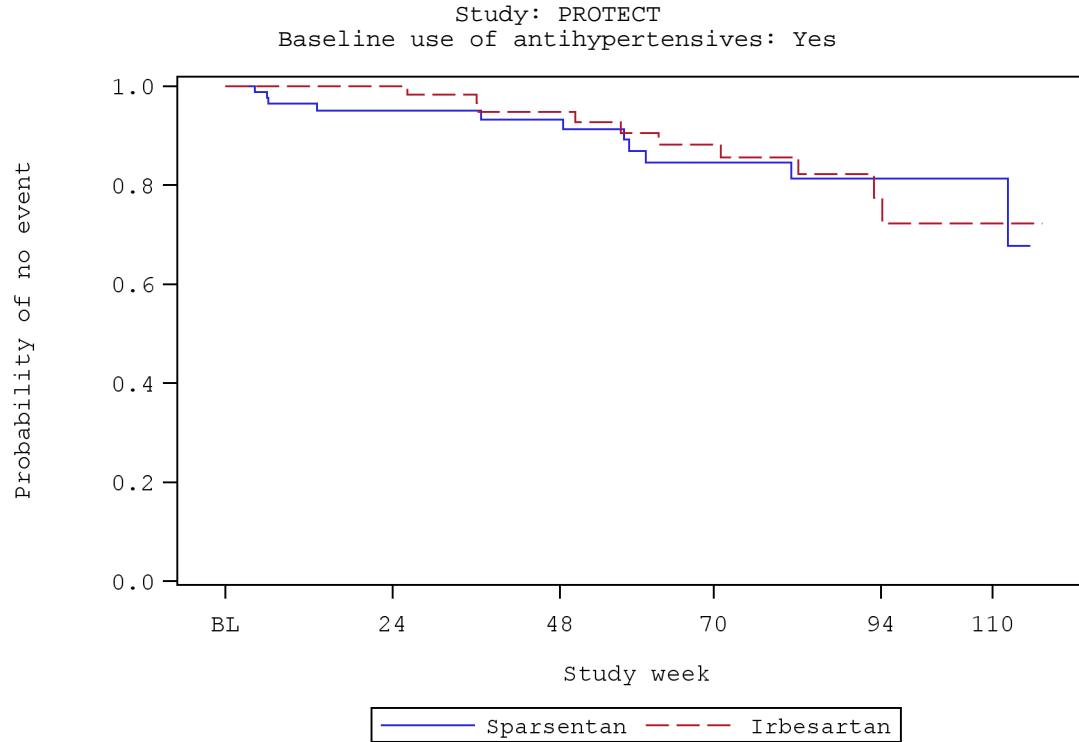
Figure PF1URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	26	18	17	10	4	2
Irbesartan	25	16	7	6	3	2

Complete remission means an urinary protein excretion of  $<0.3$  g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT1URFT\_FSTM

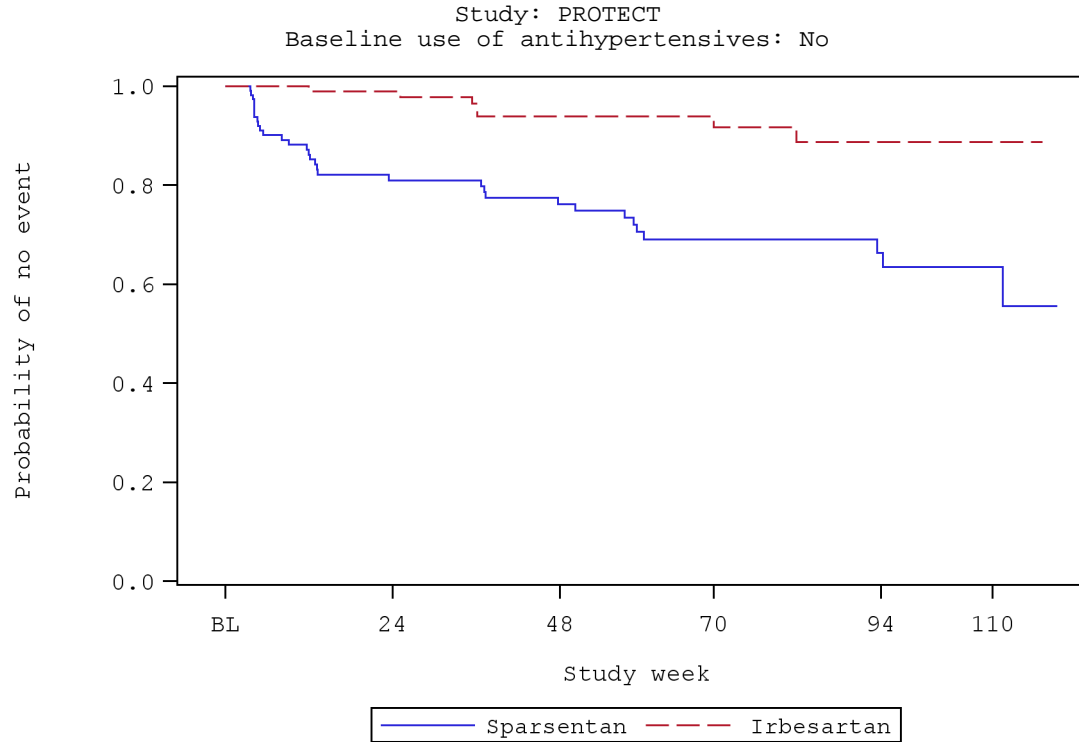
Figure PF1URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	88	60	48	34	12	7
Irbesartan	83	66	47	36	15	5

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT1URFT\_FSTM

Figure PF1URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	114	74	62	44	24	9
Irbesartan	119	86	66	42	17	7

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT1URFT\_FSTM

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Sex														
Male	eGFR	Baseline	Sparsentan	139	139 (100.0)	53.52 (22.58)	25.0	36.00	45.00	66.00	126.0			
			Irbesartan	143	143 (100.0)	57.03 (23.53)	26.0	38.00	50.00	70.00	123.0			
		Week 6	Sparsentan	139	133 (95.7)	52.58 (22.55)	21.0	35.00	48.00	65.00	121.0			
			Irbesartan	143	139 (97.2)	55.73 (23.29)	21.0	37.00	50.00	70.00	120.0			
		Week 36	Sparsentan	139	92 (66.2)	50.26 (22.05)	18.0	34.00	46.00	62.00	123.0			
			Irbesartan	143	97 (67.8)	51.90 (23.55)	7.0	34.00	48.00	64.00	125.0			
		Week 58	Sparsentan	139	64 (46.0)	47.86 (21.45)	13.0	33.00	42.00	59.50	121.0			
			Irbesartan	143	70 (49.0)	50.09 (22.58)	18.0	33.00	44.50	66.00	122.0			
			Change from baseline in eGFR	Week 6	Sparsentan	139	133 (95.7)	-0.89 (6.70)	-27.0	-4.00	-1.00	2.00	20.0	0.04 [-0.19, 0.28]
		Irbesartan			143	139 (97.2)	-1.18 (6.63)	-31.0	-5.00	-1.00	2.00	20.0		
		Week 36		Sparsentan	139	92 (66.2)	-3.24 (7.63)	-26.0	-8.00	-2.00	1.00	26.0	0.11 [-0.18, 0.39]	
				Irbesartan	143	97 (67.8)	-4.07 (8.14)	-32.0	-7.00	-4.00	0.00	15.0		
		Week 58		Sparsentan	139	64 (46.0)	-4.83 (7.63)	-33.0	-8.00	-4.00	0.00	13.0	0.17 [-0.16, 0.51]	
				Irbesartan	143	70 (49.0)	-6.19 (7.88)	-24.0	-12.00	-7.00	-1.00	12.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Female	eGFR	Baseline	Sparsentan	63	63 (100.0)	64.22 (26.67)	24.0	41.00	59.00	86.00	128.0			
			Irbesartan	59	59 (100.0)	57.19 (23.89)	27.0	39.00	50.00	70.00	123.0			
		Week 6	Sparsentan	63	60 (95.2)	63.32 (24.52)	26.0	44.00	60.00	82.00	120.0			
			Irbesartan	59	56 (94.9)	55.20 (23.83)	19.0	39.00	47.00	68.00	116.0			
		Week 36	Sparsentan	63	44 (69.8)	62.57 (23.33)	25.0	45.00	60.00	81.50	114.0			
			Irbesartan	59	36 (61.0)	47.08 (20.70)	15.0	33.50	42.00	55.50	109.0			
		Week 58	Sparsentan	63	46 (73.0)	61.89 (25.89)	23.0	43.00	56.50	85.00	133.0			
			Irbesartan	59	25 (42.4)	44.32 (20.63)	13.0	32.00	38.00	54.00	116.0			
		Change from baseline in eGFR		Week 6	Sparsentan	63	60 (95.2)	-1.67 (9.58)	-43.0	-4.00	-0.50	3.00	14.0	0.09 [-0.28, 0.45]
					Irbesartan	59	56 (94.9)	-2.39 (6.34)	-19.0	-6.50	-2.50	3.50	9.0	
				Week 36	Sparsentan	63	44 (69.8)	-5.09 (10.56)	-39.0	-9.50	-4.00	1.00	17.0	0.06 [-0.38, 0.50]
					Irbesartan	59	36 (61.0)	-5.61 (6.34)	-24.0	-9.00	-5.00	-1.00	3.0	
	Week 58			Sparsentan	63	46 (73.0)	-4.98 (10.62)	-34.0	-10.00	-5.00	2.00	24.0	0.16 [-0.33, 0.65]	
				Irbesartan	59	25 (42.4)	-6.52 (8.05)	-24.0	-12.00	-6.00	0.00	8.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age														
<= 45 years	eGFR	Baseline	Sparsentan	96	96 (100.0)	63.16 (27.44)	25.0	41.00	55.50	83.00	128.0			
			Irbesartan	99	99 (100.0)	65.52 (26.04)	26.0	42.00	63.00	80.00	123.0			
		Week 6	Sparsentan	96	93 (96.9)	61.13 (26.48)	23.0	38.00	56.00	82.00	121.0			
			Irbesartan	99	97 (98.0)	63.19 (25.41)	23.0	43.00	57.00	80.00	120.0			
		Week 36	Sparsentan	96	65 (67.7)	61.75 (26.47)	18.0	42.00	56.00	84.00	123.0			
			Irbesartan	99	65 (65.7)	55.91 (26.42)	7.0	37.00	53.00	65.00	125.0			
		Week 58	Sparsentan	96	49 (51.0)	63.96 (27.96)	13.0	42.00	58.00	85.00	133.0			
			Irbesartan	99	43 (43.4)	53.70 (26.09)	13.0	34.00	45.00	69.00	122.0			
		Change from baseline in eGFR	Week 6	Sparsentan	96	93 (96.9)	-2.28 (8.84)	-43.0	-5.00	-1.00	1.00	15.0	0.01 [-0.28, 0.29]	
				Irbesartan	99	97 (98.0)	-2.35 (6.22)	-17.0	-6.00	-2.00	2.00	14.0		
				Week 36	Sparsentan	96	65 (67.7)	-5.72 (9.89)	-39.0	-12.00	-3.00	1.00	12.0	-0.01 [-0.35, 0.34]
					Irbesartan	99	65 (65.7)	-5.66 (8.94)	-32.0	-12.00	-5.00	1.00	15.0	
				Week 58	Sparsentan	96	49 (51.0)	-6.14 (11.23)	-34.0	-15.00	-5.00	0.00	24.0	0.13 [-0.28, 0.54]
					Irbesartan	99	43 (43.4)	-7.40 (8.35)	-24.0	-13.00	-7.00	0.00	10.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 45 years	eGFR	Baseline	Sparsentan	106	106 (100.0)	51.15 (19.68)	24.0	36.00	45.00	64.00	103.0	
			Irbesartan	103	103 (100.0)	48.97 (17.56)	27.0	36.00	44.00	56.00	104.0	
		Week 6	Sparsentan	106	100 (94.3)	51.07 (19.59)	21.0	35.00	49.00	64.50	104.0	
			Irbesartan	103	98 (95.1)	48.05 (18.43)	19.0	35.00	43.00	59.00	102.0	
		Week 36	Sparsentan	106	71 (67.0)	47.37 (17.02)	24.0	34.00	45.00	62.00	97.0	
			Irbesartan	103	68 (66.0)	45.51 (17.53)	23.0	32.00	39.50	56.00	98.0	
		Week 58	Sparsentan	106	61 (57.5)	45.51 (17.14)	21.0	32.00	43.00	56.00	94.0	
			Irbesartan	103	52 (50.5)	44.33 (17.35)	18.0	31.00	38.00	55.00	88.0	
	Change from baseline in eGFR	Week 6	Sparsentan	106	100 (94.3)	-0.06 (6.30)	-21.0	-3.00	0.00	3.00	20.0	0.10 [-0.18, 0.38]
			Irbesartan	103	98 (95.1)	-0.71 (6.81)	-31.0	-4.00	0.00	3.00	20.0	
		Week 36	Sparsentan	106	71 (67.0)	-2.11 (7.06)	-22.0	-7.00	-2.00	1.00	26.0	0.19 [-0.14, 0.52]
			Irbesartan	103	68 (66.0)	-3.37 (6.15)	-25.0	-6.00	-3.00	-1.00	13.0	
		Week 58	Sparsentan	106	61 (57.5)	-3.89 (6.51)	-19.0	-8.00	-4.00	0.00	14.0	0.21 [-0.16, 0.58]
			Irbesartan	103	52 (50.5)	-5.35 (7.43)	-24.0	-10.00	-6.00	-0.50	12.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Age at IgAN diagnosis													
<= 18 years	eGFR	Baseline	Sparsentan	9	9 (100.0)	57.33 (29.45)	32.0	35.00	41.00	70.00	119.0		
			Irbesartan	5	5 (100.0)	66.60 (34.46)	35.0	39.00	66.00	72.00	121.0		
		Week 6	Sparsentan	9	9 (100.0)	50.00 (20.99)	26.0	36.00	38.00	62.00	86.0		
			Irbesartan	5	5 (100.0)	64.80 (32.64)	34.0	37.00	61.00	79.00	113.0		
		Week 36	Sparsentan	9	6 (66.7)	56.50 (22.13)	34.0	36.00	51.00	82.00	85.0		
			Irbesartan	5	2 (40.0)	45.00 (19.80)	31.0	31.00	45.00	59.00	59.0		
		Week 58	Sparsentan	9	4 (44.4)	58.50 (29.85)	33.0	34.50	52.50	82.50	96.0		
			Irbesartan	5	1 (20.0)	26.00	26.0	26.00	26.00	26.00	26.0		
		Change from baseline in eGFR	Week 6	Sparsentan	9	9 (100.0)	-7.33 (10.75)	-33.0	-8.00	-4.00	-1.00	1.0	-0.59 [-1.71, 0.52]
				Irbesartan	5	5 (100.0)	-1.80 (5.63)	-8.0	-5.00	-2.00	-1.00	7.0	
			Week 36	Sparsentan	9	6 (66.7)	-11.67 (12.82)	-34.0	-19.00	-7.00	-2.00	-1.0	-0.36 [-1.97, 1.25]
				Irbesartan	5	2 (40.0)	-7.50 (0.71)	-8.0	-8.00	-7.50	-7.00	-7.0	
			Week 58	Sparsentan	9	4 (44.4)	-11.00 (9.83)	-23.0	-19.00	-9.50	-3.00	-2.0	NE
				Irbesartan	5	1 (20.0)	-13.00	-13.0	-13.00	-13.00	-13.00	-13.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 18 to 40 years	eGFR	Baseline	Sparsentan	102	102 (100.0)	63.27 (26.69)	25.0	42.00	56.50	85.00	128.0		
			Irbesartan	109	109 (100.0)	60.08 (25.53)	26.0	40.00	52.00	71.00	123.0		
		Week 6	Sparsentan	102	98 (96.1)	61.86 (26.08)	23.0	40.00	57.00	82.00	121.0		
			Irbesartan	109	105 (96.3)	58.53 (24.53)	23.0	40.00	54.00	72.00	120.0		
		Week 36	Sparsentan	102	68 (66.7)	60.79 (26.00)	18.0	40.50	54.50	82.50	123.0		
			Irbesartan	109	74 (67.9)	51.97 (24.73)	7.0	34.00	47.50	64.00	125.0		
		Week 58	Sparsentan	102	56 (54.9)	61.07 (26.96)	13.0	41.50	53.50	84.50	133.0		
			Irbesartan	109	51 (46.8)	51.37 (25.06)	13.0	34.00	42.00	66.00	122.0		
		Change from baseline in eGFR	Week 6	Sparsentan	102	98 (96.1)	-1.44 (8.29)	-43.0	-4.00	-1.00	2.00	15.0	0.07 [-0.21, 0.34]
				Irbesartan	109	105 (96.3)	-1.92 (5.95)	-17.0	-5.00	-1.00	2.00	14.0	
			Week 36	Sparsentan	102	68 (66.7)	-4.68 (9.02)	-39.0	-10.00	-3.00	1.50	10.0	0.04 [-0.29, 0.37]
				Irbesartan	109	74 (67.9)	-5.03 (8.18)	-32.0	-10.00	-4.50	0.00	15.0	
			Week 58	Sparsentan	102	56 (54.9)	-5.48 (10.46)	-34.0	-9.50	-5.00	1.00	24.0	0.08 [-0.30, 0.46]
	Irbesartan			109	51 (46.8)	-6.22 (7.43)	-24.0	-10.00	-7.00	0.00	10.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	eGFR	Baseline	Sparsentan	91	91 (100.0)	49.62 (18.73)	24.0	35.00	45.00	64.00	101.0	
			Irbesartan	88	88 (100.0)	52.82 (19.64)	27.0	38.00	48.00	65.50	106.0	
		Week 6	Sparsentan	91	86 (94.5)	49.77 (19.05)	21.0	34.00	46.00	62.00	99.0	
			Irbesartan	88	85 (96.6)	51.39 (20.82)	19.0	36.00	46.00	69.00	108.0	
		Week 36	Sparsentan	91	62 (68.1)	46.84 (17.21)	22.0	33.00	45.00	62.00	97.0	
			Irbesartan	88	57 (64.8)	49.00 (20.47)	15.0	33.00	42.00	61.00	109.0	
	Week 58	Sparsentan	91	50 (54.9)	45.12 (17.47)	21.0	30.00	43.00	57.00	94.0		
		Irbesartan	88	43 (48.9)	45.77 (17.83)	18.0	33.00	41.00	56.00	88.0		
	Change from baseline in eGFR	Week 6	Sparsentan	91	86 (94.5)	-0.13 (6.25)	-21.0	-3.00	0.00	3.00	20.0	0.13 [-0.17, 0.43]
			Irbesartan	88	85 (96.6)	-1.02 (7.31)	-31.0	-5.00	-2.00	3.00	20.0	
		Week 36	Sparsentan	91	62 (68.1)	-2.16 (7.39)	-22.0	-7.00	-2.50	1.00	26.0	0.21 [-0.15, 0.57]
			Irbesartan	88	57 (64.8)	-3.68 (7.15)	-25.0	-6.00	-4.00	0.00	13.0	
		Week 58	Sparsentan	91	50 (54.9)	-3.74 (6.68)	-19.0	-8.00	-3.50	0.00	14.0	0.32 [-0.09, 0.73]
			Irbesartan	88	43 (48.9)	-6.19 (8.49)	-24.0	-13.00	-7.00	0.00	12.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Geographic region													
North America	eGFR	Baseline	Sparsentan	35	35 (100.0)	56.54 (25.93)	25.0	38.00	50.00	67.00	121.0		
			Irbesartan	46	46 (100.0)	55.93 (23.44)	26.0	36.00	50.50	71.00	123.0		
	Week 6	Sparsentan	35	34 (97.1)	54.41 (25.54)	21.0	33.00	51.00	67.00	120.0			
		Irbesartan	46	44 (95.7)	53.43 (22.83)	21.0	35.00	46.00	71.00	107.0			
	Week 36	Sparsentan	35	19 (54.3)	57.68 (24.72)	27.0	36.00	51.00	78.00	104.0			
		Irbesartan	46	26 (56.5)	46.23 (21.24)	23.0	32.00	39.00	60.00	123.0			
	Week 58	Sparsentan	35	14 (40.0)	60.93 (28.04)	23.0	41.00	55.50	85.00	119.0			
		Irbesartan	46	18 (39.1)	44.83 (16.47)	21.0	34.00	39.00	60.00	72.0			
		Change from baseline in eGFR	Week 6	Sparsentan	35	34 (97.1)	-2.47 (8.28)	-33.0	-4.00	-1.00	1.00	9.0	-0.08 [-0.53, 0.37]
	Irbesartan			46	44 (95.7)	-1.84 (7.80)	-17.0	-6.50	-3.00	3.50	20.0		
	Week 36		Sparsentan	35	19 (54.3)	-4.68 (9.85)	-34.0	-10.00	-2.00	3.00	8.0	-0.10 [-0.69, 0.49]	
			Irbesartan	46	26 (56.5)	-3.77 (8.81)	-19.0	-8.00	-3.00	1.00	14.0		
	Week 58		Sparsentan	35	14 (40.0)	-6.86 (13.34)	-33.0	-14.00	-5.50	-1.00	24.0	-0.16 [-0.86, 0.54]	
			Irbesartan	46	18 (39.1)	-5.06 (8.94)	-20.0	-10.00	-7.00	1.00	12.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Europe	eGFR	Baseline	Sparsentan	98	98 (100.0)	56.00 (24.15)	24.0	36.00	48.00	71.00	128.0		
			Irbesartan	115	115 (100.0)	56.89 (24.29)	27.0	38.00	50.00	69.00	123.0		
		Week 6	Sparsentan	98	92 (93.9)	55.46 (24.07)	25.0	34.50	50.00	70.00	117.0		
			Irbesartan	115	111 (96.5)	55.81 (24.15)	19.0	37.00	48.00	69.00	120.0		
		Week 36	Sparsentan	98	57 (58.2)	54.18 (24.65)	18.0	37.00	47.00	65.00	123.0		
			Irbesartan	115	77 (67.0)	51.42 (23.72)	7.0	34.00	46.00	63.00	125.0		
		Week 58	Sparsentan	98	47 (48.0)	51.98 (23.92)	13.0	36.00	45.00	66.00	133.0		
			Irbesartan	115	54 (47.0)	49.57 (23.91)	18.0	33.00	43.00	59.00	122.0		
		Change from baseline in eGFR	Week 6	Sparsentan	98	92 (93.9)	-0.49 (6.30)	-17.0	-4.00	0.00	2.50	15.0	0.15 [-0.12, 0.43]
				Irbesartan	115	111 (96.5)	-1.47 (6.36)	-31.0	-5.00	-1.00	2.00	14.0	
				Sparsentan	98	57 (58.2)	-2.54 (7.68)	-26.0	-5.00	-2.00	2.00	17.0	0.17 [-0.18, 0.51]
				Irbesartan	115	77 (67.0)	-3.78 (7.22)	-32.0	-6.00	-4.00	0.00	15.0	
	Week 36	Sparsentan	98	57 (58.2)	-2.54 (7.68)	-26.0	-5.00	-2.00	2.00	17.0	0.17 [-0.18, 0.51]		
		Irbesartan	115	77 (67.0)	-3.78 (7.22)	-32.0	-6.00	-4.00	0.00	15.0			
Week 58	Sparsentan	98	47 (48.0)	-2.96 (7.68)	-19.0	-8.00	-1.00	2.00	14.0	0.35 [-0.04, 0.74]			
	Irbesartan	115	54 (47.0)	-5.54 (7.08)	-24.0	-10.00	-6.50	-1.00	10.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Asia Pacific	eGFR	Baseline	Sparsentan	69	69 (100.0)	58.23 (24.20)	26.0	40.00	51.00	74.00	121.0			
			Irbesartan	41	41 (100.0)	58.90 (22.11)	31.0	41.00	52.00	69.00	112.0			
		Week 6	Sparsentan	69	67 (97.1)	57.31 (22.32)	26.0	38.00	54.00	70.00	121.0			
			Irbesartan	41	40 (97.6)	57.30 (22.20)	23.0	41.50	53.50	70.00	119.0			
		Week 36	Sparsentan	69	60 (87.0)	53.22 (21.33)	19.0	34.00	50.00	66.00	108.0			
			Irbesartan	41	30 (73.2)	52.27 (22.10)	15.0	38.00	44.50	70.00	101.0			
		Week 58	Sparsentan	69	49 (71.0)	53.35 (23.71)	21.0	33.00	48.00	64.00	121.0			
			Irbesartan	41	23 (56.1)	49.13 (22.13)	13.0	33.00	42.00	65.00	98.0			
		Change from baseline in eGFR	Change from baseline in eGFR	Week 6	Sparsentan	69	67 (97.1)	-1.33 (9.03)	-43.0	-4.00	-1.00	3.00	20.0	0.00 [-0.39, 0.39]
					Irbesartan	41	40 (97.6)	-1.35 (5.71)	-19.0	-4.00	-1.50	2.00	9.0	
				Week 36	Sparsentan	69	60 (87.0)	-4.80 (9.19)	-39.0	-8.50	-4.00	-1.00	26.0	0.24 [-0.19, 0.68]
					Irbesartan	41	30 (73.2)	-6.93 (7.64)	-24.0	-12.00	-5.50	-2.00	5.0	
				Week 58	Sparsentan	69	49 (71.0)	-6.18 (8.42)	-34.0	-9.00	-6.00	-1.00	13.0	0.33 [-0.17, 0.83]
					Irbesartan	41	23 (56.1)	-8.96 (8.53)	-24.0	-15.00	-9.00	-2.00	8.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline BMI														
< 27 kg/m**2	eGFR	Baseline	Sparsentan	84	84 (100.0)	58.71 (24.55)	24.0	39.00	52.50	72.00	126.0			
			Irbesartan	94	94 (100.0)	58.86 (26.11)	26.0	39.00	50.00	70.00	123.0			
		Week 6	Sparsentan	84	78 (92.9)	58.27 (23.19)	23.0	37.00	55.00	74.00	117.0			
			Irbesartan	94	91 (96.8)	56.64 (25.44)	19.0	37.00	50.00	71.00	120.0			
		Week 36	Sparsentan	84	56 (66.7)	55.96 (24.60)	19.0	34.00	50.50	72.50	123.0			
			Irbesartan	94	60 (63.8)	51.20 (27.21)	7.0	33.00	41.50	60.50	125.0			
		Week 58	Sparsentan	84	46 (54.8)	53.35 (21.52)	21.0	36.00	47.00	71.00	96.0			
			Irbesartan	94	38 (40.4)	50.89 (26.65)	13.0	33.00	41.50	60.00	122.0			
			Change from baseline in eGFR	Week 6	Sparsentan	84	78 (92.9)	-1.46 (9.46)	-43.0	-4.00	0.00	3.00	20.0	0.10 [-0.20, 0.41]
					Irbesartan	94	91 (96.8)	-2.24 (5.58)	-16.0	-5.00	-2.00	2.00	12.0	
				Week 36	Sparsentan	84	56 (66.7)	-4.18 (10.24)	-39.0	-8.00	-3.00	0.00	26.0	0.21 [-0.15, 0.58]
					Irbesartan	94	60 (63.8)	-6.10 (7.91)	-32.0	-11.00	-5.00	-2.00	15.0	
				Week 58	Sparsentan	84	46 (54.8)	-5.41 (10.04)	-34.0	-10.00	-5.50	0.00	24.0	0.28 [-0.15, 0.71]
					Irbesartan	94	38 (40.4)	-7.79 (6.39)	-24.0	-12.00	-7.00	-5.00	7.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
>= 27 kg/m**2	eGFR	Baseline	Sparsentan	118	118 (100.0)	55.53 (24.28)	25.0	38.00	48.00	69.00	128.0			
			Irbesartan	107	107 (100.0)	55.66 (21.19)	27.0	38.00	50.00	70.00	114.0			
		Week 6	Sparsentan	118	115 (97.5)	54.32 (23.92)	21.0	35.00	50.00	67.00	121.0			
			Irbesartan	107	103 (96.3)	54.81 (21.56)	21.0	39.00	48.00	69.00	119.0			
		Week 36	Sparsentan	118	80 (67.8)	53.04 (22.10)	18.0	36.00	47.00	64.00	114.0			
			Irbesartan	107	73 (68.2)	50.10 (18.68)	20.0	34.00	45.00	64.00	101.0			
		Week 58	Sparsentan	118	64 (54.2)	54.00 (26.29)	13.0	35.00	47.00	60.50	133.0			
			Irbesartan	107	57 (53.3)	47.02 (18.61)	21.0	32.00	41.00	59.00	96.0			
		Change from baseline in eGFR	Change from baseline in eGFR	Week 6	Sparsentan	118	115 (97.5)	-0.90 (6.25)	-20.0	-4.00	-1.00	2.00	15.0	-0.00 [-0.27, 0.26]
					Irbesartan	107	103 (96.3)	-0.89 (7.31)	-31.0	-4.00	0.00	3.00	20.0	
				Week 36	Sparsentan	118	80 (67.8)	-3.60 (7.47)	-26.0	-8.00	-2.50	1.50	12.0	-0.06 [-0.38, 0.26]
					Irbesartan	107	73 (68.2)	-3.16 (7.31)	-25.0	-6.00	-3.00	1.00	13.0	
				Week 58	Sparsentan	118	64 (54.2)	-4.52 (8.15)	-33.0	-8.50	-4.00	1.00	13.0	0.09 [-0.27, 0.45]
					Irbesartan	107	57 (53.3)	-5.26 (8.64)	-24.0	-12.00	-4.00	1.00	12.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Randomization strata														
eGFR Low and UP High	eGFR	Baseline	Sparsentan	71	71 (100.0)	40.34 (9.83)	25.0	31.00	39.00	48.00	67.0			
			Irbesartan	74	74 (100.0)	41.78 (9.97)	26.0	34.00	39.00	50.00	66.0			
		Week 6	Sparsentan	71	69 (97.2)	39.59 (10.79)	21.0	32.00	38.00	49.00	65.0			
			Irbesartan	74	70 (94.6)	39.64 (10.00)	21.0	32.00	38.50	46.00	64.0			
		Week 36	Sparsentan	71	49 (69.0)	38.76 (11.13)	18.0	32.00	39.00	46.00	65.0			
			Irbesartan	74	47 (63.5)	34.83 (10.69)	7.0	28.00	33.00	41.00	58.0			
		Week 58	Sparsentan	71	42 (59.2)	37.93 (10.71)	13.0	30.00	36.50	45.00	61.0			
			Irbesartan	74	34 (45.9)	33.53 (9.57)	13.0	28.00	33.50	38.00	55.0			
		Change from baseline in eGFR	Change from baseline in eGFR	Week 6	Sparsentan	71	69 (97.2)	-0.55 (5.71)	-27.0	-3.00	-1.00	1.00	15.0	0.20 [-0.14, 0.53]
					Irbesartan	74	70 (94.6)	-1.57 (4.60)	-12.0	-5.00	-2.00	2.00	9.0	
				Week 36	Sparsentan	71	49 (69.0)	-2.00 (5.48)	-13.0	-5.00	-2.00	1.00	10.0	0.59 [0.18, 1.00]
					Irbesartan	74	47 (63.5)	-5.87 (7.49)	-32.0	-8.00	-5.00	-2.00	5.0	
				Week 58	Sparsentan	71	42 (59.2)	-3.79 (5.70)	-18.0	-7.00	-3.50	0.00	7.0	0.47 [0.02, 0.93]
					Irbesartan	74	34 (45.9)	-6.68 (6.56)	-20.0	-10.00	-7.00	-2.00	7.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR Low and UP Low	eGFR	Baseline	Sparsentan	55	55 (100.0)	42.11 (9.25)	24.0	35.00	42.00	50.00	62.0			
			Irbesartan	55	55 (100.0)	44.27 (9.37)	27.0	38.00	43.00	50.00	71.0			
		Week 6	Sparsentan	55	50 (90.9)	42.20 (10.15)	26.0	34.00	39.50	50.00	70.0			
			Irbesartan	55	54 (98.2)	43.26 (9.91)	19.0	36.00	43.00	48.00	68.0			
		Week 36	Sparsentan	55	35 (63.6)	42.23 (11.93)	25.0	34.00	41.00	48.00	76.0			
			Irbesartan	55	40 (72.7)	41.28 (8.99)	28.0	34.50	40.00	47.50	66.0			
		Week 58	Sparsentan	55	28 (50.9)	40.89 (10.81)	24.0	32.50	41.50	47.50	73.0			
			Irbesartan	55	29 (52.7)	39.76 (9.36)	25.0	33.00	39.00	45.00	59.0			
		Change from baseline in eGFR		Week 6	Sparsentan	55	50 (90.9)	0.44 (6.74)	-17.0	-3.00	0.00	3.00	20.0	0.23 [-0.16, 0.61]
					Irbesartan	55	54 (98.2)	-1.09 (6.81)	-31.0	-4.00	0.00	3.00	16.0	
				Week 36	Sparsentan	55	35 (63.6)	-0.20 (7.33)	-11.0	-4.00	-1.00	3.00	26.0	0.38 [-0.08, 0.84]
					Irbesartan	55	40 (72.7)	-2.45 (4.29)	-11.0	-5.00	-3.00	0.50	13.0	
				Week 58	Sparsentan	55	28 (50.9)	-1.75 (5.31)	-11.0	-5.50	-1.50	2.00	14.0	0.28 [-0.24, 0.80]
	Irbesartan				55	29 (52.7)	-3.41 (6.51)	-19.0	-7.00	-3.00	1.00	10.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP High	eGFR	Baseline	Sparsentan	37	37 (100.0)	82.76 (20.47)	51.0	66.00	77.00	91.00	128.0			
		Week 6	Irbesartan	36	36 (100.0)	79.83 (14.94)	57.0	69.00	74.00	91.50	109.0			
			Sparsentan	37	35 (94.6)	79.23 (18.78)	56.0	62.00	74.00	90.00	121.0			
		Week 36	Irbesartan	36	34 (94.4)	77.12 (14.73)	56.0	63.00	76.00	85.00	108.0			
			Sparsentan	37	26 (70.3)	76.31 (19.95)	51.0	62.00	71.50	86.00	118.0			
		Week 58	Irbesartan	36	22 (61.1)	73.91 (19.24)	40.0	60.00	67.00	87.00	123.0			
			Sparsentan	37	20 (54.1)	81.40 (23.66)	54.0	60.00	75.50	93.50	133.0			
		Week 58	Irbesartan	36	15 (41.7)	69.67 (16.59)	37.0	55.00	72.00	78.00	98.0			
			Sparsentan	37	35 (94.6)	-3.89 (11.57)	-43.0	-8.00	-1.00	4.00	12.0	-0.08 [-0.56, 0.39]		
		Change from baseline in eGFR	eGFR	Week 6	Irbesartan	36	34 (94.4)	-3.09 (6.45)	-17.0	-6.00	-3.00	2.00	12.0	
				Week 36	Sparsentan	37	26 (70.3)	-9.46 (12.24)	-39.0	-17.00	-9.00	1.00	7.0	-0.26 [-0.83, 0.31]
					Irbesartan	36	22 (61.1)	-6.55 (10.24)	-25.0	-15.00	-6.50	0.00	14.0	
				Week 58	Sparsentan	37	20 (54.1)	-7.40 (14.33)	-34.0	-15.00	-10.50	3.00	24.0	0.23 [-0.44, 0.90]
	Irbesartan	36	15 (41.7)		-10.20 (8.54)	-23.0	-15.00	-12.00	-5.00	10.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP Low	eGFR	Baseline	Sparsentan	39	39 (100.0)	83.15 (16.11)	57.0	70.00	80.00	96.00	126.0			
			Irbesartan	37	37 (100.0)	84.57 (22.97)	50.0	66.00	79.00	104.00	123.0			
		Week 6	Sparsentan	39	39 (100.0)	81.46 (16.34)	53.0	67.00	78.00	98.00	113.0			
			Irbesartan	37	37 (100.0)	83.92 (20.66)	43.0	69.00	76.00	101.00	120.0			
		Week 36	Sparsentan	39	26 (66.7)	77.54 (17.72)	47.0	65.00	75.50	90.00	123.0			
			Irbesartan	37	24 (64.9)	75.63 (21.39)	41.0	61.50	68.50	87.50	125.0			
		Week 58	Sparsentan	39	20 (51.3)	77.20 (15.38)	58.0	62.50	74.50	88.50	106.0			
			Irbesartan	37	17 (45.9)	75.06 (22.55)	37.0	60.00	71.00	88.00	122.0			
		Change from baseline in eGFR		Week 6	Sparsentan	39	39 (100.0)	-1.69 (7.14)	-20.0	-7.00	-1.00	3.00	14.0	-0.13 [-0.58, 0.32]
					Irbesartan	37	37 (100.0)	-0.65 (9.02)	-19.0	-6.00	0.00	6.00	20.0	
				Week 36	Sparsentan	39	26 (66.7)	-6.58 (7.83)	-26.0	-12.00	-5.50	-2.00	12.0	-0.39 [-0.95, 0.17]
					Irbesartan	37	24 (64.9)	-3.29 (9.20)	-24.0	-10.00	-3.50	4.00	15.0	
				Week 58	Sparsentan	39	20 (51.3)	-9.10 (10.09)	-33.0	-16.00	-7.50	-4.00	13.0	-0.22 [-0.87, 0.43]
	Irbesartan				37	17 (45.9)	-6.88 (10.40)	-24.0	-14.00	-7.00	0.00	12.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 1														
< 60 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	127	127 (100.0)	41.08 (9.35)	24.0	33.00	41.00	49.00	59.0			
			Irbesartan	129	129 (100.0)	42.28 (8.61)	26.0	35.00	41.00	50.00	59.0			
		Week 6	Sparsentan	127	120 (94.5)	40.94 (10.91)	21.0	33.00	38.00	50.00	73.0			
			Irbesartan	129	126 (97.7)	41.75 (10.91)	19.0	34.00	41.00	47.00	76.0			
		Week 36	Sparsentan	127	86 (67.7)	40.59 (11.63)	18.0	33.00	39.50	48.00	76.0			
			Irbesartan	129	88 (68.2)	38.16 (10.97)	7.0	31.50	37.00	44.50	67.0			
		Week 58	Sparsentan	127	69 (54.3)	38.91 (10.70)	13.0	32.00	39.00	45.00	73.0			
			Irbesartan	129	66 (51.2)	37.79 (11.87)	13.0	31.00	36.00	44.00	71.0			
		Change from baseline in eGFR		Week 6	Sparsentan	127	120 (94.5)	0.15 (6.29)	-27.0	-3.00	0.00	2.00	20.0	0.12 [-0.13, 0.37]
					Irbesartan	129	126 (97.7)	-0.60 (6.14)	-31.0	-4.00	0.00	3.00	20.0	
				Week 36	Sparsentan	127	86 (67.7)	-1.05 (6.24)	-13.0	-4.00	-1.50	2.00	26.0	0.40 [0.10, 0.70]
					Irbesartan	129	88 (68.2)	-3.60 (6.61)	-32.0	-6.00	-3.00	0.00	13.0	
				Week 58	Sparsentan	127	69 (54.3)	-2.68 (5.58)	-18.0	-6.00	-2.00	1.00	14.0	0.25 [-0.09, 0.59]
					Irbesartan	129	66 (51.2)	-4.29 (7.16)	-20.0	-9.00	-5.50	1.00	12.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	49	49 (100.0)	72.71 (8.32)	60.0	66.00	71.00	79.00	89.0		
			Irbesartan	48	48 (100.0)	71.60 (7.60)	60.0	66.50	70.00	76.00	89.0		
		Week 6	Sparsentan	49	47 (95.9)	69.87 (9.06)	56.0	62.00	69.00	76.00	90.0		
			Irbesartan	48	44 (91.7)	69.11 (9.14)	54.0	61.50	69.50	75.00	85.0		
		Week 36	Sparsentan	49	30 (61.2)	66.13 (10.46)	47.0	57.00	65.00	72.00	86.0		
			Irbesartan	48	32 (66.7)	64.59 (10.63)	40.0	58.50	63.50	70.00	87.0		
		Week 58	Sparsentan	49	23 (46.9)	66.22 (10.67)	51.0	58.00	64.00	74.00	91.0		
			Irbesartan	48	19 (39.6)	61.42 (11.45)	37.0	54.00	60.00	73.00	78.0		
		Change from baseline in eGFR	Week 6	Sparsentan	49	47 (95.9)	-2.68 (7.30)	-21.0	-7.00	-2.00	3.00	12.0	0.01 [-0.40, 0.42]
				Irbesartan	48	44 (91.7)	-2.73 (7.52)	-19.0	-7.00	-3.00	3.00	14.0	
			Week 36	Sparsentan	49	30 (61.2)	-7.30 (8.27)	-26.0	-12.00	-8.50	-2.00	12.0	-0.03 [-0.53, 0.47]
				Irbesartan	48	32 (66.7)	-7.06 (8.22)	-25.0	-13.00	-6.50	-2.00	12.0	
			Week 58	Sparsentan	49	23 (46.9)	-7.70 (9.50)	-19.0	-15.00	-10.00	-4.00	24.0	0.48 [-0.14, 1.09]
	Irbesartan			48	19 (39.6)	-11.95 (8.07)	-24.0	-18.00	-14.00	-5.00	1.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	26	26 (100.0)	104.04 (11.51)	90.0	95.00	102.00	111.00	128.0		
			Irbesartan	25	25 (100.0)	105.56 (9.98)	91.0	98.00	105.00	112.00	123.0		
		Week 6	Sparsentan	26	26 (100.0)	99.81 (11.47)	77.0	92.00	99.00	104.00	121.0		
			Irbesartan	25	25 (100.0)	101.48 (10.76)	80.0	95.00	101.00	108.00	120.0		
		Week 36	Sparsentan	26	20 (76.9)	95.10 (14.20)	63.0	86.00	93.50	103.00	123.0		
			Irbesartan	25	13 (52.0)	100.31 (15.45)	80.0	89.00	98.00	109.00	125.0		
		Week 58	Sparsentan	26	18 (69.2)	94.56 (17.79)	61.0	85.00	90.00	105.00	133.0		
			Irbesartan	25	10 (40.0)	95.30 (14.34)	77.0	84.00	94.50	98.00	122.0		
		Change from baseline in eGFR	Week 6	Sparsentan	26	26 (100.0)	-4.23 (12.06)	-43.0	-5.00	-1.00	1.00	13.0	-0.02 [-0.56, 0.53]
				Irbesartan	25	25 (100.0)	-4.08 (6.00)	-16.0	-8.00	-3.00	0.00	7.0	
			Week 36	Sparsentan	26	20 (76.9)	-10.65 (12.44)	-39.0	-16.50	-10.50	-2.50	7.0	-0.53 [-1.24, 0.18]
				Irbesartan	25	13 (52.0)	-4.15 (11.72)	-24.0	-12.00	-4.00	3.00	15.0	
			Week 58	Sparsentan	26	18 (69.2)	-9.78 (14.72)	-34.0	-20.00	-9.50	2.00	13.0	-0.09 [-0.87, 0.68]
	Irbesartan			25	10 (40.0)	-8.60 (6.74)	-21.0	-12.00	-9.50	-1.00	0.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 2														
< 45 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	82	82 (100.0)	35.37 (5.60)	24.0	31.00	35.00	41.00	44.0			
			Irbesartan	80	80 (100.0)	36.58 (4.61)	26.0	33.00	37.00	40.00	44.0			
		Week 6	Sparsentan	82	79 (96.3)	36.10 (8.12)	21.0	31.00	35.00	40.00	58.0			
			Irbesartan	80	78 (97.5)	35.71 (6.41)	19.0	31.00	36.50	40.00	50.0			
		Week 36	Sparsentan	82	54 (65.9)	34.41 (7.94)	18.0	30.00	34.00	38.00	54.0			
			Irbesartan	80	56 (70.0)	32.38 (7.27)	7.0	28.50	33.00	37.00	47.0			
		Week 58	Sparsentan	82	44 (53.7)	33.05 (7.28)	13.0	29.00	33.00	37.50	48.0			
			Irbesartan	80	42 (52.5)	31.98 (7.50)	13.0	28.00	33.00	36.00	52.0			
		Change from baseline in eGFR		Week 6	Sparsentan	82	79 (96.3)	0.87 (4.98)	-10.0	-2.00	0.00	3.00	15.0	0.37 [0.05, 0.69]
					Irbesartan	80	78 (97.5)	-0.88 (4.50)	-16.0	-4.00	-0.50	2.00	10.0	
				Week 36	Sparsentan	82	54 (65.9)	-1.09 (5.34)	-13.0	-4.00	-1.00	2.00	10.0	0.51 [0.13, 0.89]
					Irbesartan	80	56 (70.0)	-4.11 (6.37)	-32.0	-6.00	-3.00	0.00	5.0	
				Week 58	Sparsentan	82	44 (53.7)	-2.77 (5.12)	-18.0	-6.00	-2.50	1.00	6.0	0.41 [-0.01, 0.84]
					Irbesartan	80	42 (52.5)	-5.05 (5.87)	-18.0	-9.00	-6.00	-1.00	8.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
45 to < 60 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	45	45 (100.0)	51.49 (4.57)	45.0	48.00	51.00	55.00	59.0		
			Irbesartan	49	49 (100.0)	51.59 (4.45)	45.0	48.00	51.00	55.00	59.0		
		Week 6	Sparsentan	45	41 (91.1)	50.27 (9.48)	23.0	46.00	50.00	54.00	73.0		
			Irbesartan	49	48 (98.0)	51.56 (9.50)	21.0	46.00	51.00	57.50	76.0		
		Week 36	Sparsentan	45	32 (71.1)	51.03 (9.16)	39.0	45.00	49.00	56.00	76.0		
			Irbesartan	49	32 (65.3)	48.28 (8.83)	33.0	41.50	48.00	54.00	67.0		
		Week 58	Sparsentan	45	25 (55.6)	49.24 (7.46)	42.0	44.00	47.00	52.00	73.0		
			Irbesartan	49	24 (49.0)	47.96 (11.31)	31.0	40.00	45.00	54.50	71.0		
		Change from baseline in eGFR	Week 6	Sparsentan	45	41 (91.1)	-1.24 (8.15)	-27.0	-4.00	-1.00	1.00	20.0	-0.13 [-0.55, 0.28]
				Irbesartan	49	48 (98.0)	-0.15 (8.17)	-31.0	-4.00	0.50	4.00	20.0	
			Week 36	Sparsentan	45	32 (71.1)	-0.97 (7.63)	-12.0	-5.00	-2.00	2.00	26.0	0.24 [-0.25, 0.73]
				Irbesartan	49	32 (65.3)	-2.72 (7.03)	-19.0	-6.00	-2.50	1.50	13.0	
			Week 58	Sparsentan	45	25 (55.6)	-2.52 (6.43)	-15.0	-7.00	-2.00	1.00	14.0	0.06 [-0.50, 0.62]
		Irbesartan		49	24 (49.0)	-2.96 (8.98)	-20.0	-9.00	-2.00	3.50	12.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	49	49 (100.0)	72.71 (8.32)	60.0	66.00	71.00	79.00	89.0		
			Irbesartan	48	48 (100.0)	71.60 (7.60)	60.0	66.50	70.00	76.00	89.0		
		Week 6	Sparsentan	49	47 (95.9)	69.87 (9.06)	56.0	62.00	69.00	76.00	90.0		
			Irbesartan	48	44 (91.7)	69.11 (9.14)	54.0	61.50	69.50	75.00	85.0		
		Week 36	Sparsentan	49	30 (61.2)	66.13 (10.46)	47.0	57.00	65.00	72.00	86.0		
			Irbesartan	48	32 (66.7)	64.59 (10.63)	40.0	58.50	63.50	70.00	87.0		
		Week 58	Sparsentan	49	23 (46.9)	66.22 (10.67)	51.0	58.00	64.00	74.00	91.0		
			Irbesartan	48	19 (39.6)	61.42 (11.45)	37.0	54.00	60.00	73.00	78.0		
		Change from baseline in eGFR	Week 6	Sparsentan	49	47 (95.9)	-2.68 (7.30)	-21.0	-7.00	-2.00	3.00	12.0	0.01 [-0.40, 0.42]
				Irbesartan	48	44 (91.7)	-2.73 (7.52)	-19.0	-7.00	-3.00	3.00	14.0	
			Week 36	Sparsentan	49	30 (61.2)	-7.30 (8.27)	-26.0	-12.00	-8.50	-2.00	12.0	-0.03 [-0.53, 0.47]
				Irbesartan	48	32 (66.7)	-7.06 (8.22)	-25.0	-13.00	-6.50	-2.00	12.0	
			Week 58	Sparsentan	49	23 (46.9)	-7.70 (9.50)	-19.0	-15.00	-10.00	-4.00	24.0	0.48 [-0.14, 1.09]
		Irbesartan		48	19 (39.6)	-11.95 (8.07)	-24.0	-18.00	-14.00	-5.00	1.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
>= 90 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	26	26 (100.0)	104.04 (11.51)	90.0	95.00	102.00	111.00	128.0			
			Irbesartan	25	25 (100.0)	105.56 (9.98)	91.0	98.00	105.00	112.00	123.0			
		Week 6	Sparsentan	26	26 (100.0)	99.81 (11.47)	77.0	92.00	99.00	104.00	121.0			
			Irbesartan	25	25 (100.0)	101.48 (10.76)	80.0	95.00	101.00	108.00	120.0			
		Week 36	Sparsentan	26	20 (76.9)	95.10 (14.20)	63.0	86.00	93.50	103.00	123.0			
			Irbesartan	25	13 (52.0)	100.31 (15.45)	80.0	89.00	98.00	109.00	125.0			
		Week 58	Sparsentan	26	18 (69.2)	94.56 (17.79)	61.0	85.00	90.00	105.00	133.0			
			Irbesartan	25	10 (40.0)	95.30 (14.34)	77.0	84.00	94.50	98.00	122.0			
		Change from baseline in eGFR	Week 6	Sparsentan	26	26 (100.0)	-4.23 (12.06)	-43.0	-5.00	-1.00	1.00	13.0	-0.02 [-0.56, 0.53]	
				Irbesartan	25	25 (100.0)	-4.08 (6.00)	-16.0	-8.00	-3.00	0.00	7.0		
				Week 36	Sparsentan	26	20 (76.9)	-10.65 (12.44)	-39.0	-16.50	-10.50	-2.50	7.0	-0.53 [-1.24, 0.18]
					Irbesartan	25	13 (52.0)	-4.15 (11.72)	-24.0	-12.00	-4.00	3.00	15.0	
				Week 58	Sparsentan	26	18 (69.2)	-9.78 (14.72)	-34.0	-20.00	-9.50	2.00	13.0	-0.09 [-0.87, 0.68]
					Irbesartan	25	10 (40.0)	-8.60 (6.74)	-21.0	-12.00	-9.50	-1.00	0.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline urine protein excretion														
<= 1.75 g/day	eGFR	Baseline	Sparsentan	98	98 (100.0)	59.05 (24.14)	24.0	40.00	53.50	77.00	126.0			
			Irbesartan	94	94 (100.0)	59.67 (25.13)	27.0	40.00	52.00	71.00	123.0			
		Week 6	Sparsentan	98	93 (94.9)	59.52 (23.68)	21.0	38.00	54.00	73.00	113.0			
			Irbesartan	94	91 (96.8)	59.05 (24.72)	19.0	41.00	54.00	72.00	120.0			
		Week 36	Sparsentan	98	70 (71.4)	57.37 (22.51)	25.0	38.00	51.00	73.00	123.0			
			Irbesartan	94	61 (64.9)	53.26 (22.50)	20.0	37.00	46.00	65.00	125.0			
		Week 58	Sparsentan	98	58 (59.2)	55.84 (22.10)	24.0	38.00	50.00	73.00	106.0			
			Irbesartan	94	36 (38.3)	53.44 (22.33)	25.0	36.50	52.50	65.00	122.0			
			Change from baseline in eGFR	Week 6	Sparsentan	98	93 (94.9)	-0.26 (6.82)	-17.0	-4.00	0.00	3.00	20.0	0.06 [-0.23, 0.35]
					Irbesartan	94	91 (96.8)	-0.69 (7.88)	-31.0	-5.00	0.00	4.00	20.0	
				Week 36	Sparsentan	98	70 (71.4)	-1.76 (7.35)	-18.0	-6.00	-2.00	2.00	26.0	0.08 [-0.26, 0.42]
					Irbesartan	94	61 (64.9)	-2.33 (7.10)	-24.0	-6.00	-3.00	2.00	15.0	
				Week 58	Sparsentan	98	58 (59.2)	-3.83 (8.91)	-33.0	-7.00	-3.00	2.00	24.0	0.02 [-0.40, 0.43]
					Irbesartan	94	36 (38.3)	-3.97 (8.37)	-24.0	-10.00	-2.00	1.50	12.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 1.75 g/day	eGFR	Baseline	Sparsentan	104	104 (100.0)	54.79 (24.54)	25.0	37.00	48.50	66.00	128.0			
			Irbesartan	108	108 (100.0)	54.82 (22.00)	26.0	37.50	48.00	68.00	116.0			
		Week 6	Sparsentan	104	100 (96.2)	52.57 (23.24)	23.0	35.00	47.50	63.50	121.0			
			Irbesartan	108	104 (96.3)	52.54 (21.82)	21.0	36.50	46.00	65.50	116.0			
		Week 36	Sparsentan	104	66 (63.5)	50.92 (23.46)	18.0	34.00	46.50	63.00	118.0			
			Irbesartan	108	72 (66.7)	48.33 (23.04)	7.0	32.00	40.00	60.00	123.0			
		Week 58	Sparsentan	104	52 (50.0)	51.37 (26.57)	13.0	33.00	44.50	59.50	133.0			
			Irbesartan	108	59 (54.6)	45.59 (21.65)	13.0	31.00	38.00	55.00	116.0			
		Change from baseline in eGFR	Change from baseline in eGFR	Week 6	Sparsentan	104	100 (96.2)	-1.94 (8.37)	-43.0	-4.00	-1.00	1.50	15.0	0.05 [-0.23, 0.32]
					Irbesartan	108	104 (96.3)	-2.26 (5.06)	-17.0	-5.00	-2.00	1.00	12.0	
				Week 36	Sparsentan	104	66 (63.5)	-6.05 (9.48)	-39.0	-10.00	-4.00	-1.00	10.0	0.03 [-0.30, 0.37]
					Irbesartan	108	72 (66.7)	-6.32 (7.76)	-32.0	-10.00	-5.00	-2.00	14.0	
				Week 58	Sparsentan	104	52 (50.0)	-6.08 (8.94)	-34.0	-10.50	-6.00	0.00	13.0	0.20 [-0.18, 0.57]
					Irbesartan	108	59 (54.6)	-7.68 (7.29)	-24.0	-12.00	-8.00	-2.00	10.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline use of antihypertensives												
Yes	eGFR	Baseline	Sparsentan	88	88 (100.0)	51.08 (21.98)	25.0	35.00	44.00	65.50	128.0	
			Irbesartan	83	83 (100.0)	50.61 (18.68)	26.0	36.00	45.00	62.00	108.0	
	Week 6	Sparsentan	88	83 (94.3)	50.78 (21.91)	25.0	33.00	44.00	60.00	121.0		
		Irbesartan	83	77 (92.8)	48.83 (18.85)	19.0	34.00	44.00	58.00	108.0		
	Week 36	Sparsentan	88	55 (62.5)	48.29 (21.94)	18.0	33.00	44.00	56.00	114.0		
		Irbesartan	83	59 (71.1)	45.69 (20.79)	7.0	32.00	38.00	54.00	109.0		
	Week 58	Sparsentan	88	44 (50.0)	48.57 (24.99)	13.0	32.50	43.00	55.00	133.0		
		Irbesartan	83	44 (53.0)	44.68 (17.82)	21.0	31.50	37.50	54.00	97.0		
	Change from baseline in eGFR	Week 6	Sparsentan	88	83 (94.3)	-0.49 (6.38)	-21.0	-3.00	-1.00	3.00	13.0	0.17 [-0.14, 0.48]
			Irbesartan	83	77 (92.8)	-1.62 (7.07)	-31.0	-4.00	0.00	2.00	13.0	
		Week 36	Sparsentan	88	55 (62.5)	-3.62 (6.28)	-22.0	-8.00	-3.00	1.00	10.0	0.03 [-0.34, 0.39]
			Irbesartan	83	59 (71.1)	-3.80 (7.01)	-32.0	-6.00	-4.00	0.00	12.0	
		Week 58	Sparsentan	88	44 (50.0)	-4.34 (5.88)	-18.0	-7.50	-4.50	0.00	6.0	0.20 [-0.22, 0.62]
			Irbesartan	83	44 (53.0)	-5.75 (8.04)	-24.0	-10.50	-6.00	1.00	12.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
No	eGFR	Baseline	Sparsentan	114	114 (100.0)	61.32 (25.28)	24.0	41.00	56.00	80.00	126.0			
			Irbesartan	119	119 (100.0)	61.59 (25.58)	28.0	41.00	55.00	76.00	123.0			
		Week 6	Sparsentan	114	110 (96.5)	59.79 (24.26)	21.0	38.00	56.50	74.00	120.0			
			Irbesartan	119	118 (99.2)	59.98 (25.03)	23.0	40.00	54.00	76.00	120.0			
		Week 36	Sparsentan	114	81 (71.1)	58.28 (23.16)	19.0	40.00	51.00	76.00	123.0			
			Irbesartan	119	74 (62.2)	54.50 (23.77)	15.0	38.00	49.50	64.00	125.0			
		Week 58	Sparsentan	114	66 (57.9)	57.17 (23.41)	21.0	41.00	52.50	73.00	119.0			
			Irbesartan	119	51 (42.9)	51.92 (24.95)	13.0	34.00	45.00	66.00	122.0			
		Change from baseline in eGFR	Change from baseline in eGFR	Week 6	Sparsentan	114	110 (96.5)	-1.61 (8.55)	-43.0	-4.00	-1.00	2.00	20.0	-0.02 [-0.28, 0.24]
					Irbesartan	119	118 (99.2)	-1.47 (6.23)	-17.0	-5.00	-2.00	3.00	20.0	
				Week 36	Sparsentan	114	81 (71.1)	-3.99 (10.03)	-39.0	-8.00	-3.00	2.00	26.0	0.11 [-0.20, 0.43]
					Irbesartan	119	74 (62.2)	-5.04 (8.22)	-25.0	-10.00	-4.00	0.00	15.0	
				Week 58	Sparsentan	114	66 (57.9)	-5.26 (10.55)	-34.0	-11.00	-4.00	1.00	24.0	0.16 [-0.21, 0.52]
					Irbesartan	119	51 (42.9)	-6.73 (7.79)	-24.0	-12.00	-7.00	-2.00	10.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Time since renal biopsy														
<= 5 years	eGFR	Baseline	Sparsentan	113	113 (100.0)	59.19 (25.65)	24.0	39.00	52.00	74.00	126.0			
			Irbesartan	127	127 (100.0)	61.37 (24.93)	28.0	41.00	56.00	73.00	123.0			
		Week 6	Sparsentan	113	106 (93.8)	58.44 (25.73)	21.0	37.00	53.50	76.00	121.0			
			Irbesartan	127	122 (96.1)	59.78 (25.18)	21.0	40.00	55.00	74.00	120.0			
		Week 36	Sparsentan	113	72 (63.7)	58.81 (25.96)	18.0	40.00	52.50	77.00	123.0			
			Irbesartan	127	80 (63.0)	54.80 (25.15)	7.0	37.50	48.00	66.00	125.0			
		Week 58	Sparsentan	113	56 (49.6)	57.84 (25.76)	13.0	41.50	54.00	72.50	121.0			
			Irbesartan	127	56 (44.1)	52.95 (23.72)	13.0	36.00	45.50	69.00	122.0			
			Change from baseline in eGFR	Week 6	Sparsentan	113	106 (93.8)	-1.31 (8.41)	-43.0	-4.00	-1.00	2.00	20.0	0.02 [-0.24, 0.28]
		Irbesartan			127	122 (96.1)	-1.47 (7.17)	-31.0	-5.00	-1.50	3.00	20.0		
		Week 36		Sparsentan	113	72 (63.7)	-4.17 (9.37)	-39.0	-9.50	-3.00	0.50	26.0	0.05 [-0.26, 0.37]	
				Irbesartan	127	80 (63.0)	-4.66 (8.95)	-32.0	-9.50	-4.00	1.00	15.0		
		Week 58		Sparsentan	113	56 (49.6)	-6.32 (9.16)	-34.0	-11.50	-6.00	-1.00	14.0	-0.03 [-0.40, 0.34]	
				Irbesartan	127	56 (44.1)	-6.02 (9.12)	-24.0	-12.50	-7.00	0.50	12.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 5 years	eGFR	Baseline	Sparsentan	89	89 (100.0)	53.90 (22.45)	28.0	36.00	44.00	66.00	128.0			
			Irbesartan	75	75 (100.0)	49.81 (19.15)	26.0	36.00	43.00	64.00	108.0			
		Week 6	Sparsentan	89	87 (97.8)	52.84 (20.55)	26.0	36.00	46.00	65.00	115.0			
			Irbesartan	75	73 (97.3)	48.56 (18.12)	19.0	35.00	43.00	59.00	95.0			
		Week 36	Sparsentan	89	64 (71.9)	49.11 (18.31)	25.0	34.50	45.00	59.50	114.0			
			Irbesartan	75	53 (70.7)	44.25 (17.18)	23.0	32.00	38.00	55.00	90.0			
		Week 58	Sparsentan	89	54 (60.7)	49.46 (22.14)	23.0	33.00	42.50	57.00	133.0			
			Irbesartan	75	39 (52.0)	42.28 (18.11)	18.0	29.00	36.00	53.00	97.0			
		Change from baseline in eGFR	Week 6	Sparsentan	89	87 (97.8)	-0.91 (6.76)	-33.0	-4.00	0.00	2.00	14.0	0.12 [-0.19, 0.43]	
				Irbesartan	75	73 (97.3)	-1.63 (5.41)	-16.0	-5.00	-2.00	2.00	13.0		
				Week 36	Sparsentan	89	64 (71.9)	-3.47 (7.91)	-34.0	-7.50	-3.00	1.50	10.0	0.11 [-0.25, 0.47]
					Irbesartan	75	53 (70.7)	-4.23 (5.37)	-18.0	-6.00	-4.00	-1.00	10.0	
	Week 58	Sparsentan	89	54 (60.7)	-3.41 (8.56)	-29.0	-8.00	-2.50	2.00	24.0	0.43 [0.01, 0.85]			
		Irbesartan	75	39 (52.0)	-6.64 (5.74)	-18.0	-10.00	-7.00	-2.00	6.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: History of hypertension														
Yes	eGFR	Baseline	Sparsentan	153	153 (100.0)	53.08 (22.63)	24.0	36.00	45.00	66.00	128.0			
			Irbesartan	157	157 (100.0)	54.81 (21.78)	26.0	38.00	50.00	68.00	123.0			
		Week 6	Sparsentan	153	144 (94.1)	52.47 (22.59)	21.0	35.00	47.50	66.00	121.0			
			Irbesartan	157	151 (96.2)	53.49 (22.08)	19.0	37.00	47.00	68.00	120.0			
		Week 36	Sparsentan	153	101 (66.0)	50.85 (22.30)	18.0	34.00	46.00	63.00	118.0			
			Irbesartan	157	105 (66.9)	50.17 (23.79)	7.0	33.00	42.00	61.00	125.0			
		Week 58	Sparsentan	153	82 (53.6)	50.68 (23.80)	13.0	33.00	45.00	59.00	133.0			
			Irbesartan	157	76 (48.4)	48.14 (22.34)	18.0	33.00	39.50	57.50	122.0			
			Change from baseline in eGFR	Week 6	Sparsentan	153	144 (94.1)	-0.63 (6.96)	-33.0	-3.50	-0.50	2.50	20.0	0.10 [-0.13, 0.33]
					Irbesartan	157	151 (96.2)	-1.32 (6.86)	-31.0	-5.00	-1.00	3.00	20.0	
				Week 36	Sparsentan	153	101 (66.0)	-2.91 (8.05)	-34.0	-7.00	-2.00	1.00	26.0	0.11 [-0.17, 0.38]
					Irbesartan	157	105 (66.9)	-3.74 (7.61)	-32.0	-7.00	-4.00	1.00	15.0	
				Week 58	Sparsentan	153	82 (53.6)	-3.73 (7.64)	-23.0	-8.00	-4.00	1.00	24.0	0.27 [-0.04, 0.59]
					Irbesartan	157	76 (48.4)	-5.87 (8.07)	-24.0	-12.00	-6.00	0.00	12.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
No	eGFR	Baseline	Sparsentan	49	49 (100.0)	68.65 (26.08)	29.0	46.00	69.00	89.00	126.0		
			Irbesartan	45	45 (100.0)	65.00 (27.83)	31.0	41.00	52.00	86.00	123.0		
		Week 6	Sparsentan	49	49 (100.0)	66.06 (23.98)	26.0	48.00	62.00	83.00	120.0		
			Irbesartan	45	44 (97.8)	62.75 (26.43)	23.0	42.50	51.00	81.00	119.0		
		Week 36	Sparsentan	49	35 (71.4)	64.03 (22.95)	32.0	46.00	63.00	82.00	123.0		
			Irbesartan	45	28 (62.2)	52.18 (19.17)	17.0	37.00	49.50	62.50	95.0		
		Week 58	Sparsentan	49	28 (57.1)	62.64 (24.00)	29.0	42.00	61.00	85.00	119.0		
			Irbesartan	45	19 (42.2)	50.26 (21.76)	13.0	33.00	48.00	69.00	98.0		
		Change from baseline in eGFR	Week 6	Sparsentan	49	49 (100.0)	-2.59 (9.46)	-43.0	-5.00	-1.00	1.00	12.0	-0.05 [-0.45, 0.36]
				Irbesartan	45	44 (97.8)	-2.23 (5.37)	-16.0	-5.50	-2.00	1.50	7.0	
			Week 36	Sparsentan	49	35 (71.4)	-6.51 (9.96)	-39.0	-12.00	-5.00	0.00	10.0	0.09 [-0.41, 0.58]
				Irbesartan	45	28 (62.2)	-7.29 (7.50)	-25.0	-12.50	-4.50	-2.50	4.0	
	Week 58		Sparsentan	49	28 (57.1)	-8.29 (11.51)	-34.0	-16.00	-6.50	-0.50	13.0	-0.04 [-0.62, 0.54]	
			Irbesartan	45	19 (42.2)	-7.89 (7.05)	-24.0	-10.00	-8.00	-5.00	8.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.135
Male	Week 6	Sparsentan	139	133 (95.7)	-0.95 (0.61)	(-2.15, 0.24)	0.15 (0.85)	(-1.53, 1.82)	0.864
		Irbesartan	143	139 (97.2)	-1.10 (0.60)	(-2.27, 0.07)			
	Week 36	Sparsentan	139	92 (66.2)	-3.27 (0.71)	(-4.67, -1.88)	0.68 (0.99)	(-1.27, 2.63)	0.494
		Irbesartan	143	97 (67.8)	-3.95 (0.69)	(-5.31, -2.60)			
	Week 58	Sparsentan	139	64 (46.0)	-4.67 (0.84)	(-6.32, -3.02)	1.13 (1.17)	(-1.16, 3.42)	0.332
		Irbesartan	143	70 (49.0)	-5.80 (0.81)	(-7.39, -4.22)			
Female	Week 6	Sparsentan	63	60 (95.2)	-1.49 (0.97)	(-3.40, 0.41)	1.01 (1.40)	(-1.74, 3.77)	0.469
		Irbesartan	59	56 (94.9)	-2.51 (1.00)	(-4.48, -0.53)			
	Week 36	Sparsentan	63	44 (69.8)	-4.63 (1.09)	(-6.77, -2.48)	1.50 (1.63)	(-1.71, 4.71)	0.358
		Irbesartan	59	36 (61.0)	-6.13 (1.21)	(-8.50, -3.76)			
	Week 58	Sparsentan	63	46 (73.0)	-4.61 (1.11)	(-6.79, -2.43)	2.53 (1.82)	(-1.04, 6.10)	0.164
		Irbesartan	59	25 (42.4)	-7.14 (1.43)	(-9.95, -4.33)			

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LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
 Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.723
<= 45 years	Week 6	Sparsentan	96	93 (96.9)	-2.40 (0.85)	(-4.07, -0.73)	-0.05 (1.19)	(-2.40, 2.29)	0.963
		Irbesartan	99	97 (98.0)	-2.35 (0.84)	(-3.99, -0.70)			
	Week 36	Sparsentan	96	65 (67.7)	-5.58 (0.98)	(-7.50, -3.65)	0.06 (1.39)	(-2.67, 2.78)	0.968
		Irbesartan	99	65 (65.7)	-5.63 (0.98)	(-7.55, -3.71)			
	Week 58	Sparsentan	96	49 (51.0)	-5.67 (1.13)	(-7.88, -3.45)	1.26 (1.64)	(-1.95, 4.47)	0.442
		Irbesartan	99	43 (43.4)	-6.93 (1.19)	(-9.25, -4.60)			
> 45 years	Week 6	Sparsentan	106	100 (94.3)	0.12 (0.61)	(-1.07, 1.31)	0.83 (0.86)	(-0.86, 2.52)	0.334
		Irbesartan	103	98 (95.1)	-0.71 (0.61)	(-1.91, 0.49)			
	Week 36	Sparsentan	106	71 (67.0)	-1.98 (0.70)	(-3.35, -0.61)	1.52 (1.00)	(-0.44, 3.48)	0.128
		Irbesartan	103	68 (66.0)	-3.50 (0.71)	(-4.90, -2.10)			
	Week 58	Sparsentan	106	61 (57.5)	-3.69 (0.76)	(-5.19, -2.20)	1.68 (1.11)	(-0.50, 3.87)	0.131
		Irbesartan	103	52 (50.5)	-5.37 (0.81)	(-6.97, -3.78)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
 Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.172
<= 18 years	Week 6	Sparsentan	9	9 (100.0)	-7.25 (3.26)	(-13.77, -0.72)	-5.36 (5.58)	(-16.52, 5.80)	0.340
		Irbesartan	5	5 (100.0)	-1.88 (4.42)	(-10.72, 6.95)			
	Week 36	Sparsentan	9	6 (66.7)	-9.57 (4.03)	(-17.62, -1.51)	-1.50 (7.92)	(-17.35, 14.35)	0.851
		Irbesartan	5	2 (40.0)	-8.07 (6.86)	(-21.79, 5.65)			
	Week 58	Sparsentan	9	4 (44.4)	-8.67 (4.99)	(-18.66, 1.31)	7.30 (10.96)	(-14.62, 29.23)	0.508
		Irbesartan	5	1 (20.0)	-15.98 (9.83)	(-35.63, 3.68)			
> 18 to 40 years	Week 6	Sparsentan	102	98 (96.1)	-1.40 (0.76)	(-2.89, 0.09)	0.66 (1.06)	(-1.41, 2.74)	0.531
		Irbesartan	109	105 (96.3)	-2.06 (0.74)	(-3.51, -0.62)			
	Week 36	Sparsentan	102	68 (66.7)	-4.33 (0.88)	(-6.05, -2.60)	0.90 (1.22)	(-1.49, 3.29)	0.460
		Irbesartan	109	74 (67.9)	-5.23 (0.84)	(-6.88, -3.57)			
	Week 58	Sparsentan	102	56 (54.9)	-4.78 (0.97)	(-6.69, -2.87)	1.23 (1.40)	(-1.51, 3.97)	0.378
		Irbesartan	109	51 (46.8)	-6.02 (1.00)	(-7.97, -4.06)			

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 40 years	Week 6	Sparsentan	91	86 (94.5)	-0.17 (0.69)	(-1.53, 1.19)	0.60 (0.98)	(-1.33, 2.53)	0.542
		Irbesartan	88	85 (96.6)	-0.77 (0.70)	(-2.14, 0.60)			
	Week 36	Sparsentan	91	62 (68.1)	-2.25 (0.79)	(-3.80, -0.70)	1.26 (1.14)	(-0.97, 3.50)	0.266
		Irbesartan	88	57 (64.8)	-3.52 (0.82)	(-5.12, -1.91)			
	Week 58	Sparsentan	91	50 (54.9)	-3.81 (0.88)	(-5.54, -2.08)	2.20 (1.29)	(-0.33, 4.73)	0.089
		Irbesartan	88	43 (48.9)	-6.01 (0.94)	(-7.85, -4.16)			

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.424
North America	Week 6	Sparsentan	35	34 (97.1)	-2.52 (1.28)	(-5.03, -0.01)	-0.86 (1.70)	(-4.20, 2.48)	0.612
		Irbesartan	46	44 (95.7)	-1.66 (1.12)	(-3.86, 0.54)			
	Week 36	Sparsentan	35	19 (54.3)	-4.36 (1.65)	(-7.61, -1.10)	-0.66 (2.19)	(-4.96, 3.64)	0.763
		Irbesartan	46	26 (56.5)	-3.70 (1.42)	(-6.49, -0.90)			
	Week 58	Sparsentan	35	14 (40.0)	-5.87 (1.94)	(-9.69, -2.05)	-0.31 (2.59)	(-5.40, 4.77)	0.903
		Irbesartan	46	18 (39.1)	-5.56 (1.69)	(-8.88, -2.23)			
Europe	Week 6	Sparsentan	98	92 (93.9)	-0.50 (0.71)	(-1.89, 0.90)	0.94 (0.97)	(-0.95, 2.84)	0.329
		Irbesartan	115	111 (96.5)	-1.44 (0.65)	(-2.72, -0.16)			
	Week 36	Sparsentan	98	57 (58.2)	-2.48 (0.87)	(-4.19, -0.78)	1.61 (1.15)	(-0.65, 3.88)	0.162
		Irbesartan	115	77 (67.0)	-4.10 (0.76)	(-5.59, -2.61)			
	Week 58	Sparsentan	98	47 (48.0)	-2.89 (0.98)	(-4.81, -0.98)	2.36 (1.33)	(-0.24, 4.97)	0.075
		Irbesartan	115	54 (47.0)	-5.26 (0.90)	(-7.02, -3.50)			

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Asia Pacific	Week 6	Sparsentan	69	67 (97.1)	-1.23 (0.92)	(-3.04, 0.57)	0.24 (1.50)	(-2.71, 3.19)	0.874
		Irbesartan	41	40 (97.6)	-1.47 (1.18)	(-3.80, 0.86)			
	Week 36	Sparsentan	69	60 (87.0)	-4.66 (0.95)	(-6.54, -2.78)	1.81 (1.62)	(-1.38, 5.00)	0.266
		Irbesartan	41	30 (73.2)	-6.47 (1.31)	(-9.05, -3.89)			
	Week 58	Sparsentan	69	49 (71.0)	-5.84 (1.03)	(-7.87, -3.82)	3.18 (1.82)	(-0.39, 6.76)	0.080
		Irbesartan	41	23 (56.1)	-9.03 (1.49)	(-11.96, -6.09)			

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
 Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline BMI	Overall		Sparsentan						Interaction:	0.106
< 27 kg/m**2	Week 6		Sparsentan	84	78 (92.9)	-1.49 (0.85)	(-3.15, 0.18)	0.80 (1.16)	(-1.49, 3.08)	0.495
			Irbesartan	94	91 (96.8)	-2.28 (0.79)	(-3.84, -0.73)			
	Week 36		Sparsentan	84	56 (66.7)	-3.78 (0.98)	(-5.70, -1.87)	2.60 (1.36)	(-0.07, 5.26)	0.056
			Irbesartan	94	60 (63.8)	-6.38 (0.94)	(-8.23, -4.54)			
	Week 58		Sparsentan	84	46 (54.8)	-5.04 (1.08)	(-7.16, -2.92)	2.72 (1.58)	(-0.39, 5.83)	0.086
			Irbesartan	94	38 (40.4)	-7.76 (1.16)	(-10.04, -5.49)			
≥ 27 kg/m**2	Week 6		Sparsentan	118	115 (97.5)	-0.78 (0.64)	(-2.04, 0.48)	0.13 (0.94)	(-1.71, 1.96)	0.893
			Irbesartan	107	103 (96.3)	-0.91 (0.68)	(-2.24, 0.43)			
	Week 36		Sparsentan	118	80 (67.8)	-3.59 (0.74)	(-5.05, -2.13)	-0.52 (1.08)	(-2.64, 1.59)	0.627
			Irbesartan	107	73 (68.2)	-3.07 (0.78)	(-4.60, -1.53)			
	Week 58		Sparsentan	118	64 (54.2)	-4.25 (0.84)	(-5.89, -2.61)	0.71 (1.22)	(-1.68, 3.10)	0.561
			Irbesartan	107	57 (53.3)	-4.96 (0.88)	(-6.69, -3.23)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Randomization strata	Overall		Sparsentan							Interaction: 0.137
eGFR Low and UP High	Week 6		Sparsentan	71	69 (97.2)	-0.58 (0.68)	(-1.93, 0.76)	1.09 (0.96)	(-0.80, 2.98)	0.258
			Irbesartan	74	70 (94.6)	-1.67 (0.67)	(-3.00, -0.34)			
	Week 36		Sparsentan	71	49 (69.0)	-2.30 (0.78)	(-3.83, -0.77)	4.07 (1.10)	(1.90, 6.24)	<0.001 *
			Irbesartan	74	47 (63.5)	-6.36 (0.78)	(-7.90, -4.83)			
Week 58		Sparsentan	71	42 (59.2)	-3.91 (0.85)	(-5.58, -2.24)	3.11 (1.25)	(0.67, 5.56)	0.013 *	
		Irbesartan	74	34 (45.9)	-7.02 (0.91)	(-8.81, -5.23)				
eGFR Low and UP Low	Week 6		Sparsentan	55	50 (90.9)	0.27 (0.80)	(-1.31, 1.85)	1.27 (1.13)	(-0.95, 3.50)	0.262
			Irbesartan	55	54 (98.2)	-1.00 (0.79)	(-2.56, 0.56)			
	Week 36		Sparsentan	55	35 (63.6)	-0.62 (0.91)	(-2.41, 1.18)	1.76 (1.27)	(-0.74, 4.26)	0.167
			Irbesartan	55	40 (72.7)	-2.38 (0.88)	(-4.11, -0.65)			
	Week 58		Sparsentan	55	28 (50.9)	-1.93 (1.03)	(-3.96, 0.10)	2.01 (1.45)	(-0.84, 4.86)	0.166
			Irbesartan	55	29 (52.7)	-3.94 (1.01)	(-5.93, -1.94)			

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Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
eGFR High and UP High	Week 6	Sparsentan	37	35 (94.6)	-3.64 (1.69)	(-6.97, -0.30)	-0.41 (2.42)	(-5.16, 4.35)	0.867	
		Irbesartan	36	34 (94.4)	-3.23 (1.72)	(-6.62, 0.16)				
	Week 36	Sparsentan	37	26 (70.3)	-8.73 (1.94)	(-12.56, -4.91)	-2.71 (2.86)	(-8.33, 2.91)	0.344	
		Irbesartan	36	22 (61.1)	-6.02 (2.09)	(-10.14, -1.91)				
	Week 58	Sparsentan	37	20 (54.1)	-6.85 (2.21)	(-11.20, -2.50)	1.60 (3.35)	(-4.99, 8.19)	0.633	
		Irbesartan	36	15 (41.7)	-8.45 (2.51)	(-13.39, -3.52)				
eGFR High and UP Low	Week 6	Sparsentan	39	39 (100.0)	-1.69 (1.31)	(-4.28, 0.90)	-1.22 (1.88)	(-4.93, 2.48)	0.516	
		Irbesartan	37	37 (100.0)	-0.47 (1.35)	(-3.13, 2.19)				
	Week 36	Sparsentan	39	26 (66.7)	-5.96 (1.56)	(-9.03, -2.89)	-2.65 (2.25)	(-7.08, 1.78)	0.240	
		Irbesartan	37	24 (64.9)	-3.31 (1.62)	(-6.50, -0.12)				
	Week 58	Sparsentan	39	20 (51.3)	-7.97 (1.76)	(-11.43, -4.50)	-1.61 (2.59)	(-6.71, 3.49)	0.535	
		Irbesartan	37	17 (45.9)	-6.36 (1.90)	(-10.10, -2.62)				

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline eGFR Group 1		Overall	Sparsentan							Interaction: 0.207
< 60 mL/min/1.73 m**2	Week 6	Sparsentan	127	120 (94.5)	0.21 (0.53)	(-0.82, 1.24)	0.89 (0.74)	(-0.56, 2.34)	0.227	
		Irbesartan	129	126 (97.7)	-0.68 (0.52)	(-1.69, 0.34)				
	Week 36	Sparsentan	127	86 (67.7)	-1.33 (0.60)	(-2.50, -0.16)	2.64 (0.84)	(0.99, 4.28)	0.002 *	
		Irbesartan	129	88 (68.2)	-3.97 (0.59)	(-5.12, -2.81)				
	Week 58	Sparsentan	127	69 (54.3)	-2.72 (0.67)	(-4.03, -1.41)	2.13 (0.95)	(0.26, 4.00)	0.025 *	
		Irbesartan	129	66 (51.2)	-4.85 (0.68)	(-6.18, -3.52)				
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	47 (95.9)	-2.78 (1.25)	(-5.23, -0.33)	-0.07 (1.79)	(-3.59, 3.46)	0.970	
		Irbesartan	48	44 (91.7)	-2.72 (1.28)	(-5.23, -0.20)				
	Week 36	Sparsentan	49	30 (61.2)	-6.84 (1.52)	(-9.82, -3.85)	-0.39 (2.13)	(-4.57, 3.79)	0.854	
		Irbesartan	48	32 (66.7)	-6.45 (1.48)	(-9.36, -3.53)				
	Week 58	Sparsentan	49	23 (46.9)	-7.96 (1.73)	(-11.36, -4.56)	2.04 (2.55)	(-2.98, 7.05)	0.425	
		Irbesartan	48	19 (39.6)	-10.00 (1.88)	(-13.69, -6.31)				

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	26 (100.0)	-4.22 (1.97)	(-8.10, -0.34)	-0.13 (2.81)	(-5.68, 5.42)	0.963
		Irbesartan	25	25 (100.0)	-4.09 (2.01)	(-8.05, -0.13)			
	Week 36	Sparsentan	26	20 (76.9)	-10.49 (2.22)	(-14.86, -6.12)	-6.21 (3.44)	(-13.00, 0.58)	0.073
		Irbesartan	25	13 (52.0)	-4.28 (2.64)	(-9.47, 0.91)			
	Week 58	Sparsentan	26	18 (69.2)	-8.95 (2.33)	(-13.55, -4.35)	-0.79 (3.84)	(-8.36, 6.77)	0.836
		Irbesartan	25	10 (40.0)	-8.15 (3.05)	(-14.16, -2.15)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.228
< 45 mL/min/1.73 m**2	Week 6	Sparsentan	82	79 (96.3)	0.84 (0.58)	(-0.29, 1.98)	1.71 (0.82)	(0.10, 3.33)	0.038 *
		Irbesartan	80	78 (97.5)	-0.87 (0.58)	(-2.02, 0.28)			
	Week 36	Sparsentan	82	54 (65.9)	-1.44 (0.66)	(-2.74, -0.15)	3.20 (0.92)	(1.38, 5.01)	<0.001 *
		Irbesartan	80	56 (70.0)	-4.64 (0.65)	(-5.92, -3.37)			
	Week 58	Sparsentan	82	44 (53.7)	-3.01 (0.73)	(-4.45, -1.57)	2.84 (1.04)	(0.79, 4.89)	0.007 *
		Irbesartan	80	42 (52.5)	-5.85 (0.74)	(-7.30, -4.40)			
45 to < 60 mL/min/1.73 m**2	Week 6	Sparsentan	45	41 (91.1)	-1.01 (1.06)	(-3.09, 1.06)	-0.66 (1.45)	(-3.51, 2.19)	0.649
		Irbesartan	49	48 (98.0)	-0.35 (0.99)	(-2.30, 1.60)			
	Week 36	Sparsentan	45	32 (71.1)	-1.20 (1.17)	(-3.49, 1.10)	1.62 (1.66)	(-1.64, 4.89)	0.328
		Irbesartan	49	32 (65.3)	-2.82 (1.17)	(-5.13, -0.51)			
	Week 58	Sparsentan	45	25 (55.6)	-2.26 (1.33)	(-4.87, 0.35)	1.01 (1.90)	(-2.73, 4.75)	0.596
		Irbesartan	49	24 (49.0)	-3.27 (1.35)	(-5.93, -0.61)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis			
						Change from Baseline		Treatment Difference	
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	47 (95.9)	-2.78 (1.25)	(-5.23, -0.33)	-0.07 (1.79)	(-3.59, 3.46)	0.970
		Irbesartan	48	44 (91.7)	-2.72 (1.28)	(-5.23, -0.20)			
	Week 36	Sparsentan	49	30 (61.2)	-6.84 (1.52)	(-9.82, -3.85)	-0.39 (2.13)	(-4.57, 3.79)	0.854
		Irbesartan	48	32 (66.7)	-6.45 (1.48)	(-9.36, -3.53)			
	Week 58	Sparsentan	49	23 (46.9)	-7.96 (1.73)	(-11.36, -4.56)	2.04 (2.55)	(-2.98, 7.05)	0.425
		Irbesartan	48	19 (39.6)	-10.00 (1.88)	(-13.69, -6.31)			
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	26 (100.0)	-4.22 (1.97)	(-8.10, -0.34)	-0.13 (2.81)	(-5.68, 5.42)	0.963
		Irbesartan	25	25 (100.0)	-4.09 (2.01)	(-8.05, -0.13)			
	Week 36	Sparsentan	26	20 (76.9)	-10.49 (2.22)	(-14.86, -6.12)	-6.21 (3.44)	(-13.00, 0.58)	0.073
		Irbesartan	25	13 (52.0)	-4.28 (2.64)	(-9.47, 0.91)			
	Week 58	Sparsentan	26	18 (69.2)	-8.95 (2.33)	(-13.55, -4.35)	-0.79 (3.84)	(-8.36, 6.77)	0.836
		Irbesartan	25	10 (40.0)	-8.15 (3.05)	(-14.16, -2.15)			

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Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline urine protein excretion		Overall	Sparsentan							Interaction: 0.825
<= 1.75 g/day	Week 6	Sparsentan	98	93 (94.9)	-0.26 (0.77)	(-1.77, 1.24)	0.33 (1.09)	(-1.82, 2.47)	0.765	
		Irbesartan	94	91 (96.8)	-0.59 (0.78)	(-2.12, 0.94)				
	Week 36	Sparsentan	98	70 (71.4)	-1.73 (0.86)	(-3.42, -0.03)	0.67 (1.26)	(-1.82, 3.15)	0.598	
		Irbesartan	94	61 (64.9)	-2.39 (0.92)	(-4.20, -0.58)				
	Week 58	Sparsentan	98	58 (59.2)	-3.69 (0.95)	(-5.56, -1.83)	0.41 (1.51)	(-2.55, 3.37)	0.785	
		Irbesartan	94	36 (38.3)	-4.11 (1.17)	(-6.40, -1.81)				
> 1.75 g/day	Week 6	Sparsentan	104	100 (96.2)	-1.90 (0.68)	(-3.23, -0.56)	0.40 (0.96)	(-1.48, 2.27)	0.679	
		Irbesartan	108	104 (96.3)	-2.29 (0.67)	(-3.61, -0.98)				
	Week 36	Sparsentan	104	66 (63.5)	-5.65 (0.81)	(-7.23, -4.06)	0.70 (1.12)	(-1.50, 2.90)	0.532	
		Irbesartan	108	72 (66.7)	-6.35 (0.77)	(-7.87, -4.83)				
	Week 58	Sparsentan	104	52 (50.0)	-5.55 (0.92)	(-7.36, -3.75)	2.06 (1.26)	(-0.42, 4.54)	0.103	
		Irbesartan	108	59 (54.6)	-7.61 (0.86)	(-9.31, -5.92)				

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Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline use of antihypertensives		Overall	Sparsentan							Interaction: 0.970
Yes	Week 6	Sparsentan	88	83 (94.3)	-0.36 (0.73)	(-1.79, 1.06)	1.49 (1.05)	(-0.56, 3.55)	0.154	
		Irbesartan	83	77 (92.8)	-1.86 (0.75)	(-3.34, -0.38)				
	Week 36	Sparsentan	88	55 (62.5)	-3.26 (0.87)	(-4.97, -1.56)	0.74 (1.21)	(-1.64, 3.13)	0.541	
		Irbesartan	83	59 (71.1)	-4.00 (0.85)	(-5.67, -2.34)				
	Week 58	Sparsentan	88	44 (50.0)	-4.17 (0.98)	(-6.10, -2.25)	1.64 (1.39)	(-1.08, 4.36)	0.237	
		Irbesartan	83	44 (53.0)	-5.81 (0.98)	(-7.73, -3.90)				
No	Week 6	Sparsentan	114	110 (96.5)	-1.69 (0.72)	(-3.10, -0.28)	-0.41 (1.00)	(-2.38, 1.56)	0.680	
		Irbesartan	119	118 (99.2)	-1.27 (0.70)	(-2.64, 0.10)				
	Week 36	Sparsentan	114	81 (71.1)	-4.06 (0.81)	(-5.65, -2.46)	0.91 (1.17)	(-1.39, 3.20)	0.438	
		Irbesartan	119	74 (62.2)	-4.96 (0.84)	(-6.61, -3.32)				
	Week 58	Sparsentan	114	66 (57.9)	-4.98 (0.90)	(-6.74, -3.22)	1.41 (1.34)	(-1.22, 4.05)	0.293	
		Irbesartan	119	51 (42.9)	-6.39 (1.00)	(-8.35, -4.44)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.289
<= 5 years	Week 6	Sparsentan	113	106 (93.8)	-1.38 (0.75)	(-2.87, 0.10)	0.07 (1.03)	(-1.96, 2.10)	0.945
		Irbesartan	127	122 (96.1)	-1.46 (0.71)	(-2.84, -0.07)			
	Week 36	Sparsentan	113	72 (63.7)	-3.90 (0.88)	(-5.62, -2.17)	0.77 (1.21)	(-1.62, 3.15)	0.528
		Irbesartan	127	80 (63.0)	-4.67 (0.84)	(-6.31, -3.02)			
	Week 58	Sparsentan	113	56 (49.6)	-5.79 (1.00)	(-7.75, -3.82)	0.11 (1.41)	(-2.66, 2.88)	0.941
		Irbesartan	127	56 (44.1)	-5.89 (0.99)	(-7.84, -3.94)			
> 5 years	Week 6	Sparsentan	89	87 (97.8)	-0.72 (0.66)	(-2.02, 0.58)	0.96 (0.98)	(-0.96, 2.89)	0.327
		Irbesartan	75	73 (97.3)	-1.68 (0.72)	(-3.10, -0.26)			
	Week 36	Sparsentan	89	64 (71.9)	-3.37 (0.76)	(-4.86, -1.89)	1.06 (1.12)	(-1.14, 3.26)	0.346
		Irbesartan	75	53 (70.7)	-4.43 (0.83)	(-6.05, -2.81)			
	Week 58	Sparsentan	89	54 (60.7)	-3.27 (0.83)	(-4.89, -1.65)	3.45 (1.26)	(0.98, 5.93)	0.006 *
		Irbesartan	75	39 (52.0)	-6.72 (0.95)	(-8.59, -4.85)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGC\_FSCM: Change from baseline in eGFR by subgroup  
 Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.848
Yes	Week 6	Sparsentan	153	144 (94.1)	-0.63 (0.57)	(-1.76, 0.49)	0.64 (0.80)	(-0.94, 2.22)	0.427
		Irbesartan	157	151 (96.2)	-1.27 (0.56)	(-2.38, -0.17)			
	Week 36	Sparsentan	153	101 (66.0)	-2.87 (0.67)	(-4.18, -1.57)	0.79 (0.93)	(-1.04, 2.62)	0.396
		Irbesartan	157	105 (66.9)	-3.67 (0.65)	(-4.95, -2.38)			
	Week 58	Sparsentan	153	82 (53.6)	-3.73 (0.74)	(-5.19, -2.28)	1.94 (1.07)	(-0.15, 4.03)	0.069
		Irbesartan	157	76 (48.4)	-5.67 (0.76)	(-7.17, -4.17)			
No	Week 6	Sparsentan	49	49 (100.0)	-2.39 (1.13)	(-4.61, -0.18)	0.11 (1.65)	(-3.13, 3.36)	0.945
		Irbesartan	45	44 (97.8)	-2.51 (1.19)	(-4.84, -0.17)			
	Week 36	Sparsentan	49	35 (71.4)	-6.00 (1.29)	(-8.53, -3.47)	2.11 (1.92)	(-1.68, 5.89)	0.274
		Irbesartan	45	28 (62.2)	-8.10 (1.42)	(-10.90, -5.31)			
	Week 58	Sparsentan	49	28 (57.1)	-7.08 (1.43)	(-9.90, -4.26)	1.27 (2.23)	(-3.12, 5.65)	0.570
		Irbesartan	45	19 (42.2)	-8.35 (1.70)	(-11.69, -5.01)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model. Only key visits up to Week 58 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Figure PF1GGC\_FSGM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Not done: No significant subgroup found.

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m<sup>2</sup>.  
eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High  $\geq$  60 ml/min/1.73m<sup>2</sup>, UP Low =  $\leq$  1.75 g/day, UP High =  $>$  1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Reference table: PT1GGC\_FSCM.

Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan					Interaction:	0.033 #
Male	Baseline to Week 58	Sparsentan	139	-4.00 (1.08)	(-6.12, -1.87)	0.37 (1.50)	(-2.58, 3.33)	0.804
		Irbesartan	143	-4.37 (1.05)	(-6.43, -2.31)			
Female	Baseline to Week 58	Sparsentan	63	-4.76 (1.57)	(-7.87, -1.66)	1.55 (2.39)	(-3.17, 6.28)	0.517
		Irbesartan	59	-6.32 (1.80)	(-9.88, -2.75)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan					Interaction:	0.529
<= 45 years	Baseline to Week 58	Sparsentan	96	-4.74 (1.51)	(-7.71, -1.76)	0.93 (2.14)	(-3.29, 5.15)	0.663
		Irbesartan	99	-5.67 (1.52)	(-8.66, -2.68)			
> 45 years	Baseline to Week 58	Sparsentan	106	-3.77 (0.99)	(-5.72, -1.82)	0.38 (1.43)	(-2.43, 3.19)	0.790
		Irbesartan	103	-4.15 (1.03)	(-6.17, -2.12)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		p-value
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Age at IgAN diagnosis	Overall	Sparsentan					Interaction:	0.008 #
<= 18 years	Baseline to Week 58	Sparsentan	9	-5.09 (3.62)	(-12.30, 2.12)	6.62 (7.00)	(-7.34, 20.58)	0.348
		Irbesartan	5	-11.71 (5.99)	(-23.65, 0.22)			
> 18 to 40 years	Baseline to Week 58	Sparsentan	102	-4.27 (1.35)	(-6.94, -1.60)	0.42 (1.89)	(-3.31, 4.14)	0.826
		Irbesartan	109	-4.68 (1.32)	(-7.28, -2.08)			
> 40 years	Baseline to Week 58	Sparsentan	91	-3.77 (1.15)	(-6.03, -1.50)	1.22 (1.66)	(-2.06, 4.50)	0.464
		Irbesartan	88	-4.99 (1.20)	(-7.36, -2.62)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan					Interaction:	0.177
North America	Baseline to Week 58	Sparsentan	35	-4.65 (2.54)	(-9.72, 0.43)	-0.93 (3.35)	(-7.62, 5.76)	0.782
		Irbesartan	46	-3.72 (2.18)	(-8.07, 0.64)			
Europe	Baseline to Week 58	Sparsentan	98	-3.21 (1.31)	(-5.79, -0.62)	1.21 (1.75)	(-2.25, 4.66)	0.493
		Irbesartan	115	-4.41 (1.17)	(-6.71, -2.11)			
Asia Pacific	Baseline to Week 58	Sparsentan	69	-5.30 (1.36)	(-7.99, -2.60)	1.59 (2.35)	(-3.05, 6.24)	0.498
		Irbesartan	41	-6.89 (1.91)	(-10.67, -3.11)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan					Interaction:	0.161
< 27 kg/m**2	Baseline to Week 58	Sparsentan	84	-3.76 (1.50)	(-6.72, -0.80)	2.87 (2.11)	(-1.29, 7.03)	0.175
		Irbesartan	94	-6.63 (1.48)	(-9.55, -3.70)			
>= 27 kg/m**2	Baseline to Week 58	Sparsentan	118	-4.56 (1.07)	(-6.68, -2.45)	-1.03 (1.56)	(-4.10, 2.05)	0.511
		Irbesartan	107	-3.54 (1.13)	(-5.77, -1.31)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan					Interaction:	0.037 #
eGFR Low and UP High	Baseline to Week 58	Sparsentan	71	-4.60 (1.40)	(-7.38, -1.82)	2.41 (1.98)	(-1.52, 6.35)	0.227
		Irbesartan	74	-7.01 (1.41)	(-9.80, -4.22)			
eGFR Low and UP Low	Baseline to Week 58	Sparsentan	55	-2.51 (1.30)	(-5.09, 0.07)	0.02 (1.82)	(-3.59, 3.64)	0.991
		Irbesartan	55	-2.53 (1.28)	(-5.06, -0.00)			
eGFR High and UP High	Baseline to Week 58	Sparsentan	37	-3.57 (2.90)	(-9.36, 2.22)	1.64 (4.28)	(-6.88, 10.16)	0.702
		Irbesartan	36	-5.21 (3.14)	(-11.46, 1.04)			
eGFR High and UP Low	Baseline to Week 58	Sparsentan	39	-7.49 (2.16)	(-11.80, -3.18)	-2.98 (3.14)	(-9.24, 3.28)	0.345
		Irbesartan	37	-4.51 (2.28)	(-9.05, 0.03)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan					Interaction:	0.154
< 60 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	127	-3.62 (0.93)	(-5.45, -1.79)	1.02 (1.31)	(-1.56, 3.60)	0.438
		Irbesartan	129	-4.64 (0.93)	(-6.47, -2.82)			
60 to < 90 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	49	-5.11 (2.10)	(-9.28, -0.93)	1.39 (3.00)	(-4.56, 7.35)	0.643
		Irbesartan	48	-6.50 (2.14)	(-10.75, -2.25)			
>= 90 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	26	-6.59 (3.36)	(-13.38, 0.19)	-3.84 (5.23)	(-14.35, 6.68)	0.467
		Irbesartan	25	-2.76 (4.00)	(-10.80, 5.28)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
				Annualized Slope		Slope Difference		
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan					Interaction:	0.097
< 45 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	82	-4.19 (1.16)	(-6.49, -1.89)	1.87 (1.63)	(-1.37, 5.11)	0.255
		Irbesartan	80	-6.06 (1.15)	(-8.34, -3.79)			
45 to < 60 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	45	-2.56 (1.70)	(-5.93, 0.81)	-0.04 (2.41)	(-4.82, 4.74)	0.987
		Irbesartan	49	-2.52 (1.71)	(-5.91, 0.87)			
60 to < 90 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	49	-5.11 (2.10)	(-9.28, -0.93)	1.39 (3.00)	(-4.56, 7.35)	0.643
		Irbesartan	48	-6.50 (2.14)	(-10.75, -2.25)			
>= 90 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	26	-6.59 (3.36)	(-13.38, 0.19)	-3.84 (5.23)	(-14.35, 6.68)	0.467
		Irbesartan	25	-2.76 (4.00)	(-10.80, 5.28)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
				Annualized Slope			Slope Difference	
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan					Interaction:	0.606
<= 1.75 g/day	Baseline to Week 58	Sparsentan	98	-3.48 (1.19)	(-5.83, -1.13)	-0.18 (1.79)	(-3.70, 3.34)	0.920
		Irbesartan	94	-3.30 (1.33)	(-5.92, -0.68)			
> 1.75 g/day	Baseline to Week 58	Sparsentan	104	-4.98 (1.30)	(-7.54, -2.42)	1.18 (1.79)	(-2.36, 4.72)	0.511
		Irbesartan	108	-6.16 (1.24)	(-8.60, -3.71)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan					Interaction:	0.764
Yes	Baseline to Week 58	Sparsentan	88	-3.69 (1.22)	(-6.10, -1.29)	0.79 (1.72)	(-2.61, 4.18)	0.648
		Irbesartan	83	-4.48 (1.21)	(-6.87, -2.09)			
No	Baseline to Week 58	Sparsentan	114	-4.73 (1.24)	(-7.18, -2.29)	0.37 (1.79)	(-3.17, 3.90)	0.838
		Irbesartan	119	-5.10 (1.30)	(-7.65, -2.55)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan					Interaction:	0.388
<= 5 years	Baseline to Week 58	Sparsentan	113	-4.51 (1.33)	(-7.14, -1.88)	-0.02 (1.85)	(-3.67, 3.63)	0.993
		Irbesartan	127	-4.49 (1.29)	(-7.03, -1.96)			
> 5 years	Baseline to Week 58	Sparsentan	89	-3.96 (1.09)	(-6.12, -1.80)	1.49 (1.63)	(-1.73, 4.71)	0.363
		Irbesartan	75	-5.45 (1.21)	(-7.84, -3.06)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan					Interaction:	0.727
Yes	Baseline to Week 58	Sparsentan	153	-3.56 (0.97)	(-5.47, -1.64)	0.84 (1.38)	(-1.87, 3.55)	0.540
		Irbesartan	157	-4.40 (0.97)	(-6.32, -2.49)			
No	Baseline to Week 58	Sparsentan	49	-6.22 (1.98)	(-10.16, -2.28)	0.36 (2.94)	(-5.50, 6.22)	0.903
		Irbesartan	45	-6.58 (2.18)	(-10.92, -2.25)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan					Interaction:	0.026 #
Male	Week 6 to Week 58	Sparsentan	139	-3.83 (1.13)	(-6.06, -1.61)	0.93 (1.57)	(-2.17, 4.02)	0.556
		Irbesartan	143	-4.76 (1.10)	(-6.91, -2.61)			
Female	Week 6 to Week 58	Sparsentan	63	-4.08 (1.65)	(-7.34, -0.82)	1.27 (2.55)	(-3.76, 6.30)	0.619
		Irbesartan	59	-5.35 (1.94)	(-9.18, -1.51)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan					Interaction:	0.493
<= 45 years	Week 6 to Week 58	Sparsentan	96	-4.07 (1.58)	(-7.17, -0.96)	1.40 (2.24)	(-3.02, 5.81)	0.533
		Irbesartan	99	-5.47 (1.59)	(-8.60, -2.33)			
> 45 years	Week 6 to Week 58	Sparsentan	106	-3.70 (1.06)	(-5.78, -1.62)	0.71 (1.53)	(-2.29, 3.72)	0.641
		Irbesartan	103	-4.41 (1.10)	(-6.58, -2.24)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan					Interaction:	0.102
<= 18 years	Week 6 to Week 58	Sparsentan	9	-6.01 (4.34)	(-16.11, 4.09)	4.46 (8.67)	(-16.48, 25.40)	0.624
		Irbesartan	5	-10.47 (7.52)	(-28.88, 7.94)			
> 18 to 40 years	Week 6 to Week 58	Sparsentan	102	-3.89 (1.41)	(-6.67, -1.11)	1.04 (1.98)	(-2.86, 4.93)	0.600
		Irbesartan	109	-4.93 (1.38)	(-7.66, -2.21)			
> 40 years	Week 6 to Week 58	Sparsentan	91	-3.55 (1.22)	(-5.96, -1.14)	1.26 (1.77)	(-2.23, 4.76)	0.477
		Irbesartan	88	-4.81 (1.28)	(-7.34, -2.28)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		p-value
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Geographic region	Overall	Sparsentan					Interaction:	0.788
North America	Week 6 to Week 58	Sparsentan	35	-3.81 (2.70)	(-9.19, 1.56)	-0.61 (3.56)	(-7.68, 6.47)	0.865
		Irbesartan	46	-3.21 (2.32)	(-7.81, 1.40)			
Europe	Week 6 to Week 58	Sparsentan	98	-3.12 (1.37)	(-5.83, -0.42)	1.37 (1.84)	(-2.25, 4.99)	0.456
		Irbesartan	115	-4.50 (1.22)	(-6.90, -2.09)			
Asia Pacific	Week 6 to Week 58	Sparsentan	69	-4.87 (1.43)	(-7.71, -2.04)	2.62 (2.50)	(-2.32, 7.55)	0.297
		Irbesartan	41	-7.49 (2.05)	(-11.53, -3.44)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan					Interaction:	0.067
< 27 kg/m**2	Week 6 to Week 58	Sparsentan	84	-3.53 (1.56)	(-6.60, -0.45)	3.04 (2.20)	(-1.31, 7.38)	0.169
		Irbesartan	94	-6.57 (1.56)	(-9.64, -3.49)			
≥ 27 kg/m**2	Week 6 to Week 58	Sparsentan	118	-4.12 (1.15)	(-6.39, -1.86)	-0.41 (1.67)	(-3.70, 2.87)	0.804
		Irbesartan	107	-3.71 (1.21)	(-6.09, -1.33)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan					Interaction:	0.036 #
eGFR Low and UP High	Week 6 to Week 58	Sparsentan	71	-4.57 (1.39)	(-7.32, -1.81)	2.70 (1.98)	(-1.22, 6.62)	0.175
		Irbesartan	74	-7.27 (1.41)	(-10.05, -4.48)			
eGFR Low and UP Low	Week 6 to Week 58	Sparsentan	55	-2.39 (1.42)	(-5.18, 0.40)	0.10 (1.98)	(-3.80, 4.01)	0.959
		Irbesartan	55	-2.49 (1.38)	(-5.22, 0.23)			
eGFR High and UP High	Week 6 to Week 58	Sparsentan	37	-1.94 (3.09)	(-8.09, 4.21)	1.05 (4.57)	(-8.03, 10.13)	0.819
		Irbesartan	36	-2.99 (3.36)	(-9.67, 3.69)			
eGFR High and UP Low	Week 6 to Week 58	Sparsentan	39	-7.34 (2.31)	(-11.92, -2.76)	-1.03 (3.37)	(-7.72, 5.65)	0.760
		Irbesartan	37	-6.31 (2.45)	(-11.17, -1.44)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline eGFR Group 1	Overall	Sparsentan					Interaction:	0.034 #	
< 60 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	127	-3.57 (0.97)	(-5.47, -1.66)	1.19 (1.37)	(-1.49, 3.88)	0.382	
		Irbesartan	129	-4.76 (0.96)	(-6.66, -2.86)				
60 to < 90 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	49	-4.18 (2.20)	(-8.55, 0.18)	1.08 (3.15)	(-5.17, 7.32)	0.734	
		Irbesartan	48	-5.26 (2.25)	(-9.72, -0.80)				
>= 90 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	26	-5.74 (3.61)	(-12.97, 1.50)	-1.23 (5.72)	(-12.68, 10.21)	0.830	
		Irbesartan	25	-4.50 (4.43)	(-13.37, 4.36)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
				Annualized Slope		Slope Difference		
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan					Interaction:	0.035 #
< 45 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	82	-4.53 (1.15)	(-6.80, -2.26)	2.01 (1.62)	(-1.18, 5.20)	0.216
		Irbesartan	80	-6.54 (1.14)	(-8.78, -4.29)			
45 to < 60 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	45	-1.89 (1.85)	(-5.55, 1.77)	0.32 (2.62)	(-4.87, 5.51)	0.902
		Irbesartan	49	-2.22 (1.86)	(-5.90, 1.46)			
60 to < 90 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	49	-4.18 (2.20)	(-8.55, 0.18)	1.08 (3.15)	(-5.17, 7.32)	0.734
		Irbesartan	48	-5.26 (2.25)	(-9.72, -0.80)			
>= 90 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	26	-5.74 (3.61)	(-12.97, 1.50)	-1.23 (5.72)	(-12.68, 10.21)	0.830
		Irbesartan	25	-4.50 (4.43)	(-13.37, 4.36)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. Source Data: alb, created on: 19FEB2024

Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model					
				Annualized Slope			Slope Difference		
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline urine protein excretion	Overall	Sparsentan					Interaction:	0.639	
<= 1.75 g/day	Week 6 to Week 58	Sparsentan	98	-3.62 (1.26)	(-6.11, -1.14)	-0.07 (1.90)	(-3.82, 3.68)	0.970	
		Irbesartan	94	-3.55 (1.42)	(-6.36, -0.75)				
> 1.75 g/day	Week 6 to Week 58	Sparsentan	104	-4.18 (1.37)	(-6.87, -1.49)	1.82 (1.88)	(-1.90, 5.53)	0.336	
		Irbesartan	108	-6.00 (1.30)	(-8.55, -3.44)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model					
				Annualized Slope			Slope Difference		
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline use of antihypertensives	Overall	Sparsentan					Interaction:	0.355	
Yes	Week 6 to Week 58	Sparsentan	88	-4.03 (1.29)	(-6.57, -1.50)	0.35 (1.81)	(-3.22, 3.93)	0.847	
		Irbesartan	83	-4.39 (1.28)	(-6.90, -1.87)				
No	Week 6 to Week 58	Sparsentan	114	-3.85 (1.31)	(-6.42, -1.28)	1.42 (1.90)	(-2.31, 5.15)	0.454	
		Irbesartan	119	-5.28 (1.38)	(-7.98, -2.57)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan					Interaction:	0.339
<= 5 years	Week 6 to Week 58	Sparsentan	113	-4.11 (1.40)	(-6.88, -1.35)	0.42 (1.96)	(-3.43, 4.27)	0.829
		Irbesartan	127	-4.54 (1.36)	(-7.22, -1.86)			
> 5 years	Week 6 to Week 58	Sparsentan	89	-3.67 (1.15)	(-5.93, -1.40)	1.78 (1.72)	(-1.60, 5.17)	0.300
		Irbesartan	75	-5.45 (1.27)	(-7.96, -2.94)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan					Interaction:	0.550
Yes	Week 6 to Week 58	Sparsentan	153	-3.63 (1.03)	(-5.65, -1.60)	0.87 (1.46)	(-1.99, 3.74)	0.550
		Irbesartan	157	-4.50 (1.03)	(-6.53, -2.47)			
No	Week 6 to Week 58	Sparsentan	49	-4.75 (2.05)	(-8.82, -0.69)	1.69 (3.07)	(-4.39, 7.77)	0.582
		Irbesartan	45	-6.45 (2.28)	(-10.97, -1.93)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.039 #
Male	Double-blind period	Sparsentan	139	7 (5.0)	0.424 [0.181, 0.990]	0.393 [0.158, 0.980]	-6.9 [-14.0, 0.3]	0.053
		Irbesartan	143	17 (11.9)				
Female	Double-blind period	Sparsentan	63	7 (11.1)	2.185 [0.593, 8.059]	2.333 [0.574, 9.484]	6.0 [-5.2, 17.2]	0.326
		Irbesartan	59	3 (5.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.416
<= 45 years	Double-blind period	Sparsentan	96	8 (8.3)	0.917 [0.369, 2.277]	0.909 [0.336, 2.463]	-0.8 [-9.7, 8.2]	1.000
		Irbesartan	99	9 (9.1)				
> 45 years	Double-blind period	Sparsentan	106	6 (5.7)	0.530 [0.204, 1.380]	0.502 [0.178, 1.412]	-5.0 [-13.4, 3.3]	0.213
		Irbesartan	103	11 (10.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024



Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.673
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	1.111 [0.131, 9.416]	1.143 [0.077, 16.947]	2.2 [-57.7, 62.1]	1.000
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	7 (6.9)	0.831 [0.321, 2.149]	0.819 [0.293, 2.286]	-1.4 [-9.5, 6.7]	0.798
		Irbesartan	109	9 (8.3)				
> 40 years	Double-blind period	Sparsentan	91	5 (5.5)	0.484 [0.172, 1.358]	0.453 [0.148, 1.385]	-5.9 [-15.1, 3.4]	0.185
		Irbesartan	88	10 (11.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.253
North America	Double-blind period	Sparsentan	35	5 (14.3)	0.548 [0.213, 1.411]	0.472 [0.149, 1.496]	-11.8 [-31.5, 7.9]	0.273
		Irbesartan	46	12 (26.1)				
Europe	Double-blind period	Sparsentan	98	6 (6.1)	1.760 [0.511, 6.059]	1.810 [0.496, 6.607]	2.6 [-4.1, 9.4]	0.518
		Irbesartan	115	4 (3.5)				
Asia Pacific	Double-blind period	Sparsentan	69	3 (4.3)	0.446 [0.105, 1.893]	0.420 [0.089, 1.981]	-5.4 [-17.6, 6.8]	0.421
		Irbesartan	41	4 (9.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.430
< 27 kg/m**2	Double-blind period	Sparsentan	84	6 (7.1)	0.959 [0.336, 2.741]	0.956 [0.308, 2.967]	-0.3 [-9.1, 8.5]	1.000
		Irbesartan	94	7 (7.4)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	8 (6.8)	0.558 [0.241, 1.294]	0.526 [0.209, 1.323]	-5.4 [-13.9, 3.2]	0.178
		Irbesartan	107	13 (12.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.154
eGFR Low and UP High	Double-blind period	Sparsentan	71	3 (4.2)	0.261 [0.077, 0.885]	0.228 [0.061, 0.846]	-12.0 [-23.0, -1.0]	0.027 *
		Irbesartan	74	12 (16.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	5 (9.1)	2.500 [0.506, 12.341]	2.650 [0.492, 14.286]	5.5 [-5.4, 16.3]	0.438
		Irbesartan	55	2 (3.6)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)	0.973 [0.210, 4.507]	0.971 [0.183, 5.158]	-0.2 [-15.6, 15.1]	1.000
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	3 (7.7)	0.949 [0.204, 4.407]	0.944 [0.178, 5.005]	-0.4 [-15.2, 14.4]	1.000
		Irbesartan	37	3 (8.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 28FEB2024

Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.371
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	8 (6.3)	0.508 [0.225, 1.145]	0.475 [0.196, 1.153]	-6.1 [-14.0, 1.8]	0.132
		Irbesartan	129	16 (12.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	3 (6.1)	1.469 [0.257, 8.408]	1.500 [0.239, 9.400]	2.0 [-8.9, 12.8]	1.000
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	3 (11.5)	1.442 [0.263, 7.918]	1.500 [0.229, 9.832]	3.5 [-16.6, 23.7]	1.000
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.551
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	4 (4.9)	0.434 [0.139, 1.351]	0.405 [0.119, 1.372]	-6.4 [-16.0, 3.2]	0.158
		Irbesartan	80	9 (11.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	0.622 [0.195, 1.985]	0.585 [0.159, 2.151]	-5.4 [-20.4, 9.6]	0.528
		Irbesartan	49	7 (14.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	3 (6.1)	1.469 [0.257, 8.408]	1.500 [0.239, 9.400]	2.0 [-8.9, 12.8]	1.000
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	3 (11.5)	1.442 [0.263, 7.918]	1.500 [0.229, 9.832]	3.5 [-16.6, 23.7]	1.000
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.058
<= 1.75 g/day	Double-blind period	Sparsentan	98	9 (9.2)	1.439 [0.533, 3.886]	1.483 [0.507, 4.342]	2.8 [-5.8, 11.4]	0.593
		Irbesartan	94	6 (6.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	5 (4.8)	0.371 [0.139, 0.993]	0.339 [0.118, 0.978]	-8.2 [-16.7, 0.3]	0.053
		Irbesartan	108	14 (13.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.253
Yes	Double-blind period	Sparsentan	88	4 (4.5)	0.419 [0.134, 1.309]	0.392 [0.116, 1.324]	-6.3 [-15.4, 2.9]	0.153
		Irbesartan	83	9 (10.8)				
No	Double-blind period	Sparsentan	114	10 (8.8)	0.949 [0.419, 2.148]	0.944 [0.385, 2.317]	-0.5 [-8.7, 7.7]	1.000
		Irbesartan	119	11 (9.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024



Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.012 #
<= 5 years	Double-blind period	Sparsentan	113	5 (4.4)	0.331 [0.126, 0.867]	0.300 [0.107, 0.841]	-9.0 [-16.8, -1.1]	0.023 *
		Irbesartan	127	17 (13.4)				
> 5 years	Double-blind period	Sparsentan	89	9 (10.1)	2.528 [0.710, 9.002]	2.700 [0.704, 10.362]	6.1 [-2.8, 15.0]	0.228
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.413
Yes	Double-blind period	Sparsentan	153	11 (7.2)	0.627 [0.306, 1.284]	0.598 [0.273, 1.312]	-4.3 [-11.4, 2.8]	0.243
		Irbesartan	157	18 (11.5)				
No	Double-blind period	Sparsentan	49	3 (6.1)	1.378 [0.241, 7.871]	1.402 [0.223, 8.801]	1.7 [-9.5, 12.8]	1.000
		Irbesartan	45	2 (4.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.052
Male	Sparsentan	139	7 (5.0)	NE		0.408	(0.169, 0.985)	0.046 *
	Irbesartan	143	17 (11.9)	NE				
Female	Sparsentan	63	7 (11.1)	NE		1.782	(0.449, 7.070)	0.411
	Irbesartan	59	3 (5.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff\_tte, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.448
<= 45 years	Sparsentan	96	8 (8.3)	NE		0.857	(0.330, 2.223)	0.751
	Irbesartan	99	9 (9.1)	NE				
> 45 years	Sparsentan	106	6 (5.7)	NE		0.528	(0.195, 1.430)	0.209
	Irbesartan	103	11 (10.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff\_tte, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: 0.754
<= 18 years	Sparsentan	9	2 (22.2)	NE		0.785	(0.070, 8.812)	0.845
	Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Sparsentan	102	7 (6.9)	NE		0.845	(0.313, 2.286)	0.741
	Irbesartan	109	9 (8.3)	NE				
> 40 years	Sparsentan	91	5 (5.5)	NE		0.464	(0.158, 1.366)	0.163
	Irbesartan	88	10 (11.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff\_tte, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.151
North America	Sparsentan	35	5 (14.3)	NE		0.501	(0.176, 1.424)	0.194
	Irbesartan	46	12 (26.1)	NE				
Europe	Sparsentan	98	6 (6.1)	NE		2.092	(0.590, 7.422)	0.253
	Irbesartan	115	4 (3.5)	NE				
Asia Pacific	Sparsentan	69	3 (4.3)	NE		0.293	(0.063, 1.361)	0.117
	Irbesartan	41	4 (9.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff\_tte, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.460
< 27 kg/m**2	Sparsentan	84	6 (7.1)	NE		0.860	(0.287, 2.574)	0.787
	Irbesartan	94	7 (7.4)	NE				
≥ 27 kg/m**2	Sparsentan	118	8 (6.8)	NE		0.543	(0.224, 1.311)	0.175
	Irbesartan	107	13 (12.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff\_tte, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.081
eGFR Low and UP High	Sparsentan	71	3 (4.2)	NE		0.226	(0.064, 0.803)	0.021 *
	Irbesartan	74	12 (16.2)	NE				
eGFR Low and UP Low	Sparsentan	55	5 (9.1)	NE		2.642	(0.512, 13.625)	0.246
	Irbesartan	55	2 (3.6)	NE				
eGFR High and UP High	Sparsentan	37	3 (8.1)	NE		0.882	(0.178, 4.375)	0.878
	Irbesartan	36	3 (8.3)	NE				
eGFR High and UP Low	Sparsentan	39	3 (7.7)	NE		0.947	(0.191, 4.699)	0.947
	Irbesartan	37	3 (8.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff\_tte, created on: 28FEB2024



Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.446
< 60 mL/min/1.73 m**2	Sparsentan	127	8 (6.3)	NE		0.498	(0.212, 1.172)	0.110
	Irbesartan	129	16 (12.4)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	3 (6.1)	NE		1.427	(0.238, 8.552)	0.697
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	3 (11.5)	NE		1.095	(0.181, 6.613)	0.921
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff\_tte, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.621
< 45 mL/min/1.73 m**2	Sparsentan	82	4 (4.9)	NE		0.426	(0.131, 1.385)	0.156
	Irbesartan	80	9 (11.3)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	4 (8.9)	NE		0.584	(0.167, 2.046)	0.401
	Irbesartan	49	7 (14.3)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	3 (6.1)	NE		1.427	(0.238, 8.552)	0.697
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	3 (11.5)	NE		1.095	(0.181, 6.613)	0.921
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff\_tte, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.059
<= 1.75 g/day	Sparsentan	98	9 (9.2)	NE		1.348	(0.479, 3.797)	0.572
	Irbesartan	94	6 (6.4)	NE				
> 1.75 g/day	Sparsentan	104	5 (4.8)	NE		0.353	(0.127, 0.984)	0.047 *
	Irbesartan	108	14 (13.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff\_tte, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.350
Yes	Sparsentan	88	4 (4.5)	NE		0.465	(0.142, 1.520)	0.205
	Irbesartan	83	9 (10.8)	NE				
No	Sparsentan	114	10 (8.8)	NE		0.877	(0.372, 2.068)	0.764
	Irbesartan	119	11 (9.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment  
 was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff\_tte, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.005 #
<= 5 years	Sparsentan	113	5 (4.4)	NE		0.276	(0.102, 0.751)	0.012 *
	Irbesartan	127	17 (13.4)	NE				
> 5 years	Sparsentan	89	9 (10.1)	NE		2.656	(0.718, 9.825)	0.143
	Irbesartan	75	3 (4.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff\_tte, created on: 28FEB2024

Table PT1MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.533
Yes	Sparsentan	153	11 (7.2)	NE		0.625	(0.295, 1.325)	0.220
	Irbesartan	157	18 (11.5)	NE				
No	Sparsentan	49	3 (6.1)	NE		1.241	(0.196, 7.864)	0.819
	Irbesartan	45	2 (4.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

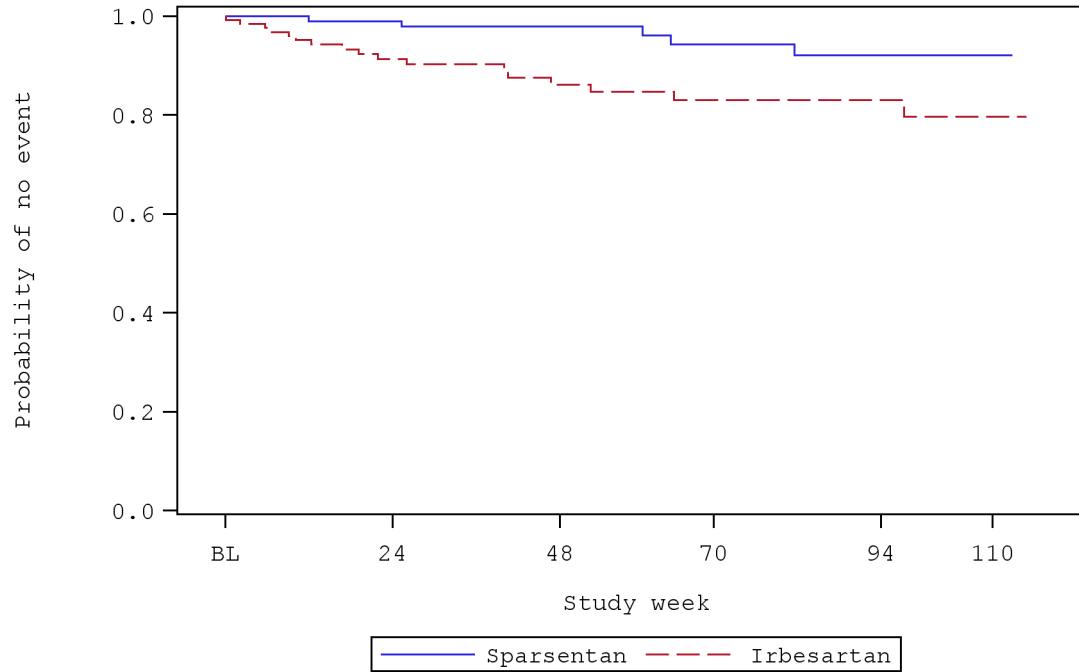
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff\_tte, created on: 28FEB2024

Figure PF1MIST\_FSKM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

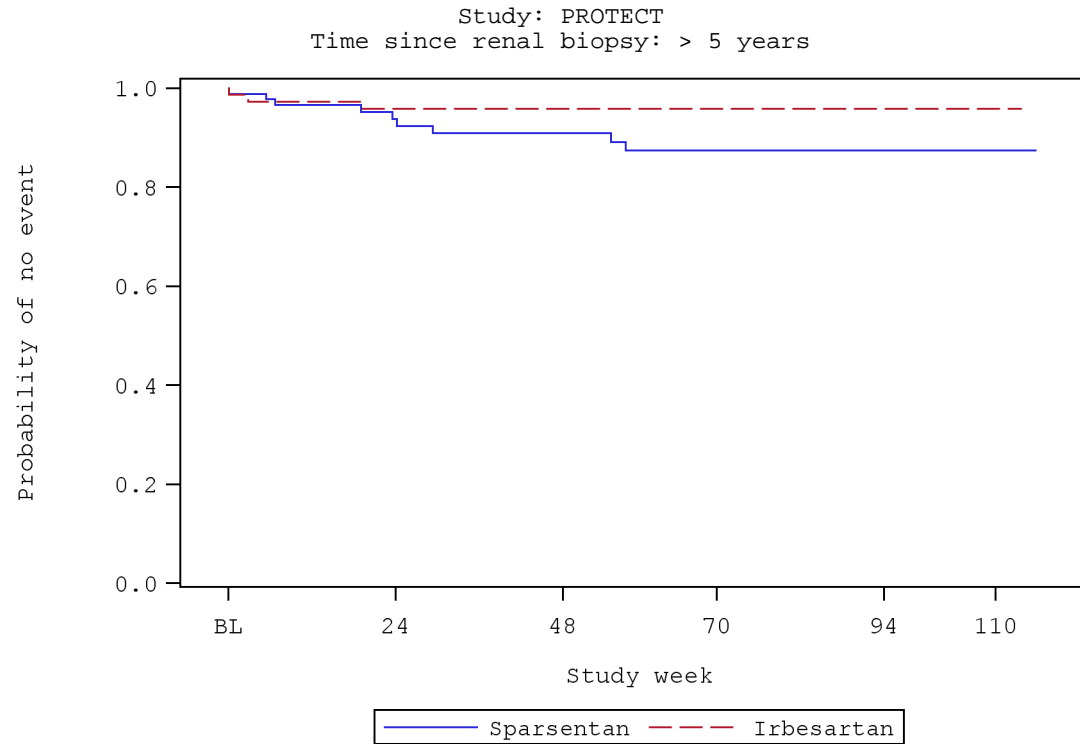
Study: PROTECT  
 Time since renal biopsy: <= 5 years



Sparsentan	113	90	66	52	28	7
Irbesartan	127	88	61	48	28	10

Reference table: PT1MIST\_FSTM

Figure PF1MIST\_FSKM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set



Sparsentan	89	66	55	45	22	9
Irbesartan	75	61	49	40	18	7

Reference table: PT1MIST\_FSTM



Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: NE
Male	Double-blind period	Sparsentan	139	1 (0.7)				NE
		Irbesartan	143	8 (5.6)				
Female	Double-blind period	Sparsentan	63	1 (1.6)				NE
		Irbesartan	59	1 (1.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: NE
<= 45 years	Double-blind period	Sparsentan	96	2 (2.1) all n<10				NE
		Irbesartan	99	5 (5.1)				
> 45 years	Double-blind period	Sparsentan	106	0 (0.0) all n<10				NE
		Irbesartan	103	4 (3.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: NE
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2) all n<10				NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	0 (0.0) all n<10				NE
		Irbesartan	109	5 (4.6)				
> 40 years	Double-blind period	Sparsentan	91	0 (0.0) all n<10				NE
		Irbesartan	88	4 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 28FEB2024

Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: NE
North America	Double-blind period	Sparsentan	35	2 (5.7) all n<10				NE
		Irbesartan	46	3 (6.5)				
Europe	Double-blind period	Sparsentan	98	0 (0.0) all n<10				NE
		Irbesartan	115	3 (2.6)				
Asia Pacific	Double-blind period	Sparsentan	69	0 (0.0) all n<10				NE
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: NE
< 27 kg/m**2	Double-blind period	Sparsentan	84	2 (2.4) all n<10				NE
		Irbesartan	94	5 (5.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	0 (0.0) all n<10				NE
		Irbesartan	107	4 (3.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	0 (0.0) all n<10				NE
		Irbesartan	74	6 (8.1)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	0 (0.0) all n<10				NE
		Irbesartan	55	1 (1.8)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4) all n<10				NE
		Irbesartan	36	2 (5.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0) all n<10				NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: NE
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	0 (0.0) all n<10				NE
		Irbesartan	129	7 (5.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	1 (2.0) all n<10				NE
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8) all n<10				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	0 (0.0) all n<10				NE
		Irbesartan	80	5 (6.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	0 (0.0) all n<10				NE
		Irbesartan	49	2 (4.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	1 (2.0) all n<10				NE
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8) all n<10				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024



Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication  
 (renal indication)

Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.505
<= 1.75 g/day	Double-blind period	Sparsentan	98	0 (0.0)	0.960 + [0.019, 47.874]	0.959 + [0.019, 48.843]	0.0 [NE, NE]	NE
		Irbesartan	94	0 (0.0)				
> 1.75 g/day	Double-blind period	Sparsentan	104	2 (1.9)	0.231 [0.051, 1.043]	0.216 [0.045, 1.023]	-6.4 [-13.2, 0.4]	0.059
		Irbesartan	108	9 (8.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 28FEB2024

Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: NE
Yes	Double-blind period	Sparsentan	88	0 (0.0) all n<10				NE
		Irbesartan	83	4 (4.8)				
No	Double-blind period	Sparsentan	114	2 (1.8) all n<10				NE
		Irbesartan	119	5 (4.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: NE
<= 5 years	Double-blind period	Sparsentan	113	0 (0.0) all n<10				NE
		Irbesartan	127	7 (5.5)				
> 5 years	Double-blind period	Sparsentan	89	2 (2.2) all n<10				NE
		Irbesartan	75	2 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: NE
Yes	Double-blind period	Sparsentan	153	1 (0.7) all n<10				NE
		Irbesartan	157	8 (5.1)				
No	Double-blind period	Sparsentan	49	1 (2.0) all n<10				NE
		Irbesartan	45	1 (2.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Not done. Less than 10 patients with events in every subgroup level.

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeфф\_tte, created on: 28FEB2024

Figure PF1MIKT\_FSKM: Time to systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Not done. No significant subgroups.

Reference table: PT1MIKT\_FSTM

Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.621
Male	Double-blind period	Sparsentan	139	15 (10.8)	1.102 [0.553, 2.197]	1.115 [0.517, 2.405]	1.0 [-6.8, 8.8]	0.846
		Irbesartan	143	14 (9.8)				
Female	Double-blind period	Sparsentan	63	7 (11.1)	0.819 [0.317, 2.119]	0.797 [0.270, 2.354]	-2.4 [-15.8, 10.9]	0.785
		Irbesartan	59	8 (13.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.672
<= 45 years	Double-blind period	Sparsentan	96	11 (11.5)	1.134 [0.505, 2.548]	1.152 [0.465, 2.851]	1.4 [-8.4, 11.1]	0.820
		Irbesartan	99	10 (10.1)				
> 45 years	Double-blind period	Sparsentan	106	11 (10.4)	0.891 [0.412, 1.927]	0.878 [0.369, 2.090]	-1.3 [-10.7, 8.2]	0.827
		Irbesartan	103	12 (11.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024



Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.394
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	3.000 + [0.171, 52.527]	3.667 + [0.145, 92.653]	22.2 [-20.5, 64.9]	0.505
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	10 (9.8)	1.336 [0.549, 3.252]	1.372 [0.519, 3.626]	2.5 [-6.1, 11.0]	0.624
		Irbesartan	109	8 (7.3)				
> 40 years	Double-blind period	Sparsentan	91	10 (11.0)	0.691 [0.324, 1.472]	0.653 [0.273, 1.558]	-4.9 [-16.0, 6.2]	0.384
		Irbesartan	88	14 (15.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.987
North America	Double-blind period	Sparsentan	35	4 (11.4)	0.876 [0.268, 2.870]	0.860 [0.223, 3.316]	-1.6 [-18.5, 15.2]	1.000
		Irbesartan	46	6 (13.0)				
Europe	Double-blind period	Sparsentan	98	8 (8.2)	0.939 [0.386, 2.286]	0.933 [0.353, 2.466]	-0.5 [-9.0, 7.9]	1.000
		Irbesartan	115	10 (8.7)				
Asia Pacific	Double-blind period	Sparsentan	69	10 (14.5)	0.990 [0.389, 2.524]	0.989 [0.331, 2.956]	-0.1 [-15.7, 15.4]	1.000
		Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.961
< 27 kg/m**2	Double-blind period	Sparsentan	84	10 (11.9)	1.017 [0.455, 2.274]	1.020 [0.410, 2.538]	0.2 [-10.4, 10.8]	1.000
		Irbesartan	94	11 (11.7)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	12 (10.2)	0.989 [0.456, 2.148]	0.988 [0.417, 2.343]	-0.1 [-8.9, 8.7]	1.000
		Irbesartan	107	11 (10.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

Hospitalizations								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.280
eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	0.695 [0.302, 1.599]	0.656 [0.251, 1.715]	-4.9 [-17.5, 7.6]	0.473
		Irbesartan	74	12 (16.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	8 (14.5)	1.600 [0.558, 4.586]	1.702 [0.520, 5.574]	5.5 [-8.4, 19.3]	0.556
		Irbesartan	55	5 (9.1)				
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	3.892 [0.457, 33.169]	4.242 [0.451, 39.943]	8.0 [-6.1, 22.1]	0.358
		Irbesartan	36	1 (2.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)	0.474 [0.092, 2.437]	0.446 [0.077, 2.595]	-5.7 [-20.5, 9.1]	0.425
		Irbesartan	37	4 (10.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.541
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	16 (12.6)	0.855 [0.461, 1.587]	0.835 [0.408, 1.707]	-2.1 [-11.3, 7.1]	0.717
		Irbesartan	129	19 (14.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	1.959 [0.519, 7.389]	2.093 [0.492, 8.901]	6.0 [-7.5, 19.5]	0.487
		Irbesartan	48	3 (6.3)				
>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 28FEB2024

Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

Hospitalizations								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.679
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	11 (13.4)	0.976 [0.449, 2.122]	0.972 [0.396, 2.388]	-0.3 [-12.1, 11.5]	1.000
		Irbesartan	80	11 (13.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	5 (11.1)	0.681 [0.240, 1.928]	0.641 [0.193, 2.125]	-5.2 [-21.2, 10.8]	0.557
		Irbesartan	49	8 (16.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	1.959 [0.519, 7.389]	2.093 [0.492, 8.901]	6.0 [-7.5, 19.5]	0.487
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.149
<= 1.75 g/day	Double-blind period	Sparsentan	98	13 (13.3)	1.559 [0.677, 3.589]	1.644 [0.648, 4.168]	4.8 [-5.1, 14.6]	0.358
		Irbesartan	94	8 (8.5)				
> 1.75 g/day	Double-blind period	Sparsentan	104	9 (8.7)	0.668 [0.302, 1.475]	0.636 [0.263, 1.541]	-4.3 [-13.6, 5.0]	0.380
		Irbesartan	108	14 (13.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.618
Yes	Double-blind period	Sparsentan	88	9 (10.2)	0.849 [0.363, 1.984]	0.832 [0.320, 2.161]	-1.8 [-12.4, 8.8]	0.809
		Irbesartan	83	10 (12.0)				
No	Double-blind period	Sparsentan	114	13 (11.4)	1.131 [0.539, 2.373]	1.148 [0.500, 2.633]	1.3 [-7.5, 10.1]	0.833
		Irbesartan	119	12 (10.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024



Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.019 #
<= 5 years	Double-blind period	Sparsentan	113	9 (8.0)	0.562 [0.263, 1.200]	0.524 [0.225, 1.219]	-6.2 [-14.9, 2.5]	0.154
		Irbesartan	127	18 (14.2)				
> 5 years	Double-blind period	Sparsentan	89	13 (14.6)	2.739 [0.932, 8.046]	3.036 [0.946, 9.748]	9.3 [-0.9, 19.4]	0.071
		Irbesartan	75	4 (5.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.882
Yes	Double-blind period	Sparsentan	153	18 (11.8)	1.026 [0.555, 1.897]	1.030 [0.514, 2.063]	0.3 [-7.5, 8.1]	1.000
		Irbesartan	157	18 (11.5)				
No	Double-blind period	Sparsentan	49	4 (8.2)	0.918 [0.244, 3.457]	0.911 [0.214, 3.881]	-0.7 [-14.2, 12.7]	1.000
		Irbesartan	45	4 (8.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1HNS\_FSNM: Number of hospitalizations by subgroup  
Full Analysis Set

Number of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Sex	Sparsentan								NE
Male	Sparsentan	139	139 (100.0)	175.4	17	0.10	1.155	(0.564, 2.366)	0.693
	Irbesartan	143	143 (100.0)	194.0	16	0.08			
Female	Sparsentan	63	63 (100.0)	90.4	7	0.08	0.617	(0.229, 1.663)	0.340
	Irbesartan	59	59 (100.0)	68.9	10	0.15			
Age	Sparsentan								0.764
<= 45 years	Sparsentan	96	96 (100.0)	121.4	11	0.09	1.104	(0.486, 2.505)	0.813
	Irbesartan	99	99 (100.0)	129.7	12	0.09			
> 45 years	Sparsentan	106	106 (100.0)	144.4	13	0.09	0.917	(0.410, 2.051)	0.833
	Irbesartan	103	103 (100.0)	133.2	14	0.11			
Age at IgAN diagnosis	Sparsentan								NE
<= 18 years	Sparsentan	9	9 (100.0)	9.9	2	0.20			NE
	Irbesartan	5	5 (100.0)	3.5	0	0.00			
> 18 to 40 years	Sparsentan	102	102 (100.0)	133.1	11	0.08	1.342	(0.569, 3.168)	0.501
	Irbesartan	109	109 (100.0)	142.4	10	0.07			
> 40 years	Sparsentan	91	91 (100.0)	122.8	11	0.09	0.666	(0.305, 1.455)	0.308
	Irbesartan	88	88 (100.0)	117.0	16	0.14			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.

Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.

\* = significant treatment effect. # = significant interaction.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1HNS\_FSNM: Number of hospitalizations by subgroup  
Full Analysis Set

Number of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Geographic region	Sparsentan								NE
	North America	35	35 (100.0)	38.8	4	0.10	0.693	(0.205, 2.339)	0.554
Europe	Sparsentan	46	46 (100.0)	54.5	8	0.15	0.956	(0.384, 2.378)	0.923
	Irbesartan	98	98 (100.0)	116.1	8	0.07			
Asia Pacific	Sparsentan	115	115 (100.0)	152.1	11	0.07	0.797	(0.301, 2.108)	0.648
	Irbesartan	69	69 (100.0)	111.0	12	0.11			
Baseline BMI < 27 kg/m**2	Sparsentan	41	41 (100.0)	56.3	7	0.12	0.997	(0.446, 2.229)	0.907
	Sparsentan	84	84 (100.0)	109.3	11	0.10			0.994
>= 27 kg/m**2	Sparsentan	94	94 (100.0)	118.5	13	0.11	0.908	(0.411, 2.010)	0.813
	Irbesartan	118	118 (100.0)	156.5	13	0.08			
Randomization strata eGFR Low and UP High	Sparsentan	107	107 (100.0)	142.4	13	0.09	0.659	(0.284, 1.528)	NE
	Sparsentan	71	71 (100.0)	96.1	9	0.09			0.331
eGFR Low and UP Low	Irbesartan	74	74 (100.0)	98.4	14	0.14	1.794	(0.601, 5.354)	0.295
	Sparsentan	55	55 (100.0)	72.9	9	0.12			
eGFR High and UP High	Sparsentan	55	55 (100.0)	72.7	5	0.07	3.917	(0.438, 35.048)	0.222
	Irbesartan	37	37 (100.0)	47.0	4	0.09			
eGFR High and UP Low	Sparsentan	36	36 (100.0)	46.0	1	0.02	0.329	(0.054, 1.997)	0.227
	Irbesartan	39	39 (100.0)	49.8	2	0.04			
	Irbesartan	37	37 (100.0)	45.7	6	0.13			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 28FEB2024

Table PT1HNS\_FSNM: Number of hospitalizations by subgroup  
Full Analysis Set

Number of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline eGFR Group 1 < 60 mL/min/1.73 m**2	Sparsentan								NE
	Sparsentan	127	127 (100.0)	171.0	18	0.11	0.879	(0.461, 1.673)	0.693
	Irbesartan	129	129 (100.0)	174.1	23	0.13			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	59.8	6	0.10	2.081	(0.512, 8.464)	0.306
	Irbesartan	48	48 (100.0)	60.9	3	0.05			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	35.0	0	0.00			NE
	Irbesartan	25	25 (100.0)	27.9	0	0.00			
Baseline eGFR Group 2 < 45 mL/min/1.73 m**2	Sparsentan								NE
	Sparsentan	82	82 (100.0)	110.0	13	0.12	1.016	(0.456, 2.264)	0.969
	Irbesartan	80	80 (100.0)	110.6	13	0.12			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	45 (100.0)	61.0	5	0.08	0.653	(0.215, 1.980)	0.452
	Irbesartan	49	49 (100.0)	63.4	10	0.16			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	59.8	6	0.10	2.081	(0.512, 8.464)	0.306
	Irbesartan	48	48 (100.0)	60.9	3	0.05			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	35.0	0	0.00			NE
	Irbesartan	25	25 (100.0)	27.9	0	0.00			
Baseline urine protein excretion <= 1.75 g/day	Sparsentan								0.158
	Sparsentan	98	98 (100.0)	137.3	14	0.10	1.433	(0.600, 3.421)	0.418
	Irbesartan	94	94 (100.0)	114.9	8	0.07			
	Sparsentan	104	104 (100.0)	128.5	10	0.08	0.608	(0.280, 1.320)	0.208
> 1.75 g/day	Sparsentan	104	104 (100.0)	128.5	10	0.08			
	Irbesartan	108	108 (100.0)	148.0	18	0.12			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 28FEB2024

Table PT1HNS\_FSNM: Number of hospitalizations by subgroup  
 Full Analysis Set

Number of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan								0.757
Yes	Sparsentan	88	88 (100.0)	110.2	10	0.09	0.943	(0.385, 2.309)	0.899
	Irbesartan	83	83 (100.0)	115.1	12	0.10			
No	Sparsentan	114	114 (100.0)	155.6	14	0.09	1.056	(0.502, 2.221)	0.886
	Irbesartan	119	119 (100.0)	147.8	14	0.09			
Time since renal biopsy	Sparsentan								0.017 #
<= 5 years	Sparsentan	113	113 (100.0)	144.6	10	0.07	0.535	(0.246, 1.162)	0.114
	Irbesartan	127	127 (100.0)	162.0	21	0.13			
> 5 years	Sparsentan	89	89 (100.0)	121.2	14	0.12	2.429	(0.870, 6.780)	0.090
	Irbesartan	75	75 (100.0)	100.8	5	0.05			
History of hypertension	Sparsentan								NE
Yes	Sparsentan	153	153 (100.0)	199.2	19	0.10	0.974	(0.519, 1.827)	0.934
	Irbesartan	157	157 (100.0)	207.9	20	0.10			
No	Sparsentan	49	49 (100.0)	66.6	5	0.08	0.753	(0.226, 2.503)	0.643
	Irbesartan	45	45 (100.0)	55.0	6	0.11			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
 NE = not evaluable.  
 Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
 A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
 \* = significant treatment effect. # = significant interaction.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024

Table PT1HDS\_FSNM: Duration of hospitalizations by subgroup  
Full Analysis Set

Duration of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Sex	Sparsentan								0.990
Male	Sparsentan	139	139 (100.0)	175.4	75	0.43	0.812	(0.237, 2.777)	0.740
	Irbesartan	143	143 (100.0)	194.0	134	0.69			
Female	Sparsentan	63	63 (100.0)	90.4	30	0.33	0.975	(0.200, 4.763)	0.975
	Irbesartan	59	59 (100.0)	68.9	44	0.64			
Age	Sparsentan								0.849
<= 45 years	Sparsentan	96	96 (100.0)	121.4	46	0.38	1.400	(0.321, 6.096)	0.654
	Irbesartan	99	99 (100.0)	129.7	76	0.59			
> 45 years	Sparsentan	106	106 (100.0)	144.4	59	0.41	0.840	(0.227, 3.104)	0.794
	Irbesartan	103	103 (100.0)	133.2	102	0.77			
Age at IgAN diagnosis	Sparsentan								NE
<= 18 years	Sparsentan	9	9 (100.0)	9.9	2	0.20			NE
	Irbesartan	5	5 (100.0)	3.5	0	0.00			
> 18 to 40 years	Sparsentan	102	102 (100.0)	133.1	53	0.40	1.369	(0.304, 6.162)	0.683
	Irbesartan	109	109 (100.0)	142.4	54	0.38			
> 40 years	Sparsentan	91	91 (100.0)	122.8	50	0.41	0.501	(0.137, 1.836)	0.297
	Irbesartan	88	88 (100.0)	117.0	124	1.06			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.

Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.

\* = significant treatment effect. # = significant interaction.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 28FEB2024

Table PT1HDS\_FSNM: Duration of hospitalizations by subgroup  
 Full Analysis Set

Duration of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Geographic region	Sparsentan								0.954
	North America								
Europe	Sparsentan	35	35 (100.0)	38.8	15	0.39	0.840	(0.126, 5.583)	0.857
	Irbesartan	46	46 (100.0)	54.5	38	0.70			
Asia Pacific	Sparsentan	98	98 (100.0)	116.1	41	0.35	0.684	(0.083, 5.624)	0.724
	Irbesartan	115	115 (100.0)	152.1	94	0.62			
Baseline BMI	Sparsentan	69	69 (100.0)	111.0	49	0.44	0.837	(0.176, 3.983)	0.824
	Irbesartan	41	41 (100.0)	56.3	46	0.82			
< 27 kg/m**2	Sparsentan								0.529
	Sparsentan	84	84 (100.0)	109.3	48	0.44	1.405	(0.395, 4.999)	0.600
>= 27 kg/m**2	Irbesartan	94	94 (100.0)	118.5	68	0.57			
	Sparsentan	118	118 (100.0)	156.5	57	0.36	0.438	(0.108, 1.773)	0.247
Randomization strata	Irbesartan	107	107 (100.0)	142.4	110	0.77			
	eGFR Low and UP High								0.219
eGFR Low and UP Low	Sparsentan	71	71 (100.0)	96.1	32	0.33	0.401	(0.109, 1.481)	0.170
	Irbesartan	74	74 (100.0)	98.4	101	1.03			
eGFR High and UP High	Sparsentan	55	55 (100.0)	72.9	53	0.73	2.204	(0.447, 10.863)	0.331
	Irbesartan	55	55 (100.0)	72.7	24	0.33			
eGFR High and UP Low	Sparsentan	37	37 (100.0)	47.0	9	0.19	2.824	(0.364, 21.921)	0.321
	Irbesartan	36	36 (100.0)	46.0	3	0.07			
	Sparsentan	39	39 (100.0)	49.8	11	0.22	0.277	(0.021, 3.729)	0.333
	Irbesartan	37	37 (100.0)	45.7	50	1.09			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
 NE = not evaluable.  
 Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
 A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
 \* = significant treatment effect. # = significant interaction.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 28FEB2024



Table PT1HDS\_FSNM: Duration of hospitalizations by subgroup  
Full Analysis Set

Duration of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline eGFR Group 1 < 60 mL/min/1.73 m**2	Sparsentan								NE
	Sparsentan	127	127 (100.0)	171.0	85	0.50	0.838	(0.287, 2.450)	0.747
	Irbesartan	129	129 (100.0)	174.1	157	0.90			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	59.8	20	0.33	3.023	(0.470, 19.456)	0.244
	Irbesartan	48	48 (100.0)	60.9	21	0.34			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	35.0	0	0.00			NE
	Irbesartan	25	25 (100.0)	27.9	0	0.00			
Baseline eGFR Group 2 < 45 mL/min/1.73 m**2	Sparsentan								NE
	Sparsentan	82	82 (100.0)	110.0	68	0.62	0.918	(0.199, 4.227)	0.912
	Irbesartan	80	80 (100.0)	110.6	76	0.69			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	45 (100.0)	61.0	17	0.28	0.574	(0.110, 2.999)	0.510
	Irbesartan	49	49 (100.0)	63.4	81	1.28			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	59.8	20	0.33	3.023	(0.470, 19.456)	0.244
	Irbesartan	48	48 (100.0)	60.9	21	0.34			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	35.0	0	0.00			NE
	Irbesartan	25	25 (100.0)	27.9	0	0.00			
Baseline urine protein excretion ≤ 1.75 g/day	Sparsentan								0.339
	Sparsentan	98	98 (100.0)	137.3	68	0.50	1.076	(0.267, 4.340)	0.918
	Irbesartan	94	94 (100.0)	114.9	59	0.51			
	Irbesartan	94	94 (100.0)	114.9	59	0.51			
> 1.75 g/day	Sparsentan	104	104 (100.0)	128.5	37	0.29	0.512	(0.143, 1.825)	0.302
	Irbesartan	108	108 (100.0)	148.0	119	0.80			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 28FEB2024

Table PT1HDS\_FSNM: Duration of hospitalizations by subgroup  
Full Analysis Set

Duration of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan								0.488
Yes	Sparsentan	88	88 (100.0)	110.2	40	0.36	0.844	(0.217, 3.286)	0.807
	Irbesartan	83	83 (100.0)	115.1	85	0.74			
No	Sparsentan	114	114 (100.0)	155.6	65	0.42	1.978	(0.496, 7.889)	0.334
	Irbesartan	119	119 (100.0)	147.8	93	0.63			
Time since renal biopsy	Sparsentan								0.320
<= 5 years	Sparsentan	113	113 (100.0)	144.6	36	0.25	0.486	(0.157, 1.505)	0.211
	Irbesartan	127	127 (100.0)	162.0	117	0.72			
> 5 years	Sparsentan	89	89 (100.0)	121.2	69	0.57	2.375	(0.338, 16.667)	0.384
	Irbesartan	75	75 (100.0)	100.8	61	0.61			
History of hypertension	Sparsentan								0.910
Yes	Sparsentan	153	153 (100.0)	199.2	84	0.42	0.765	(0.255, 2.294)	0.632
	Irbesartan	157	157 (100.0)	207.9	150	0.72			
No	Sparsentan	49	49 (100.0)	66.6	21	0.32	1.258	(0.091, 17.336)	0.864
	Irbesartan	45	45 (100.0)	55.0	28	0.51			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 28FEB2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Sex												
Male	EQ-5D VAS	Baseline	Sparsentan	139	132 (95.0)	81.11 (11.21)	40.0	74.50	81.00	90.00	100.0	
			Irbesartan	143	131 (91.6)	79.58 (14.73)	19.0	75.00	80.00	90.00	100.0	
		Week 24	Sparsentan	139	91 (65.5)	83.32 (10.98)	50.0	76.00	84.00	91.00	100.0	
			Irbesartan	143	83 (58.0)	80.06 (18.01)	4.0	73.00	82.00	91.00	100.0	
		Week 48	Sparsentan	139	75 (54.0)	82.76 (9.78)	49.0	80.00	83.00	90.00	100.0	
			Irbesartan	143	61 (42.7)	83.44 (12.33)	40.0	79.00	85.00	91.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	139	91 (65.5)	2.46 (11.91)	-31.0	-4.00	2.00	8.00	44.0	0.12 [-0.18, 0.41]
			Irbesartan	143	83 (58.0)	0.66 (18.83)	-90.0	-6.00	0.00	11.00	52.0	
		Week 48	Sparsentan	139	75 (54.0)	2.61 (10.45)	-30.0	-2.00	1.00	7.00	41.0	-0.07 [-0.41, 0.27]
			Irbesartan	143	61 (42.7)	3.36 (10.16)	-35.0	-1.00	3.00	9.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Female	EQ-5D VAS	Baseline	Sparsentan	63	59 (93.7)	79.07 (19.44)	7.0	75.00	82.00	90.00	100.0	
			Irbesartan	59	53 (89.8)	82.30 (12.78)	37.0	75.00	83.00	91.00	100.0	
		Week 24	Sparsentan	63	41 (65.1)	82.95 (10.66)	51.0	78.00	83.00	90.00	100.0	
			Irbesartan	59	32 (54.2)	83.19 (10.21)	48.0	80.00	81.00	90.00	99.0	
		Week 48	Sparsentan	63	38 (60.3)	80.82 (12.07)	49.0	74.00	80.00	90.00	100.0	
			Irbesartan	59	22 (37.3)	80.91 (13.86)	50.0	80.00	82.00	90.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	63	41 (65.1)	4.85 (20.39)	-25.0	-9.00	0.00	12.00	75.0	0.26 [-0.20, 0.73]
			Irbesartan	59	32 (54.2)	0.34 (12.17)	-37.0	-3.50	0.50	7.00	23.0	
		Week 48	Sparsentan	63	38 (60.3)	3.29 (22.68)	-29.0	-11.00	0.00	10.00	87.0	0.25 [-0.28, 0.77]
			Irbesartan	59	22 (37.3)	-1.50 (11.66)	-31.0	-8.00	0.50	5.00	23.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age												
<= 45 years	EQ-5D VAS	Baseline	Sparsentan	96	89 (92.7)	79.93 (14.66)	7.0	75.00	81.00	90.00	100.0	
			Irbesartan	99	92 (92.9)	81.66 (14.94)	19.0	75.00	82.00	90.50	100.0	
	Week 24	Sparsentan	96	62 (64.6)	84.79 (10.90)	59.0	78.00	87.50	91.00	100.0		
		Irbesartan	99	58 (58.6)	79.76 (18.45)	10.0	71.00	83.50	91.00	100.0		
	Week 48	Sparsentan	96	54 (56.3)	82.39 (11.84)	49.0	79.00	82.50	91.00	100.0		
		Irbesartan	99	41 (41.4)	82.44 (14.21)	40.0	80.00	83.00	90.00	100.0		
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	96	62 (64.6)	4.77 (16.93)	-31.0	-4.00	1.50	13.00	75.0	0.41 [0.04, 0.77]
			Irbesartan	99	58 (58.6)	-2.21 (17.48)	-90.0	-9.00	-1.00	9.00	23.0	
		Week 48	Sparsentan	96	54 (56.3)	2.15 (15.73)	-30.0	-6.00	1.00	10.00	64.0	0.19 [-0.22, 0.59]
			Irbesartan	99	41 (41.4)	-0.49 (11.86)	-35.0	-5.00	0.00	5.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 45 years	EQ-5D VAS	Baseline	Sparsentan	106	102 (96.2)	80.96 (13.92)	11.0	75.00	82.00	90.00	100.0	
			Irbesartan	103	92 (89.3)	79.07 (13.40)	34.0	74.00	80.00	89.00	100.0	
		Week 24	Sparsentan	106	70 (66.0)	81.80 (10.67)	50.0	75.00	80.50	90.00	100.0	
			Irbesartan	103	57 (55.3)	82.12 (13.69)	4.0	79.00	81.00	90.00	100.0	
		Week 48	Sparsentan	106	59 (55.7)	81.85 (9.41)	49.0	79.00	81.00	89.00	99.0	
			Irbesartan	103	42 (40.8)	83.10 (11.24)	52.0	80.00	82.50	90.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	106	70 (66.0)	1.81 (13.07)	-29.0	-5.00	0.50	7.00	44.0	-0.11 [-0.46, 0.24]
			Irbesartan	103	57 (55.3)	3.40 (16.56)	-71.0	-1.00	1.00	10.00	52.0	
		Week 48	Sparsentan	106	59 (55.7)	3.47 (15.51)	-20.0	-3.00	0.00	7.00	87.0	-0.08 [-0.48, 0.31]
			Irbesartan	103	42 (40.8)	4.57 (8.93)	-19.0	0.00	5.00	9.00	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age at IgAN diagnosis												
<= 18 years	EQ-5D VAS	Baseline	Sparsentan	9	9 (100.0)	74.67 (15.03)	51.0	62.00	79.00	82.00	95.0	
			Irbesartan	5	5 (100.0)	76.40 (11.59)	60.0	73.00	74.00	86.00	89.0	
	Week 24	Sparsentan	9	3 (33.3)	86.67 (23.09)	60.0	60.00	100.00	100.00	100.0		
		Irbesartan	5	2 (40.0)	50.00 (26.87)	31.0	31.00	50.00	69.00	69.0		
	Week 48	Sparsentan	9	5 (55.6)	83.00 (14.25)	66.0	71.00	86.00	92.00	100.0		
		Irbesartan	5	1 (20.0)	66.00	66.0	66.00	66.00	66.00	66.0		
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	9	3 (33.3)	9.33 (10.69)	0.0	0.00	7.00	21.00	21.0	1.14 [-0.78, 3.07]
			Irbesartan	5	2 (40.0)	-16.50 (36.06)	-42.0	-42.00	-16.50	9.00	9.0	
		Week 48	Sparsentan	9	5 (55.6)	6.80 (9.28)	-9.0	7.00	10.00	11.00	15.0	NE
			Irbesartan	5	1 (20.0)	6.00	6.0	6.00	6.00	6.00	6.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 18 to 40 years	EQ-5D VAS	Baseline	Sparsentan	102	95 (93.1)	80.85 (14.44)	7.0	75.00	82.00	90.00	100.0	
			Irbesartan	109	100 (91.7)	81.14 (15.01)	19.0	75.00	82.00	90.00	100.0	
		Week 24	Sparsentan	102	66 (64.7)	84.33 (10.13)	59.0	78.00	85.50	91.00	100.0	
			Irbesartan	109	63 (57.8)	81.75 (16.66)	10.0	78.00	85.00	91.00	100.0	
		Week 48	Sparsentan	102	58 (56.9)	82.16 (10.83)	49.0	79.00	82.00	90.00	100.0	
			Irbesartan	109	45 (41.3)	82.29 (14.05)	40.0	80.00	84.00	90.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	102	66 (64.7)	4.62 (17.29)	-31.0	-4.00	1.50	12.00	75.0	0.28 [-0.06, 0.63]
			Irbesartan	109	63 (57.8)	-0.22 (16.72)	-90.0	-7.00	-1.00	11.00	23.0	
		Week 48	Sparsentan	102	58 (56.9)	2.07 (16.14)	-30.0	-6.00	1.00	9.00	64.0	0.12 [-0.27, 0.51]
			Irbesartan	109	45 (41.3)	0.38 (12.09)	-35.0	-5.00	0.00	7.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024



Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	EQ-5D VAS	Baseline	Sparsentan	91	87 (95.6)	80.68 (13.98)	11.0	72.00	81.00	90.00	100.0	
			Irbesartan	88	79 (89.8)	79.63 (13.37)	40.0	73.00	80.00	90.00	100.0	
		Week 24	Sparsentan	91	63 (69.2)	81.86 (10.95)	50.0	75.00	80.00	90.00	100.0	
			Irbesartan	88	50 (56.8)	81.14 (14.40)	4.0	79.00	80.00	90.00	100.0	
		Week 48	Sparsentan	91	50 (54.9)	81.96 (10.17)	49.0	79.00	81.00	90.00	100.0	
			Irbesartan	88	37 (42.0)	83.81 (10.86)	56.0	80.00	82.00	90.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	91	63 (69.2)	1.43 (12.35)	-29.0	-6.00	0.00	7.00	40.0	-0.06 [-0.43, 0.31]
			Irbesartan	88	50 (56.8)	2.26 (17.07)	-71.0	-1.00	0.50	10.00	52.0	
		Week 48	Sparsentan	91	50 (54.9)	3.34 (15.52)	-20.0	-3.00	0.00	5.00	87.0	-0.05 [-0.48, 0.37]
			Irbesartan	88	37 (42.0)	4.03 (8.69)	-19.0	0.00	4.00	9.00	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Geographic region													
North America	EQ-5D VAS	Baseline	Sparsentan	35	33 (94.3)	79.61 (18.57)	7.0	75.00	85.00	93.00	100.0		
			Irbesartan	46	44 (95.7)	81.18 (15.14)	34.0	75.00	82.50	92.50	100.0		
	Week 24	Sparsentan	35	15 (42.9)	85.73 (10.79)	70.0	75.00	85.00	95.00	100.0			
		Irbesartan	46	28 (60.9)	78.96 (23.68)	4.0	73.50	86.00	93.50	100.0			
	Week 48	Sparsentan	35	13 (37.1)	80.23 (13.59)	49.0	71.00	80.00	90.00	99.0			
		Irbesartan	46	18 (39.1)	83.33 (12.37)	56.0	75.00	82.00	96.00	100.0			
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	35	15 (42.9)	6.40 (22.37)	-21.0	-2.00	0.00	10.00	75.0	0.39 [-0.24, 1.02]	
				46	28 (60.9)	-3.00 (24.84)	-90.0	-4.00	0.50	9.00	23.0		
			Week 48	Sparsentan	35	13 (37.1)	6.62 (20.44)	-15.0	-4.00	-1.00	15.00	64.0	0.26 [-0.45, 0.98]
				Irbesartan	46	18 (39.1)	2.72 (8.99)	-19.0	0.00	3.00	7.00	24.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Europe	EQ-5D VAS	Baseline	Sparsentan	98	91 (92.9)	80.81 (13.55)	11.0	75.00	81.00	90.00	100.0	
			Irbesartan	115	99 (86.1)	81.11 (13.68)	40.0	75.00	81.00	90.00	100.0	
		Week 24	Sparsentan	98	60 (61.2)	81.92 (11.89)	50.0	75.50	81.00	90.00	100.0	
			Irbesartan	115	52 (45.2)	82.73 (12.81)	31.0	78.50	85.00	91.00	100.0	
		Week 48	Sparsentan	98	45 (45.9)	83.20 (10.61)	50.0	79.00	83.00	91.00	100.0	
			Irbesartan	115	40 (34.8)	84.05 (14.15)	40.0	80.00	86.50	92.50	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	98	60 (61.2)	1.35 (13.57)	-31.0	-5.00	0.50	7.00	40.0	0.03 [-0.34, 0.40]
			Irbesartan	115	52 (45.2)	0.90 (14.23)	-42.0	-7.00	-0.50	10.50	52.0	
		Week 48	Sparsentan	98	45 (45.9)	2.98 (17.14)	-30.0	-2.00	1.00	7.00	87.0	0.05 [-0.37, 0.48]
			Irbesartan	115	40 (34.8)	2.20 (10.94)	-35.0	-1.00	1.00	7.50	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Asia Pacific	EQ-5D VAS	Baseline	Sparsentan	69	67 (97.1)	80.46 (12.87)	40.0	73.00	81.00	90.00	100.0	
			Irbesartan	41	41 (100.0)	77.68 (14.49)	19.0	72.00	80.00	85.00	100.0	
		Week 24	Sparsentan	69	57 (82.6)	83.89 (9.64)	60.0	78.00	85.00	90.00	100.0	
			Irbesartan	41	35 (85.4)	79.83 (13.52)	30.0	72.00	80.00	90.00	100.0	
		Week 48	Sparsentan	69	55 (79.7)	81.65 (9.89)	49.0	80.00	82.00	89.00	100.0	
			Irbesartan	41	25 (61.0)	80.32 (10.48)	54.0	75.00	81.00	89.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	69	57 (82.6)	4.32 (14.18)	-25.0	-3.00	5.00	12.00	44.0	0.10 [-0.32, 0.52]
			Irbesartan	41	35 (85.4)	2.94 (13.42)	-37.0	-2.00	0.00	10.00	34.0	
		Week 48	Sparsentan	69	55 (79.7)	1.84 (12.87)	-29.0	-6.00	0.00	10.00	46.0	0.03 [-0.44, 0.51]
			Irbesartan	41	25 (61.0)	1.40 (11.84)	-31.0	-4.00	5.00	10.00	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline BMI														
< 27 kg/m**2	EQ-5D VAS	Baseline	Sparsentan	84	78 (92.9)	78.91 (14.89)	11.0	72.00	81.00	89.00	100.0			
			Irbesartan	94	86 (91.5)	84.01 (10.64)	52.0	77.00	83.00	91.00	100.0			
		Week 24	Sparsentan	84	54 (64.3)	81.33 (11.56)	51.0	72.00	80.00	90.00	100.0			
			Irbesartan	94	56 (59.6)	82.14 (13.66)	31.0	78.50	81.50	91.00	100.0			
		Week 48	Sparsentan	84	49 (58.3)	79.43 (11.27)	49.0	75.00	80.00	87.00	98.0			
			Irbesartan	94	39 (41.5)	83.67 (13.53)	40.0	80.00	82.00	92.00	100.0			
		Change from baseline in EQ-5D VAS	Week 24	Sparsentan	84	54 (64.3)	1.89 (13.44)	-25.0	-6.00	0.00	7.00	40.0	0.23 [-0.15, 0.60]	
				Irbesartan	94	56 (59.6)	-0.95 (11.50)	-42.0	-5.50	-1.00	7.00	19.0		
				Week 48	Sparsentan	84	49 (58.3)	1.04 (18.60)	-30.0	-8.00	-1.00	4.00	87.0	0.10 [-0.32, 0.52]
					Irbesartan	94	39 (41.5)	-0.49 (11.05)	-35.0	-5.00	0.00	6.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 27 kg/m**2	EQ-5D VAS	Baseline	Sparsentan	118	113 (95.8)	81.57 (13.74)	7.0	77.00	82.00	91.00	100.0	
			Irbesartan	107	97 (90.7)	77.55 (15.74)	19.0	73.00	80.00	88.00	100.0	
		Week 24	Sparsentan	118	78 (66.1)	84.50 (10.20)	50.0	80.00	85.50	91.00	100.0	
			Irbesartan	107	59 (55.1)	79.78 (18.40)	4.0	74.00	81.00	91.00	100.0	
		Week 48	Sparsentan	118	64 (54.2)	84.16 (9.64)	51.0	80.00	84.50	90.50	100.0	
			Irbesartan	107	44 (41.1)	81.98 (12.05)	50.0	77.00	83.50	90.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	118	78 (66.1)	4.12 (16.05)	-31.0	-4.00	3.00	10.00	75.0	0.11 [-0.22, 0.45]
			Irbesartan	107	59 (55.1)	2.02 (21.23)	-90.0	-5.00	1.00	13.00	52.0	
		Week 48	Sparsentan	118	64 (54.2)	4.22 (12.74)	-20.0	-1.00	1.50	10.00	64.0	-0.01 [-0.39, 0.37]
			Irbesartan	107	44 (41.1)	4.34 (10.00)	-20.0	0.00	5.00	10.00	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Randomization strata												
eGFR Low and UP High	EQ-5D VAS	Baseline	Sparsentan	71	66 (93.0)	82.14 (10.50)	51.0	79.00	81.50	90.00	100.0	
			Irbesartan	74	65 (87.8)	78.74 (15.05)	37.0	75.00	80.00	87.00	100.0	
	Week 24	Sparsentan	71	45 (63.4)	83.67 (11.19)	50.0	76.00	85.00	91.00	100.0		
		Irbesartan	74	38 (51.4)	76.74 (18.94)	4.0	70.00	80.00	90.00	100.0		
	Week 48	Sparsentan	71	39 (54.9)	84.67 (8.90)	60.0	80.00	84.00	91.00	100.0		
		Irbesartan	74	25 (33.8)	84.08 (12.35)	54.0	80.00	85.00	90.00	100.0		
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	71	45 (63.4)	0.11 (11.89)	-31.0	-5.00	0.00	7.00	35.0	0.29 [-0.15, 0.72]
			Irbesartan	74	38 (51.4)	-4.55 (20.40)	-71.0	-11.00	-2.00	2.00	52.0	
		Week 48	Sparsentan	71	39 (54.9)	1.69 (8.25)	-13.0	-2.00	0.00	5.00	28.0	0.05 [-0.46, 0.55]
			Irbesartan	74	25 (33.8)	1.24 (11.66)	-31.0	-1.00	2.00	5.00	24.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR Low and UP Low	EQ-5D VAS	Baseline	Sparsentan	55	52 (94.5)	79.10 (17.39)	11.0	71.00	81.00	90.00	100.0		
			Irbesartan	55	51 (92.7)	81.61 (13.00)	34.0	74.00	80.00	90.00	100.0		
		Week 24	Sparsentan	55	40 (72.7)	81.88 (10.79)	51.0	76.50	81.00	90.00	100.0		
			Irbesartan	55	35 (63.6)	82.71 (14.89)	10.0	80.00	83.00	90.00	100.0		
		Week 48	Sparsentan	55	28 (50.9)	81.32 (10.99)	49.0	80.00	81.50	90.00	98.0		
			Irbesartan	55	26 (47.3)	83.00 (13.03)	50.0	80.00	83.00	92.00	100.0		
		Change from baseline in EQ-5D VAS	Week 24	Sparsentan	55	40 (72.7)	4.25 (14.75)	-25.0	-3.50	0.00	10.50	44.0	0.20 [-0.26, 0.65]
			Week 48	Irbesartan	55	35 (63.6)	0.91 (18.70)	-90.0	-2.00	1.00	10.00	23.0	
		Sparsentan		55	28 (50.9)	5.04 (20.42)	-20.0	-4.00	0.50	10.50	87.0	0.18 [-0.35, 0.72]	
		Irbesartan	55	26 (47.3)	2.04 (9.96)	-20.0	-5.00	2.00	9.00	23.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024



Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR High and UP High	EQ-5D VAS	Baseline	Sparsentan	37	35 (94.6)	78.66 (18.53)	7.0	70.00	82.00	91.00	100.0		
			Irbesartan	36	32 (88.9)	81.06 (11.59)	50.0	72.50	80.00	90.00	100.0		
		Week 24	Sparsentan	37	22 (59.5)	81.59 (11.67)	60.0	71.00	81.00	90.00	100.0		
			Irbesartan	36	20 (55.6)	85.90 (10.30)	69.0	76.00	90.00	91.50	99.0		
		Week 48	Sparsentan	37	22 (59.5)	77.50 (10.98)	50.0	71.00	79.00	86.00	90.0		
			Irbesartan	36	13 (36.1)	79.62 (15.16)	40.0	80.00	81.00	90.00	99.0		
	Change from baseline in EQ-5D VAS		Week 24	Sparsentan	37	22 (59.5)	5.86 (21.28)	-25.0	-4.00	5.00	8.00	75.0	-0.14 [-0.75, 0.46]
				Irbesartan	36	20 (55.6)	8.30 (10.34)	-11.0	-1.00	9.00	15.00	34.0	
			Week 48	Sparsentan	37	22 (59.5)	2.36 (21.95)	-30.0	-12.00	-1.00	11.00	64.0	-0.04 [-0.72, 0.65]
				Irbesartan	36	13 (36.1)	3.08 (14.51)	-35.0	0.00	6.00	9.00	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP Low	EQ-5D VAS	Baseline	Sparsentan	39	38 (97.4)	81.18 (10.21)	50.0	75.00	80.50	90.00	100.0			
			Irbesartan	37	36 (97.3)	80.92 (16.54)	19.0	75.00	84.00	90.00	100.0			
		Week 24	Sparsentan	39	25 (64.1)	85.92 (9.52)	70.0	80.00	88.00	91.00	100.0			
			Irbesartan	37	22 (59.5)	80.82 (16.82)	30.0	78.00	80.00	91.00	100.0			
		Week 48	Sparsentan	39	24 (61.5)	83.08 (11.43)	49.0	80.00	84.50	91.00	100.0			
			Irbesartan	37	19 (51.4)	82.89 (11.62)	60.0	75.00	80.00	90.00	100.0			
		Change from baseline in EQ-5D VAS	EQ-5D VAS	Week 24	Sparsentan	39	25 (64.1)	4.76 (13.93)	-20.0	-3.00	3.00	16.00	29.0	0.24 [-0.34, 0.81]
					Irbesartan	37	22 (59.5)	1.86 (10.03)	-10.0	-5.00	0.00	9.00	30.0	
				Week 48	Sparsentan	39	24 (61.5)	2.58 (11.37)	-29.0	-3.00	3.00	10.00	24.0	0.01 [-0.60, 0.61]
					Irbesartan	37	19 (51.4)	2.53 (7.92)	-10.0	-1.00	0.00	5.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline eGFR Group 1												
< 60 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	127	119 (93.7)	80.61 (13.81)	11.0	75.00	81.00	90.00	100.0	
			Irbesartan	129	117 (90.7)	79.89 (13.65)	34.0	75.00	80.00	89.00	100.0	
	Week 24	Sparsentan	127	87 (68.5)	82.74 (10.73)	50.0	76.00	82.00	91.00	100.0		
		Irbesartan	129	75 (58.1)	79.52 (17.07)	4.0	74.00	81.00	90.00	100.0		
	Week 48	Sparsentan	127	67 (52.8)	83.21 (9.72)	49.0	80.00	82.00	90.00	100.0		
		Irbesartan	129	53 (41.1)	83.11 (12.42)	50.0	80.00	83.00	91.00	100.0		
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	127	87 (68.5)	2.29 (13.35)	-31.0	-5.00	0.00	10.00	44.0	0.19 [-0.12, 0.50]
			Irbesartan	129	75 (58.1)	-0.93 (19.64)	-90.0	-9.00	0.00	10.00	52.0	
		Week 48	Sparsentan	127	67 (52.8)	3.43 (14.61)	-20.0	-2.00	0.00	7.00	87.0	0.11 [-0.25, 0.47]
			Irbesartan	129	53 (41.1)	1.96 (10.50)	-31.0	-1.00	2.00	7.00	24.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	49	47 (95.9)	80.83 (16.08)	7.0	75.00	82.00	90.00	100.0	
			Irbesartan	48	43 (89.6)	80.44 (13.89)	37.0	71.00	81.00	90.00	100.0	
		Week 24	Sparsentan	49	29 (59.2)	84.55 (11.57)	60.0	75.00	87.00	91.00	100.0	
			Irbesartan	48	27 (56.3)	84.96 (10.01)	65.0	80.00	82.00	91.00	100.0	
		Week 48	Sparsentan	49	29 (59.2)	81.07 (10.59)	49.0	77.00	82.00	87.00	100.0	
			Irbesartan	48	20 (41.7)	82.20 (14.64)	40.0	80.00	85.00	90.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	49	29 (59.2)	4.00 (19.09)	-25.0	-8.00	3.00	8.00	75.0	0.04 [-0.48, 0.56]
			Irbesartan	48	27 (56.3)	3.37 (11.73)	-11.0	-5.00	0.00	9.00	34.0	
		Week 48	Sparsentan	49	29 (59.2)	1.10 (16.78)	-29.0	-6.00	0.00	5.00	64.0	-0.08 [-0.65, 0.49]
			Irbesartan	48	20 (41.7)	2.25 (11.98)	-35.0	-2.50	5.00	9.50	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	26	25 (96.2)	79.20 (13.04)	40.0	71.00	81.00	89.00	97.0	
		Week 24	Irbesartan	25	24 (96.0)	82.54 (17.58)	19.0	78.00	86.00	92.50	100.0	
			Sparsentan	26	16 (61.5)	83.31 (10.60)	60.0	78.00	84.00	90.50	100.0	
	Week 48	Irbesartan	25	13 (52.0)	80.69 (21.01)	30.0	70.00	90.00	92.00	100.0		
		Sparsentan	26	17 (65.4)	79.53 (13.62)	50.0	77.00	80.00	89.00	96.0		
		Irbesartan	25	10 (40.0)	82.10 (11.35)	66.0	75.00	80.00	90.00	100.0		
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	26	16 (61.5)	6.75 (15.80)	-20.0	-2.00	5.50	17.00	40.0	0.25 [-0.49, 0.98]
			Irbesartan	25	13 (52.0)	3.46 (9.42)	-11.0	-4.00	0.00	11.00	21.0	
		Week 48	Sparsentan	26	17 (65.4)	3.47 (17.75)	-30.0	-6.00	4.00	11.00	46.0	0.08 [-0.71, 0.86]
			Irbesartan	25	10 (40.0)	2.30 (10.39)	-14.0	-1.00	0.00	5.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline eGFR Group 2												
< 45 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	82	75 (91.5)	79.11 (15.04)	11.0	71.00	81.00	88.00	100.0	
			Irbesartan	80	70 (87.5)	77.90 (14.78)	34.0	73.00	80.00	85.00	100.0	
		Week 24	Sparsentan	82	54 (65.9)	82.91 (11.28)	50.0	80.00	83.00	91.00	100.0	
			Irbesartan	80	48 (60.0)	77.54 (17.12)	4.0	71.00	80.00	90.00	99.0	
		Week 48	Sparsentan	82	44 (53.7)	84.55 (9.72)	49.0	80.00	84.00	90.50	100.0	
			Irbesartan	80	32 (40.0)	81.59 (13.61)	50.0	79.50	82.50	89.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	82	54 (65.9)	3.37 (12.98)	-29.0	-4.00	0.00	10.00	44.0	0.26 [-0.13, 0.65]
			Irbesartan	80	48 (60.0)	-0.96 (19.60)	-71.0	-9.00	0.00	10.00	52.0	
		Week 48	Sparsentan	82	44 (53.7)	6.07 (16.34)	-20.0	-1.00	1.50	9.50	87.0	0.31 [-0.15, 0.77]
			Irbesartan	80	32 (40.0)	1.47 (12.25)	-31.0	-0.50	3.50	7.50	24.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
45 to < 60 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	45	44 (97.8)	83.18 (11.11)	50.0	78.50	81.50	91.00	100.0	
			Irbesartan	49	47 (95.9)	82.85 (11.28)	50.0	75.00	82.00	90.00	100.0	
			Sparsentan	45	33 (73.3)	82.45 (9.92)	59.0	74.00	82.00	90.00	100.0	
		Week 24	Irbesartan	49	27 (55.1)	83.04 (16.71)	10.0	80.00	85.00	91.00	100.0	
			Sparsentan	45	23 (51.1)	80.65 (9.39)	55.0	75.00	80.00	90.00	95.0	
			Irbesartan	49	21 (42.9)	85.43 (10.22)	65.0	80.00	83.00	92.00	100.0	
		Week 48	Sparsentan	45	23 (51.1)	80.65 (9.39)	55.0	75.00	80.00	90.00	95.0	
			Irbesartan	49	21 (42.9)	85.43 (10.22)	65.0	80.00	83.00	92.00	100.0	
			Sparsentan	45	33 (73.3)	0.52 (13.95)	-31.0	-9.00	0.00	8.00	35.0	0.08 [-0.43, 0.59]
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	45	33 (73.3)	0.52 (13.95)	-31.0	-9.00	0.00	8.00	35.0	0.08 [-0.43, 0.59]
Irbesartan			49	27 (55.1)	-0.89 (20.08)	-90.0	-5.00	0.00	9.00	22.0		
Sparsentan			45	23 (51.1)	-1.61 (8.82)	-20.0	-8.00	0.00	1.00	13.0	-0.53 [-1.13, 0.07]	
Week 48	Sparsentan	45	23 (51.1)	-1.61 (8.82)	-20.0	-8.00	0.00	1.00	13.0	-0.53 [-1.13, 0.07]		
	Irbesartan	49	21 (42.9)	2.71 (7.27)	-8.0	-1.00	1.00	5.00	19.0			
	Irbesartan	49	21 (42.9)	2.71 (7.27)	-8.0	-1.00	1.00	5.00	19.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	49	47 (95.9)	80.83 (16.08)	7.0	75.00	82.00	90.00	100.0	
			Irbesartan	48	43 (89.6)	80.44 (13.89)	37.0	71.00	81.00	90.00	100.0	
		Week 24	Sparsentan	49	29 (59.2)	84.55 (11.57)	60.0	75.00	87.00	91.00	100.0	
			Irbesartan	48	27 (56.3)	84.96 (10.01)	65.0	80.00	82.00	91.00	100.0	
		Week 48	Sparsentan	49	29 (59.2)	81.07 (10.59)	49.0	77.00	82.00	87.00	100.0	
			Irbesartan	48	20 (41.7)	82.20 (14.64)	40.0	80.00	85.00	90.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	49	29 (59.2)	4.00 (19.09)	-25.0	-8.00	3.00	8.00	75.0	0.04 [-0.48, 0.56]
			Irbesartan	48	27 (56.3)	3.37 (11.73)	-11.0	-5.00	0.00	9.00	34.0	
		Week 48	Sparsentan	49	29 (59.2)	1.10 (16.78)	-29.0	-6.00	0.00	5.00	64.0	-0.08 [-0.65, 0.49]
			Irbesartan	48	20 (41.7)	2.25 (11.98)	-35.0	-2.50	5.00	9.50	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	26	25 (96.2)	79.20 (13.04)	40.0	71.00	81.00	89.00	97.0	
		Week 24	Irbesartan	25	24 (96.0)	82.54 (17.58)	19.0	78.00	86.00	92.50	100.0	
			Sparsentan	26	16 (61.5)	83.31 (10.60)	60.0	78.00	84.00	90.50	100.0	
	Week 48	Irbesartan	25	13 (52.0)	80.69 (21.01)	30.0	70.00	90.00	92.00	100.0		
		Sparsentan	26	17 (65.4)	79.53 (13.62)	50.0	77.00	80.00	89.00	96.0		
		Irbesartan	25	10 (40.0)	82.10 (11.35)	66.0	75.00	80.00	90.00	100.0		
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	26	16 (61.5)	6.75 (15.80)	-20.0	-2.00	5.50	17.00	40.0	0.25 [-0.49, 0.98]
			Irbesartan	25	13 (52.0)	3.46 (9.42)	-11.0	-4.00	0.00	11.00	21.0	
		Week 48	Sparsentan	26	17 (65.4)	3.47 (17.75)	-30.0	-6.00	4.00	11.00	46.0	0.08 [-0.71, 0.86]
			Irbesartan	25	10 (40.0)	2.30 (10.39)	-14.0	-1.00	0.00	5.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline urine protein excretion													
<= 1.75 g/day	EQ-5D VAS	Baseline	Sparsentan	98	94 (95.9)	79.36 (16.33)	7.0	73.00	82.00	90.00	100.0		
			Irbesartan	94	84 (89.4)	81.11 (14.51)	19.0	74.50	81.00	90.00	100.0		
	Week 24	Sparsentan	98	69 (70.4)	82.28 (10.73)	51.0	75.00	81.00	90.00	100.0			
		Irbesartan	94	48 (51.1)	82.77 (12.93)	30.0	79.50	81.00	90.50	100.0			
	Week 48	Sparsentan	98	61 (62.2)	82.21 (11.02)	49.0	79.00	83.00	90.00	100.0			
		Irbesartan	94	36 (38.3)	85.25 (9.00)	66.0	80.00	84.00	91.50	100.0			
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	98	69 (70.4)	4.38 (16.88)	-25.0	-5.00	0.00	11.00	75.0	0.20 [-0.16, 0.57]	
				Irbesartan	94	48 (51.1)	1.48 (8.90)	-20.0	-4.50	0.00	9.50	21.0	
			Week 48	Sparsentan	98	61 (62.2)	4.38 (17.79)	-29.0	-2.00	1.00	10.00	87.0	0.08 [-0.33, 0.49]
				Irbesartan	94	36 (38.3)	3.19 (9.21)	-20.0	-1.00	3.50	9.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 1.75 g/day	EQ-5D VAS	Baseline	Sparsentan	104	97 (93.3)	81.57 (11.86)	40.0	75.00	81.00	91.00	100.0	
			Irbesartan	108	100 (92.6)	79.74 (14.00)	34.0	75.00	80.00	88.50	100.0	
		Week 24	Sparsentan	104	63 (60.6)	84.22 (10.97)	50.0	78.00	86.00	91.00	100.0	
			Irbesartan	108	67 (62.0)	79.61 (18.22)	4.0	72.00	82.00	91.00	100.0	
		Week 48	Sparsentan	104	52 (50.0)	81.98 (10.18)	50.0	78.50	81.50	89.00	100.0	
			Irbesartan	108	47 (43.5)	80.87 (14.77)	40.0	74.00	82.00	90.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	104	63 (60.6)	1.92 (12.70)	-31.0	-4.00	4.00	8.00	40.0	0.11 [-0.23, 0.46]
			Irbesartan	108	67 (62.0)	-0.07 (21.28)	-90.0	-7.00	0.00	11.00	52.0	
		Week 48	Sparsentan	104	52 (50.0)	1.04 (12.39)	-30.0	-6.00	0.00	7.00	46.0	-0.01 [-0.41, 0.38]
			Irbesartan	108	47 (43.5)	1.21 (11.78)	-35.0	-1.00	2.00	8.00	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline use of antihypertensives												
Yes	EQ-5D VAS	Baseline	Sparsentan	88	80 (90.9)	78.55 (17.02)	7.0	71.00	81.00	90.00	100.0	
			Irbesartan	83	74 (89.2)	78.82 (13.68)	37.0	72.00	80.00	89.00	100.0	
	Week 24	Sparsentan	88	55 (62.5)	84.27 (11.04)	51.0	78.00	89.00	91.00	100.0		
		Irbesartan	83	45 (54.2)	79.71 (14.94)	4.0	72.00	80.00	90.00	99.0		
	Week 48	Sparsentan	88	44 (50.0)	82.73 (12.17)	49.0	76.00	84.00	92.00	100.0		
		Irbesartan	83	32 (38.6)	81.88 (12.73)	50.0	74.50	83.50	90.00	100.0		
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	88	55 (62.5)	6.89 (16.18)	-25.0	0.00	7.00	11.00	75.0	0.28 [-0.11, 0.68]
				Irbesartan	83	45 (54.2)	2.09 (18.01)	-71.0	-6.00	0.00	10.00	52.0
Week 48			Sparsentan	88	44 (50.0)	6.98 (18.40)	-19.0	-1.00	1.50	9.00	87.0	0.19 [-0.27, 0.64]
			Irbesartan	83	32 (38.6)	4.06 (10.35)	-19.0	0.00	5.00	9.50	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	EQ-5D VAS	Baseline	Sparsentan	114	111 (97.4)	81.87 (11.73)	40.0	77.00	81.00	90.00	100.0	
			Irbesartan	119	110 (92.4)	81.40 (14.53)	19.0	75.00	83.00	90.00	100.0	
		Week 24	Sparsentan	114	77 (67.5)	82.44 (10.71)	50.0	75.00	81.00	90.00	100.0	
			Irbesartan	119	70 (58.8)	81.71 (17.08)	10.0	80.00	85.00	91.00	100.0	
		Week 48	Sparsentan	114	69 (60.5)	81.71 (9.53)	49.0	80.00	82.00	88.00	100.0	
			Irbesartan	119	51 (42.9)	83.33 (12.81)	40.0	80.00	83.00	91.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	114	77 (67.5)	0.57 (13.64)	-31.0	-6.00	0.00	5.00	40.0	0.06 [-0.26, 0.39]
			Irbesartan	119	70 (58.8)	-0.40 (16.69)	-90.0	-5.00	0.00	11.00	23.0	
		Week 48	Sparsentan	114	69 (60.5)	0.20 (12.91)	-30.0	-6.00	0.00	9.00	46.0	-0.05 [-0.41, 0.31]
			Irbesartan	119	51 (42.9)	0.82 (10.87)	-35.0	-4.00	1.00	7.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Time since renal biopsy												
<= 5 years	EQ-5D VAS	Baseline	Sparsentan	113	107 (94.7)	80.36 (13.96)	11.0	74.00	81.00	90.00	100.0	
			Irbesartan	127	119 (93.7)	80.81 (14.66)	19.0	75.00	80.00	90.00	100.0	
	Week 24	Sparsentan	113	76 (67.3)	83.50 (11.55)	50.0	75.00	86.00	91.00	100.0		
		Irbesartan	127	75 (59.1)	80.56 (17.80)	4.0	79.00	82.00	91.00	100.0		
	Week 48	Sparsentan	113	62 (54.9)	81.29 (11.79)	49.0	77.00	82.00	91.00	100.0		
		Irbesartan	127	55 (43.3)	82.13 (13.78)	40.0	75.00	82.00	90.00	100.0		
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	113	76 (67.3)	4.25 (13.90)	-29.0	-4.00	3.50	11.00	40.0	0.26 [-0.06, 0.58]
			Irbesartan	127	75 (59.1)	0.00 (18.78)	-90.0	-5.00	0.00	10.00	52.0	
		Week 48	Sparsentan	113	62 (54.9)	2.56 (16.44)	-30.0	-6.00	1.00	9.00	87.0	0.13 [-0.24, 0.49]
			Irbesartan	127	55 (43.3)	0.75 (11.09)	-35.0	-2.00	1.00	7.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 5 years	EQ-5D VAS	Baseline	Sparsentan	89	84 (94.4)	80.63 (14.68)	7.0	77.50	82.00	90.00	100.0	
			Irbesartan	75	65 (86.7)	79.55 (13.43)	34.0	73.00	80.00	89.00	100.0	
		Week 24	Sparsentan	89	56 (62.9)	82.80 (9.90)	59.0	78.00	82.00	90.00	100.0	
			Irbesartan	75	40 (53.3)	81.63 (12.97)	31.0	73.00	81.00	90.00	100.0	
		Week 48	Sparsentan	89	51 (57.3)	83.10 (8.94)	51.0	79.00	82.00	90.00	100.0	
			Irbesartan	75	28 (37.3)	84.04 (10.44)	52.0	80.00	85.00	90.50	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	89	56 (62.9)	1.79 (16.44)	-31.0	-6.00	0.00	7.50	75.0	0.01 [-0.40, 0.41]
			Irbesartan	75	40 (53.3)	1.65 (13.88)	-42.0	-4.50	0.00	10.00	34.0	
		Week 48	Sparsentan	89	51 (57.3)	3.18 (14.58)	-20.0	-3.00	0.00	7.00	64.0	-0.11 [-0.58, 0.35]
			Irbesartan	75	28 (37.3)	4.68 (9.63)	-20.0	-1.00	5.00	9.50	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: History of hypertension												
Yes	EQ-5D VAS	Baseline	Sparsentan	153	143 (93.5)	79.99 (13.66)	11.0	71.00	81.00	90.00	100.0	
			Irbesartan	157	140 (89.2)	79.76 (13.77)	34.0	74.50	80.00	90.00	100.0	
	Week 24	Sparsentan	153	100 (65.4)	83.44 (10.97)	50.0	76.00	83.50	91.00	100.0		
		Irbesartan	157	86 (54.8)	80.38 (16.54)	4.0	74.00	81.00	90.00	100.0		
	Week 48	Sparsentan	153	84 (54.9)	82.61 (11.06)	49.0	79.00	82.00	90.00	100.0		
		Irbesartan	157	62 (39.5)	82.55 (12.76)	40.0	80.00	84.50	90.00	100.0		
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	153	100 (65.4)	3.46 (13.43)	-29.0	-4.00	2.00	10.00	44.0	0.16 [-0.12, 0.45]
				Irbesartan	157	86 (54.8)	0.80 (18.78)	-90.0	-6.00	0.00	11.00	52.0
Week 48			Sparsentan	153	84 (54.9)	3.85 (14.67)	-30.0	-1.50	1.00	10.00	87.0	0.10 [-0.23, 0.43]
			Irbesartan	157	62 (39.5)	2.56 (11.01)	-35.0	-1.00	2.50	9.00	27.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	EQ-5D VAS	Baseline	Sparsentan	49	48 (98.0)	81.96 (15.91)	7.0	80.00	83.50	90.00	100.0	
			Irbesartan	45	44 (97.8)	82.30 (15.55)	19.0	75.00	85.00	91.00	100.0	
		Week 24	Sparsentan	49	32 (65.3)	82.47 (10.58)	59.0	76.00	83.50	90.50	100.0	
			Irbesartan	45	29 (64.4)	82.55 (15.44)	30.0	80.00	82.00	91.00	100.0	
		Week 48	Sparsentan	49	29 (59.2)	80.66 (9.13)	49.0	77.00	80.00	87.00	95.0	
			Irbesartan	45	21 (46.7)	83.43 (12.87)	54.0	80.00	81.00	94.00	100.0	
	Change from baseline in EQ-5D VAS	Week 24	Sparsentan	49	32 (65.3)	2.41 (19.41)	-31.0	-7.50	0.00	8.00	75.0	0.16 [-0.35, 0.66]
			Irbesartan	45	29 (64.4)	-0.10 (11.43)	-37.0	-3.00	0.00	5.00	22.0	
		Week 48	Sparsentan	49	29 (59.2)	-0.07 (17.85)	-29.0	-8.00	-2.00	4.00	64.0	-0.05 [-0.61, 0.52]
			Irbesartan	45	21 (46.7)	0.62 (9.95)	-31.0	-4.00	0.00	7.00	19.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.907
Male	Week 24	Sparsentan	139	91 (65.5)	2.88 (1.26)	(0.40, 5.36)	2.64 (1.83)	(-0.95, 6.24)	0.149
		Irbesartan	143	83 (58.0)	0.23 (1.32)	(-2.36, 2.83)			
	Week 48	Sparsentan	139	75 (54.0)	2.50 (1.38)	(-0.21, 5.21)	-0.77 (2.06)	(-4.82, 3.27)	0.707
		Irbesartan	143	61 (42.7)	3.27 (1.52)	(0.27, 6.27)			
Female	Week 24	Sparsentan	63	41 (65.1)	3.30 (1.76)	(-0.18, 6.78)	0.39 (2.66)	(-4.87, 5.64)	0.885
		Irbesartan	59	32 (54.2)	2.91 (1.98)	(-1.02, 6.84)			
	Week 48	Sparsentan	63	38 (60.3)	1.40 (1.81)	(-2.19, 4.99)	0.63 (3.01)	(-5.33, 6.60)	0.834
		Irbesartan	59	22 (37.3)	0.77 (2.39)	(-3.96, 5.50)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.069
<= 45 years	Week 24	Sparsentan	96	62 (64.6)	4.01 (1.65)	(0.75, 7.27)	5.60 (2.38)	(0.92, 10.29)	0.019 *
		Irbesartan	99	58 (58.6)	-1.59 (1.70)	(-4.95, 1.77)			
	Week 48	Sparsentan	96	54 (56.3)	1.51 (1.76)	(-1.96, 4.98)	0.96 (2.68)	(-4.32, 6.24)	0.719
		Irbesartan	99	41 (41.4)	0.55 (2.02)	(-3.43, 4.52)			
> 45 years	Week 24	Sparsentan	106	70 (66.0)	2.53 (1.28)	(0.00, 5.05)	-0.70 (1.92)	(-4.48, 3.08)	0.715
		Irbesartan	103	57 (55.3)	3.23 (1.43)	(0.42, 6.04)			
	Week 48	Sparsentan	106	59 (55.7)	2.66 (1.38)	(-0.07, 5.39)	-1.83 (2.15)	(-6.06, 2.41)	0.396
		Irbesartan	103	42 (40.8)	4.49 (1.64)	(1.26, 7.72)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.006 #
<= 18 years	Week 24	Sparsentan	9	3 (33.3)	4.69 (10.12)	(-21.57, 30.96)	23.09 (18.12)	(-25.95, 72.13)	0.267
		Irbesartan	5	2 (40.0)	-18.40 (14.14)	(-57.79, 20.99)			
	Week 48	Sparsentan	9	5 (55.6)	9.09 (9.00)	(-15.68, 33.87)	28.95 (18.65)	(-20.43, 78.33)	0.187
		Irbesartan	5	1 (20.0)	-19.86 (15.76)	(-60.86, 21.14)			
> 18 to 40 years	Week 24	Sparsentan	102	66 (64.7)	3.89 (1.52)	(0.89, 6.88)	3.36 (2.18)	(-0.94, 7.65)	0.125
		Irbesartan	109	63 (57.8)	0.53 (1.56)	(-2.54, 3.61)			
	Week 48	Sparsentan	102	58 (56.9)	1.68 (1.62)	(-1.52, 4.87)	0.60 (2.46)	(-4.25, 5.46)	0.807
		Irbesartan	109	45 (41.3)	1.07 (1.84)	(-2.55, 4.70)			
> 40 years	Week 24	Sparsentan	91	63 (69.2)	2.13 (1.38)	(-0.58, 4.85)	0.29 (2.07)	(-3.79, 4.37)	0.889
		Irbesartan	88	50 (56.8)	1.84 (1.54)	(-1.20, 4.89)			
	Week 48	Sparsentan	91	50 (54.9)	2.40 (1.53)	(-0.62, 5.41)	-1.99 (2.35)	(-6.61, 2.64)	0.398
		Irbesartan	88	37 (42.0)	4.38 (1.78)	(0.88, 7.89)			

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LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.490
North America	Week 24	Sparsentan	35	15 (42.9)	5.72 (4.35)	(-2.98, 14.41)	7.40 (5.42)	(-3.44, 18.24)	0.177
		Irbesartan	46	28 (60.9)	-1.69 (3.22)	(-8.11, 4.74)			
	Week 48	Sparsentan	35	13 (37.1)	3.29 (4.76)	(-6.21, 12.80)	-1.90 (6.24)	(-14.37, 10.57)	0.762
		Irbesartan	46	18 (39.1)	5.19 (3.97)	(-2.74, 13.13)			
Europe	Week 24	Sparsentan	98	60 (61.2)	1.10 (1.46)	(-1.78, 3.97)	-0.14 (2.14)	(-4.35, 4.08)	0.949
		Irbesartan	115	52 (45.2)	1.24 (1.57)	(-1.86, 4.33)			
	Week 48	Sparsentan	98	45 (45.9)	2.64 (1.69)	(-0.68, 5.97)	0.32 (2.46)	(-4.54, 5.18)	0.897
		Irbesartan	115	40 (34.8)	2.32 (1.79)	(-1.21, 5.86)			
Asia Pacific	Week 24	Sparsentan	69	57 (82.6)	5.08 (1.33)	(2.46, 7.71)	3.68 (2.19)	(-0.65, 8.01)	0.095
		Irbesartan	41	35 (85.4)	1.41 (1.72)	(-2.00, 4.81)			
	Week 48	Sparsentan	69	55 (79.7)	2.05 (1.35)	(-0.61, 4.71)	2.17 (2.36)	(-2.50, 6.84)	0.360
		Irbesartan	41	25 (61.0)	-0.12 (1.92)	(-3.93, 3.68)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.180
< 27 kg/m**2	Week 24	Sparsentan	84	54 (64.3)	0.95 (1.55)	(-2.11, 4.02)	0.75 (2.19)	(-3.57, 5.06)	0.733
		Irbesartan	94	56 (59.6)	0.21 (1.53)	(-2.81, 3.23)			
	Week 48	Sparsentan	84	49 (58.3)	-0.58 (1.63)	(-3.80, 2.64)	-1.66 (2.46)	(-6.52, 3.20)	
		Irbesartan	94	39 (41.5)	1.08 (1.82)	(-2.52, 4.67)			
>= 27 kg/m**2	Week 24	Sparsentan	118	78 (66.1)	4.99 (1.38)	(2.26, 7.71)	4.02 (2.11)	(-0.14, 8.18)	0.058
		Irbesartan	107	59 (55.1)	0.97 (1.59)	(-2.17, 4.10)			
	Week 48	Sparsentan	118	64 (54.2)	4.57 (1.52)	(1.58, 7.55)	1.06 (2.38)	(-3.62, 5.74)	
		Irbesartan	107	44 (41.1)	3.50 (1.83)	(-0.10, 7.10)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan						Interaction:	0.281
eGFR Low and UP High	Week 24	Sparsentan	71	45 (63.4)	0.77 (1.88)	(-2.95, 4.49)	6.27 (2.81)	(0.70, 11.85)	0.028 *
		Irbesartan	74	38 (51.4)	-5.50 (2.09)	(-9.64, -1.37)			
	Week 48	Sparsentan	71	39 (54.9)	1.41 (1.97)	(-2.48, 5.30)	-1.44 (3.09)	(-7.55, 4.68)	0.643
		Irbesartan	74	25 (33.8)	2.85 (2.38)	(-1.87, 7.57)			
eGFR Low and UP Low	Week 24	Sparsentan	55	40 (72.7)	3.17 (1.88)	(-0.54, 6.89)	0.27 (2.76)	(-5.19, 5.74)	0.922
		Irbesartan	55	35 (63.6)	2.90 (2.01)	(-1.08, 6.89)			
	Week 48	Sparsentan	55	28 (50.9)	3.00 (2.26)	(-1.47, 7.47)	-0.39 (3.26)	(-6.85, 6.06)	0.904
		Irbesartan	55	26 (47.3)	3.40 (2.34)	(-1.23, 8.02)			
eGFR High and UP High	Week 24	Sparsentan	37	22 (59.5)	5.79 (2.40)	(1.00, 10.57)	-3.81 (3.50)	(-10.79, 3.17)	0.280
		Irbesartan	36	20 (55.6)	9.60 (2.55)	(4.51, 14.68)			
	Week 48	Sparsentan	37	22 (59.5)	2.11 (2.40)	(-2.68, 6.89)	-0.55 (3.90)	(-8.32, 7.22)	0.889
		Irbesartan	36	13 (36.1)	2.65 (3.07)	(-3.47, 8.77)			
eGFR High and UP Low	Week 24	Sparsentan	39	25 (64.1)	5.18 (1.93)	(1.33, 9.02)	3.80 (2.83)	(-1.83, 9.42)	0.183
		Irbesartan	37	22 (59.5)	1.38 (2.06)	(-2.72, 5.48)			
	Week 48	Sparsentan	39	24 (61.5)	2.54 (1.97)	(-1.39, 6.46)	0.15 (2.97)	(-5.75, 6.05)	0.960
		Irbesartan	37	19 (51.4)	2.39 (2.22)	(-2.02, 6.80)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.698
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	87 (68.5)	2.23 (1.31)	(-0.35, 4.82)	3.35 (1.93)	(-0.45, 7.15)	0.084
		Irbesartan	129	75 (58.1)	-1.12 (1.42)	(-3.90, 1.67)			
	Week 48	Sparsentan	127	67 (52.8)	2.86 (1.48)	(-0.06, 5.79)	0.38 (2.24)	(-4.04, 4.80)	0.865
		Irbesartan	129	53 (41.1)	2.48 (1.67)	(-0.81, 5.78)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	29 (59.2)	4.60 (1.98)	(0.67, 8.53)	0.64 (2.86)	(-5.04, 6.32)	0.823
		Irbesartan	48	27 (56.3)	3.96 (2.04)	(-0.10, 8.01)			
	Week 48	Sparsentan	49	29 (59.2)	1.22 (1.98)	(-2.72, 5.15)	0.05 (3.12)	(-6.14, 6.24)	0.987
		Irbesartan	48	20 (41.7)	1.16 (2.37)	(-3.55, 5.87)			
≥ 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	16 (61.5)	6.24 (3.06)	(0.07, 12.42)	2.53 (4.57)	(-6.69, 11.75)	0.584
		Irbesartan	25	13 (52.0)	3.72 (3.39)	(-3.12, 10.56)			
	Week 48	Sparsentan	26	17 (65.4)	3.13 (3.00)	(-2.93, 9.19)	-0.11 (4.87)	(-9.92, 9.69)	0.981
		Irbesartan	25	10 (40.0)	3.24 (3.81)	(-4.41, 10.90)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024



Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.280
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	54 (65.9)	3.44 (1.71)	(0.07, 6.81)	4.85 (2.49)	(-0.07, 9.76)	0.053
		Irbesartan	80	48 (60.0)	-1.41 (1.81)	(-4.99, 2.18)			
	Week 48	Sparsentan	82	44 (53.7)	5.24 (1.87)	(1.55, 8.93)	2.73 (2.87)	(-2.94, 8.40)	0.344
		Irbesartan	80	32 (40.0)	2.51 (2.18)	(-1.79, 6.82)			
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	33 (73.3)	0.01 (2.05)	(-4.06, 4.09)	0.46 (3.09)	(-5.68, 6.60)	0.882
		Irbesartan	49	27 (55.1)	-0.45 (2.30)	(-5.02, 4.12)			
	Week 48	Sparsentan	45	23 (51.1)	-1.76 (2.47)	(-6.66, 3.15)	-4.25 (3.63)	(-11.45, 2.95)	0.244
		Irbesartan	49	21 (42.9)	2.49 (2.62)	(-2.72, 7.70)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	29 (59.2)	4.60 (1.98)	(0.67, 8.53)	0.64 (2.86)	(-5.04, 6.32)	0.823
		Irbesartan	48	27 (56.3)	3.96 (2.04)	(-0.10, 8.01)			
	Week 48	Sparsentan	49	29 (59.2)	1.22 (1.98)	(-2.72, 5.15)	0.05 (3.12)	(-6.14, 6.24)	0.987
		Irbesartan	48	20 (41.7)	1.16 (2.37)	(-3.55, 5.87)			
≥ 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	16 (61.5)	6.24 (3.06)	(0.07, 12.42)	2.53 (4.57)	(-6.69, 11.75)	0.584
		Irbesartan	25	13 (52.0)	3.72 (3.39)	(-3.12, 10.56)			
	Week 48	Sparsentan	26	17 (65.4)	3.13 (3.00)	(-2.93, 9.19)	-0.11 (4.87)	(-9.92, 9.69)	0.981
		Irbesartan	25	10 (40.0)	3.24 (3.81)	(-4.41, 10.90)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
 Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.329
<= 1.75 g/day	Week 24	Sparsentan	98	69 (70.4)	3.46 (1.18)	(1.13, 5.78)	0.68 (1.84)	(-2.95, 4.31)	0.712
		Irbesartan	94	48 (51.1)	2.77 (1.42)	(-0.02, 5.57)			
	Week 48	Sparsentan	98	61 (62.2)	3.21 (1.25)	(0.75, 5.68)	-1.54 (2.06)	(-5.59, 2.52)	0.455
		Irbesartan	94	36 (38.3)	4.75 (1.62)	(1.55, 7.95)			
> 1.75 g/day	Week 24	Sparsentan	104	63 (60.6)	2.92 (1.72)	(-0.46, 6.30)	3.70 (2.40)	(-1.04, 8.44)	0.125
		Irbesartan	108	67 (62.0)	-0.78 (1.67)	(-4.07, 2.51)			
	Week 48	Sparsentan	104	52 (50.0)	1.03 (1.89)	(-2.69, 4.75)	0.16 (2.77)	(-5.29, 5.61)	0.954
		Irbesartan	108	47 (43.5)	0.87 (1.99)	(-3.05, 4.78)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
 Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan							Interaction: 0.224
	Yes	Sparsentan	88	55 (62.5)	6.87 (1.60)	(3.72, 10.03)	4.42 (2.38)	(-0.28, 9.12)	0.065
		Irbesartan	83	45 (54.2)	2.45 (1.77)	(-1.04, 5.94)			
	Week 48	Sparsentan	88	44 (50.0)	5.96 (1.77)	(2.46, 9.46)	0.91 (2.73)	(-4.48, 6.30)	0.740
		Irbesartan	83	32 (38.6)	5.05 (2.07)	(0.97, 9.14)			
No	Week 24	Sparsentan	114	77 (67.5)	0.74 (1.34)	(-1.91, 3.39)	1.00 (1.95)	(-2.83, 4.84)	0.606
		Irbesartan	119	70 (58.8)	-0.27 (1.41)	(-3.04, 2.51)			
	Week 48	Sparsentan	114	69 (60.5)	-0.26 (1.42)	(-3.04, 2.53)	-1.19 (2.17)	(-5.46, 3.08)	0.584
		Irbesartan	119	51 (42.9)	0.93 (1.65)	(-2.31, 4.17)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.549
<= 5 years	Week 24	Sparsentan	113	76 (67.3)	3.76 (1.48)	(0.84, 6.68)	3.47 (2.10)	(-0.67, 7.61)	0.100
		Irbesartan	127	75 (59.1)	0.29 (1.49)	(-2.65, 3.23)			
	Week 48	Sparsentan	113	62 (54.9)	1.51 (1.64)	(-1.72, 4.75)	-0.13 (2.39)	(-4.84, 4.58)	0.958
		Irbesartan	127	55 (43.3)	1.64 (1.74)	(-1.79, 5.07)			
> 5 years	Week 24	Sparsentan	89	56 (62.9)	2.67 (1.33)	(0.04, 5.31)	0.96 (2.09)	(-3.17, 5.10)	0.645
		Irbesartan	75	40 (53.3)	1.71 (1.62)	(-1.49, 4.91)			
	Week 48	Sparsentan	89	51 (57.3)	2.79 (1.38)	(0.07, 5.52)	-0.68 (2.28)	(-5.19, 3.83)	0.767
		Irbesartan	75	28 (37.3)	3.47 (1.81)	(-0.11, 7.05)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 04APR2024

Table PT1VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
 Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.407
Yes	Week 24	Sparsentan	153	100 (65.4)	3.66 (1.22)	(1.26, 6.06)	2.87 (1.79)	(-0.65, 6.39)	0.110
		Irbesartan	157	86 (54.8)	0.79 (1.31)	(-1.80, 3.37)			
	Week 48	Sparsentan	153	84 (54.9)	3.28 (1.32)	(0.67, 5.89)	0.49 (2.03)	(-3.51, 4.49)	0.810
		Irbesartan	157	62 (39.5)	2.79 (1.54)	(-0.24, 5.83)			
No	Week 24	Sparsentan	49	32 (65.3)	1.72 (1.92)	(-2.10, 5.54)	1.10 (2.81)	(-4.49, 6.69)	0.697
		Irbesartan	45	29 (64.4)	0.62 (2.04)	(-3.44, 4.68)			
	Week 48	Sparsentan	49	29 (59.2)	-1.19 (1.99)	(-5.14, 2.75)	-2.32 (3.06)	(-8.41, 3.76)	0.450
		Irbesartan	45	21 (46.7)	1.13 (2.32)	(-3.47, 5.73)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

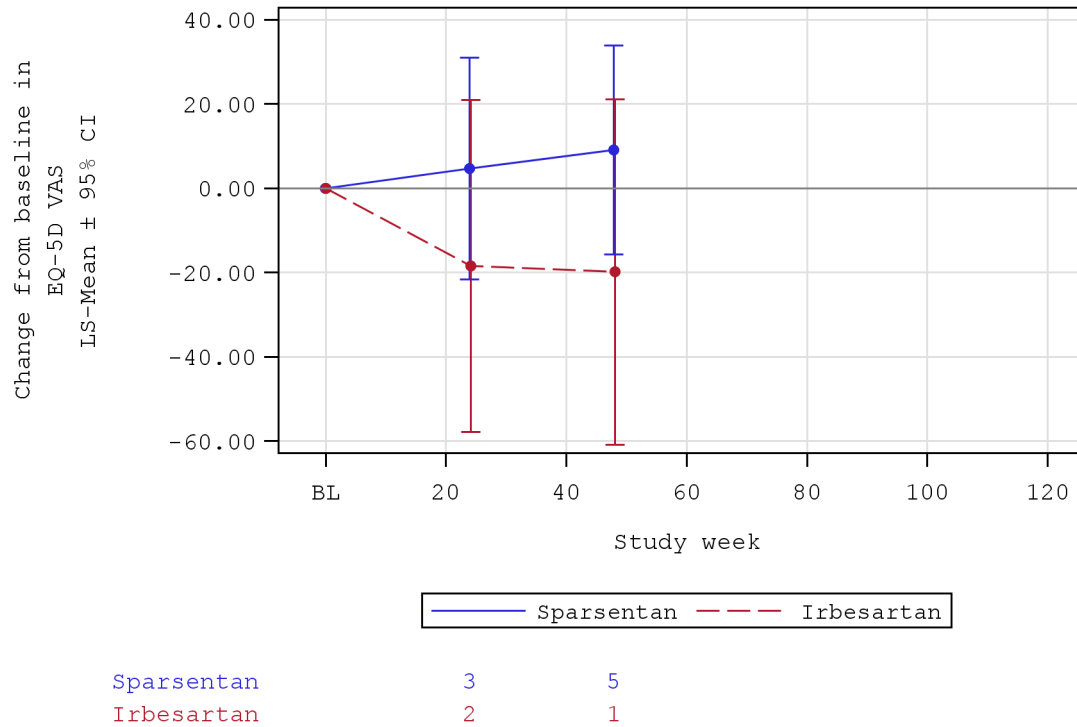
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

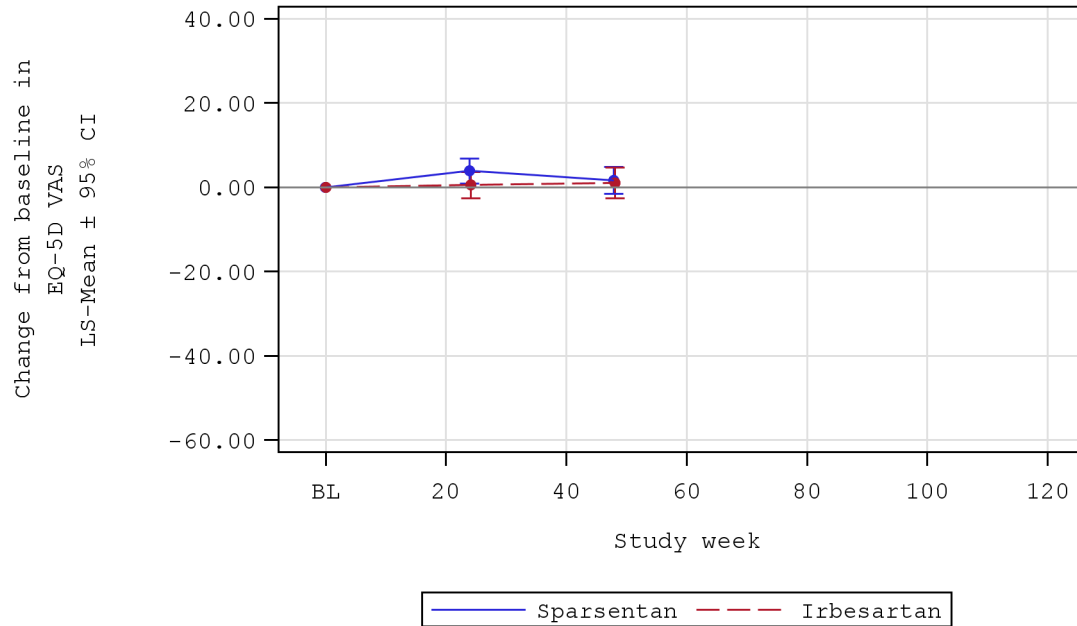
Source Data: aqs, created on: 04APR2024

Figure PF1VSC\_FSGM: Change from baseline in EQ-5D VAS by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Age at IgAN diagnosis: <= 18 years



Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
 95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1VSC\_FSCM.

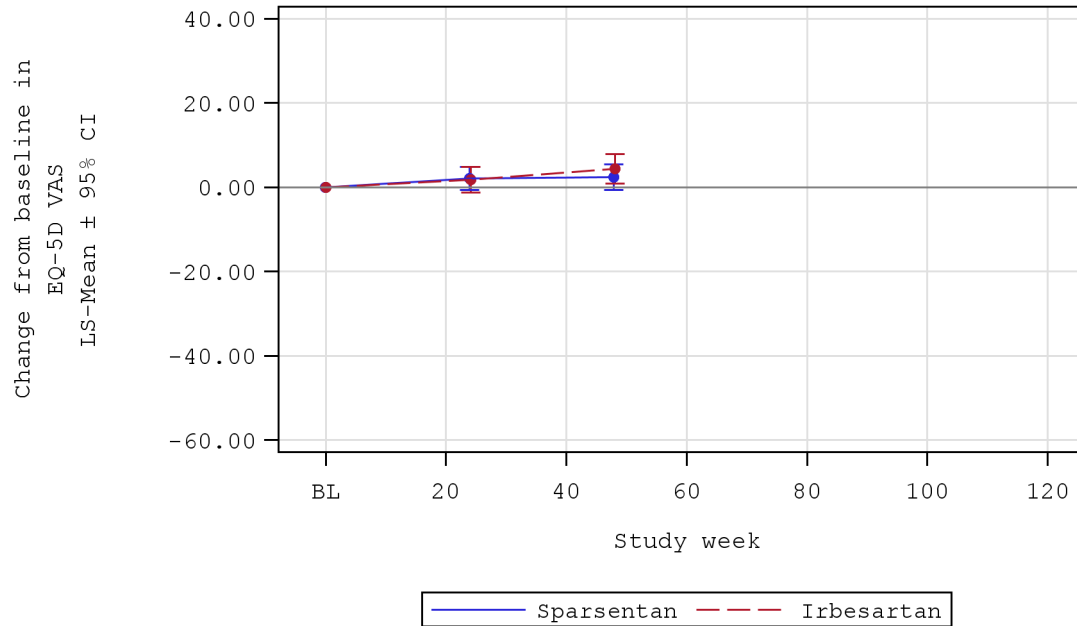
Figure PF1VSC\_FSGM: Change from baseline in EQ-5D VAS by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Age at IgAN diagnosis: > 18 to 40 years



Sparsentan	66	58
Irbesartan	63	45

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
 95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1VSC\_FSCM.

Figure PF1VSC\_FSGM: Change from baseline in EQ-5D VAS by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Age at IgAN diagnosis: > 40 years



Sparsentan	63	50
Irbesartan	50	37

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
 95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1VSC\_FSCM.



Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.974
Male	Sparsentan	139	17 (12.2)	NE		0.949	(0.481, 1.872)	0.880
	Irbesartan	143	21 (14.7)	NE				
Female	Sparsentan	63	12 (19.0)	NE		1.178	(0.338, 4.110)	0.797
	Irbesartan	59	5 (8.5)	110.0	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.070
<= 45 years	Sparsentan	96	16 (16.7)	NE		1.572	(0.688, 3.593)	0.284
	Irbesartan	99	11 (11.1)	NE				
> 45 years	Sparsentan	106	13 (12.3)	NE		0.517	(0.234, 1.139)	0.101
	Irbesartan	103	15 (14.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	2 (22.2) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	15 (14.7)	NE		1.201	(0.553, 2.609)	0.644
	Irbesartan	109	14 (12.8)	110.0	(94.9, NE)			
> 40 years	Sparsentan	91	12 (13.2)	NE		0.662	(0.282, 1.558)	0.345
	Irbesartan	88	12 (13.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.134
North America	Sparsentan	35	5 (14.3)	NE		0.580	(0.125, 2.690)	0.486
	Irbesartan	46	7 (15.2)	NE				
Europe	Sparsentan	98	12 (12.2)	108.0	(95.0, NE)	0.735	(0.318, 1.696)	0.470
	Irbesartan	115	13 (11.3)	NE				
Asia Pacific	Sparsentan	69	12 (17.4)	NE		2.818	(0.805, 9.861)	0.105
	Irbesartan	41	6 (14.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.332
< 27 kg/m**2	Sparsentan	84	14 (16.7)	108.0	(95.0, NE)	1.190	(0.452, 3.128)	0.725
	Irbesartan	94	7 (7.4)	NE				
≥ 27 kg/m**2	Sparsentan	118	15 (12.7)	NE		0.771	(0.375, 1.586)	0.480
	Irbesartan	107	18 (16.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan							Interaction test: 0.007 #
eGFR Low and UP High	Sparsentan	71	6 (8.5)	NE		1.052	(0.302, 3.663)	0.937
	Irbesartan	74	8 (10.8)	NE				
eGFR Low and UP Low	Sparsentan	55	9 (16.4)	NE		0.659	(0.221, 1.962)	0.453
	Irbesartan	55	8 (14.5)	NE				
eGFR High and UP High	Sparsentan	37	7 (18.9)	NE		0.444	(0.141, 1.393)	0.164
	Irbesartan	36	7 (19.4)	71.4	(25.1, NE)			
eGFR High and UP Low	Sparsentan	39	7 (17.9)	NE		7.790	(1.072, 56.617)	0.042 *
	Irbesartan	37	3 (8.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.191
< 60 mL/min/1.73 m**2	Sparsentan	127	16 (12.6)	NE		0.814	(0.402, 1.647)	0.567
	Irbesartan	129	18 (14.0)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	7 (14.3)	NE		0.853	(0.253, 2.870)	0.797
	Irbesartan	48	5 (10.4)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	6 (23.1)	NE		2.564	(0.452, 14.546)	0.288
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.240
< 45 mL/min/1.73 m**2	Sparsentan	82	10 (12.2)	NE		0.710	(0.297, 1.695)	0.440
	Irbesartan	80	13 (16.3)	110.0	(94.9, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	6 (13.3)	NE		1.132	(0.272, 4.710)	0.865
	Irbesartan	49	5 (10.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	7 (14.3)	NE		0.853	(0.253, 2.870)	0.797
	Irbesartan	48	5 (10.4)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	6 (23.1)	NE		2.564	(0.452, 14.546)	0.288
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.429
<= 1.75 g/day	Sparsentan	98	16 (16.3)	NE		1.718	(0.699, 4.224)	0.238
	Irbesartan	94	9 (9.6)	110.0	(94.9, NE)			
> 1.75 g/day	Sparsentan	104	13 (12.5)	NE		0.908	(0.414, 1.992)	0.810
	Irbesartan	108	17 (15.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.157
Yes	Sparsentan	88	12 (13.6)	NE		0.677	(0.292, 1.568)	0.363
	Irbesartan	83	13 (15.7)	110.0	(94.9, NE)			
No	Sparsentan	114	17 (14.9)	NE		1.393	(0.634, 3.064)	0.409
	Irbesartan	119	13 (10.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.081
<= 5 years	Sparsentan	113	18 (15.9)	NE		1.420	(0.707, 2.850)	0.324
	Irbesartan	127	17 (13.4)	NE				
> 5 years	Sparsentan	89	11 (12.4)	NE		0.515	(0.195, 1.356)	0.179
	Irbesartan	75	9 (12.0)	NE				

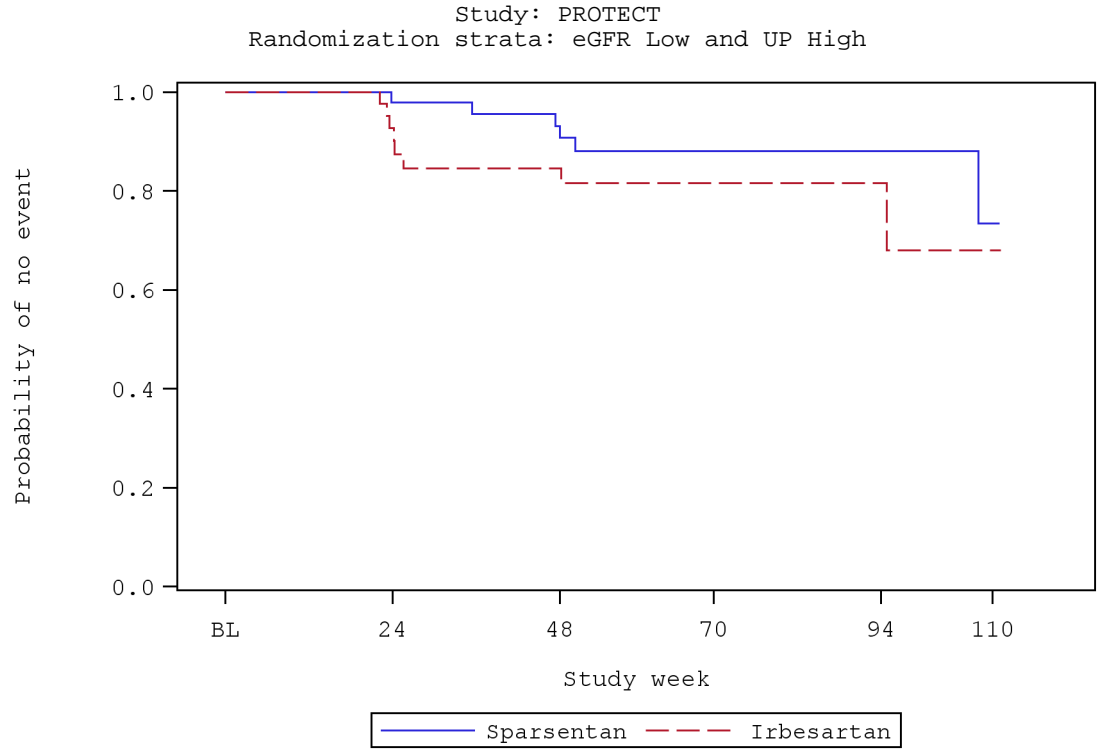
N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.197
Yes	Sparsentan	153	23 (15.0)	NE		0.788	(0.434, 1.433)	0.436
	Irbesartan	157	24 (15.3)	NE				
No	Sparsentan	49	6 (12.2)	NE		3.535	(0.641, 19.492)	0.147
	Irbesartan	45	2 (4.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

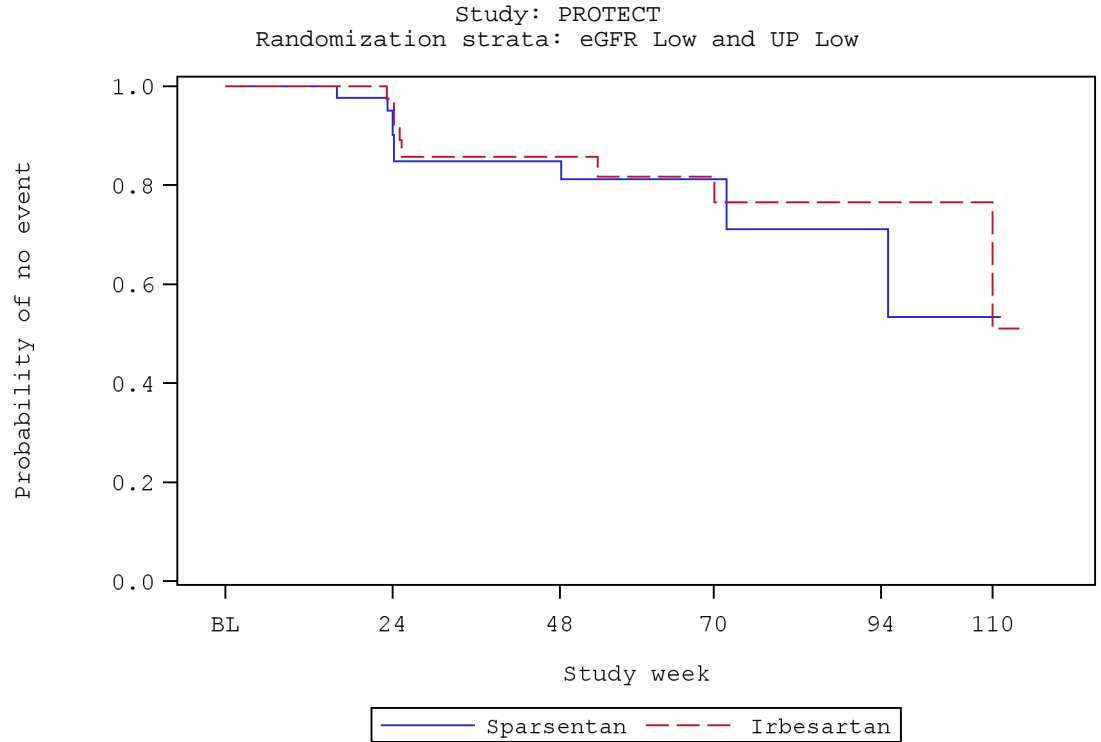
Figure PF1VSIT\_FSKM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	71	47	39	31	16	4
Irbesartan	74	36	28	17	9	2

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1VSIT\_FSTM

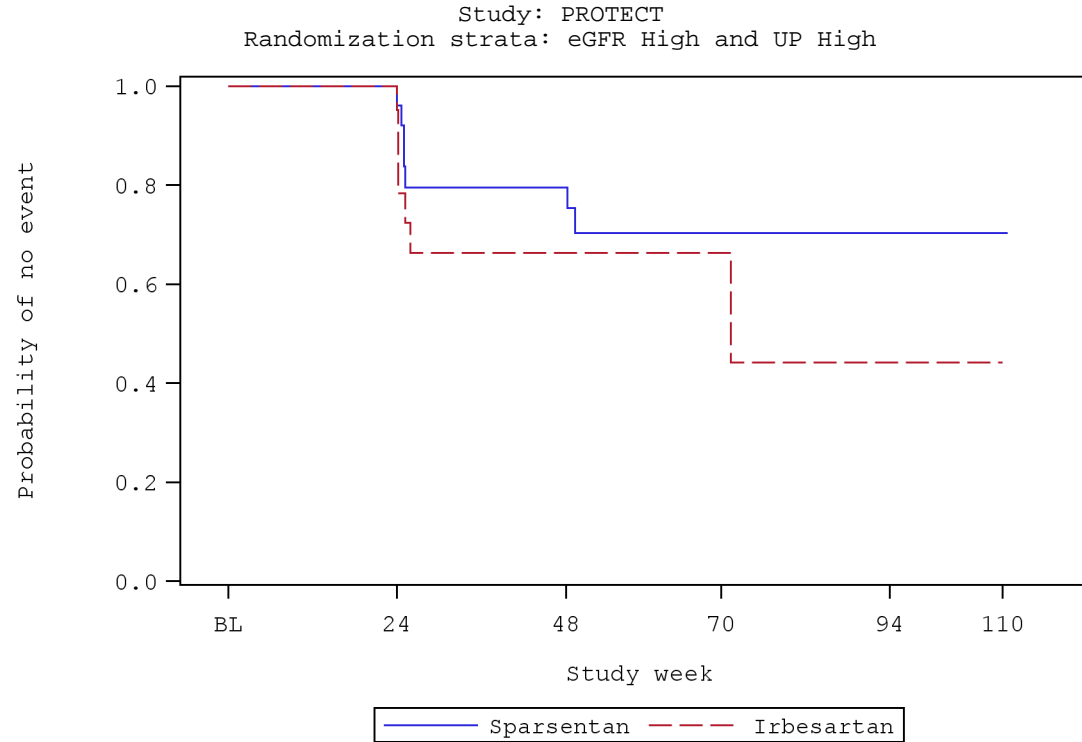
Figure PF1VSIT\_FSKM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	55	38	24	15	6	3
Irbesartan	55	37	26	16	8	3

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1VSIT\_FSTM

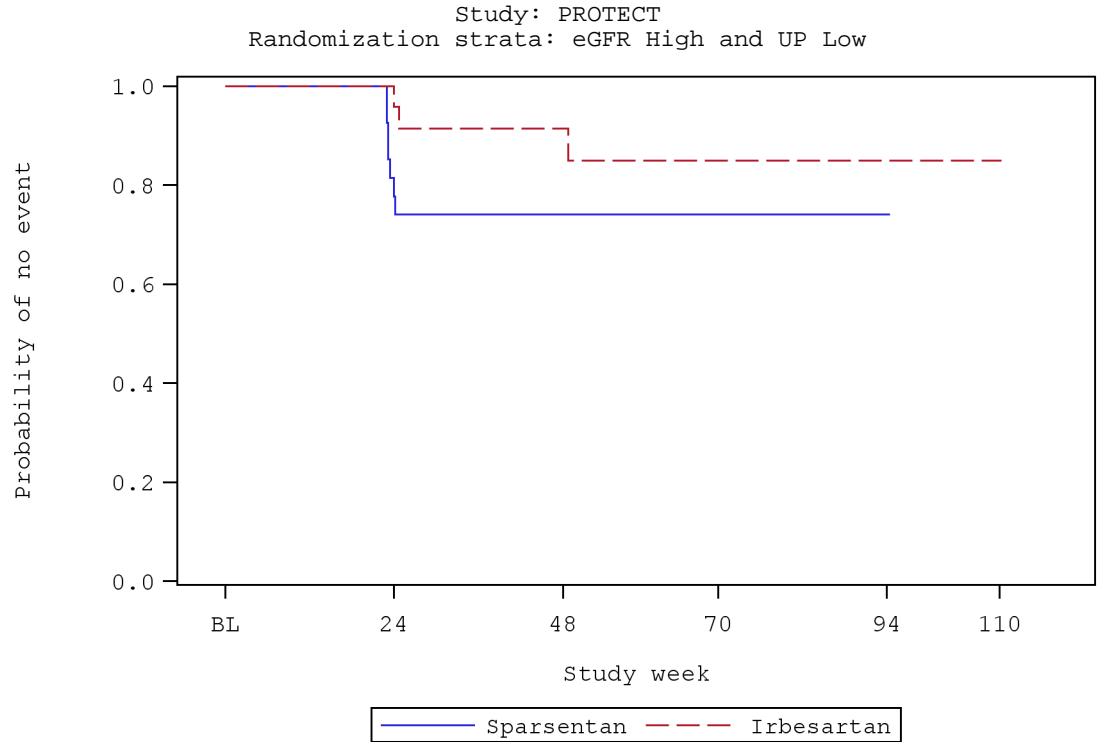
Figure PF1VSIT\_FSKM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	37	26	19	12	4	1
Irbesartan	36	21	9	5	1	1

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1VSIT\_FSTM

Figure PF1VSIT\_FSKM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	39	22	16	11	3	0
Irbesartan	37	24	17	12	2	1

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1VSIT\_FSTM



Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.543
Male	Sparsentan	139	13 (9.4)	NE		1.013	(0.437, 2.350)	0.976
	Irbesartan	143	10 (7.0)	NE				
Female	Sparsentan	63	13 (20.6)	112.1	(112.1, NE)	1.612	(0.594, 4.372)	0.348
	Irbesartan	59	6 (10.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.783
<= 45 years	Sparsentan	96	14 (14.6)	NE		1.305	(0.559, 3.047)	0.538
	Irbesartan	99	9 (9.1)	NE				
> 45 years	Sparsentan	106	12 (11.3)	112.1	(111.0, NE)	1.078	(0.413, 2.813)	0.878
	Irbesartan	103	7 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Sparsentan	102	15 (14.7)	NE		1.521	(0.655, 3.534)	0.329
	Irbesartan	109	9 (8.3)	NE				
> 40 years	Sparsentan	91	11 (12.1)	112.1	(111.0, NE)	1.072	(0.388, 2.963)	0.893
	Irbesartan	88	6 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
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 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.679
North America	Sparsentan	35	4 (11.4)	NE		1.370	(0.326, 5.761)	0.668
	Irbesartan	46	4 (8.7)	NE				
Europe	Sparsentan	98	11 (11.2)	112.1	(112.1, NE)	0.930	(0.384, 2.251)	0.872
	Irbesartan	115	10 (8.7)	NE				
Asia Pacific	Sparsentan	69	11 (15.9)	NE		3.090	(0.618, 15.452)	0.170
	Irbesartan	41	2 (4.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan							Interaction test: 0.029 #
< 27 kg/m**2	Sparsentan	84	14 (16.7)	112.1	(112.1, NE)	2.575	(0.955, 6.944)	0.062
	Irbesartan	94	6 (6.4)	NE				
≥ 27 kg/m**2	Sparsentan	118	12 (10.2)	NE		0.654	(0.272, 1.573)	0.343
	Irbesartan	107	10 (9.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	NE
eGFR Low and UP High	Sparsentan	71	7 (9.9)	111.0	(111.0, NE)	0.437	(0.165, 1.160)	0.097
	Irbesartan	74	10 (13.5)	NE				
eGFR Low and UP Low	Sparsentan	55	8 (14.5)	112.1	(NE, NE)	2.002	(0.578, 6.939)	0.274
	Irbesartan	55	4 (7.3)	NE				
eGFR High and UP High	Sparsentan	37	6 (16.2)	NE		2.081	(0.413, 10.480)	0.374
	Irbesartan	36	2 (5.6)	NE				
eGFR High and UP Low	Sparsentan	39	5 (12.8)	NE		NE		NE
	Irbesartan	37	No events in 1 group 0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	NE
< 60 mL/min/1.73 m**2	Sparsentan	127	15 (11.8)	112.1	(111.0, 112.1)	0.770	(0.365, 1.625)	0.493
	Irbesartan	129	14 (10.9)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	6 (12.2)	NE		2.239	(0.445, 11.270)	0.328
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	5 (19.2) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: NE
< 45 mL/min/1.73 m**2	Sparsentan	82	6 (7.3)	NE		0.337	(0.126, 0.902)	0.030 *
	Irbesartan	80	13 (16.3)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	9 (20.0)	112.1	(94.0, 112.1)	8.124	(0.986, 66.937)	0.052
	Irbesartan	49	1 (2.0)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	6 (12.2)	NE		2.239	(0.445, 11.270)	0.328
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	5 (19.2)	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.003 #
<= 1.75 g/day	Sparsentan	98	16 (16.3)	112.1	(NE, NE)	7.300	(1.610, 33.094)	0.010 *
	Irbesartan	94	2 (2.1)	NE				
> 1.75 g/day	Sparsentan	104	10 (9.6)	NE		0.590	(0.257, 1.355)	0.213
	Irbesartan	108	14 (13.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.280
Yes	Sparsentan	88	7 (8.0)	NE		0.747	(0.239, 2.336)	0.616
	Irbesartan	83	6 (7.2)	NE				
No	Sparsentan	114	19 (16.7)	112.1	(112.1, NE)	1.535	(0.705, 3.342)	0.280
	Irbesartan	119	10 (8.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.202
<= 5 years	Sparsentan	113	17 (15.0)	111.0	(111.0, NE)	1.699	(0.749, 3.857)	0.205
	Irbesartan	127	9 (7.1)	NE				
> 5 years	Sparsentan	89	9 (10.1)	NE		0.778	(0.282, 2.141)	0.626
	Irbesartan	75	7 (9.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

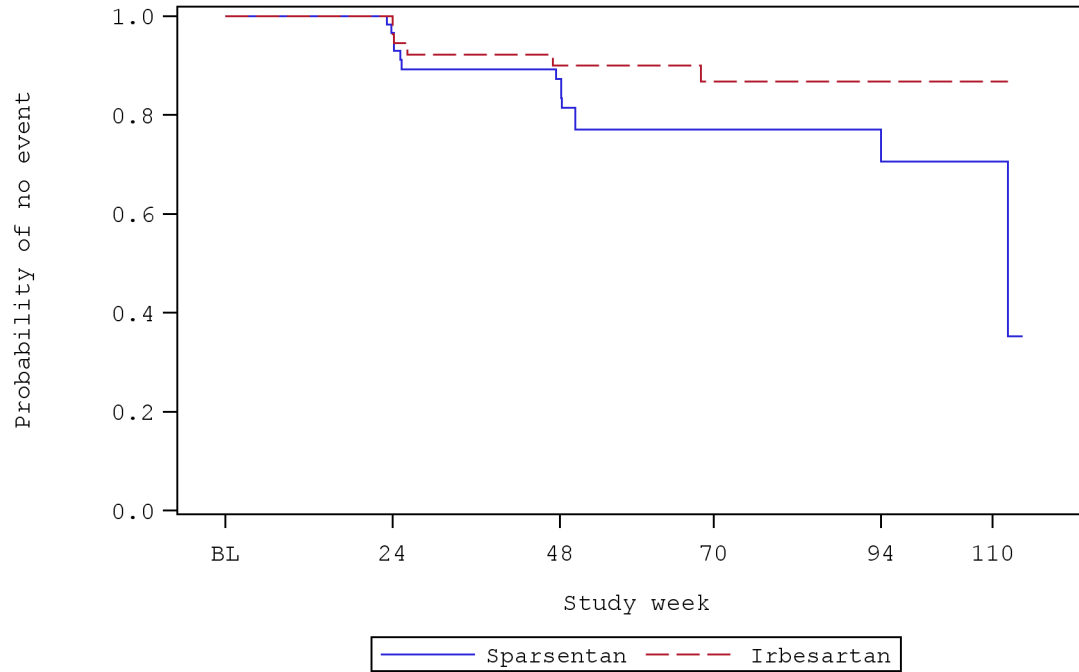
Table PT1VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension		Sparsentan				Interaction test: 0.240		
Yes	Sparsentan	153	17 (11.1)	112.1	(111.0, NE)	0.992	(0.467, 2.108)	0.983
	Irbesartan	157	12 (7.6)	NE				
No	Sparsentan	49	9 (18.4)	NE		2.657	(0.748, 9.437)	0.131
	Irbesartan	45	4 (8.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Figure PF1VSDT\_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

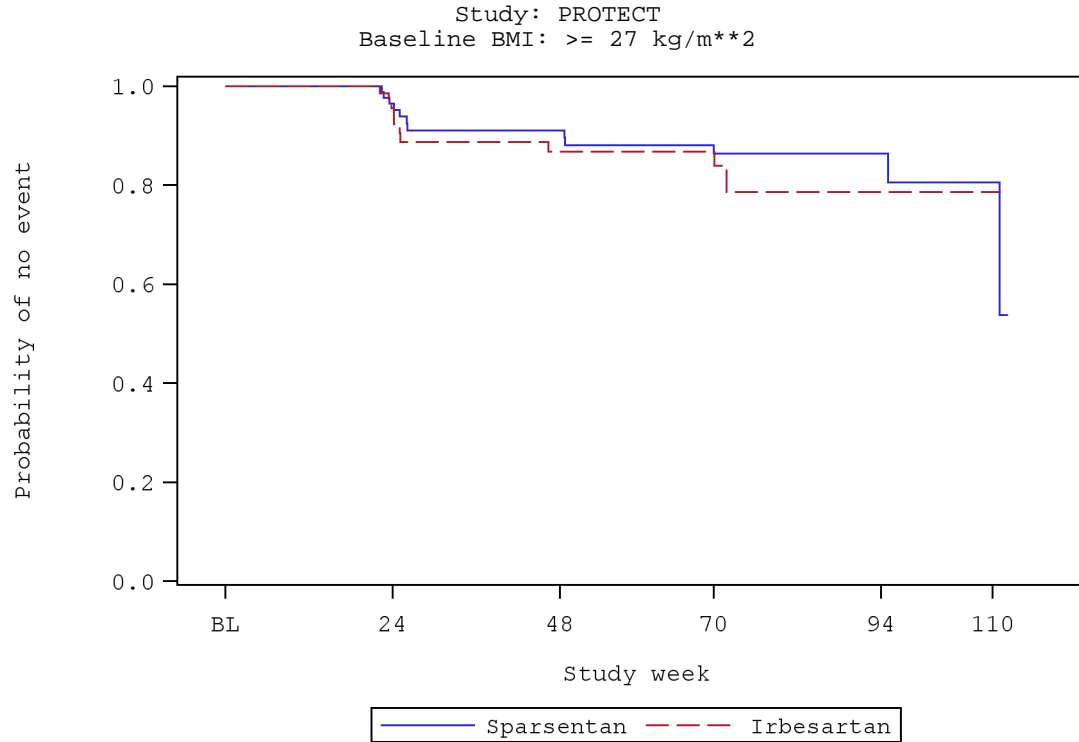
Study: PROTECT  
 Baseline BMI: < 27 kg/m\*\*2



Sparsentan	84	56	45	30	12	5
Irbesartan	94	55	39	24	9	4

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1VSDT\_FSTM

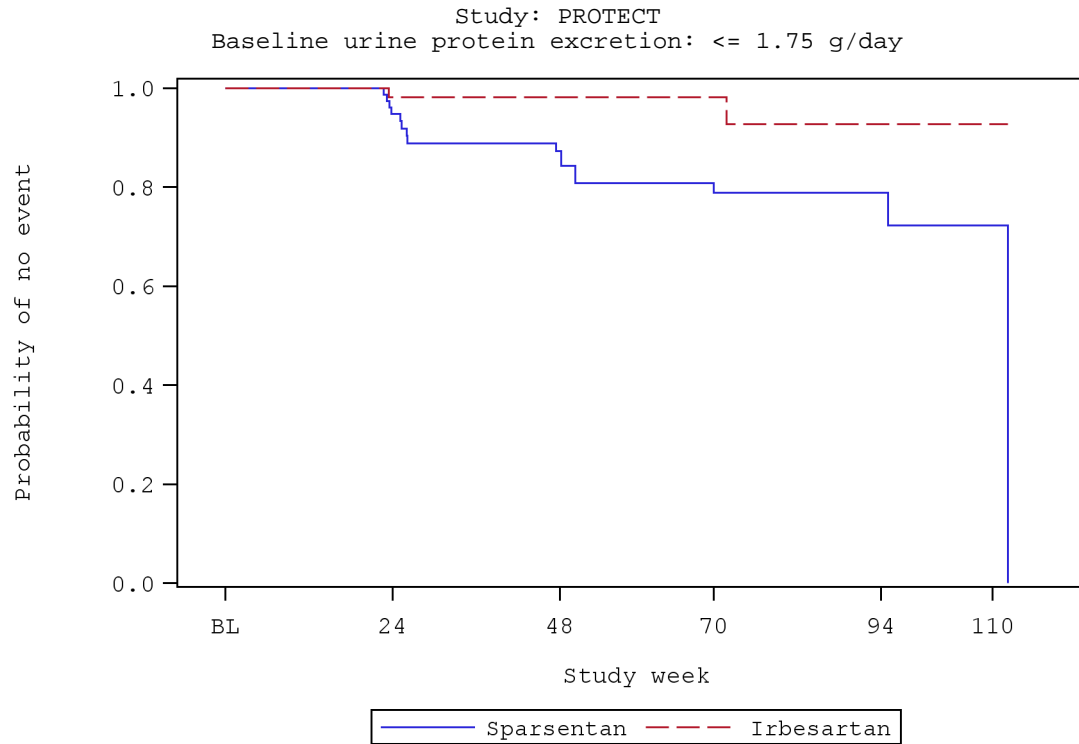
Figure PF1VSDT\_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	118	81	64	49	23	8
Irbesartan	107	63	42	30	11	4

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1VSDT\_FSTM

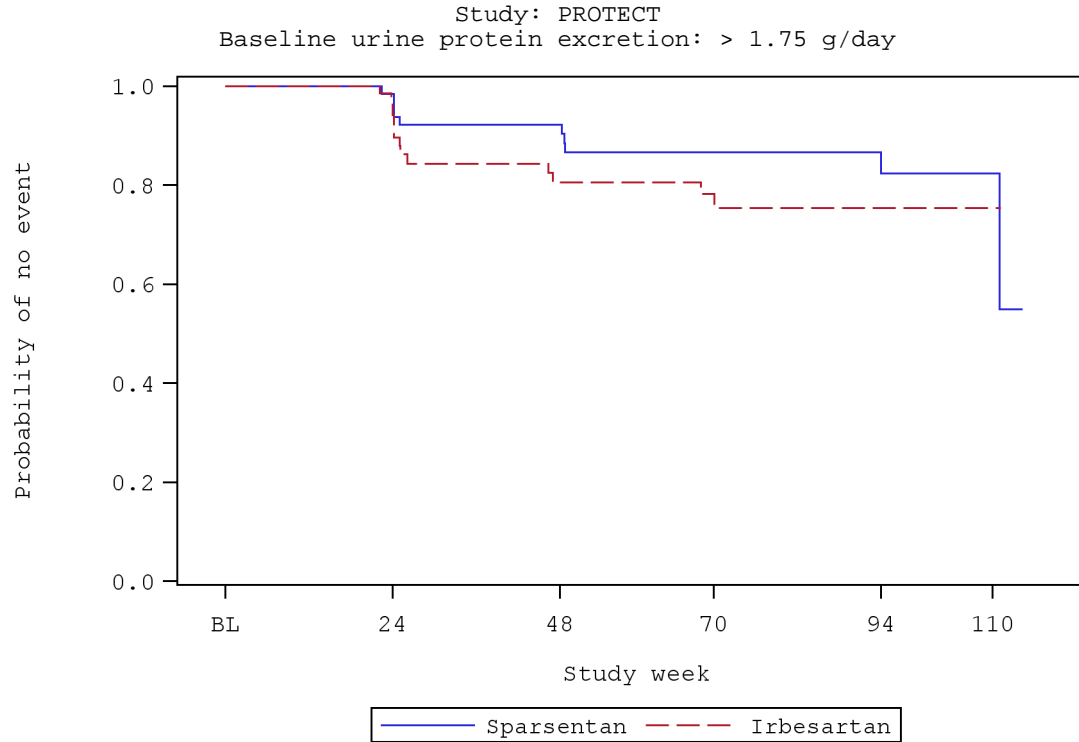
Figure PF1VSDT\_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	98	73	58	40	15	8
Irbesartan	94	53	40	27	10	5

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1VSDT\_FSTM

Figure PF1VSDT\_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	104	64	51	39	20	5
Irbesartan	108	66	42	28	11	3

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1VSDT\_FSTM



Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Sex												
Male	KDQOL: burden of kidney disease	Baseline	Sparsentan	139	130 (93.5)	71.68 (23.02)	6.3	62.50	75.00	87.50	100.0	
			Irbesartan	143	128 (89.5)	79.25 (22.24)	6.3	68.75	87.50	100.00	100.0	
		Week 24	Sparsentan	139	87 (62.6)	75.86 (21.70)	12.5	68.75	81.25	93.75	100.0	
			Irbesartan	143	73 (51.0)	78.00 (22.99)	0.0	68.75	81.25	100.00	100.0	
		Week 48	Sparsentan	139	72 (51.8)	78.04 (20.34)	6.3	71.88	81.25	93.75	100.0	
			Irbesartan	143	55 (38.5)	83.75 (18.66)	25.0	75.00	93.75	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	139	87 (62.6)	6.11 (21.80)	-50.0	-6.25	6.25	18.75	62.5	0.43 [0.12, 0.75]
			Irbesartan	143	73 (51.0)	-2.65 (18.40)	-50.0	-12.50	0.00	6.25	50.0	
		Week 48	Sparsentan	139	72 (51.8)	7.55 (19.87)	-62.5	0.00	6.25	25.00	43.8	0.27 [-0.08, 0.62]
			Irbesartan	143	55 (38.5)	2.50 (16.99)	-43.8	-6.25	0.00	6.25	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Female	KDQOL: burden of kidney disease	Baseline	Sparsentan	63	56 (88.9)	64.51 (30.01)	0.0	46.88	68.75	93.75	100.0	
			Irbesartan	59	51 (86.4)	70.83 (23.31)	18.8	56.25	75.00	87.50	100.0	
		Week 24	Sparsentan	63	38 (60.3)	74.84 (22.95)	18.8	62.50	81.25	93.75	100.0	
			Irbesartan	59	31 (52.5)	80.85 (20.22)	25.0	75.00	81.25	100.00	100.0	
		Week 48	Sparsentan	63	38 (60.3)	75.99 (22.21)	31.3	62.50	81.25	93.75	100.0	
			Irbesartan	59	22 (37.3)	73.30 (22.59)	6.3	56.25	75.00	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	63	38 (60.3)	11.51 (25.27)	-81.3	0.00	6.25	31.25	75.0	0.26 [-0.22, 0.73]
			Irbesartan	59	31 (52.5)	6.05 (14.57)	-18.8	0.00	0.00	18.75	56.3	
		Week 48	Sparsentan	63	38 (60.3)	12.50 (21.36)	-25.0	0.00	6.25	25.00	62.5	0.44 [-0.09, 0.97]
			Irbesartan	59	22 (37.3)	3.69 (17.75)	-50.0	0.00	0.00	12.50	37.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age												
<= 45 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	96	87 (90.6)	66.31 (27.25)	0.0	50.00	68.75	87.50	100.0	
			Irbesartan	99	92 (92.9)	71.20 (25.00)	6.3	53.13	75.00	93.75	100.0	
		Week 24	Sparsentan	96	57 (59.4)	74.45 (23.66)	18.8	62.50	81.25	93.75	100.0	
			Irbesartan	99	53 (53.5)	74.29 (26.19)	0.0	62.50	81.25	93.75	100.0	
		Week 48	Sparsentan	96	53 (55.2)	72.76 (24.02)	6.3	56.25	75.00	93.75	100.0	
			Irbesartan	99	40 (40.4)	78.75 (24.83)	6.3	62.50	87.50	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	96	57 (59.4)	9.87 (27.95)	-81.3	0.00	6.25	25.00	75.0	0.37 [-0.01, 0.75]
			Irbesartan	99	53 (53.5)	0.71 (20.79)	-50.0	-12.50	0.00	12.50	56.3	
		Week 48	Sparsentan	96	53 (55.2)	8.96 (21.60)	-62.5	0.00	6.25	25.00	50.0	0.29 [-0.13, 0.70]
			Irbesartan	99	40 (40.4)	3.28 (17.39)	-43.8	-6.25	0.00	9.38	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 45 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	106	99 (93.4)	72.35 (23.56)	6.3	56.25	75.00	87.50	100.0	
			Irbesartan	103	87 (84.5)	82.83 (18.56)	18.8	75.00	87.50	100.00	100.0	
		Week 24	Sparsentan	106	68 (64.2)	76.47 (20.63)	12.5	68.75	81.25	93.75	100.0	
			Irbesartan	103	51 (49.5)	83.58 (15.86)	43.8	75.00	87.50	100.00	100.0	
		Week 48	Sparsentan	106	57 (53.8)	81.58 (16.68)	37.5	75.00	81.25	93.75	100.0	
			Irbesartan	103	37 (35.9)	82.94 (13.79)	50.0	75.00	81.25	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	106	68 (64.2)	5.97 (17.72)	-43.8	-3.13	0.00	18.75	43.8	0.42 [0.05, 0.79]
			Irbesartan	103	51 (49.5)	-0.86 (14.03)	-31.3	-12.50	0.00	6.25	37.5	
		Week 48	Sparsentan	106	57 (53.8)	9.54 (19.48)	-50.0	0.00	6.25	25.00	62.5	0.39 [-0.03, 0.80]
			Irbesartan	103	37 (35.9)	2.36 (17.01)	-50.0	-6.25	0.00	12.50	43.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age at IgAN diagnosis												
<= 18 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	9	9 (100.0)	58.33 (30.94)	6.3	37.50	62.50	68.75	100.0	
			Irbesartan	5	5 (100.0)	72.50 (31.12)	18.8	75.00	81.25	93.75	93.8	
		Week 24	Sparsentan	9	3 (33.3)	95.83 (7.22)	87.5	87.50	100.00	100.00	100.0	
			Irbesartan	5	2 (40.0)	56.25 (35.36)	31.3	31.25	56.25	81.25	81.3	
		Week 48	Sparsentan	9	5 (55.6)	61.25 (40.84)	6.3	31.25	75.00	93.75	100.0	
			Irbesartan	5	1 (20.0)	81.25	81.3	81.25	81.25	81.25	81.3	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	9	3 (33.3)	27.08 (25.26)	0.0	0.00	31.25	50.00	50.0	0.98 [-0.91, 2.87]
			Irbesartan	5	2 (40.0)	6.25 (8.84)	0.0	0.00	6.25	12.50	12.5	
		Week 48	Sparsentan	9	5 (55.6)	-1.25 (38.63)	-62.5	-6.25	0.00	25.00	37.5	NE
			Irbesartan	5	1 (20.0)	0.00	0.0	0.00	0.00	0.00	0.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 18 to 40 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	102	93 (91.2)	69.76 (25.21)	0.0	62.50	75.00	87.50	100.0	
			Irbesartan	109	99 (90.8)	73.23 (24.39)	6.3	56.25	75.00	93.75	100.0	
		Week 24	Sparsentan	102	62 (60.8)	74.70 (22.67)	18.8	62.50	81.25	93.75	100.0	
			Irbesartan	109	59 (54.1)	76.48 (24.87)	0.0	62.50	81.25	100.00	100.0	
		Week 48	Sparsentan	102	57 (55.9)	75.00 (21.26)	25.0	62.50	81.25	93.75	100.0	
			Irbesartan	109	44 (40.4)	78.84 (24.01)	6.3	65.63	87.50	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	102	62 (60.8)	8.57 (26.64)	-81.3	0.00	6.25	25.00	75.0	0.35 [-0.01, 0.71]
			Irbesartan	109	59 (54.1)	0.32 (19.85)	-50.0	-6.25	0.00	6.25	56.3	
		Week 48	Sparsentan	102	57 (55.9)	8.44 (21.05)	-50.0	0.00	0.00	25.00	50.0	0.25 [-0.15, 0.64]
			Irbesartan	109	44 (40.4)	3.69 (16.74)	-43.8	-6.25	0.00	12.50	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	91	84 (92.3)	70.46 (25.18)	6.3	50.00	75.00	87.50	100.0	
			Irbesartan	88	75 (85.2)	81.92 (19.13)	18.8	75.00	87.50	100.00	100.0	
		Week 24	Sparsentan	91	60 (65.9)	75.42 (21.49)	12.5	68.75	78.13	93.75	100.0	
			Irbesartan	88	43 (48.9)	83.14 (16.34)	43.8	75.00	81.25	100.00	100.0	
		Week 48	Sparsentan	91	48 (52.7)	81.77 (16.75)	37.5	75.00	84.38	96.88	100.0	
			Irbesartan	88	32 (36.4)	83.40 (13.99)	50.0	75.00	81.25	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	91	60 (65.9)	5.94 (18.10)	-43.8	-6.25	0.00	18.75	43.8	0.40 [0.01, 0.80]
			Irbesartan	88	43 (48.9)	-0.87 (14.91)	-31.3	-12.50	0.00	6.25	37.5	
		Week 48	Sparsentan	91	48 (52.7)	11.33 (17.23)	-31.3	0.00	6.25	25.00	62.5	0.54 [0.09, 1.00]
			Irbesartan	88	32 (36.4)	1.76 (18.05)	-50.0	-6.25	0.00	9.38	43.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Geographic region												
North America	KDQOL: burden of kidney disease	Baseline	Sparsentan	35	32 (91.4)	70.70 (26.91)	6.3	59.38	75.00	93.75	100.0	
			Irbesartan	46	43 (93.5)	82.99 (17.11)	43.8	68.75	81.25	100.00	100.0	
		Week 24	Sparsentan	35	15 (42.9)	80.42 (19.32)	25.0	75.00	81.25	93.75	100.0	
			Irbesartan	46	26 (56.5)	80.53 (23.07)	6.3	68.75	87.50	100.00	100.0	
		Week 48	Sparsentan	35	13 (37.1)	76.44 (23.41)	31.3	68.75	81.25	93.75	100.0	
			Irbesartan	46	17 (37.0)	80.51 (20.24)	25.0	68.75	81.25	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	35	15 (42.9)	2.50 (19.45)	-37.5	-12.50	0.00	12.50	31.3	0.35 [-0.29, 0.99]
			Irbesartan	46	26 (56.5)	-3.61 (16.02)	-37.5	-12.50	-6.25	6.25	31.3	
		Week 48	Sparsentan	35	13 (37.1)	11.54 (14.62)	-12.5	0.00	12.50	25.00	37.5	0.92 [0.16, 1.67]
			Irbesartan	46	17 (37.0)	-3.68 (17.96)	-43.8	-6.25	0.00	6.25	37.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Europe	KDQOL: burden of kidney disease	Baseline	Sparsentan	98	87 (88.8)	75.86 (21.97)	6.3	68.75	81.25	93.75	100.0	
			Irbesartan	115	96 (83.5)	75.20 (24.19)	18.8	56.25	87.50	93.75	100.0	
		Week 24	Sparsentan	98	53 (54.1)	78.18 (21.25)	12.5	75.00	81.25	93.75	100.0	
			Irbesartan	115	47 (40.9)	78.99 (19.04)	31.3	68.75	81.25	93.75	100.0	
		Week 48	Sparsentan	98	44 (44.9)	79.12 (19.94)	6.3	75.00	81.25	93.75	100.0	
			Irbesartan	115	38 (33.0)	82.40 (18.77)	37.5	68.75	87.50	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	98	53 (54.1)	3.54 (26.02)	-81.3	-6.25	0.00	12.50	75.0	0.16 [-0.23, 0.55]
			Irbesartan	115	47 (40.9)	-0.13 (19.17)	-50.0	-12.50	0.00	12.50	56.3	
		Week 48	Sparsentan	98	44 (44.9)	5.11 (20.85)	-62.5	-3.13	0.00	18.75	50.0	0.04 [-0.40, 0.47]
			Irbesartan	115	38 (33.0)	4.44 (15.44)	-50.0	0.00	0.00	12.50	31.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Asia Pacific	KDQOL: burden of kidney disease	Baseline	Sparsentan	69	67 (97.1)	60.73 (26.76)	0.0	43.75	62.50	87.50	100.0	
			Irbesartan	41	40 (97.6)	74.22 (24.03)	6.3	62.50	75.00	93.75	100.0	
		Week 24	Sparsentan	69	57 (82.6)	71.82 (23.06)	12.5	56.25	75.00	93.75	100.0	
			Irbesartan	41	31 (75.6)	77.22 (26.05)	0.0	68.75	81.25	100.00	100.0	
		Week 48	Sparsentan	69	53 (76.8)	76.06 (21.40)	25.0	62.50	75.00	93.75	100.0	
			Irbesartan	41	22 (53.7)	78.13 (23.29)	6.3	75.00	81.25	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	69	57 (82.6)	13.05 (19.74)	-25.0	0.00	6.25	31.25	62.5	0.53 [0.09, 0.98]
			Irbesartan	41	31 (75.6)	3.02 (16.76)	-37.5	0.00	0.00	6.25	50.0	
		Week 48	Sparsentan	69	53 (76.8)	12.15 (21.03)	-50.0	0.00	6.25	25.00	62.5	0.35 [-0.16, 0.85]
			Irbesartan	41	22 (53.7)	5.11 (18.66)	-25.0	0.00	0.00	6.25	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline BMI												
< 27 kg/m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	84	76 (90.5)	64.47 (25.88)	6.3	46.88	68.75	87.50	100.0	
			Irbesartan	94	84 (89.4)	72.77 (22.37)	18.8	56.25	75.00	93.75	100.0	
		Week 24	Sparsentan	84	52 (61.9)	73.08 (22.91)	12.5	68.75	75.00	87.50	100.0	
			Irbesartan	94	50 (53.2)	76.38 (23.09)	6.3	68.75	81.25	93.75	100.0	
		Week 48	Sparsentan	84	47 (56.0)	74.07 (21.69)	25.0	62.50	75.00	93.75	100.0	
			Irbesartan	94	36 (38.3)	75.17 (22.28)	6.3	65.63	81.25	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	84	52 (61.9)	9.25 (22.47)	-43.8	-3.13	6.25	21.88	62.5	0.23 [-0.16, 0.62]
			Irbesartan	94	50 (53.2)	4.38 (19.28)	-37.5	-12.50	0.00	12.50	56.3	
		Week 48	Sparsentan	84	47 (56.0)	10.90 (22.02)	-50.0	0.00	6.25	25.00	62.5	0.34 [-0.09, 0.78]
			Irbesartan	94	36 (38.3)	3.65 (20.07)	-50.0	-6.25	0.00	15.63	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 27 kg/m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	118	110 (93.2)	73.01 (24.68)	0.0	62.50	75.00	93.75	100.0	
			Irbesartan	107	94 (87.9)	81.12 (21.88)	6.3	68.75	87.50	100.00	100.0	
		Week 24	Sparsentan	118	73 (61.9)	77.31 (21.31)	12.5	68.75	81.25	93.75	100.0	
			Irbesartan	107	54 (50.5)	81.13 (21.19)	0.0	68.75	87.50	100.00	100.0	
		Week 48	Sparsentan	118	63 (53.4)	79.76 (20.17)	6.3	75.00	81.25	100.00	100.0	
			Irbesartan	107	41 (38.3)	85.67 (17.13)	31.3	75.00	93.75	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	118	73 (61.9)	6.68 (23.37)	-81.3	0.00	6.25	18.75	75.0	0.53 [0.18, 0.89]
			Irbesartan	107	54 (50.5)	-4.17 (15.21)	-50.0	-12.50	0.00	6.25	31.3	
		Week 48	Sparsentan	118	63 (53.4)	8.04 (19.26)	-62.5	0.00	6.25	25.00	50.0	0.34 [-0.06, 0.73]
			Irbesartan	107	41 (38.3)	2.13 (14.23)	-31.3	-6.25	0.00	6.25	43.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Randomization strata												
eGFR Low and UP High	KDQOL: burden of kidney disease	Baseline	Sparsentan	71	63 (88.7)	69.64 (25.02)	6.3	50.00	75.00	87.50	100.0	
			Irbesartan	74	63 (85.1)	72.82 (24.13)	18.8	56.25	75.00	93.75	100.0	
		Week 24	Sparsentan	71	44 (62.0)	75.71 (22.20)	12.5	65.63	81.25	93.75	100.0	
			Irbesartan	74	32 (43.2)	72.85 (25.48)	6.3	59.38	78.13	96.88	100.0	
		Week 48	Sparsentan	71	37 (52.1)	77.53 (19.79)	25.0	68.75	75.00	93.75	100.0	
			Irbesartan	74	22 (29.7)	79.55 (23.00)	6.3	75.00	81.25	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	71	44 (62.0)	6.96 (23.78)	-50.0	-6.25	6.25	25.00	62.5	0.54 [0.08, 1.01]
			Irbesartan	74	32 (43.2)	-3.91 (13.16)	-37.5	-12.50	-3.13	0.00	31.3	
		Week 48	Sparsentan	71	37 (52.1)	7.94 (19.52)	-50.0	-6.25	6.25	25.00	43.8	0.42 [-0.11, 0.96]
			Irbesartan	74	22 (29.7)	0.57 (13.21)	-25.0	-6.25	0.00	6.25	31.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR Low and UP Low	KDQOL: burden of kidney disease	Baseline	Sparsentan	55	51 (92.7)	66.91 (25.47)	0.0	50.00	68.75	87.50	100.0	
			Irbesartan	55	49 (89.1)	78.44 (22.28)	25.0	62.50	81.25	100.00	100.0	
		Week 24	Sparsentan	55	38 (69.1)	72.70 (24.07)	12.5	56.25	78.13	93.75	100.0	
			Irbesartan	55	33 (60.0)	83.52 (15.92)	50.0	68.75	87.50	100.00	100.0	
		Week 48	Sparsentan	55	28 (50.9)	82.37 (17.52)	37.5	75.00	81.25	100.00	100.0	
			Irbesartan	55	25 (45.5)	80.75 (18.83)	31.3	75.00	81.25	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	55	38 (69.1)	6.25 (24.10)	-81.3	0.00	0.00	18.75	62.5	0.11 [-0.36, 0.57]
			Irbesartan	55	33 (60.0)	3.98 (17.94)	-31.3	-6.25	0.00	12.50	50.0	
		Week 48	Sparsentan	55	28 (50.9)	12.95 (18.32)	-25.0	0.00	6.25	25.00	62.5	0.34 [-0.20, 0.88]
			Irbesartan	55	25 (45.5)	6.25 (21.19)	-50.0	-6.25	0.00	18.75	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP High	KDQOL: burden of kidney disease	Baseline	Sparsentan	37	35 (94.6)	67.68 (27.47)	6.3	50.00	75.00	87.50	100.0	
			Irbesartan	36	31 (86.1)	77.22 (21.32)	31.3	62.50	81.25	93.75	100.0	
		Week 24	Sparsentan	37	19 (51.4)	73.68 (19.83)	25.0	68.75	75.00	87.50	100.0	
			Irbesartan	36	18 (50.0)	77.78 (20.81)	31.3	68.75	81.25	93.75	100.0	
		Week 48	Sparsentan	37	22 (59.5)	63.92 (25.07)	6.3	50.00	71.88	81.25	100.0	
			Irbesartan	36	11 (30.6)	80.68 (18.64)	43.8	68.75	81.25	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	37	19 (51.4)	13.49 (25.28)	-31.3	-6.25	6.25	25.00	75.0	0.68 [0.01, 1.34]
			Irbesartan	36	18 (50.0)	-2.43 (21.56)	-50.0	-6.25	-3.13	6.25	50.0	
		Week 48	Sparsentan	37	22 (59.5)	5.40 (25.75)	-62.5	0.00	0.00	25.00	50.0	-0.04 [-0.76, 0.69]
			Irbesartan	36	11 (30.6)	6.25 (13.69)	-6.3	0.00	0.00	12.50	37.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP Low	KDQOL: burden of kidney disease	Baseline	Sparsentan	39	37 (94.9)	74.66 (24.47)	0.0	62.50	81.25	93.75	100.0	
			Irbesartan	37	36 (97.3)	81.42 (22.08)	6.3	75.00	90.63	96.88	100.0	
		Week 24	Sparsentan	39	24 (61.5)	81.25 (19.94)	25.0	71.88	84.38	100.00	100.0	
			Irbesartan	37	21 (56.8)	81.55 (25.35)	0.0	75.00	93.75	100.00	100.0	
		Week 48	Sparsentan	39	23 (59.0)	83.70 (17.24)	37.5	81.25	87.50	93.75	100.0	
			Irbesartan	37	19 (51.4)	82.24 (21.17)	25.0	68.75	93.75	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	39	24 (61.5)	7.03 (17.71)	-18.8	-6.25	3.13	15.63	43.8	0.30 [-0.29, 0.89]
			Irbesartan	37	21 (56.8)	1.49 (19.56)	-31.3	-12.50	0.00	6.25	56.3	
		Week 48	Sparsentan	39	23 (59.0)	10.60 (19.07)	-31.3	0.00	6.25	25.00	50.0	0.64 [0.02, 1.26]
			Irbesartan	37	19 (51.4)	-0.99 (16.83)	-43.8	-6.25	0.00	6.25	31.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline eGFR Group 1												
< 60 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	127	115 (90.6)	69.24 (25.44)	0.0	50.00	75.00	87.50	100.0	
			Irbesartan	129	113 (87.6)	77.21 (22.84)	18.8	62.50	81.25	100.00	100.0	
		Week 24	Sparsentan	127	83 (65.4)	74.32 (22.87)	12.5	62.50	81.25	93.75	100.0	
			Irbesartan	129	68 (52.7)	78.31 (21.74)	6.3	68.75	81.25	100.00	100.0	
		Week 48	Sparsentan	127	65 (51.2)	79.52 (18.90)	25.0	68.75	81.25	93.75	100.0	
			Irbesartan	129	49 (38.0)	80.99 (20.41)	6.3	75.00	81.25	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	127	83 (65.4)	5.80 (23.70)	-81.3	-6.25	0.00	18.75	62.5	0.33 [0.01, 0.66]
			Irbesartan	129	68 (52.7)	-1.01 (15.60)	-37.5	-12.50	0.00	6.25	50.0	
		Week 48	Sparsentan	127	65 (51.2)	9.33 (18.92)	-50.0	0.00	6.25	25.00	62.5	0.37 [-0.01, 0.74]
			Irbesartan	129	49 (38.0)	2.55 (18.04)	-50.0	-6.25	0.00	12.50	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	49	47 (95.9)	72.61 (23.91)	0.0	62.50	75.00	87.50	100.0	
			Irbesartan	48	43 (89.6)	77.91 (21.45)	18.8	62.50	87.50	93.75	100.0	
		Week 24	Sparsentan	49	27 (55.1)	75.23 (20.47)	25.0	68.75	81.25	87.50	100.0	
			Irbesartan	48	24 (50.0)	84.11 (17.77)	37.5	75.00	90.63	100.00	100.0	
		Week 48	Sparsentan	49	29 (59.2)	75.22 (23.89)	6.3	75.00	81.25	87.50	100.0	
			Irbesartan	48	19 (39.6)	78.29 (20.66)	25.0	68.75	81.25	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	49	27 (55.1)	8.33 (19.69)	-31.3	-6.25	6.25	25.00	50.0	0.29 [-0.26, 0.84]
			Irbesartan	48	24 (50.0)	2.86 (18.06)	-25.0	-9.38	0.00	15.63	50.0	
		Week 48	Sparsentan	49	29 (59.2)	8.62 (23.41)	-62.5	0.00	6.25	25.00	50.0	0.30 [-0.28, 0.88]
			Irbesartan	48	19 (39.6)	2.30 (17.46)	-43.8	0.00	0.00	12.50	37.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	26	24 (92.3)	64.84 (28.66)	6.3	50.00	75.00	87.50	100.0	
			Irbesartan	25	23 (92.0)	73.10 (25.59)	6.3	56.25	75.00	93.75	100.0	
		Week 24	Sparsentan	26	15 (57.7)	82.92 (19.40)	31.3	75.00	87.50	100.00	100.0	
			Irbesartan	25	12 (48.0)	71.35 (30.44)	0.0	50.00	81.25	93.75	100.0	
		Week 48	Sparsentan	26	16 (61.5)	72.27 (23.16)	31.3	56.25	75.00	93.75	100.0	
	Irbesartan		25	9 (36.0)	84.72 (20.28)	43.8	75.00	93.75	100.00	100.0		
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	26	15 (57.7)	17.50 (22.93)	-6.3	6.25	6.25	31.25	75.0	0.72 [-0.06, 1.50]
			Irbesartan	25	12 (48.0)	-0.52 (27.50)	-50.0	-9.38	-6.25	12.50	56.3	
		Week 48	Sparsentan	26	16 (61.5)	10.16 (21.99)	-31.3	-3.13	6.25	21.88	50.0	0.24 [-0.58, 1.06]
			Irbesartan	25	9 (36.0)	5.56 (11.46)	-6.3	0.00	6.25	6.25	31.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline eGFR Group 2												
< 45 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	82	71 (86.6)	66.55 (24.96)	6.3	50.00	68.75	87.50	100.0	
			Irbesartan	80	69 (86.3)	73.82 (24.05)	18.8	62.50	81.25	93.75	100.0	
		Week 24	Sparsentan	82	53 (64.6)	73.70 (24.46)	12.5	62.50	81.25	93.75	100.0	
			Irbesartan	80	45 (56.3)	77.36 (23.70)	6.3	62.50	81.25	100.00	100.0	
		Week 48	Sparsentan	82	42 (51.2)	79.32 (20.26)	25.0	75.00	81.25	93.75	100.0	
			Irbesartan	80	29 (36.3)	80.39 (22.21)	6.3	75.00	81.25	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	82	53 (64.6)	6.37 (23.50)	-81.3	0.00	0.00	12.50	62.5	0.28 [-0.12, 0.68]
			Irbesartan	80	45 (56.3)	0.83 (14.75)	-37.5	-6.25	0.00	12.50	37.5	
		Week 48	Sparsentan	82	42 (51.2)	10.57 (20.82)	-50.0	0.00	6.25	25.00	62.5	0.36 [-0.12, 0.83]
			Irbesartan	80	29 (36.3)	4.09 (13.50)	-18.8	-6.25	0.00	6.25	43.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
45 to < 60 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	45	44 (97.8)	73.58 (25.89)	0.0	56.25	81.25	93.75	100.0	
			Irbesartan	49	44 (89.8)	82.53 (19.93)	25.0	68.75	90.63	100.00	100.0	
		Week 24	Sparsentan	45	30 (66.7)	75.42 (20.10)	25.0	62.50	75.00	93.75	100.0	
			Irbesartan	49	23 (46.9)	80.16 (17.64)	31.3	68.75	81.25	100.00	100.0	
		Week 48	Sparsentan	45	23 (51.1)	79.89 (16.53)	50.0	68.75	75.00	100.00	100.0	
			Irbesartan	49	20 (40.8)	81.88 (18.01)	31.3	75.00	81.25	96.88	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	45	30 (66.7)	4.79 (24.44)	-50.0	-6.25	0.00	25.00	50.0	0.44 [-0.11, 0.99]
			Irbesartan	49	23 (46.9)	-4.62 (16.88)	-37.5	-12.50	0.00	0.00	50.0	
		Week 48	Sparsentan	45	23 (51.1)	7.07 (14.99)	-25.0	0.00	0.00	25.00	37.5	0.35 [-0.25, 0.95]
			Irbesartan	49	20 (40.8)	0.31 (23.34)	-50.0	-6.25	0.00	12.50	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	49	47 (95.9)	72.61 (23.91)	0.0	62.50	75.00	87.50	100.0	
			Irbesartan	48	43 (89.6)	77.91 (21.45)	18.8	62.50	87.50	93.75	100.0	
		Week 24	Sparsentan	49	27 (55.1)	75.23 (20.47)	25.0	68.75	81.25	87.50	100.0	
			Irbesartan	48	24 (50.0)	84.11 (17.77)	37.5	75.00	90.63	100.00	100.0	
		Week 48	Sparsentan	49	29 (59.2)	75.22 (23.89)	6.3	75.00	81.25	87.50	100.0	
			Irbesartan	48	19 (39.6)	78.29 (20.66)	25.0	68.75	81.25	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	49	27 (55.1)	8.33 (19.69)	-31.3	-6.25	6.25	25.00	50.0	0.29 [-0.26, 0.84]
			Irbesartan	48	24 (50.0)	2.86 (18.06)	-25.0	-9.38	0.00	15.63	50.0	
		Week 48	Sparsentan	49	29 (59.2)	8.62 (23.41)	-62.5	0.00	6.25	25.00	50.0	0.30 [-0.28, 0.88]
			Irbesartan	48	19 (39.6)	2.30 (17.46)	-43.8	0.00	0.00	12.50	37.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	26	24 (92.3)	64.84 (28.66)	6.3	50.00	75.00	87.50	100.0	
			Irbesartan	25	23 (92.0)	73.10 (25.59)	6.3	56.25	75.00	93.75	100.0	
		Week 24	Sparsentan	26	15 (57.7)	82.92 (19.40)	31.3	75.00	87.50	100.00	100.0	
			Irbesartan	25	12 (48.0)	71.35 (30.44)	0.0	50.00	81.25	93.75	100.0	
		Week 48	Sparsentan	26	16 (61.5)	72.27 (23.16)	31.3	56.25	75.00	93.75	100.0	
			Irbesartan	25	9 (36.0)	84.72 (20.28)	43.8	75.00	93.75	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	26	15 (57.7)	17.50 (22.93)	-6.3	6.25	6.25	31.25	75.0	0.72 [-0.06, 1.50]
			Irbesartan	25	12 (48.0)	-0.52 (27.50)	-50.0	-9.38	-6.25	12.50	56.3	
		Week 48	Sparsentan	26	16 (61.5)	10.16 (21.99)	-31.3	-3.13	6.25	21.88	50.0	0.24 [-0.58, 1.06]
			Irbesartan	25	9 (36.0)	5.56 (11.46)	-6.3	0.00	6.25	6.25	31.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline urine protein excretion												
<= 1.75 g/day	KDQOL: burden of kidney disease	Baseline	Sparsentan	98	92 (93.9)	69.43 (24.97)	0.0	53.13	75.00	87.50	100.0	
			Irbesartan	94	82 (87.2)	78.35 (21.77)	6.3	68.75	87.50	93.75	100.0	
		Week 24	Sparsentan	98	66 (67.3)	75.38 (23.89)	12.5	68.75	81.25	93.75	100.0	
			Irbesartan	94	43 (45.7)	82.99 (21.23)	0.0	68.75	93.75	100.00	100.0	
		Week 48	Sparsentan	98	59 (60.2)	81.14 (18.51)	31.3	75.00	81.25	93.75	100.0	
			Irbesartan	94	34 (36.2)	81.80 (19.96)	25.0	75.00	84.38	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	98	66 (67.3)	8.14 (23.28)	-81.3	0.00	6.25	25.00	62.5	0.27 [-0.12, 0.65]
			Irbesartan	94	43 (45.7)	2.47 (17.47)	-31.3	-6.25	0.00	12.50	50.0	
		Week 48	Sparsentan	98	59 (60.2)	11.12 (18.75)	-31.3	0.00	6.25	25.00	62.5	0.43 [-0.00, 0.85]
			Irbesartan	94	34 (36.2)	2.94 (20.08)	-50.0	0.00	0.00	12.50	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 1.75 g/day	KDQOL: burden of kidney disease	Baseline	Sparsentan	104	94 (90.4)	69.61 (26.06)	6.3	50.00	75.00	87.50	100.0	
			Irbesartan	108	97 (89.8)	75.58 (23.68)	18.8	62.50	81.25	100.00	100.0	
		Week 24	Sparsentan	104	59 (56.7)	75.74 (19.87)	12.5	68.75	81.25	87.50	100.0	
			Irbesartan	108	61 (56.5)	75.92 (22.47)	6.3	62.50	81.25	93.75	100.0	
		Week 48	Sparsentan	104	51 (49.0)	72.92 (22.80)	6.3	56.25	75.00	93.75	100.0	
			Irbesartan	108	43 (39.8)	79.94 (20.71)	6.3	68.75	81.25	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	104	59 (56.7)	7.31 (22.75)	-50.0	-6.25	6.25	18.75	75.0	0.45 [0.09, 0.81]
			Irbesartan	108	61 (56.5)	-1.84 (17.84)	-50.0	-12.50	0.00	6.25	56.3	
		Week 48	Sparsentan	104	51 (49.0)	7.11 (22.22)	-62.5	0.00	6.25	25.00	50.0	0.23 [-0.18, 0.63]
			Irbesartan	108	43 (39.8)	2.76 (14.58)	-31.3	-6.25	0.00	6.25	37.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline use of antihypertensives												
Yes	KDQOL: burden of kidney disease	Baseline	Sparsentan	88	77 (87.5)	69.81 (25.01)	6.3	56.25	75.00	87.50	100.0	
			Irbesartan	83	70 (84.3)	77.50 (21.87)	18.8	62.50	84.38	100.00	100.0	
		Week 24	Sparsentan	88	50 (56.8)	79.13 (19.63)	12.5	68.75	81.25	93.75	100.0	
			Irbesartan	83	39 (47.0)	79.17 (23.58)	6.3	62.50	87.50	100.00	100.0	
		Week 48	Sparsentan	88	41 (46.6)	80.03 (16.55)	31.3	75.00	81.25	93.75	100.0	
			Irbesartan	83	28 (33.7)	77.90 (19.43)	31.3	71.88	81.25	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	88	50 (56.8)	10.25 (21.86)	-31.3	0.00	3.13	25.00	75.0	0.58 [0.15, 1.00]
			Irbesartan	83	39 (47.0)	-0.96 (15.94)	-37.5	-12.50	0.00	6.25	37.5	
		Week 48	Sparsentan	88	41 (46.6)	12.35 (19.08)	-50.0	0.00	6.25	25.00	50.0	0.58 [0.09, 1.07]
			Irbesartan	83	28 (33.7)	1.12 (19.55)	-50.0	-6.25	0.00	9.38	43.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL: burden of kidney disease	Baseline	Sparsentan	114	109 (95.6)	69.32 (25.89)	0.0	50.00	75.00	87.50	100.0	
			Irbesartan	119	109 (91.6)	76.43 (23.47)	6.3	62.50	81.25	93.75	100.0	
		Week 24	Sparsentan	114	75 (65.8)	73.17 (23.27)	12.5	56.25	75.00	93.75	100.0	
			Irbesartan	119	65 (54.6)	78.65 (21.42)	0.0	68.75	81.25	93.75	100.0	
		Week 48	Sparsentan	114	69 (60.5)	75.72 (23.10)	6.3	62.50	81.25	93.75	100.0	
			Irbesartan	119	49 (41.2)	82.40 (20.76)	6.3	75.00	87.50	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	114	75 (65.8)	6.08 (23.63)	-81.3	-6.25	6.25	18.75	62.5	0.26 [-0.07, 0.59]
			Irbesartan	119	65 (54.6)	0.48 (18.82)	-50.0	-12.50	0.00	6.25	56.3	
		Week 48	Sparsentan	114	69 (60.5)	7.43 (21.12)	-62.5	0.00	0.00	25.00	62.5	0.19 [-0.18, 0.56]
			Irbesartan	119	49 (41.2)	3.83 (15.67)	-43.8	-6.25	0.00	12.50	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Time since renal biopsy												
<= 5 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	113	104 (92.0)	67.25 (26.79)	0.0	50.00	75.00	87.50	100.0	
			Irbesartan	127	116 (91.3)	75.54 (22.97)	6.3	56.25	81.25	93.75	100.0	
		Week 24	Sparsentan	113	70 (61.9)	75.27 (22.05)	12.5	68.75	78.13	93.75	100.0	
			Irbesartan	127	67 (52.8)	78.36 (21.69)	0.0	62.50	81.25	93.75	100.0	
		Week 48	Sparsentan	113	61 (54.0)	75.41 (20.09)	25.0	62.50	75.00	93.75	100.0	
			Irbesartan	127	51 (40.2)	78.06 (22.86)	6.3	62.50	81.25	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	113	70 (61.9)	11.96 (21.38)	-50.0	0.00	6.25	31.25	62.5	0.52 [0.18, 0.86]
			Irbesartan	127	67 (52.8)	1.49 (18.66)	-50.0	-12.50	0.00	12.50	56.3	
		Week 48	Sparsentan	113	61 (54.0)	11.48 (18.48)	-31.3	0.00	6.25	25.00	50.0	0.42 [0.05, 0.80]
			Irbesartan	127	51 (40.2)	3.43 (19.82)	-50.0	-6.25	0.00	12.50	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 5 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	89	82 (92.1)	72.41 (23.51)	0.0	62.50	75.00	87.50	100.0	
			Irbesartan	75	63 (84.0)	79.27 (22.47)	18.8	68.75	87.50	93.75	100.0	
		Week 24	Sparsentan	89	55 (61.8)	75.91 (22.13)	18.8	68.75	81.25	93.75	100.0	
			Irbesartan	75	37 (49.3)	79.73 (23.22)	6.3	68.75	81.25	100.00	100.0	
		Week 48	Sparsentan	89	49 (55.1)	79.72 (21.89)	6.3	75.00	87.50	93.75	100.0	
			Irbesartan	75	26 (34.7)	86.06 (12.66)	56.3	75.00	87.50	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	89	55 (61.8)	2.39 (23.92)	-81.3	-6.25	0.00	12.50	75.0	0.25 [-0.17, 0.67]
			Irbesartan	75	37 (49.3)	-2.87 (15.77)	-37.5	-6.25	0.00	0.00	37.5	
		Week 48	Sparsentan	89	49 (55.1)	6.51 (22.53)	-62.5	0.00	0.00	25.00	62.5	0.25 [-0.23, 0.73]
			Irbesartan	75	26 (34.7)	1.68 (10.09)	-25.0	-6.25	0.00	6.25	18.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: History of hypertension												
Yes	KDQOL: burden of kidney disease	Baseline	Sparsentan	153	138 (90.2)	68.98 (26.23)	0.0	50.00	75.00	87.50	100.0	
			Irbesartan	157	135 (86.0)	78.10 (21.82)	18.8	62.50	81.25	100.00	100.0	
		Week 24	Sparsentan	153	94 (61.4)	76.40 (21.22)	12.5	68.75	81.25	93.75	100.0	
			Irbesartan	157	76 (48.4)	79.93 (21.00)	6.3	68.75	81.25	100.00	100.0	
		Week 48	Sparsentan	153	81 (52.9)	77.93 (21.56)	6.3	68.75	81.25	93.75	100.0	
			Irbesartan	157	57 (36.3)	81.03 (18.60)	31.3	68.75	81.25	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	153	94 (61.4)	8.44 (22.11)	-43.8	-6.25	6.25	25.00	75.0	0.39 [0.08, 0.69]
			Irbesartan	157	76 (48.4)	0.49 (18.08)	-50.0	-9.38	0.00	12.50	56.3	
		Week 48	Sparsentan	153	81 (52.9)	9.57 (19.49)	-62.5	0.00	6.25	25.00	62.5	0.35 [0.01, 0.69]
			Irbesartan	157	57 (36.3)	3.07 (16.71)	-50.0	-6.25	0.00	12.50	43.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL: burden of kidney disease	Baseline	Sparsentan	49	48 (98.0)	71.09 (23.30)	12.5	53.13	75.00	87.50	100.0	
			Irbesartan	45	44 (97.8)	73.01 (25.46)	6.3	56.25	75.00	93.75	100.0	
		Week 24	Sparsentan	49	31 (63.3)	72.98 (24.39)	12.5	68.75	75.00	87.50	100.0	
			Irbesartan	45	28 (62.2)	75.89 (25.16)	0.0	68.75	78.13	96.88	100.0	
		Week 48	Sparsentan	49	29 (59.2)	75.65 (19.29)	31.3	68.75	75.00	93.75	100.0	
			Irbesartan	45	20 (44.4)	80.00 (24.97)	6.3	75.00	84.38	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	49	31 (63.3)	5.65 (25.58)	-81.3	0.00	6.25	18.75	56.3	0.33 [-0.19, 0.84]
			Irbesartan	45	28 (62.2)	-1.56 (16.98)	-31.3	-12.50	-3.13	0.00	50.0	
		Week 48	Sparsentan	49	29 (59.2)	8.41 (23.22)	-50.0	0.00	12.50	25.00	50.0	0.29 [-0.28, 0.86]
			Irbesartan	45	20 (44.4)	2.19 (18.62)	-43.8	-6.25	0.00	9.38	56.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.890
Male	Week 24	Sparsentan	139	87 (62.6)	3.81 (1.84)	(0.18, 7.43)	4.06 (2.75)	(-1.36, 9.48)	0.142
		Irbesartan	143	73 (51.0)	-0.25 (2.01)	(-4.20, 3.70)			
	Week 48	Sparsentan	139	72 (51.8)	5.79 (1.99)	(1.87, 9.72)	1.51 (3.06)	(-4.51, 7.53)	0.622
		Irbesartan	143	55 (38.5)	4.28 (2.29)	(-0.22, 8.79)			
Female	Week 24	Sparsentan	63	38 (60.3)	8.94 (2.63)	(3.73, 14.14)	-0.82 (3.97)	(-8.69, 7.05)	0.837
		Irbesartan	59	31 (52.5)	9.76 (2.94)	(3.92, 15.59)			
	Week 48	Sparsentan	63	38 (60.3)	10.51 (2.60)	(5.35, 15.67)	5.35 (4.32)	(-3.20, 13.90)	0.218
		Irbesartan	59	22 (37.3)	5.16 (3.42)	(-1.60, 11.93)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024



Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.879
<= 45 years	Week 24	Sparsentan	96	57 (59.4)	7.18 (2.57)	(2.12, 12.25)	4.04 (3.73)	(-3.32, 11.40)	0.280
		Irbesartan	99	53 (53.5)	3.14 (2.67)	(-2.13, 8.41)			
	Week 48	Sparsentan	96	53 (55.2)	6.57 (2.64)	(1.37, 11.78)	1.65 (4.01)	(-6.27, 9.57)	0.682
		Irbesartan	99	40 (40.4)	4.92 (3.00)	(-0.99, 10.84)			
> 45 years	Week 24	Sparsentan	106	68 (64.2)	3.43 (1.68)	(0.12, 6.75)	0.16 (2.64)	(-5.05, 5.36)	0.953
		Irbesartan	103	51 (49.5)	3.28 (1.98)	(-0.63, 7.18)			
	Week 48	Sparsentan	106	57 (53.8)	7.61 (1.82)	(4.02, 11.20)	3.11 (2.94)	(-2.68, 8.90)	0.291
		Irbesartan	103	37 (35.9)	4.50 (2.27)	(0.02, 8.99)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

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A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.937
<= 18 years	Week 24	Sparsentan	9	3 (33.3)	13.10 (15.77)	(-30.17, 56.37)	12.51 (30.95)	(-73.20, 98.22)	0.707
		Irbesartan	5	2 (40.0)	0.59 (26.30)	(-72.51, 73.68)			
	Week 48	Sparsentan	9	5 (55.6)	3.45 (15.59)	(-39.81, 46.70)	2.79 (31.02)	(-82.91, 88.49)	0.933
		Irbesartan	5	1 (20.0)	0.66 (26.53)	(-72.42, 73.74)			
> 18 to 40 years	Week 24	Sparsentan	102	62 (60.8)	6.55 (2.32)	(1.97, 11.12)	3.90 (3.36)	(-2.72, 10.51)	0.247
		Irbesartan	109	59 (54.1)	2.65 (2.40)	(-2.08, 7.38)			
	Week 48	Sparsentan	102	57 (55.9)	6.83 (2.41)	(2.07, 11.58)	2.58 (3.66)	(-4.65, 9.80)	0.483
		Irbesartan	109	44 (40.4)	4.25 (2.73)	(-1.14, 9.63)			
> 40 years	Week 24	Sparsentan	91	60 (65.9)	3.58 (1.80)	(0.02, 7.14)	0.30 (2.83)	(-5.29, 5.89)	0.916
		Irbesartan	88	43 (48.9)	3.28 (2.14)	(-0.94, 7.50)			
	Week 48	Sparsentan	91	48 (52.7)	8.97 (1.99)	(5.05, 12.90)	4.06 (3.18)	(-2.21, 10.34)	0.203
		Irbesartan	88	32 (36.4)	4.91 (2.45)	(0.07, 9.75)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Geographic region	Overall		Sparsentan						Interaction:	0.545
North America	Week 24		Sparsentan	35	15 (42.9)	2.32 (4.19)	(-6.07, 10.71)	5.04 (5.34)	(-5.67, 15.75)	0.350
			Irbesartan	46	26 (56.5)	-2.72 (3.23)	(-9.19, 3.76)			
	Week 48		Sparsentan	35	13 (37.1)	9.31 (4.71)	(-0.11, 18.73)	12.35 (6.26)	(-0.15, 24.86)	0.053
			Irbesartan	46	17 (37.0)	-3.05 (3.92)	(-10.87, 4.78)			
Europe	Week 24		Sparsentan	98	53 (54.1)	2.57 (2.33)	(-2.03, 7.18)	1.05 (3.41)	(-5.67, 7.78)	0.757
			Irbesartan	115	47 (40.9)	1.52 (2.48)	(-3.38, 6.42)			
	Week 48		Sparsentan	98	44 (44.9)	3.86 (2.56)	(-1.20, 8.92)	-0.96 (3.78)	(-8.42, 6.50)	0.800
			Irbesartan	115	38 (33.0)	4.82 (2.76)	(-0.63, 10.27)			
Asia Pacific	Week 24		Sparsentan	69	57 (82.6)	10.88 (2.24)	(6.45, 15.31)	5.77 (3.86)	(-1.87, 13.42)	0.138
			Irbesartan	41	31 (75.6)	5.11 (3.07)	(-0.97, 11.19)			
	Week 48		Sparsentan	69	53 (76.8)	12.25 (2.29)	(7.72, 16.77)	5.91 (4.24)	(-2.47, 14.30)	0.165
			Irbesartan	41	22 (53.7)	6.33 (3.53)	(-0.64, 13.30)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.850
< 27 kg/m**2	Week 24	Sparsentan	84	52 (61.9)	7.80 (2.44)	(2.98, 12.62)	1.08 (3.51)	(-5.86, 8.02)	0.759
		Irbesartan	94	50 (53.2)	6.72 (2.50)	(1.78, 11.67)			
	Week 48	Sparsentan	84	47 (56.0)	8.87 (2.56)	(3.82, 13.93)	4.02 (3.91)	(-3.68, 11.73)	0.304
		Irbesartan	94	36 (38.3)	4.85 (2.92)	(-0.91, 10.61)			
≥ 27 kg/m**2	Week 24	Sparsentan	118	73 (61.9)	3.11 (1.89)	(-0.61, 6.84)	3.55 (2.97)	(-2.30, 9.40)	0.233
		Irbesartan	107	54 (50.5)	-0.43 (2.22)	(-4.81, 3.94)			
	Week 48	Sparsentan	118	63 (53.4)	5.93 (1.99)	(2.01, 9.86)	1.50 (3.20)	(-4.80, 7.81)	0.639
		Irbesartan	107	41 (38.3)	4.43 (2.47)	(-0.43, 9.29)			

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Randomization strata	Overall		Sparsentan						Interaction:	0.506
eGFR Low and UP High	Week 24		Sparsentan	71	44 (62.0)	5.61 (2.52)	(0.63, 10.60)	7.71 (3.90)	(-0.02, 15.44)	0.051
			Irbesartan	74	32 (43.2)	-2.10 (2.96)	(-7.96, 3.77)			
	Week 48		Sparsentan	71	37 (52.1)	6.50 (2.73)	(1.10, 11.89)	4.30 (4.47)	(-4.55, 13.16)	0.338
			Irbesartan	74	22 (29.7)	2.19 (3.53)	(-4.79, 9.18)			
eGFR Low and UP Low	Week 24		Sparsentan	55	38 (69.1)	3.25 (2.62)	(-1.94, 8.45)	-4.37 (3.90)	(-12.11, 3.36)	0.265
			Irbesartan	55	33 (60.0)	7.63 (2.83)	(2.02, 13.23)			
	Week 48		Sparsentan	55	28 (50.9)	11.89 (2.99)	(5.96, 17.82)	4.44 (4.38)	(-4.23, 13.11)	0.313
			Irbesartan	55	25 (45.5)	7.46 (3.17)	(1.17, 13.74)			
eGFR High and UP High	Week 24		Sparsentan	37	19 (51.4)	9.10 (4.25)	(0.59, 17.61)	5.33 (6.38)	(-7.44, 18.10)	0.407
			Irbesartan	36	18 (50.0)	3.77 (4.58)	(-5.42, 12.96)			
	Week 48		Sparsentan	37	22 (59.5)	1.05 (4.02)	(-7.01, 9.11)	-10.01 (6.92)	(-23.84, 3.82)	0.153
			Irbesartan	36	11 (30.6)	11.05 (5.51)	(0.05, 22.06)			
eGFR High and UP Low	Week 24		Sparsentan	39	24 (61.5)	5.93 (3.19)	(-0.41, 12.28)	2.75 (4.68)	(-6.56, 12.06)	0.558
			Irbesartan	37	21 (56.8)	3.18 (3.40)	(-3.60, 9.96)			
	Week 48		Sparsentan	39	23 (59.0)	9.10 (3.26)	(2.61, 15.59)	8.33 (4.88)	(-1.38, 18.04)	0.092
			Irbesartan	37	19 (51.4)	0.77 (3.59)	(-6.38, 7.92)			

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LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.603
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	83 (65.4)	3.53 (1.80)	(-0.03, 7.08)	2.10 (2.72)	(-3.26, 7.45)	0.441
		Irbesartan	129	68 (52.7)	1.43 (2.01)	(-2.53, 5.39)			
	Week 48	Sparsentan	127	65 (51.2)	7.79 (2.01)	(3.83, 11.74)	3.77 (3.09)	(-2.31, 9.85)	0.223
		Irbesartan	129	49 (38.0)	4.02 (2.32)	(-0.56, 8.59)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	27 (55.1)	6.11 (3.38)	(-0.61, 12.83)	-1.11 (5.05)	(-11.17, 8.95)	0.826
		Irbesartan	48	24 (50.0)	7.23 (3.64)	(-0.02, 14.47)			
	Week 48	Sparsentan	49	29 (59.2)	6.96 (3.29)	(0.41, 13.50)	3.79 (5.26)	(-6.68, 14.25)	0.474
		Irbesartan	48	19 (39.6)	3.17 (3.97)	(-4.72, 11.06)			
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	15 (57.7)	16.09 (4.47)	(7.10, 25.08)	15.15 (6.72)	(1.62, 28.67)	0.029 *
		Irbesartan	25	12 (48.0)	0.94 (5.00)	(-9.13, 11.02)			
	Week 48	Sparsentan	26	16 (61.5)	8.25 (4.36)	(-0.54, 17.04)	-1.17 (7.47)	(-16.20, 13.87)	0.877
		Irbesartan	25	9 (36.0)	9.41 (5.93)	(-2.53, 21.35)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline eGFR Group 2		Overall	Sparsentan							Interaction: 0.794
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	53 (64.6)	4.59 (2.30)	(0.04, 9.14)	2.08 (3.42)	(-4.68, 8.85)	0.544	
		Irbesartan	80	45 (56.3)	2.51 (2.51)	(-2.46, 7.47)				
	Week 48	Sparsentan	82	42 (51.2)	9.46 (2.54)	(4.45, 14.48)	4.95 (3.97)	(-2.89, 12.80)	0.214	
		Irbesartan	80	29 (36.3)	4.51 (3.04)	(-1.49, 10.51)				
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	30 (66.7)	1.33 (2.92)	(-4.47, 7.13)	1.30 (4.55)	(-7.76, 10.35)	0.776	
		Irbesartan	49	23 (46.9)	0.03 (3.40)	(-6.74, 6.80)				
	Week 48	Sparsentan	45	23 (51.1)	4.78 (3.31)	(-1.80, 11.37)	1.61 (4.95)	(-8.23, 11.45)	0.746	
		Irbesartan	49	20 (40.8)	3.17 (3.62)	(-4.02, 10.37)				
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	27 (55.1)	6.11 (3.38)	(-0.61, 12.83)	-1.11 (5.05)	(-11.17, 8.95)	0.826	
		Irbesartan	48	24 (50.0)	7.23 (3.64)	(-0.02, 14.47)				
	Week 48	Sparsentan	49	29 (59.2)	6.96 (3.29)	(0.41, 13.50)	3.79 (5.26)	(-6.68, 14.25)	0.474	
		Irbesartan	48	19 (39.6)	3.17 (3.97)	(-4.72, 11.06)				
≥ 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	15 (57.7)	16.09 (4.47)	(7.10, 25.08)	15.15 (6.72)	(1.62, 28.67)	0.029 *	
		Irbesartan	25	12 (48.0)	0.94 (5.00)	(-9.13, 11.02)				
	Week 48	Sparsentan	26	16 (61.5)	8.25 (4.36)	(-0.54, 17.04)	-1.17 (7.47)	(-16.20, 13.87)	0.877	
		Irbesartan	25	9 (36.0)	9.41 (5.93)	(-2.53, 21.35)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.951
<= 1.75 g/day	Week 24	Sparsentan	98	66 (67.3)	5.73 (2.12)	(1.55, 9.92)	0.28 (3.43)	(-6.49, 7.06)	0.934
		Irbesartan	94	43 (45.7)	5.45 (2.66)	(0.21, 10.69)			
	Week 48	Sparsentan	98	59 (60.2)	10.33 (2.21)	(5.96, 14.69)	5.57 (3.69)	(-1.71, 12.85)	0.133
		Irbesartan	94	34 (36.2)	4.76 (2.92)	(-1.00, 10.51)			
> 1.75 g/day	Week 24	Sparsentan	104	59 (56.7)	5.51 (2.15)	(1.27, 9.76)	4.91 (3.06)	(-1.12, 10.95)	0.110
		Irbesartan	108	61 (56.5)	0.60 (2.13)	(-3.60, 4.80)			
	Week 48	Sparsentan	104	51 (49.0)	4.22 (2.33)	(-0.37, 8.82)	-0.13 (3.50)	(-7.04, 6.77)	0.969
		Irbesartan	108	43 (39.8)	4.36 (2.53)	(-0.64, 9.35)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

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Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.066
Yes	Week 24	Sparsentan	88	50 (56.8)	8.46 (2.27)	(3.97, 12.96)	6.60 (3.48)	(-0.28, 13.48)	0.060
		Irbesartan	83	39 (47.0)	1.86 (2.60)	(-3.28, 7.00)			
	Week 48	Sparsentan	88	41 (46.6)	8.91 (2.47)	(4.02, 13.80)	7.47 (3.89)	(-0.23, 15.16)	0.057
		Irbesartan	83	28 (33.7)	1.44 (2.97)	(-4.43, 7.31)			
No	Week 24	Sparsentan	114	75 (65.8)	3.46 (1.99)	(-0.47, 7.39)	0.35 (2.95)	(-5.46, 6.16)	0.905
		Irbesartan	119	65 (54.6)	3.11 (2.15)	(-1.12, 7.34)			
	Week 48	Sparsentan	114	69 (60.5)	5.93 (2.06)	(1.87, 9.99)	-0.06 (3.22)	(-6.39, 6.28)	0.986
		Irbesartan	119	49 (41.2)	5.98 (2.45)	(1.16, 10.81)			

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

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Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.704
<= 5 years	Week 24	Sparsentan	113	70 (61.9)	9.33 (1.95)	(5.48, 13.18)	4.68 (2.83)	(-0.89, 10.25)	0.099
		Irbesartan	127	67 (52.8)	4.65 (2.01)	(0.70, 8.61)			
	Week 48	Sparsentan	113	61 (54.0)	8.08 (2.07)	(4.01, 12.16)	2.48 (3.07)	(-3.57, 8.53)	0.420
		Irbesartan	127	51 (40.2)	5.60 (2.25)	(1.18, 10.03)			
> 5 years	Week 24	Sparsentan	89	55 (61.8)	0.47 (2.37)	(-4.21, 5.15)	0.40 (3.80)	(-7.11, 7.91)	0.917
		Irbesartan	75	37 (49.3)	0.07 (2.95)	(-5.76, 5.91)			
	Week 48	Sparsentan	89	49 (55.1)	6.16 (2.51)	(1.20, 11.12)	3.00 (4.30)	(-5.50, 11.50)	0.486
		Irbesartan	75	26 (34.7)	3.16 (3.45)	(-3.65, 9.97)			

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LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

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Table PT1KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.998
Yes	Week 24	Sparsentan	153	94 (61.4)	5.77 (1.63)	(2.56, 8.97)	2.48 (2.46)	(-2.36, 7.31)	0.315
		Irbesartan	157	76 (48.4)	3.29 (1.82)	(-0.29, 6.87)			
	Week 48	Sparsentan	153	81 (52.9)	7.65 (1.73)	(4.24, 11.06)	3.09 (2.71)	(-2.24, 8.42)	0.255
		Irbesartan	157	57 (36.3)	4.56 (2.06)	(0.50, 8.62)			
No	Week 24	Sparsentan	49	31 (63.3)	4.14 (3.55)	(-2.91, 11.18)	3.09 (5.23)	(-7.29, 13.47)	0.556
		Irbesartan	45	28 (62.2)	1.05 (3.77)	(-6.44, 8.53)			
	Week 48	Sparsentan	49	29 (59.2)	6.32 (3.66)	(-0.94, 13.59)	2.32 (5.83)	(-9.26, 13.89)	0.692
		Irbesartan	45	20 (44.4)	4.00 (4.46)	(-4.84, 12.85)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Figure PF1KBUC\_FSGM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

Not done: No significant subgroup found.

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KBUC\_FSCM.

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.149
Male	Sparsentan	139	38 (27.3)	95.9	(70.1, NE)	1.991	(1.105, 3.588)	0.022 *
	Irbesartan	143	17 (11.9)	NE				
Female	Sparsentan	63	23 (36.5)	50.1	(46.3, NE)	1.054	(0.478, 2.327)	0.896
	Irbesartan	59	12 (20.3)	71.9	(46.3, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan							Interaction test: 0.935
<= 45 years	Sparsentan	96	29 (30.2)	87.7	(48.7, NE)	1.509	(0.798, 2.854)	0.205
	Irbesartan	99	17 (17.2)	NE				
> 45 years	Sparsentan	106	32 (30.2)	75.3	(49.1, NE)	1.156	(0.571, 2.343)	0.687
	Irbesartan	103	12 (11.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: NE
<= 18 years	Sparsentan	9	3 (33.3) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	31 (30.4)	75.3	(49.7, NE)	1.686	(0.926, 3.072)	0.088
	Irbesartan	109	18 (16.5)	NE				
> 40 years	Sparsentan	91	27 (29.7)	111.0	(48.7, NE)	1.105	(0.519, 2.352)	0.795
	Irbesartan	88	11 (12.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.370
North America	Sparsentan	35	8 (22.9)	87.7	(44.4, NE)	0.886	(0.251, 3.120)	0.850
	Irbesartan	46	6 (13.0)	NE				
Europe	Sparsentan	98	18 (18.4)	NE		1.922	(0.908, 4.066)	0.087
	Irbesartan	115	15 (13.0)	NE				
Asia Pacific	Sparsentan	69	35 (50.7)	50.1	(48.1, 111.0)	1.918	(0.881, 4.179)	0.101
	Irbesartan	41	8 (19.5)	95.1	(70.6, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.299
< 27 kg/m**2	Sparsentan	84	29 (34.5)	50.1	(48.1, NE)	1.093	(0.577, 2.070)	0.785
	Irbesartan	94	18 (19.1)	95.1	(71.0, NE)			
≥ 27 kg/m**2	Sparsentan	118	32 (27.1)	95.9	(72.0, 111.0)	1.778	(0.867, 3.649)	0.117
	Irbesartan	107	11 (10.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.193
eGFR Low and UP High	Sparsentan	71	21 (29.6)	75.3	(48.4, 111.0)	2.813	(1.191, 6.643)	0.018 *
	Irbesartan	74	8 (10.8)	NE				
eGFR Low and UP Low	Sparsentan	55	15 (27.3)	NE		0.819	(0.370, 1.809)	0.621
	Irbesartan	55	12 (21.8)	73.0	(50.1, NE)			
eGFR High and UP High	Sparsentan	37	13 (35.1)	87.7	(25.7, NE)	1.472	(0.365, 5.932)	0.587
	Irbesartan	36	3 (8.3)	NE				
eGFR High and UP Low	Sparsentan	39	12 (30.8)	50.1	(48.1, NE)	1.703	(0.628, 4.624)	0.296
	Irbesartan	37	6 (16.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.696
< 60 mL/min/1.73 m**2	Sparsentan	127	35 (27.6)	111.0	(52.0, NE)	1.658	(0.946, 2.908)	0.078
	Irbesartan	129	20 (15.5)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	15 (30.6)	70.1	(44.4, NE)	1.307	(0.441, 3.870)	0.629
	Irbesartan	48	6 (12.5)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	11 (42.3)	70.0	(24.3, 95.9)	2.398	(0.528, 10.900)	0.257
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.819
< 45 mL/min/1.73 m**2	Sparsentan	82	23 (28.0)	75.3	(50.1, NE)	1.621	(0.810, 3.244)	0.172
	Irbesartan	80	13 (16.3)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	12 (26.7)	NE		1.452	(0.511, 4.124)	0.484
	Irbesartan	49	7 (14.3)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	15 (30.6)	70.1	(44.4, NE)	1.307	(0.441, 3.870)	0.629
	Irbesartan	48	6 (12.5)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	11 (42.3)	70.0	(24.3, 95.9)	2.398	(0.528, 10.900)	0.257
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.947
<= 1.75 g/day	Sparsentan	98	32 (32.7)	NE		1.832	(0.961, 3.491)	0.066
	Irbesartan	94	14 (14.9)	NE				
> 1.75 g/day	Sparsentan	104	29 (27.9)	72.1	(49.3, 111.0)	1.505	(0.770, 2.940)	0.232
	Irbesartan	108	15 (13.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.287
Yes	Sparsentan	88	25 (28.4)	72.1	(49.1, NE)	2.377	(1.102, 5.128)	0.027 *
	Irbesartan	83	10 (12.0)	NE				
No	Sparsentan	114	36 (31.6)	95.9	(49.1, NE)	1.297	(0.733, 2.298)	0.372
	Irbesartan	119	19 (16.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.211
<= 5 years	Sparsentan	113	36 (31.9)	72.1	(48.7, NE)	1.291	(0.762, 2.188)	0.342
	Irbesartan	127	24 (18.9)	NE				
> 5 years	Sparsentan	89	25 (28.1)	NE		3.550	(1.301, 9.689)	0.013 *
	Irbesartan	75	5 (6.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.056
Yes	Sparsentan	153	42 (27.5)	95.9	(70.1, NE)	1.193	(0.706, 2.015)	0.510
	Irbesartan	157	24 (15.3)	NE				
No	Sparsentan	49	19 (38.8)	50.1	(48.1, NE)	3.578	(1.307, 9.792)	0.013 *
	Irbesartan	45	5 (11.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Figure PF1KBUIT\_FSKM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_   
An increase reflects an improvement of the status of the patient.  
Reference table: PT1KBUIT\_FSTM

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.507
Male	Sparsentan	139	19 (13.7)	NE		0.816	(0.437, 1.525)	0.524
	Irbesartan	143	25 (17.5)	NE				
Female	Sparsentan	63	6 (9.5)	112.1	(112.1, NE)	0.466	(0.141, 1.534)	0.209
	Irbesartan	59	8 (13.6)	94.6	(70.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.588
<= 45 years	Sparsentan	96	12 (12.5)	NE		0.819	(0.379, 1.773)	0.613
	Irbesartan	99	16 (16.2)	NE				
> 45 years	Sparsentan	106	13 (12.3)	112.1	(112.1, NE)	0.619	(0.286, 1.339)	0.223
	Irbesartan	103	17 (16.5)	95.1	(94.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	1 (11.1) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	13 (12.7)	NE		0.861	(0.404, 1.833)	0.697
	Irbesartan	109	17 (15.6)	NE				
> 40 years	Sparsentan	91	11 (12.1)	112.1	(112.1, NE)	0.542	(0.240, 1.227)	0.142
	Irbesartan	88	16 (18.2)	96.1	(70.9, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.860
North America	Sparsentan	35	4 (11.4)	95.0	(30.4, NE)	0.478	(0.144, 1.589)	0.229
	Irbesartan	46	13 (28.3)	70.1	(48.6, 96.1)			
Europe	Sparsentan	98	12 (12.2)	112.1	(112.1, NE)	0.869	(0.393, 1.924)	0.730
	Irbesartan	115	14 (12.2)	NE				
Asia Pacific	Sparsentan	69	9 (13.0)	NE		0.543	(0.184, 1.602)	0.269
	Irbesartan	41	6 (14.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.191
< 27 kg/m**2	Sparsentan	84	13 (15.5)	112.1	(112.1, NE)	0.989	(0.441, 2.218)	0.979
	Irbesartan	94	12 (12.8)	95.3	(95.1, NE)			
≥ 27 kg/m**2	Sparsentan	118	12 (10.2)	NE		0.505	(0.239, 1.065)	0.073
	Irbesartan	107	21 (19.6)	94.6	(93.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan							Interaction test: 0.954
eGFR Low and UP High	Sparsentan	71	12 (16.9)	NE		0.647	(0.291, 1.439)	0.286
	Irbesartan	74	13 (17.6)	95.1	(70.9, NE)			
eGFR Low and UP Low	Sparsentan	55	5 (9.1)	112.1	(NE, NE)	0.482	(0.151, 1.541)	0.218
	Irbesartan	55	10 (18.2)	NE				
eGFR High and UP High	Sparsentan	37	4 (10.8)	NE		0.567	(0.129, 2.487)	0.452
	Irbesartan	36	4 (11.1)	NE				
eGFR High and UP Low	Sparsentan	39	4 (10.3)	NE		0.755	(0.200, 2.843)	0.678
	Irbesartan	37	6 (16.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.886
< 60 mL/min/1.73 m**2	Sparsentan	127	18 (14.2)	112.1	(95.0, 112.1)	0.668	(0.353, 1.263)	0.214
	Irbesartan	129	24 (18.6)	95.3	(94.1, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	6 (12.2)	NE		0.587	(0.181, 1.906)	0.375
	Irbesartan	48	7 (14.6)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	1 (3.8)	NE		0.362	(0.020, 6.400)	0.488
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.929
< 45 mL/min/1.73 m**2	Sparsentan	82	9 (11.0)	NE		0.554	(0.238, 1.289)	0.170
	Irbesartan	80	15 (18.8)	95.1	(93.0, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	9 (20.0)	112.1	(93.0, 112.1)	0.747	(0.274, 2.036)	0.569
	Irbesartan	49	9 (18.4)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	6 (12.2)	NE		0.587	(0.181, 1.906)	0.375
	Irbesartan	48	7 (14.6)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	1 (3.8)	NE		0.362	(0.020, 6.400)	0.488
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan						Interaction test:	0.722
<= 1.75 g/day	Sparsentan	98	12 (12.2)	112.1	(95.0, 112.1)	0.552	(0.238, 1.279)	0.166
	Irbesartan	94	13 (13.8)	NE				
> 1.75 g/day	Sparsentan	104	13 (12.5)	NE		0.702	(0.335, 1.471)	0.348
	Irbesartan	108	20 (18.5)	95.1	(94.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.422
Yes	Sparsentan	88	8 (9.1)	NE		0.493	(0.203, 1.197)	0.118
	Irbesartan	83	14 (16.9)	96.1	(94.1, NE)			
No	Sparsentan	114	17 (14.9)	112.1	(112.1, NE)	0.782	(0.398, 1.535)	0.475
	Irbesartan	119	19 (16.0)	95.3	(70.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.333
<= 5 years	Sparsentan	113	11 (9.7)	112.1	(112.1, NE)	0.501	(0.237, 1.060)	0.071
	Irbesartan	127	21 (16.5)	96.1	(95.1, NE)			
> 5 years	Sparsentan	89	14 (15.7)	NE		0.795	(0.358, 1.765)	0.572
	Irbesartan	75	12 (16.0)	94.6	(70.9, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							
							Interaction test:	0.141
Yes	Sparsentan	153	17 (11.1)	112.1	(112.1, NE)	0.523	(0.278, 0.985)	0.045 *
	Irbesartan	157	25 (15.9)	96.1	(94.1, NE)			
No	Sparsentan	49	8 (16.3)	NE		0.939	(0.309, 2.855)	0.911
	Irbesartan	45	8 (17.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KBUDT\_FSKM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_ reflects a worsening of the status of the patient.  
Reference table: PT1KBUDT\_FSTM

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Sex												
Male	KDQOL: effect of kidney disease	Baseline	Sparsentan	139	130 (93.5)	89.93 (10.99)	28.1	84.38	93.75	96.88	100.0	
			Irbesartan	143	128 (89.5)	89.62 (13.07)	37.5	85.94	93.75	100.00	100.0	
		Week 24	Sparsentan	139	87 (62.6)	89.30 (14.85)	9.4	84.38	93.75	100.00	100.0	
			Irbesartan	143	73 (51.0)	88.78 (15.46)	28.1	84.38	96.88	100.00	100.0	
		Week 48	Sparsentan	139	72 (51.8)	89.80 (14.69)	0.0	87.50	93.75	100.00	100.0	
			Irbesartan	143	55 (38.5)	94.77 (8.92)	50.0	93.75	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	139	87 (62.6)	-0.61 (13.39)	-75.0	-3.13	0.00	6.25	21.9	0.11 [-0.20, 0.42]
			Irbesartan	143	73 (51.0)	-1.97 (11.95)	-53.1	-3.13	0.00	3.13	31.3	
		Week 48	Sparsentan	139	72 (51.8)	-0.39 (14.80)	-100.0	-3.13	0.00	6.25	15.6	-0.23 [-0.58, 0.12]
			Irbesartan	143	55 (38.5)	2.44 (8.04)	-21.9	0.00	0.00	6.25	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Female	KDQOL: effect of kidney disease	Baseline	Sparsentan	63	56 (88.9)	84.21 (18.01)	28.1	73.44	90.63	98.44	100.0	
			Irbesartan	59	51 (86.4)	87.01 (12.88)	46.9	78.13	90.63	100.00	100.0	
		Week 24	Sparsentan	63	38 (60.3)	90.54 (9.51)	59.4	87.50	93.75	96.88	100.0	
			Irbesartan	59	31 (52.5)	88.41 (20.29)	0.0	87.50	93.75	100.00	100.0	
		Week 48	Sparsentan	63	38 (60.3)	91.04 (9.56)	62.5	87.50	93.75	96.88	100.0	
			Irbesartan	59	22 (37.3)	87.78 (13.46)	40.6	81.25	89.06	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	63	38 (60.3)	6.58 (13.84)	-15.6	-3.13	4.69	15.63	34.4	0.47 [-0.01, 0.95]
			Irbesartan	59	31 (52.5)	-1.31 (20.09)	-100.0	-3.13	0.00	9.38	18.8	
		Week 48	Sparsentan	63	38 (60.3)	6.74 (16.70)	-37.5	-3.13	3.13	9.38	56.3	0.55 [0.01, 1.08]
			Irbesartan	59	22 (37.3)	-1.42 (11.28)	-25.0	-9.38	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age												
<= 45 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	96	87 (90.6)	86.03 (16.11)	28.1	84.38	90.63	96.88	100.0	
			Irbesartan	99	92 (92.9)	86.58 (14.49)	37.5	79.69	90.63	96.88	100.0	
		Week 24	Sparsentan	96	57 (59.4)	87.50 (16.17)	9.4	84.38	93.75	100.00	100.0	
			Irbesartan	99	53 (53.5)	85.79 (20.16)	0.0	81.25	93.75	96.88	100.0	
		Week 48	Sparsentan	96	53 (55.2)	88.38 (17.00)	0.0	87.50	93.75	100.00	100.0	
			Irbesartan	99	40 (40.4)	91.33 (13.23)	40.6	89.06	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	96	57 (59.4)	0.55 (16.88)	-75.0	-6.25	3.13	6.25	34.4	0.17 [-0.20, 0.55]
			Irbesartan	99	53 (53.5)	-2.36 (17.10)	-100.0	-3.13	0.00	3.13	31.3	
		Week 48	Sparsentan	96	53 (55.2)	2.42 (21.00)	-100.0	0.00	3.13	6.25	56.3	0.06 [-0.35, 0.47]
			Irbesartan	99	40 (40.4)	1.33 (9.47)	-21.9	-3.13	0.00	4.69	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 45 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	106	99 (93.4)	90.12 (10.89)	50.0	84.38	93.75	100.00	100.0	
			Irbesartan	103	87 (84.5)	91.31 (10.85)	46.9	87.50	93.75	100.00	100.0	
		Week 24	Sparsentan	106	68 (64.2)	91.50 (10.37)	50.0	87.50	93.75	100.00	100.0	
			Irbesartan	103	51 (49.5)	91.67 (12.26)	34.4	87.50	96.88	100.00	100.0	
		Week 48	Sparsentan	106	57 (53.8)	91.94 (7.77)	75.0	87.50	93.75	100.00	100.0	
			Irbesartan	103	37 (35.9)	94.34 (7.21)	75.0	90.63	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	106	68 (64.2)	2.44 (10.80)	-37.5	-3.13	0.00	6.25	31.3	0.32 [-0.05, 0.68]
			Irbesartan	103	51 (49.5)	-1.16 (11.96)	-53.1	-3.13	0.00	6.25	18.8	
		Week 48	Sparsentan	106	57 (53.8)	1.75 (8.66)	-21.9	-3.13	0.00	6.25	31.3	0.05 [-0.37, 0.46]
			Irbesartan	103	37 (35.9)	1.35 (8.98)	-25.0	0.00	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age at IgAN diagnosis												
<= 18 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	9	9 (100.0)	76.74 (22.54)	28.1	71.88	84.38	90.63	96.9	
			Irbesartan	5	5 (100.0)	86.88 (23.01)	46.9	87.50	100.00	100.00	100.0	
		Week 24	Sparsentan	9	3 (33.3)	65.63 (49.11)	9.4	9.38	87.50	100.00	100.0	
			Irbesartan	5	2 (40.0)	79.69 (24.31)	62.5	62.50	79.69	96.88	96.9	
		Week 48	Sparsentan	9	5 (55.6)	89.38 (8.73)	75.0	87.50	93.75	93.75	96.9	
			Irbesartan	5	1 (20.0)	96.88	96.9	96.88	96.88	96.88	96.9	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	9	3 (33.3)	-17.71 (49.84)	-75.0	-75.00	6.25	15.63	15.6	-0.58 [-2.40, 1.25]
			Irbesartan	5	2 (40.0)	6.25 (13.26)	-3.1	-3.13	6.25	15.63	15.6	
		Week 48	Sparsentan	9	5 (55.6)	13.13 (19.94)	0.0	0.00	3.13	15.63	46.9	NE
			Irbesartan	5	1 (20.0)	-3.13	-3.1	-3.13	-3.13	-3.13	-3.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 18 to 40 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	102	93 (91.2)	89.01 (12.76)	34.4	87.50	93.75	96.88	100.0	
			Irbesartan	109	99 (90.8)	87.31 (13.87)	37.5	84.38	90.63	96.88	100.0	
		Week 24	Sparsentan	102	62 (60.8)	89.67 (12.04)	56.3	84.38	93.75	100.00	100.0	
			Irbesartan	109	59 (54.1)	87.18 (19.70)	0.0	84.38	93.75	100.00	100.0	
		Week 48	Sparsentan	102	57 (55.9)	89.25 (16.58)	0.0	87.50	93.75	100.00	100.0	
			Irbesartan	109	44 (40.4)	91.76 (12.75)	40.6	90.63	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	102	62 (60.8)	1.41 (12.99)	-37.5	-6.25	0.00	6.25	34.4	0.24 [-0.11, 0.60]
			Irbesartan	109	59 (54.1)	-2.22 (16.80)	-100.0	-3.13	0.00	3.13	31.3	
		Week 48	Sparsentan	102	57 (55.9)	0.99 (19.47)	-100.0	-3.13	3.13	6.25	56.3	-0.07 [-0.46, 0.33]
			Irbesartan	109	44 (40.4)	2.06 (9.51)	-21.9	0.00	0.00	6.25	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	91	84 (92.3)	88.54 (13.18)	28.1	84.38	93.75	96.88	100.0	
			Irbesartan	88	75 (85.2)	91.08 (10.81)	46.9	84.38	93.75	100.00	100.0	
		Week 24	Sparsentan	91	60 (65.9)	90.89 (10.78)	50.0	87.50	93.75	100.00	100.0	
			Irbesartan	88	43 (48.9)	91.13 (11.85)	34.4	84.38	93.75	100.00	100.0	
		Week 48	Sparsentan	91	48 (52.7)	91.47 (7.89)	75.0	87.50	93.75	100.00	100.0	
			Irbesartan	88	32 (36.4)	94.04 (7.55)	75.0	89.06	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	91	60 (65.9)	2.71 (11.22)	-37.5	-3.13	3.13	6.25	31.3	0.37 [-0.02, 0.77]
			Irbesartan	88	43 (48.9)	-1.53 (11.66)	-53.1	-6.25	0.00	6.25	18.8	
		Week 48	Sparsentan	91	48 (52.7)	2.21 (8.79)	-21.9	-1.56	1.56	6.25	31.3	0.20 [-0.25, 0.64]
			Irbesartan	88	32 (36.4)	0.49 (8.88)	-25.0	0.00	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Geographic region												
North America	KDQOL: effect of kidney disease	Baseline	Sparsentan	35	32 (91.4)	86.52 (18.37)	28.1	84.38	93.75	96.88	100.0	
			Irbesartan	46	43 (93.5)	91.28 (10.84)	46.9	87.50	96.88	100.00	100.0	
		Week 24	Sparsentan	35	15 (42.9)	87.92 (22.53)	9.4	87.50	93.75	96.88	100.0	
			Irbesartan	46	26 (56.5)	87.86 (20.08)	0.0	84.38	93.75	100.00	100.0	
		Week 48	Sparsentan	35	13 (37.1)	92.55 (7.39)	75.0	90.63	93.75	100.00	100.0	
			Irbesartan	46	17 (37.0)	92.65 (12.45)	50.0	87.50	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	35	15 (42.9)	-3.75 (21.33)	-75.0	-6.25	0.00	6.25	18.8	-0.01 [-0.65, 0.62]
			Irbesartan	46	26 (56.5)	-3.49 (21.46)	-100.0	-3.13	0.00	6.25	15.6	
		Week 48	Sparsentan	35	13 (37.1)	6.01 (13.41)	-6.3	0.00	3.13	6.25	46.9	0.35 [-0.38, 1.08]
			Irbesartan	46	17 (37.0)	2.02 (9.44)	-21.9	0.00	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Europe	KDQOL: effect of kidney disease	Baseline	Sparsentan	98	87 (88.8)	90.16 (11.32)	43.8	84.38	93.75	96.88	100.0	
			Irbesartan	115	96 (83.5)	86.95 (13.93)	43.8	78.13	90.63	96.88	100.0	
		Week 24	Sparsentan	98	53 (54.1)	91.39 (9.80)	56.3	87.50	93.75	100.00	100.0	
			Irbesartan	115	47 (40.9)	88.50 (15.57)	34.4	84.38	93.75	100.00	100.0	
		Week 48	Sparsentan	98	44 (44.9)	92.54 (8.95)	56.3	87.50	93.75	100.00	100.0	
			Irbesartan	115	38 (33.0)	94.16 (8.13)	65.6	90.63	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	98	53 (54.1)	1.89 (11.57)	-25.0	-3.13	0.00	6.25	31.3	0.26 [-0.14, 0.65]
			Irbesartan	115	47 (40.9)	-1.40 (14.09)	-53.1	-3.13	0.00	6.25	31.3	
		Week 48	Sparsentan	98	44 (44.9)	2.77 (13.36)	-40.6	-3.13	1.56	6.25	56.3	0.06 [-0.37, 0.50]
			Irbesartan	115	38 (33.0)	2.06 (8.67)	-15.6	0.00	0.00	6.25	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Asia Pacific	KDQOL: effect of kidney disease	Baseline	Sparsentan	69	67 (97.1)	86.47 (13.82)	34.4	81.25	90.63	96.88	100.0	
			Irbesartan	41	40 (97.6)	90.94 (12.50)	37.5	87.50	93.75	100.00	100.0	
		Week 24	Sparsentan	69	57 (82.6)	88.54 (13.34)	50.0	84.38	93.75	100.00	100.0	
			Irbesartan	41	31 (75.6)	89.62 (16.60)	28.1	84.38	96.88	100.00	100.0	
		Week 48	Sparsentan	69	53 (76.8)	87.74 (16.42)	0.0	81.25	93.75	96.88	100.0	
			Irbesartan	41	22 (53.7)	90.48 (13.38)	40.6	84.38	93.75	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	69	57 (82.6)	2.69 (13.39)	-37.5	-3.13	3.13	9.38	34.4	0.31 [-0.13, 0.75]
			Irbesartan	41	31 (75.6)	-0.91 (7.40)	-15.6	-6.25	0.00	3.13	12.5	
		Week 48	Sparsentan	69	53 (76.8)	0.53 (18.05)	-100.0	-3.13	3.13	6.25	34.4	0.06 [-0.44, 0.56]
			Irbesartan	41	22 (53.7)	-0.43 (9.99)	-25.0	-3.13	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline BMI												
< 27 kg/m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	84	76 (90.5)	85.94 (13.34)	28.1	78.13	90.63	96.88	100.0	
			Irbesartan	94	84 (89.4)	88.50 (11.42)	46.9	84.38	90.63	100.00	100.0	
		Week 24	Sparsentan	84	52 (61.9)	88.46 (16.32)	9.4	87.50	93.75	98.44	100.0	
			Irbesartan	94	50 (53.2)	86.50 (18.71)	0.0	84.38	93.75	96.88	100.0	
		Week 48	Sparsentan	84	47 (56.0)	88.63 (11.26)	50.0	84.38	90.63	96.88	100.0	
			Irbesartan	94	36 (38.3)	89.93 (13.53)	40.6	84.38	93.75	98.44	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	84	52 (61.9)	1.98 (16.36)	-75.0	-4.69	1.56	9.38	34.4	0.16 [-0.23, 0.55]
			Irbesartan	94	50 (53.2)	-0.75 (17.55)	-100.0	-3.13	0.00	6.25	31.3	
		Week 48	Sparsentan	84	47 (56.0)	3.26 (12.97)	-40.6	-3.13	3.13	6.25	46.9	0.12 [-0.31, 0.56]
			Irbesartan	94	36 (38.3)	1.82 (9.22)	-21.9	0.00	0.00	6.25	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 27 kg/m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	118	110 (93.2)	89.77 (13.78)	28.1	87.50	93.75	100.00	100.0	
			Irbesartan	107	94 (87.9)	89.39 (14.33)	37.5	87.50	96.88	100.00	100.0	
		Week 24	Sparsentan	118	73 (61.9)	90.54 (10.95)	56.3	84.38	93.75	100.00	100.0	
			Irbesartan	107	54 (50.5)	90.68 (15.01)	28.1	87.50	96.88	100.00	100.0	
		Week 48	Sparsentan	118	63 (53.4)	91.42 (14.32)	0.0	90.63	93.75	100.00	100.0	
			Irbesartan	107	41 (38.3)	95.27 (6.92)	75.0	93.75	100.00	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	118	73 (61.9)	1.28 (11.91)	-37.5	-3.13	0.00	6.25	31.3	0.34 [-0.02, 0.69]
			Irbesartan	107	54 (50.5)	-2.72 (11.66)	-53.1	-3.13	0.00	3.13	15.6	
		Week 48	Sparsentan	118	63 (53.4)	1.19 (17.64)	-100.0	-3.13	3.13	6.25	56.3	0.02 [-0.37, 0.41]
			Irbesartan	107	41 (38.3)	0.91 (9.22)	-25.0	-3.13	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Randomization strata												
eGFR Low and UP High	KDQOL: effect of kidney disease	Baseline	Sparsentan	71	63 (88.7)	87.85 (13.81)	28.1	84.38	93.75	96.88	100.0	
			Irbesartan	74	63 (85.1)	85.66 (15.38)	43.8	81.25	90.63	96.88	100.0	
		Week 24	Sparsentan	71	44 (62.0)	89.42 (12.02)	56.3	84.38	93.75	98.44	100.0	
			Irbesartan	74	32 (43.2)	84.28 (15.84)	40.6	78.13	87.50	96.88	100.0	
		Week 48	Sparsentan	71	37 (52.1)	90.63 (10.72)	50.0	87.50	93.75	100.00	100.0	
			Irbesartan	74	22 (29.7)	90.34 (14.17)	40.6	84.38	95.31	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	71	44 (62.0)	0.36 (8.48)	-25.0	-3.13	3.13	6.25	12.5	0.47 [0.01, 0.93]
			Irbesartan	74	32 (43.2)	-3.91 (9.95)	-25.0	-12.50	-1.56	3.13	15.6	
		Week 48	Sparsentan	71	37 (52.1)	0.76 (6.35)	-15.6	-3.13	0.00	6.25	12.5	0.22 [-0.31, 0.75]
			Irbesartan	74	22 (29.7)	-0.71 (7.34)	-15.6	-6.25	0.00	3.13	15.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR Low and UP Low	KDQOL: effect of kidney disease	Baseline	Sparsentan	55	51 (92.7)	86.89 (13.13)	43.8	78.13	90.63	96.88	100.0	
			Irbesartan	55	49 (89.1)	91.39 (8.85)	71.9	87.50	93.75	100.00	100.0	
		Week 24	Sparsentan	55	38 (69.1)	89.23 (12.70)	50.0	81.25	93.75	100.00	100.0	
			Irbesartan	55	33 (60.0)	94.98 (9.76)	50.0	93.75	100.00	100.00	100.0	
		Week 48	Sparsentan	55	28 (50.9)	89.29 (19.18)	0.0	87.50	93.75	100.00	100.0	
			Irbesartan	55	25 (45.5)	93.50 (7.54)	75.0	87.50	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	55	38 (69.1)	3.45 (13.07)	-37.5	-3.13	1.56	9.38	31.3	0.13 [-0.33, 0.60]
			Irbesartan	55	33 (60.0)	1.89 (9.76)	-37.5	0.00	0.00	6.25	18.8	
		Week 48	Sparsentan	55	28 (50.9)	2.34 (24.02)	-100.0	0.00	3.13	7.81	56.3	0.07 [-0.47, 0.61]
			Irbesartan	55	25 (45.5)	1.00 (9.79)	-25.0	0.00	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP High	KDQOL: effect of kidney disease	Baseline	Sparsentan	37	35 (94.6)	89.73 (14.15)	28.1	84.38	93.75	100.00	100.0	
			Irbesartan	36	31 (86.1)	88.31 (11.86)	56.3	84.38	90.63	96.88	100.0	
		Week 24	Sparsentan	37	19 (51.4)	89.47 (20.36)	9.4	87.50	93.75	100.00	100.0	
			Irbesartan	36	18 (50.0)	88.02 (15.02)	34.4	84.38	92.19	96.88	100.0	
		Week 48	Sparsentan	37	22 (59.5)	89.20 (11.88)	56.3	87.50	92.19	96.88	100.0	
			Irbesartan	36	11 (30.6)	93.75 (6.56)	81.3	90.63	93.75	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	37	19 (51.4)	-0.66 (21.34)	-75.0	-6.25	3.13	9.38	34.4	0.04 [-0.61, 0.68]
			Irbesartan	36	18 (50.0)	-1.39 (17.22)	-53.1	-3.13	0.00	9.38	31.3	
		Week 48	Sparsentan	37	22 (59.5)	1.14 (18.51)	-40.6	-3.13	0.00	6.25	46.9	-0.33 [-1.05, 0.40]
			Irbesartan	36	11 (30.6)	6.53 (11.22)	-6.3	-3.13	3.13	15.63	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP Low	KDQOL: effect of kidney disease	Baseline	Sparsentan	39	37 (94.9)	89.19 (14.20)	34.4	87.50	93.75	100.00	100.0	
			Irbesartan	37	36 (97.3)	91.58 (13.48)	37.5	90.63	96.88	100.00	100.0	
		Week 24	Sparsentan	39	24 (61.5)	91.02 (10.72)	59.4	89.06	93.75	100.00	100.0	
			Irbesartan	37	21 (56.8)	86.01 (25.30)	0.0	87.50	96.88	100.00	100.0	
		Week 48	Sparsentan	39	23 (59.0)	91.71 (8.56)	65.6	90.63	93.75	96.88	100.0	
			Irbesartan	37	19 (51.4)	94.08 (12.19)	50.0	90.63	100.00	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	39	24 (61.5)	2.60 (16.08)	-37.5	-4.69	0.00	14.06	31.3	0.37 [-0.22, 0.96]
			Irbesartan	37	21 (56.8)	-4.61 (22.89)	-100.0	-3.13	0.00	3.13	12.5	
		Week 48	Sparsentan	39	23 (59.0)	4.76 (10.98)	-21.9	-3.13	3.13	9.38	31.3	0.36 [-0.25, 0.97]
			Irbesartan	37	19 (51.4)	1.15 (8.54)	-21.9	0.00	3.13	6.25	15.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline eGFR Group 1												
< 60 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	127	115 (90.6)	87.55 (13.46)	28.1	84.38	93.75	96.88	100.0	
			Irbesartan	129	113 (87.6)	89.13 (12.29)	43.8	84.38	93.75	100.00	100.0	
		Week 24	Sparsentan	127	83 (65.4)	88.82 (12.65)	50.0	81.25	93.75	100.00	100.0	
			Irbesartan	129	68 (52.7)	90.17 (13.79)	40.6	84.38	96.88	100.00	100.0	
		Week 48	Sparsentan	127	65 (51.2)	90.19 (14.89)	0.0	87.50	93.75	100.00	100.0	
			Irbesartan	129	49 (38.0)	92.22 (11.03)	40.6	87.50	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	127	83 (65.4)	1.20 (11.70)	-37.5	-3.13	0.00	6.25	31.3	0.16 [-0.16, 0.48]
			Irbesartan	129	68 (52.7)	-0.55 (10.01)	-37.5	-3.13	0.00	6.25	18.8	
		Week 48	Sparsentan	127	65 (51.2)	1.54 (16.32)	-100.0	0.00	3.13	6.25	56.3	0.07 [-0.30, 0.44]
			Irbesartan	129	49 (38.0)	0.57 (8.65)	-25.0	0.00	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	49	47 (95.9)	90.69 (12.66)	34.4	87.50	93.75	100.00	100.0	
			Irbesartan	48	43 (89.6)	87.79 (13.89)	46.9	78.13	90.63	100.00	100.0	
		Week 24	Sparsentan	49	27 (55.1)	88.66 (17.79)	9.4	87.50	93.75	96.88	100.0	
			Irbesartan	48	24 (50.0)	89.06 (13.85)	34.4	85.94	92.19	96.88	100.0	
		Week 48	Sparsentan	49	29 (59.2)	90.63 (9.81)	62.5	87.50	93.75	96.88	100.0	
			Irbesartan	48	19 (39.6)	92.76 (11.74)	50.0	90.63	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	49	27 (55.1)	-0.81 (19.47)	-75.0	-6.25	0.00	9.38	31.3	-0.01 [-0.56, 0.54]
			Irbesartan	48	24 (50.0)	-0.65 (14.69)	-53.1	-3.13	0.00	6.25	31.3	
		Week 48	Sparsentan	49	29 (59.2)	1.62 (11.92)	-37.5	0.00	3.13	6.25	31.3	-0.14 [-0.72, 0.44]
			Irbesartan	48	19 (39.6)	3.29 (11.29)	-21.9	-3.13	0.00	9.38	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	26	24 (92.3)	86.46 (16.50)	28.1	84.38	90.63	98.44	100.0	
			Irbesartan	25	23 (92.0)	89.67 (15.29)	37.5	84.38	96.88	100.00	100.0	
		Week 24	Sparsentan	26	15 (57.7)	96.25 (4.29)	87.5	93.75	96.88	100.00	100.0	
			Irbesartan	25	12 (48.0)	79.43 (31.94)	0.0	79.69	93.75	96.88	100.0	
		Week 48	Sparsentan	26	16 (61.5)	89.65 (11.11)	56.3	87.50	93.75	96.88	100.0	
			Irbesartan	25	9 (36.0)	95.83 (7.49)	78.1	93.75	100.00	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	26	15 (57.7)	7.92 (12.04)	-12.5	0.00	6.25	12.50	34.4	0.87 [0.08, 1.67]
			Irbesartan	25	12 (48.0)	-10.94 (29.51)	-100.0	-12.50	-1.56	3.13	12.5	
		Week 48	Sparsentan	26	16 (61.5)	5.08 (19.89)	-40.6	-4.69	0.00	14.06	46.9	0.22 [-0.60, 1.04]
			Irbesartan	25	9 (36.0)	1.39 (7.18)	-12.5	0.00	0.00	3.13	15.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 2													
< 45 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	82	71 (86.6)	85.96 (13.24)	43.8	78.13	90.63	93.75	100.0		
			Irbesartan	80	69 (86.3)	87.14 (13.91)	43.8	81.25	90.63	96.88	100.0		
		Week 24	Sparsentan	82	53 (64.6)	89.21 (12.20)	50.0	84.38	93.75	100.00	100.0		
			Irbesartan	80	45 (56.3)	88.06 (15.80)	40.6	84.38	96.88	100.00	100.0		
		Week 48	Sparsentan	82	42 (51.2)	88.62 (17.11)	0.0	87.50	92.19	100.00	100.0		
			Irbesartan	80	29 (36.3)	90.63 (13.23)	40.6	84.38	96.88	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	82	53 (64.6)	2.42 (10.59)	-37.5	-3.13	3.13	6.25	28.1	0.31 [-0.09, 0.71]
				Irbesartan	80	45 (56.3)	-0.97 (11.28)	-37.5	-3.13	0.00	6.25	18.8	
			Week 48	Sparsentan	82	42 (51.2)	0.97 (19.54)	-100.0	-3.13	3.13	6.25	56.3	0.11 [-0.36, 0.59]
				Irbesartan	80	29 (36.3)	-0.86 (8.26)	-25.0	-3.13	0.00	3.13	18.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
45 to < 60 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	45	44 (97.8)	90.13 (13.58)	28.1	87.50	93.75	100.00	100.0	
			Irbesartan	49	44 (89.8)	92.26 (8.43)	71.9	87.50	95.31	100.00	100.0	
		Week 24	Sparsentan	45	30 (66.7)	88.13 (13.59)	56.3	81.25	93.75	96.88	100.0	
			Irbesartan	49	23 (46.9)	94.29 (7.28)	78.1	90.63	96.88	100.00	100.0	
		Week 48	Sparsentan	45	23 (51.1)	93.07 (9.28)	68.8	87.50	96.88	100.00	100.0	
			Irbesartan	49	20 (40.8)	94.53 (6.32)	81.3	89.06	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	45	30 (66.7)	-0.94 (13.36)	-37.5	-6.25	0.00	6.25	31.3	-0.11 [-0.65, 0.43]
			Irbesartan	49	23 (46.9)	0.27 (7.05)	-18.8	0.00	0.00	3.13	12.5	
		Week 48	Sparsentan	45	23 (51.1)	2.58 (7.86)	-9.4	0.00	0.00	6.25	31.3	-0.01 [-0.61, 0.59]
			Irbesartan	49	20 (40.8)	2.66 (8.97)	-15.6	0.00	3.13	6.25	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	49	47 (95.9)	90.69 (12.66)	34.4	87.50	93.75	100.00	100.0	
			Irbesartan	48	43 (89.6)	87.79 (13.89)	46.9	78.13	90.63	100.00	100.0	
		Week 24	Sparsentan	49	27 (55.1)	88.66 (17.79)	9.4	87.50	93.75	96.88	100.0	
			Irbesartan	48	24 (50.0)	89.06 (13.85)	34.4	85.94	92.19	96.88	100.0	
		Week 48	Sparsentan	49	29 (59.2)	90.63 (9.81)	62.5	87.50	93.75	96.88	100.0	
			Irbesartan	48	19 (39.6)	92.76 (11.74)	50.0	90.63	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	49	27 (55.1)	-0.81 (19.47)	-75.0	-6.25	0.00	9.38	31.3	-0.01 [-0.56, 0.54]
			Irbesartan	48	24 (50.0)	-0.65 (14.69)	-53.1	-3.13	0.00	6.25	31.3	
		Week 48	Sparsentan	49	29 (59.2)	1.62 (11.92)	-37.5	0.00	3.13	6.25	31.3	-0.14 [-0.72, 0.44]
			Irbesartan	48	19 (39.6)	3.29 (11.29)	-21.9	-3.13	0.00	9.38	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	26	24 (92.3)	86.46 (16.50)	28.1	84.38	90.63	98.44	100.0	
			Irbesartan	25	23 (92.0)	89.67 (15.29)	37.5	84.38	96.88	100.00	100.0	
		Week 24	Sparsentan	26	15 (57.7)	96.25 (4.29)	87.5	93.75	96.88	100.00	100.0	
			Irbesartan	25	12 (48.0)	79.43 (31.94)	0.0	79.69	93.75	96.88	100.0	
		Week 48	Sparsentan	26	16 (61.5)	89.65 (11.11)	56.3	87.50	93.75	96.88	100.0	
			Irbesartan	25	9 (36.0)	95.83 (7.49)	78.1	93.75	100.00	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	26	15 (57.7)	7.92 (12.04)	-12.5	0.00	6.25	12.50	34.4	0.87 [0.08, 1.67]
			Irbesartan	25	12 (48.0)	-10.94 (29.51)	-100.0	-12.50	-1.56	3.13	12.5	
		Week 48	Sparsentan	26	16 (61.5)	5.08 (19.89)	-40.6	-4.69	0.00	14.06	46.9	0.22 [-0.60, 1.04]
			Irbesartan	25	9 (36.0)	1.39 (7.18)	-12.5	0.00	0.00	3.13	15.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline urine protein excretion												
<= 1.75 g/day	KDQOL: effect of kidney disease	Baseline	Sparsentan	98	92 (93.9)	87.70 (13.75)	34.4	84.38	90.63	96.88	100.0	
			Irbesartan	94	82 (87.2)	89.90 (13.14)	37.5	84.38	95.31	100.00	100.0	
		Week 24	Sparsentan	98	66 (67.3)	88.64 (12.50)	50.0	84.38	93.75	96.88	100.0	
			Irbesartan	94	43 (45.7)	90.41 (19.64)	0.0	90.63	96.88	100.00	100.0	
		Week 48	Sparsentan	98	59 (60.2)	90.25 (15.08)	0.0	87.50	93.75	100.00	100.0	
			Irbesartan	94	34 (36.2)	94.21 (10.18)	50.0	90.63	98.44	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	98	66 (67.3)	2.18 (13.89)	-37.5	-6.25	0.00	9.38	31.3	0.24 [-0.15, 0.62]
			Irbesartan	94	43 (45.7)	-1.53 (17.92)	-100.0	0.00	3.13	6.25	18.8	
		Week 48	Sparsentan	98	59 (60.2)	2.38 (18.45)	-100.0	0.00	3.13	9.38	56.3	0.11 [-0.31, 0.53]
			Irbesartan	94	34 (36.2)	0.64 (9.32)	-25.0	0.00	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 1.75 g/day	KDQOL: effect of kidney disease	Baseline	Sparsentan	104	94 (90.4)	88.70 (13.70)	28.1	84.38	93.75	96.88	100.0	
			Irbesartan	108	97 (89.8)	88.02 (12.95)	46.9	84.38	90.63	96.88	100.0	
		Week 24	Sparsentan	104	59 (56.7)	90.84 (14.41)	9.4	87.50	96.88	100.00	100.0	
			Irbesartan	108	61 (56.5)	87.45 (14.80)	34.4	81.25	93.75	96.88	100.0	
		Week 48	Sparsentan	104	51 (49.0)	90.20 (10.55)	50.0	87.50	93.75	100.00	100.0	
			Irbesartan	108	43 (39.8)	91.64 (11.26)	40.6	87.50	93.75	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	104	59 (56.7)	0.90 (13.95)	-75.0	-3.13	3.13	6.25	34.4	0.22 [-0.14, 0.58]
			Irbesartan	108	61 (56.5)	-1.95 (12.19)	-53.1	-3.13	0.00	3.13	31.3	
		Week 48	Sparsentan	104	51 (49.0)	1.72 (12.16)	-40.6	-3.13	0.00	6.25	46.9	-0.02 [-0.42, 0.39]
			Irbesartan	108	43 (39.8)	1.89 (9.13)	-15.6	-3.13	0.00	6.25	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline use of antihypertensives												
Yes	KDQOL: effect of kidney disease	Baseline	Sparsentan	88	77 (87.5)	89.16 (12.93)	28.1	84.38	93.75	96.88	100.0	
			Irbesartan	83	70 (84.3)	88.97 (12.06)	43.8	81.25	93.75	100.00	100.0	
		Week 24	Sparsentan	88	50 (56.8)	92.38 (10.64)	50.0	90.63	96.88	100.00	100.0	
			Irbesartan	83	39 (47.0)	89.74 (14.32)	50.0	81.25	96.88	100.00	100.0	
		Week 48	Sparsentan	88	41 (46.6)	92.00 (6.85)	78.1	87.50	90.63	100.00	100.0	
			Irbesartan	83	28 (33.7)	94.98 (7.95)	65.6	93.75	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	88	50 (56.8)	2.56 (10.62)	-37.5	-3.13	3.13	6.25	31.3	0.37 [-0.05, 0.79]
			Irbesartan	83	39 (47.0)	-1.44 (10.92)	-37.5	-3.13	0.00	6.25	18.8	
		Week 48	Sparsentan	88	41 (46.6)	1.75 (8.67)	-21.9	-3.13	0.00	6.25	31.3	-0.06 [-0.54, 0.42]
			Irbesartan	83	28 (33.7)	2.23 (8.02)	-9.4	0.00	0.00	4.69	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024



Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL: effect of kidney disease	Baseline	Sparsentan	114	109 (95.6)	87.53 (14.24)	28.1	84.38	90.63	96.88	100.0	
			Irbesartan	119	109 (91.6)	88.82 (13.68)	37.5	84.38	93.75	100.00	100.0	
		Week 24	Sparsentan	114	75 (65.8)	87.88 (14.79)	9.4	81.25	93.75	96.88	100.0	
			Irbesartan	119	65 (54.6)	88.03 (18.41)	0.0	84.38	93.75	96.88	100.0	
		Week 48	Sparsentan	114	69 (60.5)	89.18 (15.66)	0.0	87.50	93.75	100.00	100.0	
			Irbesartan	119	49 (41.2)	91.52 (12.04)	40.6	87.50	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	114	75 (65.8)	0.92 (15.72)	-75.0	-6.25	0.00	9.38	34.4	0.18 [-0.15, 0.51]
			Irbesartan	119	65 (54.6)	-1.97 (16.71)	-100.0	-3.13	0.00	3.13	31.3	
		Week 48	Sparsentan	114	69 (60.5)	2.26 (18.83)	-100.0	0.00	3.13	6.25	56.3	0.09 [-0.28, 0.46]
			Irbesartan	119	49 (41.2)	0.83 (9.82)	-25.0	-3.13	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Time since renal biopsy												
<= 5 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	113	104 (92.0)	87.41 (13.71)	28.1	84.38	90.63	96.88	100.0	
			Irbesartan	127	116 (91.3)	88.39 (13.59)	37.5	82.81	93.75	100.00	100.0	
		Week 24	Sparsentan	113	70 (61.9)	91.43 (10.91)	50.0	87.50	93.75	100.00	100.0	
			Irbesartan	127	67 (52.8)	87.13 (19.18)	0.0	84.38	93.75	100.00	100.0	
		Week 48	Sparsentan	113	61 (54.0)	90.93 (10.44)	50.0	87.50	93.75	100.00	100.0	
			Irbesartan	127	51 (40.2)	92.03 (12.41)	40.6	87.50	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	113	70 (61.9)	5.00 (12.91)	-37.5	0.00	3.13	12.50	34.4	0.54 [0.19, 0.88]
			Irbesartan	127	67 (52.8)	-2.89 (16.44)	-100.0	-6.25	0.00	3.13	31.3	
		Week 48	Sparsentan	113	61 (54.0)	4.20 (11.47)	-40.6	0.00	3.13	9.38	34.4	0.29 [-0.08, 0.67]
			Irbesartan	127	51 (40.2)	1.04 (9.92)	-25.0	0.00	0.00	3.13	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 5 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	89	82 (92.1)	89.21 (13.70)	28.1	87.50	93.75	100.00	100.0	
			Irbesartan	75	63 (84.0)	89.78 (12.00)	46.9	84.38	93.75	100.00	100.0	
		Week 24	Sparsentan	89	55 (61.8)	87.44 (15.89)	9.4	81.25	90.63	100.00	100.0	
			Irbesartan	75	37 (49.3)	91.47 (11.59)	50.0	87.50	96.88	100.00	100.0	
		Week 48	Sparsentan	89	49 (55.1)	89.35 (15.91)	0.0	87.50	93.75	100.00	100.0	
			Irbesartan	75	26 (34.7)	94.23 (6.60)	78.1	90.63	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	89	55 (61.8)	-2.78 (13.96)	-75.0	-6.25	0.00	6.25	28.1	-0.24 [-0.65, 0.18]
			Irbesartan	75	37 (49.3)	0.25 (10.96)	-37.5	-3.13	0.00	6.25	18.8	
		Week 48	Sparsentan	89	49 (55.1)	-0.57 (19.70)	-100.0	-3.13	0.00	6.25	56.3	-0.15 [-0.63, 0.33]
			Irbesartan	75	26 (34.7)	1.92 (7.66)	-9.4	0.00	0.00	6.25	18.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: History of hypertension												
Yes	KDQOL: effect of kidney disease	Baseline	Sparsentan	153	138 (90.2)	87.45 (14.01)	28.1	84.38	92.19	96.88	100.0	
			Irbesartan	157	135 (86.0)	88.82 (12.27)	43.8	84.38	93.75	100.00	100.0	
		Week 24	Sparsentan	153	94 (61.4)	91.16 (10.98)	56.3	87.50	93.75	100.00	100.0	
			Irbesartan	157	76 (48.4)	91.00 (11.74)	50.0	87.50	96.88	100.00	100.0	
		Week 48	Sparsentan	153	81 (52.9)	89.74 (14.31)	0.0	87.50	93.75	100.00	100.0	
			Irbesartan	157	57 (36.3)	94.63 (7.30)	65.6	93.75	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	153	94 (61.4)	3.42 (9.43)	-25.0	-3.13	3.13	6.25	31.3	0.31 [0.01, 0.61]
			Irbesartan	157	76 (48.4)	0.41 (10.11)	-37.5	-3.13	0.00	6.25	31.3	
		Week 48	Sparsentan	153	81 (52.9)	1.74 (15.60)	-100.0	0.00	3.13	6.25	46.9	-0.10 [-0.44, 0.24]
			Irbesartan	157	57 (36.3)	3.07 (8.45)	-12.5	0.00	0.00	6.25	25.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL: effect of kidney disease	Baseline	Sparsentan	49	48 (98.0)	90.36 (12.65)	43.8	87.50	93.75	100.00	100.0	
			Irbesartan	45	44 (97.8)	89.06 (15.30)	37.5	87.50	95.31	100.00	100.0	
		Week 24	Sparsentan	49	31 (63.3)	85.18 (18.54)	9.4	81.25	87.50	100.00	100.0	
			Irbesartan	45	28 (62.2)	82.37 (25.60)	0.0	76.56	93.75	98.44	100.0	
		Week 48	Sparsentan	49	29 (59.2)	91.59 (9.04)	62.5	87.50	93.75	100.00	100.0	
			Irbesartan	45	20 (44.4)	87.50 (16.44)	40.6	82.81	92.19	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	49	31 (63.3)	-4.03 (21.85)	-75.0	-12.50	-3.13	9.38	34.4	0.17 [-0.35, 0.68]
			Irbesartan	45	28 (62.2)	-7.70 (22.27)	-100.0	-10.94	-1.56	3.13	15.6	
		Week 48	Sparsentan	49	29 (59.2)	3.02 (16.50)	-37.5	-3.13	0.00	6.25	56.3	0.47 [-0.11, 1.05]
			Irbesartan	45	20 (44.4)	-3.59 (9.58)	-25.0	-9.38	0.00	1.56	12.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.035 #
Male	Week 24	Sparsentan	139	87 (62.6)	-0.61 (1.32)	(-3.21, 2.00)	1.32 (1.96)	(-2.54, 5.18)	0.501
		Irbesartan	143	73 (51.0)	-1.93 (1.44)	(-4.76, 0.90)			
	Week 48	Sparsentan	139	72 (51.8)	-0.81 (1.44)	(-3.65, 2.02)	-2.85 (2.20)	(-7.18, 1.48)	0.197
		Irbesartan	143	55 (38.5)	2.03 (1.66)	(-1.23, 5.30)			
Female	Week 24	Sparsentan	63	38 (60.3)	5.39 (2.12)	(1.17, 9.61)	5.24 (3.28)	(-1.29, 11.76)	0.114
		Irbesartan	59	31 (52.5)	0.16 (2.46)	(-4.75, 5.06)			
	Week 48	Sparsentan	63	38 (60.3)	4.91 (2.11)	(0.71, 9.11)	7.30 (3.45)	(0.46, 14.15)	0.037 *
		Irbesartan	59	22 (37.3)	-2.39 (2.69)	(-7.73, 2.95)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.967
<= 45 years	Week 24	Sparsentan	96	57 (59.4)	0.24 (2.08)	(-3.86, 4.33)	1.77 (3.00)	(-4.14, 7.68)	0.555
		Irbesartan	99	53 (53.5)	-1.53 (2.15)	(-5.78, 2.71)			
	Week 48	Sparsentan	96	53 (55.2)	0.85 (2.14)	(-3.38, 5.08)	-0.12 (3.26)	(-6.55, 6.32)	0.971
		Irbesartan	99	40 (40.4)	0.97 (2.45)	(-3.87, 5.81)			
> 45 years	Week 24	Sparsentan	106	68 (64.2)	1.54 (1.11)	(-0.65, 3.73)	1.89 (1.73)	(-1.51, 5.30)	0.274
		Irbesartan	103	51 (49.5)	-0.35 (1.30)	(-2.93, 2.22)			
	Week 48	Sparsentan	106	57 (53.8)	0.88 (1.18)	(-1.44, 3.20)	-0.37 (1.88)	(-4.09, 3.35)	0.843
		Irbesartan	103	37 (35.9)	1.25 (1.46)	(-1.63, 4.13)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Age at IgAN diagnosis	Overall		Sparsentan							Interaction: 0.786
<= 18 years	Week 24		Sparsentan	9	3 (33.3)	1.14 (16.71)	(-44.84, 47.12)	-0.44 (32.18)	(-89.71, 88.83)	0.990
			Irbesartan	5	2 (40.0)	1.58 (27.60)	(-75.23, 78.40)			
	Week 48		Sparsentan	9	5 (55.6)	-1.62 (16.55)	(-47.59, 44.36)	-3.31 (32.28)	(-92.56, 85.94)	0.923
			Irbesartan	5	1 (20.0)	1.69 (27.85)	(-75.11, 78.48)			
> 18 to 40 years	Week 24		Sparsentan	102	62 (60.8)	1.40 (1.81)	(-2.17, 4.96)	3.26 (2.60)	(-1.87, 8.38)	0.211
			Irbesartan	109	59 (54.1)	-1.86 (1.86)	(-5.54, 1.81)			
	Week 48		Sparsentan	102	57 (55.9)	0.71 (1.89)	(-3.01, 4.42)	-0.14 (2.86)	(-5.78, 5.50)	0.961
			Irbesartan	109	44 (40.4)	0.85 (2.14)	(-3.38, 5.07)			
> 40 years	Week 24		Sparsentan	91	60 (65.9)	1.43 (1.19)	(-0.92, 3.78)	1.91 (1.84)	(-1.73, 5.55)	0.301
			Irbesartan	88	43 (48.9)	-0.48 (1.39)	(-3.23, 2.27)			
	Week 48		Sparsentan	91	48 (52.7)	0.96 (1.28)	(-1.57, 3.50)	-0.17 (2.05)	(-4.20, 3.87)	0.935
			Irbesartan	88	32 (36.4)	1.13 (1.58)	(-1.98, 4.24)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024



Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Geographic region	Overall	Sparsentan							Interaction:	0.889
North America	Week 24	Sparsentan	35	15 (42.9)	-2.41 (4.72)	(-11.93, 7.11)	0.08 (6.08)	(-12.19, 12.34)	0.990	
		Irbesartan	46	26 (56.5)	-2.48 (3.77)	(-10.09, 5.13)				
	Week 48	Sparsentan	35	13 (37.1)	0.13 (4.88)	(-9.69, 9.94)	1.04 (6.39)	(-11.80, 13.89)	0.871	
		Irbesartan	46	17 (37.0)	-0.92 (4.05)	(-9.04, 7.21)				
Europe	Week 24	Sparsentan	98	53 (54.1)	1.61 (1.37)	(-1.09, 4.31)	3.32 (1.99)	(-0.61, 7.26)	0.097	
		Irbesartan	115	47 (40.9)	-1.71 (1.45)	(-4.58, 1.16)				
	Week 48	Sparsentan	98	44 (44.9)	2.49 (1.48)	(-0.44, 5.42)	1.36 (2.18)	(-2.95, 5.67)	0.535	
		Irbesartan	115	38 (33.0)	1.13 (1.60)	(-2.02, 4.28)				
Asia Pacific	Week 24	Sparsentan	69	57 (82.6)	2.20 (1.72)	(-1.19, 5.59)	2.04 (2.93)	(-3.74, 7.82)	0.486	
		Irbesartan	41	31 (75.6)	0.16 (2.34)	(-4.47, 4.78)				
	Week 48	Sparsentan	69	53 (76.8)	0.05 (1.78)	(-3.46, 3.56)	-0.08 (3.30)	(-6.59, 6.44)	0.982	
		Irbesartan	41	22 (53.7)	0.12 (2.76)	(-5.33, 5.58)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.840
< 27 kg/m**2	Week 24	Sparsentan	84	52 (61.9)	2.15 (2.02)	(-1.86, 6.16)	2.06 (2.92)	(-3.72, 7.84)	0.482
		Irbesartan	94	50 (53.2)	0.09 (2.09)	(-4.04, 4.23)			
	Week 48	Sparsentan	84	47 (56.0)	0.80 (2.07)	(-3.29, 4.89)	0.81 (3.07)	(-5.26, 6.88)	0.792
		Irbesartan	94	36 (38.3)	-0.01 (2.25)	(-4.46, 4.44)			
≥ 27 kg/m**2	Week 24	Sparsentan	118	73 (61.9)	0.04 (1.37)	(-2.65, 2.73)	1.58 (2.11)	(-2.57, 5.73)	0.455
		Irbesartan	107	54 (50.5)	-1.54 (1.58)	(-4.66, 1.58)			
	Week 48	Sparsentan	118	63 (53.4)	0.46 (1.46)	(-2.41, 3.34)	-1.70 (2.35)	(-6.33, 2.92)	0.469
		Irbesartan	107	41 (38.3)	2.17 (1.83)	(-1.44, 5.77)			

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LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Randomization strata	Overall		Sparsentan							Interaction: 0.339
eGFR Low and UP High	Week 24	Sparsentan	71	44 (62.0)	0.40 (1.23)	(-2.05, 2.84)	4.31 (1.91)	(0.54, 8.09)	0.025 *	
		Irbesartan	74	32 (43.2)	-3.92 (1.45)	(-6.79, -1.04)				
	Week 48	Sparsentan	71	37 (52.1)	0.54 (1.34)	(-2.11, 3.19)	1.04 (2.19)	(-3.29, 5.36)	0.635	
		Irbesartan	74	22 (29.7)	-0.50 (1.73)	(-3.92, 2.92)				
eGFR Low and UP Low	Week 24	Sparsentan	55	38 (69.1)	0.93 (2.09)	(-3.21, 5.07)	-3.67 (3.11)	(-9.83, 2.48)	0.240	
		Irbesartan	55	33 (60.0)	4.60 (2.24)	(0.17, 9.04)				
	Week 48	Sparsentan	55	28 (50.9)	0.58 (2.42)	(-4.21, 5.37)	-2.61 (3.55)	(-9.64, 4.42)	0.464	
		Irbesartan	55	25 (45.5)	3.19 (2.57)	(-1.90, 8.27)				
eGFR High and UP High	Week 24	Sparsentan	37	19 (51.4)	1.63 (3.54)	(-5.48, 8.74)	2.65 (5.20)	(-7.82, 13.12)	0.613	
		Irbesartan	36	18 (50.0)	-1.02 (3.81)	(-8.71, 6.67)				
	Week 48	Sparsentan	37	22 (59.5)	-1.81 (3.39)	(-8.64, 5.02)	-3.73 (5.59)	(-14.91, 7.46)	0.507	
		Irbesartan	36	11 (30.6)	1.92 (4.44)	(-6.95, 10.78)				
eGFR High and UP Low	Week 24	Sparsentan	39	24 (61.5)	2.21 (3.12)	(-4.06, 8.48)	5.31 (4.54)	(-3.78, 14.41)	0.246	
		Irbesartan	37	21 (56.8)	-3.10 (3.27)	(-9.66, 3.45)				
	Week 48	Sparsentan	39	23 (59.0)	3.37 (3.16)	(-2.96, 9.71)	4.84 (4.65)	(-4.45, 14.14)	0.301	
		Irbesartan	37	19 (51.4)	-1.47 (3.37)	(-8.22, 5.27)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline eGFR Group 1		Overall	Sparsentan							Interaction: 0.035 #
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	83 (65.4)	0.51 (1.25)	(-1.96, 2.97)	0.63 (1.87)	(-3.06, 4.32)	0.737	
		Irbesartan	129	68 (52.7)	-0.12 (1.39)	(-2.86, 2.61)				
	Week 48	Sparsentan	127	65 (51.2)	1.05 (1.40)	(-1.72, 3.81)	0.12 (2.16)	(-4.12, 4.37)	0.954	
		Irbesartan	129	49 (38.0)	0.92 (1.63)	(-2.28, 4.13)				
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	27 (55.1)	0.15 (2.64)	(-5.12, 5.41)	0.24 (3.86)	(-7.48, 7.96)	0.951	
		Irbesartan	48	24 (50.0)	-0.09 (2.80)	(-5.68, 5.50)				
	Week 48	Sparsentan	49	29 (59.2)	-0.18 (2.59)	(-5.37, 5.00)	-0.91 (4.00)	(-8.88, 7.06)	0.820	
		Irbesartan	48	19 (39.6)	0.73 (2.99)	(-5.22, 6.68)				
≥ 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	15 (57.7)	7.75 (4.89)	(-2.26, 17.76)	15.68 (7.37)	(0.59, 30.76)	0.042 *	
		Irbesartan	25	12 (48.0)	-7.93 (5.42)	(-19.01, 3.15)				
	Week 48	Sparsentan	26	16 (61.5)	2.19 (4.87)	(-7.78, 12.16)	6.01 (7.71)	(-9.69, 21.70)	0.442	
		Irbesartan	25	9 (36.0)	-3.82 (5.83)	(-15.64, 8.00)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.  
LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.  
For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.  
A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.056
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	53 (64.6)	1.83 (1.73)	(-1.59, 5.26)	2.63 (2.55)	(-2.42, 7.67)	0.305
		Irbesartan	80	45 (56.3)	-0.79 (1.88)	(-4.50, 2.91)			
	Week 48	Sparsentan	82	42 (51.2)	0.47 (1.93)	(-3.34, 4.28)	0.58 (3.03)	(-5.40, 6.56)	0.849
		Irbesartan	80	29 (36.3)	-0.11 (2.33)	(-4.71, 4.49)			
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	30 (66.7)	-1.94 (1.62)	(-5.17, 1.29)	-3.52 (2.54)	(-8.57, 1.53)	0.169
		Irbesartan	49	23 (46.9)	1.58 (1.90)	(-2.20, 5.36)			
	Week 48	Sparsentan	45	23 (51.1)	2.08 (1.84)	(-1.57, 5.74)	-0.29 (2.75)	(-5.75, 5.17)	0.917
		Irbesartan	49	20 (40.8)	2.37 (2.01)	(-1.63, 6.37)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	27 (55.1)	0.15 (2.64)	(-5.12, 5.41)	0.24 (3.86)	(-7.48, 7.96)	0.951
		Irbesartan	48	24 (50.0)	-0.09 (2.80)	(-5.68, 5.50)			
	Week 48	Sparsentan	49	29 (59.2)	-0.18 (2.59)	(-5.37, 5.00)	-0.91 (4.00)	(-8.88, 7.06)	0.820
		Irbesartan	48	19 (39.6)	0.73 (2.99)	(-5.22, 6.68)			
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	15 (57.7)	7.75 (4.89)	(-2.26, 17.76)	15.68 (7.37)	(0.59, 30.76)	0.042 *
		Irbesartan	25	12 (48.0)	-7.93 (5.42)	(-19.01, 3.15)			
	Week 48	Sparsentan	26	16 (61.5)	2.19 (4.87)	(-7.78, 12.16)	6.01 (7.71)	(-9.69, 21.70)	0.442
		Irbesartan	25	9 (36.0)	-3.82 (5.83)	(-15.64, 8.00)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.614
<= 1.75 g/day	Week 24	Sparsentan	98	66 (67.3)	0.71 (1.74)	(-2.72, 4.14)	0.83 (2.79)	(-4.68, 6.34)	0.767
		Irbesartan	94	43 (45.7)	-0.12 (2.16)	(-4.38, 4.15)			
	Week 48	Sparsentan	98	59 (60.2)	1.42 (1.82)	(-2.18, 5.02)	-0.50 (3.06)	(-6.54, 5.53)	0.869
		Irbesartan	94	34 (36.2)	1.93 (2.42)	(-2.85, 6.71)			
> 1.75 g/day	Week 24	Sparsentan	104	59 (56.7)	1.69 (1.51)	(-1.28, 4.66)	3.87 (2.13)	(-0.35, 8.09)	0.072
		Irbesartan	108	61 (56.5)	-2.18 (1.50)	(-5.14, 0.78)			
	Week 48	Sparsentan	104	51 (49.0)	0.60 (1.59)	(-2.53, 3.74)	0.52 (2.35)	(-4.12, 5.16)	0.825
		Irbesartan	108	43 (39.8)	0.08 (1.70)	(-3.28, 3.44)			

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LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

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Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.935
Yes	Week 24	Sparsentan	88	50 (56.8)	2.32 (1.31)	(-0.28, 4.92)	3.23 (1.98)	(-0.70, 7.16)	0.106
		Irbesartan	83	39 (47.0)	-0.92 (1.48)	(-3.86, 2.03)			
	Week 48	Sparsentan	88	41 (46.6)	0.71 (1.38)	(-2.03, 3.44)	-0.84 (2.14)	(-5.09, 3.40)	0.694
		Irbesartan	83	28 (33.7)	1.55 (1.63)	(-1.68, 4.77)			
No	Week 24	Sparsentan	114	75 (65.8)	0.28 (1.67)	(-3.01, 3.58)	1.44 (2.46)	(-3.40, 6.28)	0.559
		Irbesartan	119	65 (54.6)	-1.16 (1.80)	(-4.70, 2.39)			
	Week 48	Sparsentan	114	69 (60.5)	0.92 (1.73)	(-2.49, 4.33)	0.38 (2.69)	(-4.92, 5.68)	0.888
		Irbesartan	119	49 (41.2)	0.54 (2.06)	(-3.51, 4.59)			

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Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.020 #
<= 5 years	Week 24	Sparsentan	113	70 (61.9)	4.13 (1.47)	(1.24, 7.02)	5.94 (2.10)	(1.80, 10.08)	0.005 *
		Irbesartan	127	67 (52.8)	-1.81 (1.50)	(-4.76, 1.14)			
	Week 48	Sparsentan	113	61 (54.0)	2.91 (1.54)	(-0.12, 5.93)	2.37 (2.26)	(-2.08, 6.83)	0.295
		Irbesartan	127	51 (40.2)	0.53 (1.66)	(-2.73, 3.80)			
> 5 years	Week 24	Sparsentan	89	55 (61.8)	-2.90 (1.78)	(-6.41, 0.61)	-3.56 (2.81)	(-9.12, 1.99)	0.207
		Irbesartan	75	37 (49.3)	0.66 (2.19)	(-3.67, 5.00)			
	Week 48	Sparsentan	89	49 (55.1)	-1.11 (1.90)	(-4.86, 2.63)	-3.30 (3.23)	(-9.69, 3.08)	0.308
		Irbesartan	75	26 (34.7)	2.19 (2.60)	(-2.95, 7.33)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024



Table PT1KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.121
Yes	Week 24	Sparsentan	153	94 (61.4)	2.84 (1.01)	(0.85, 4.83)	1.86 (1.51)	(-1.12, 4.83)	0.221
		Irbesartan	157	76 (48.4)	0.98 (1.12)	(-1.23, 3.19)			
	Week 48	Sparsentan	153	81 (52.9)	1.05 (1.08)	(-1.08, 3.18)	-2.58 (1.70)	(-5.92, 0.76)	0.129
		Irbesartan	157	57 (36.3)	3.63 (1.30)	(1.07, 6.19)			
No	Week 24	Sparsentan	49	31 (63.3)	-3.77 (3.43)	(-10.62, 3.08)	2.70 (5.03)	(-7.34, 12.75)	0.593
		Irbesartan	45	28 (62.2)	-6.48 (3.66)	(-13.78, 0.83)			
	Week 48	Sparsentan	49	29 (59.2)	-0.70 (3.48)	(-7.63, 6.24)	8.12 (5.29)	(-2.41, 18.66)	0.129
		Irbesartan	45	20 (44.4)	-8.82 (3.96)	(-16.69, -0.94)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

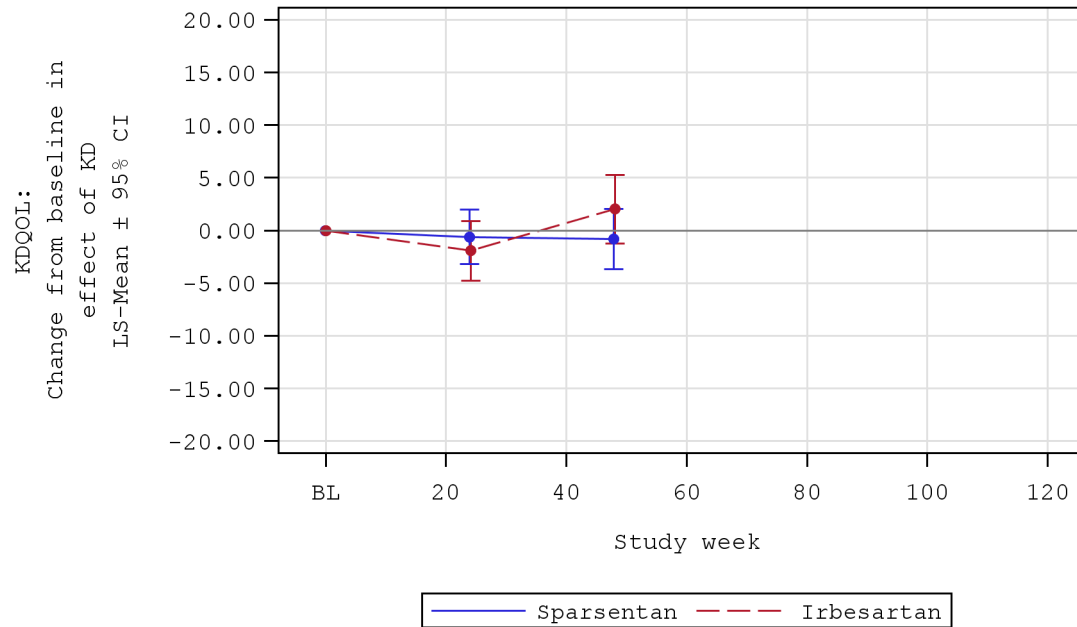
A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

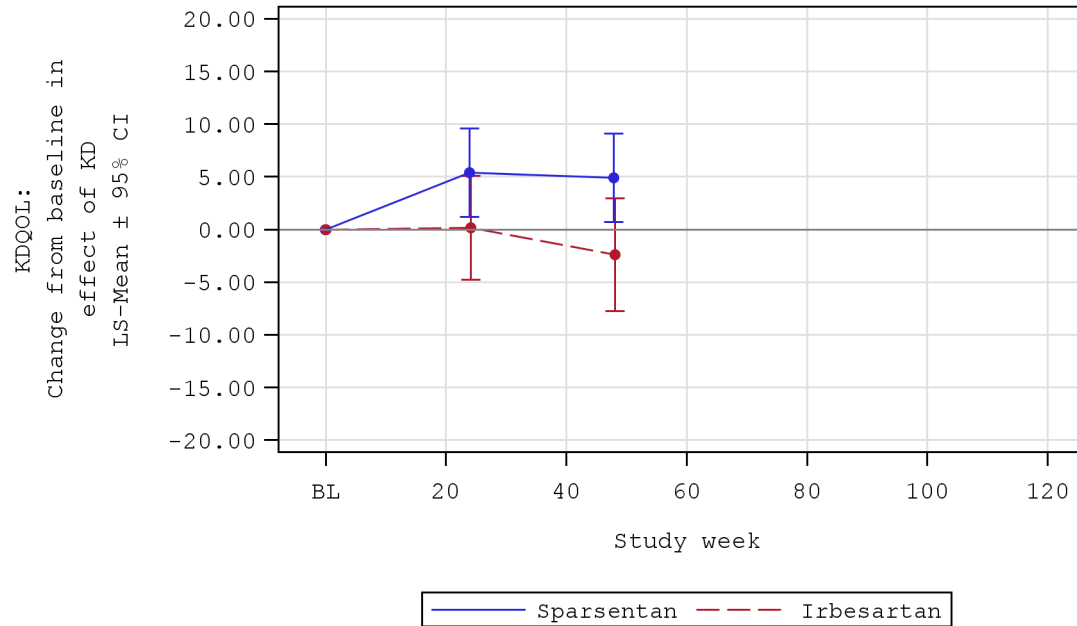
Figure PF1KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Sex: Male



Sparsentan	87	72
Irbesartan	73	55

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
 95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KEFC\_FSCM.

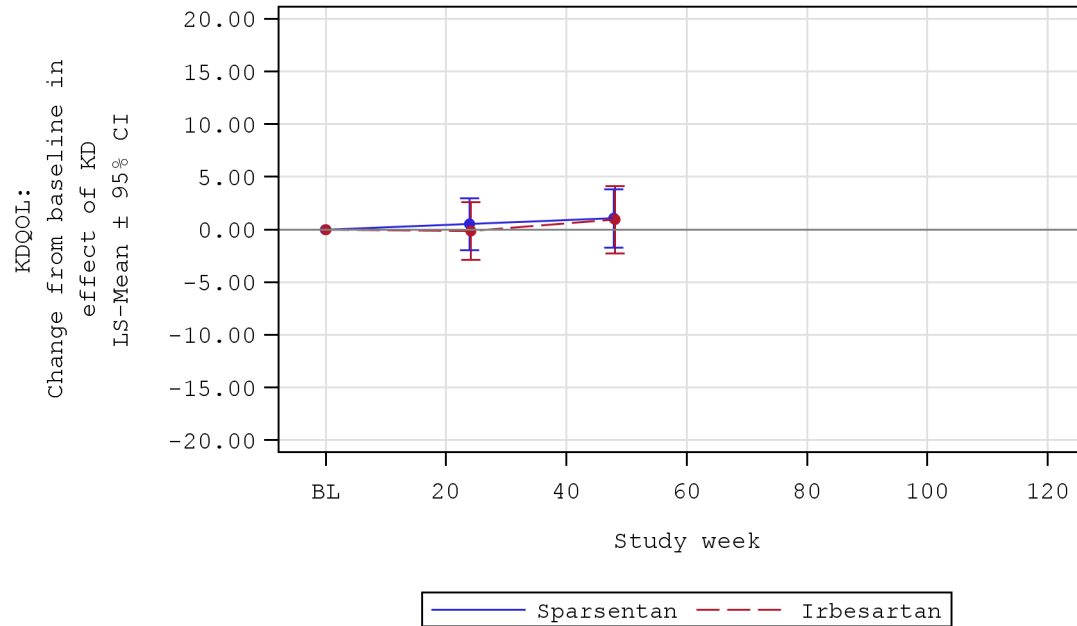
Figure PF1KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Sex: Female



Sparsentan	38	38
Irbesartan	31	22

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
 95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KEFC\_FSCM.

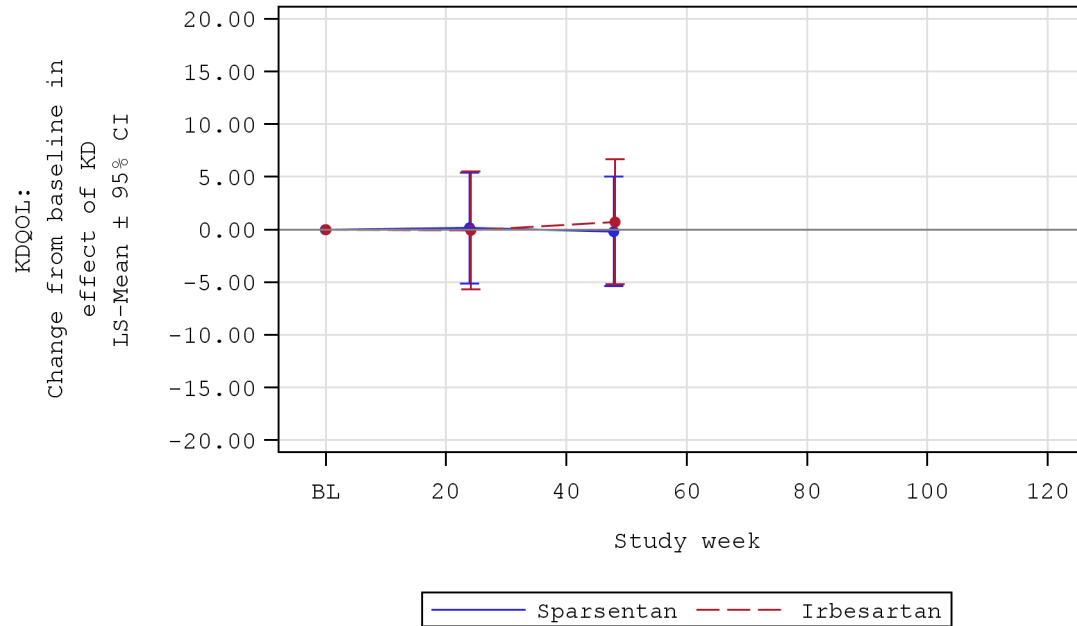
Figure PF1KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1: < 60 mL/min/1.73 m\*\*2



Sparsentan	83	65
Irbesartan	68	49

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
 95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KEFC\_FSCM.

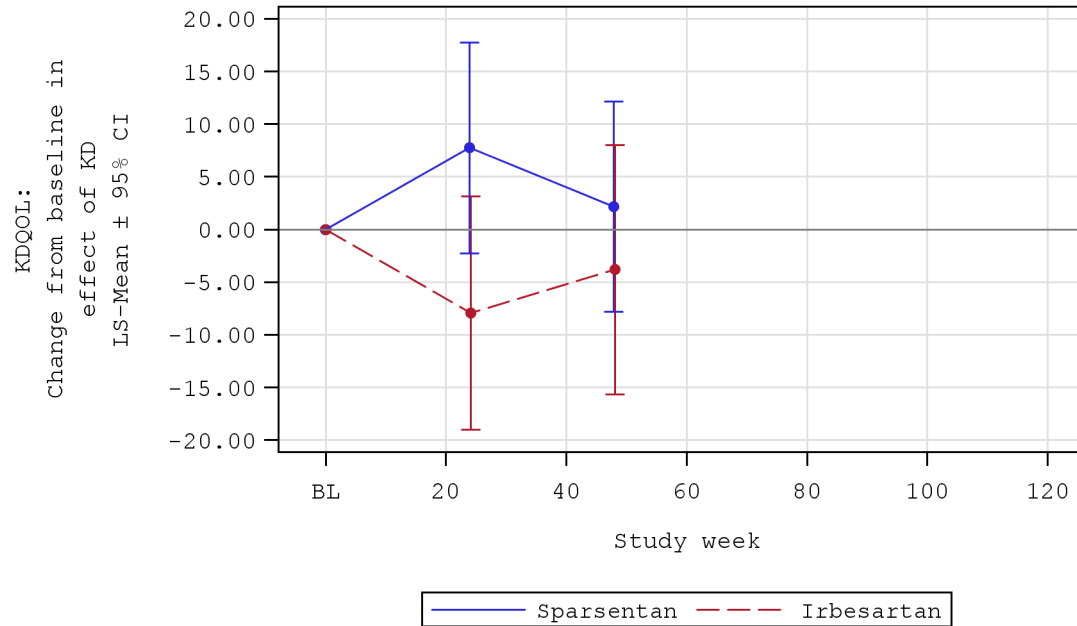
Figure PF1KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1: 60 to < 90 mL/min/1.73 m\*\*2



Sparsentan	27	29
Irbesartan	24	19

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
 95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KEFC\_FSCM.

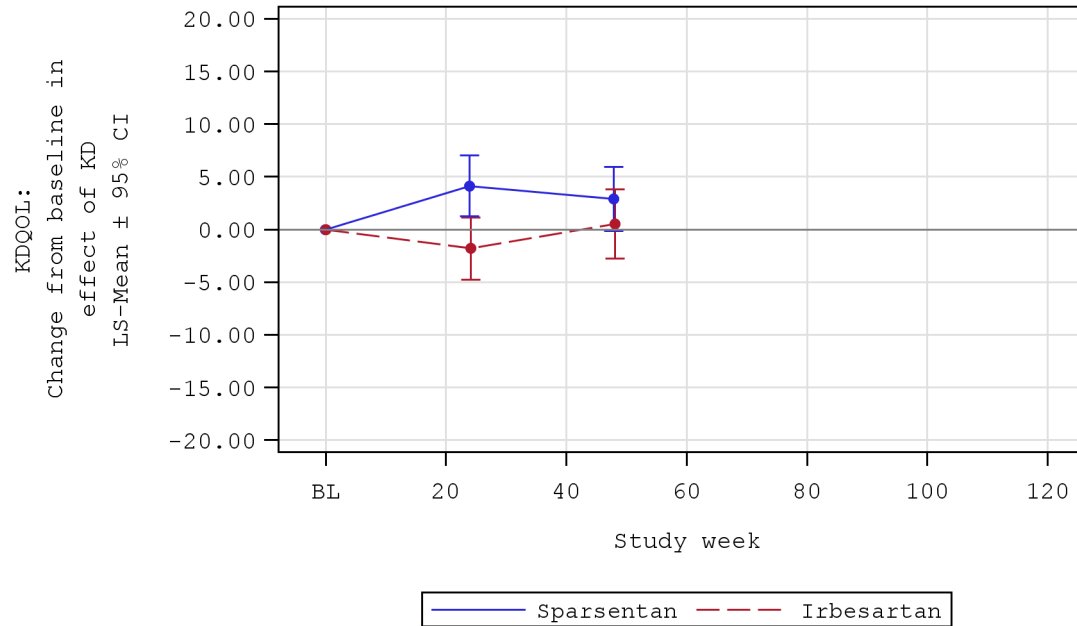
Figure PF1KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1:  $\geq 90$  mL/min/1.73 m<sup>2</sup>



Sparsentan	15	16
Irbesartan	12	9

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
 95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
 eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High  $\geq 60$  ml/min/1.73m<sup>2</sup>, UP Low =  $\leq 1.75$  g/day, UP High =  $> 1.75$  g/day.  
 IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KEFC\_FSCM.

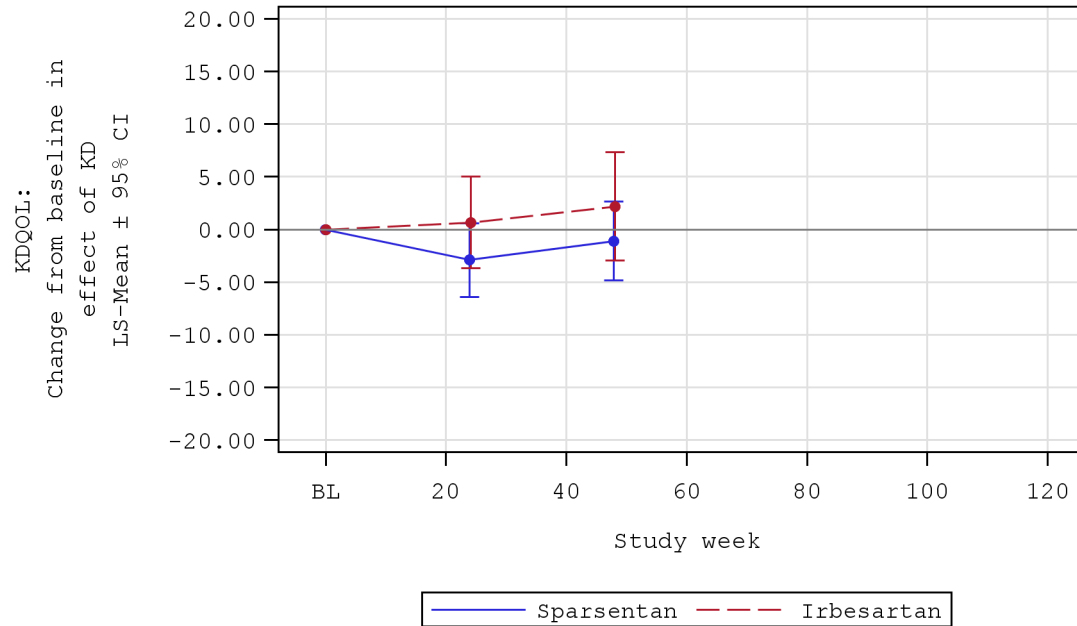
Figure PF1KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Time since renal biopsy: <= 5 years



Sparsentan	70	61
Irbesartan	67	51

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
 95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KEFC\_FSCM.

Figure PF1KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Time since renal biopsy: > 5 years



Sparsentan	55	49
Irbesartan	37	26

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
 95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KEFC\_FSCM.



Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.229
Male	Sparsentan	139	11 (7.9)	NE		1.979	(0.707, 5.539)	0.194
	Irbesartan	143	9 (6.3)	NE				
Female	Sparsentan	63	12 (19.0)	NE		0.744	(0.191, 2.901)	0.671
	Irbesartan	59	5 (8.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan							Interaction test: 0.851
<= 45 years	Sparsentan	96	13 (13.5)	NE		1.613	(0.585, 4.446)	0.355
	Irbesartan	99	7 (7.1)	NE				
> 45 years	Sparsentan	106	10 (9.4)	NE		1.354	(0.428, 4.280)	0.606
	Irbesartan	103	7 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: <0.001 #
<= 18 years	Sparsentan	9	2 (22.2)	NE		0.191	(0.005, 6.762)	0.363
	Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Sparsentan	102	11 (10.8)	NE		1.821	(0.698, 4.750)	0.221
	Irbesartan	109	9 (8.3)	NE				
> 40 years	Sparsentan	91	10 (11.0)	NE		1.489	(0.411, 5.392)	0.545
	Irbesartan	88	4 (4.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan							
							Interaction test:	0.009 #
North America	Sparsentan	35	3 (8.6)	NE		0.729	(0.101, 5.236)	0.753
	Irbesartan	46	3 (6.5)	NE				
Europe	Sparsentan	98	7 (7.1)	NE		0.723	(0.223, 2.343)	0.589
	Irbesartan	115	9 (7.8)	NE				
Asia Pacific	Sparsentan	69	13 (18.8)	NE		4.884	(0.941, 25.347)	0.059
	Irbesartan	41	2 (4.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.860
< 27 kg/m**2	Sparsentan	84	14 (16.7)	NE		1.392	(0.538, 3.602)	0.495
	Irbesartan	94	8 (8.5)	NE				
>= 27 kg/m**2	Sparsentan	118	9 (7.6)	NE		1.322	(0.395, 4.431)	0.651
	Irbesartan	107	6 (5.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.008 #
eGFR Low and UP High	Sparsentan	71	3 (4.2)	NE		0.316	(0.052, 1.923)	0.211
	Irbesartan	74	3 (4.1)	NE				
eGFR Low and UP Low	Sparsentan	55	9 (16.4)	NE		0.998	(0.246, 4.050)	0.997
	Irbesartan	55	4 (7.3)	NE				
eGFR High and UP High	Sparsentan	37	5 (13.5)	NE		0.542	(0.146, 2.015)	0.360
	Irbesartan	36	5 (13.9)	NE				
eGFR High and UP Low	Sparsentan	39	6 (15.4)	NE		11.943	(1.281, 111.381)	0.029 *
	Irbesartan	37	2 (5.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.761
< 60 mL/min/1.73 m**2	Sparsentan	127	12 (9.4)	NE		0.812	(0.286, 2.306)	0.696
	Irbesartan	129	7 (5.4)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	6 (12.2)	NE		1.919	(0.458, 8.045)	0.372
	Irbesartan	48	6 (12.5)	93.3	(70.1, NE)			
≥ 90 mL/min/1.73 m**2	Sparsentan	26	5 (19.2)	NE		3.022	(0.301, 30.297)	0.347
	Irbesartan	25	1 (4.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.906
< 45 mL/min/1.73 m**2	Sparsentan	82	7 (8.5)	NE		0.667	(0.153, 2.914)	0.590
	Irbesartan	80	4 (5.0)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	5 (11.1)	NE		0.354	(0.046, 2.713)	0.318
	Irbesartan	49	3 (6.1)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	6 (12.2)	NE		1.919	(0.458, 8.045)	0.372
	Irbesartan	48	6 (12.5)	93.3	(70.1, NE)			
≥ 90 mL/min/1.73 m**2	Sparsentan	26	5 (19.2)	NE		3.022	(0.301, 30.297)	0.347
	Irbesartan	25	1 (4.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.024 #
<= 1.75 g/day	Sparsentan	98	15 (15.3)	NE		3.533	(1.085, 11.498)	0.036 *
	Irbesartan	94	5 (5.3)	NE				
> 1.75 g/day	Sparsentan	104	8 (7.7)	NE		0.569	(0.198, 1.637)	0.296
	Irbesartan	108	9 (8.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.329
Yes	Sparsentan	88	8 (9.1)	NE		1.549	(0.395, 6.065)	0.530
	Irbesartan	83	4 (4.8)	NE				
No	Sparsentan	114	15 (13.2)	NE		1.073	(0.451, 2.554)	0.874
	Irbesartan	119	10 (8.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: <0.001 #
<= 5 years	Sparsentan	113	18 (15.9)	NE		4.893	(1.782, 13.436)	0.002 *
	Irbesartan	127	6 (4.7)	NE				
> 5 years	Sparsentan	89	5 (5.6)	NE		0.035	(0.004, 0.308)	0.003 *
	Irbesartan	75	8 (10.7)	108.1	(108.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
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 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

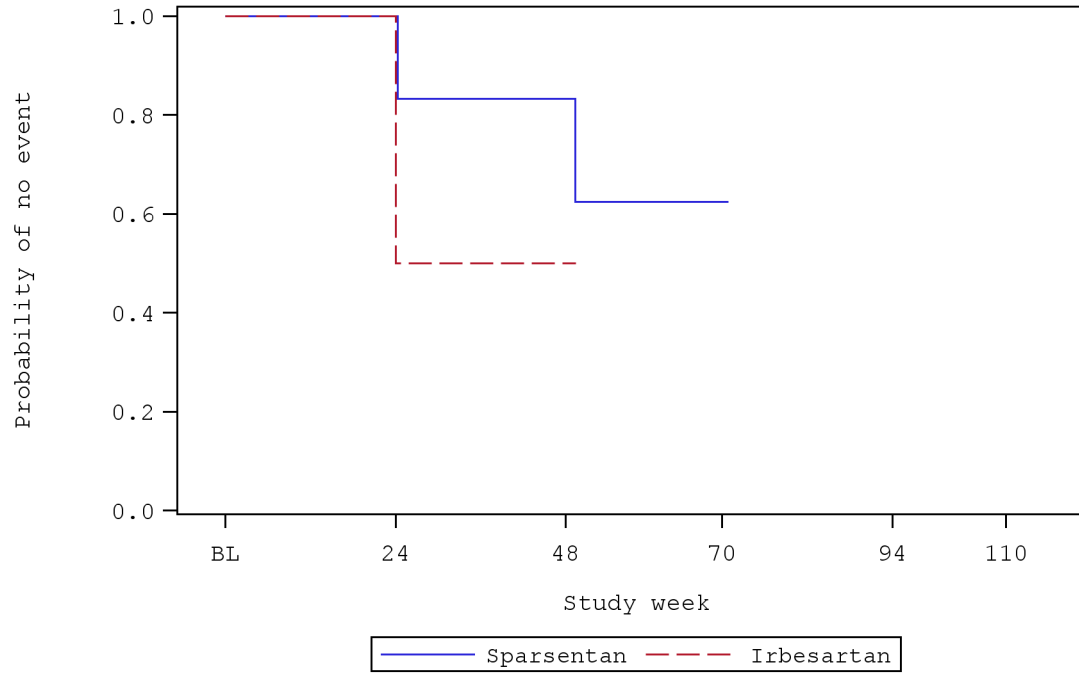
Table PT1KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.020 #
Yes	Sparsentan	153	17 (11.1)	NE		0.692	(0.302, 1.584)	0.383
	Irbesartan	157	12 (7.6)	NE				
No	Sparsentan	49	6 (12.2)	NE		15.899	(1.260, 200.530)	0.032 *
	Irbesartan	45	2 (4.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

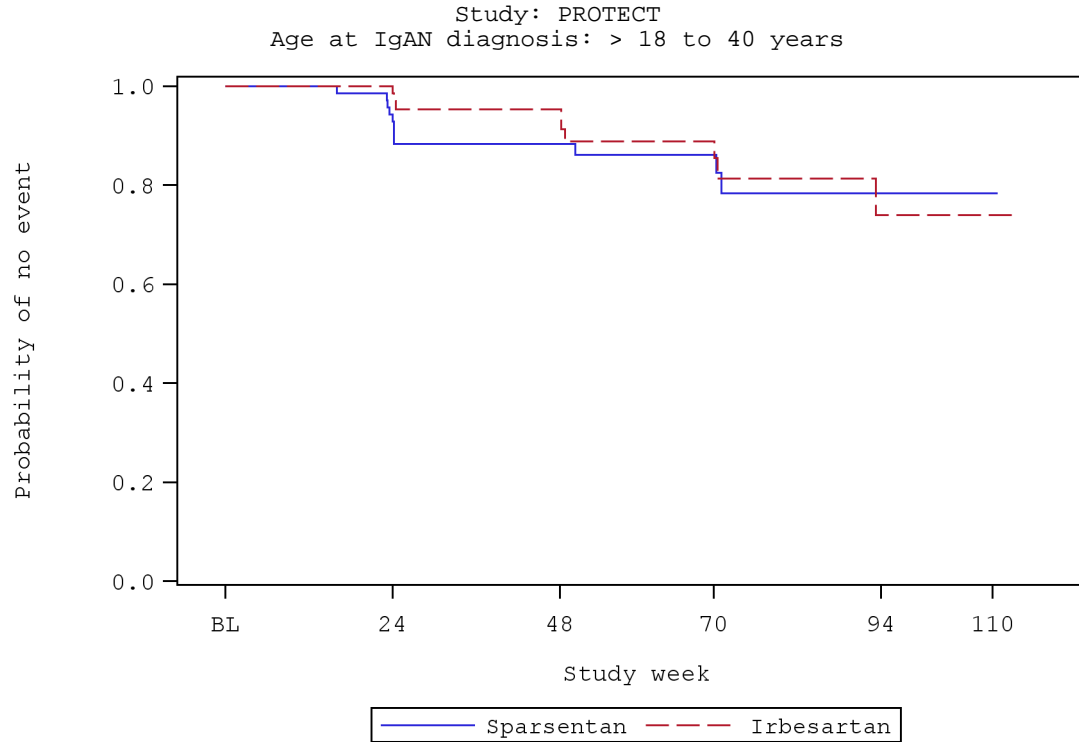
Study: PROTECT  
 Age at IgAN diagnosis: <= 18 years



Sparsentan	9	6	4	1	0
Irbesartan	5	2	1	0	

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

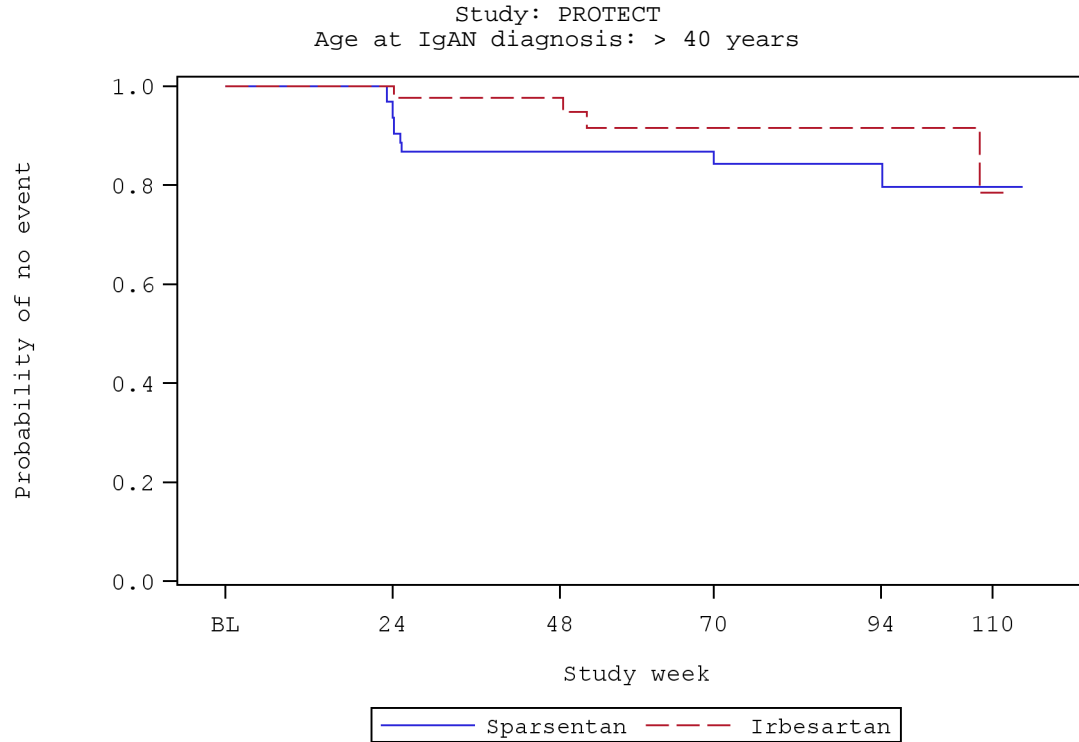
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	102	65	50	33	10	3
Irbesartan	109	68	47	27	9	5

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

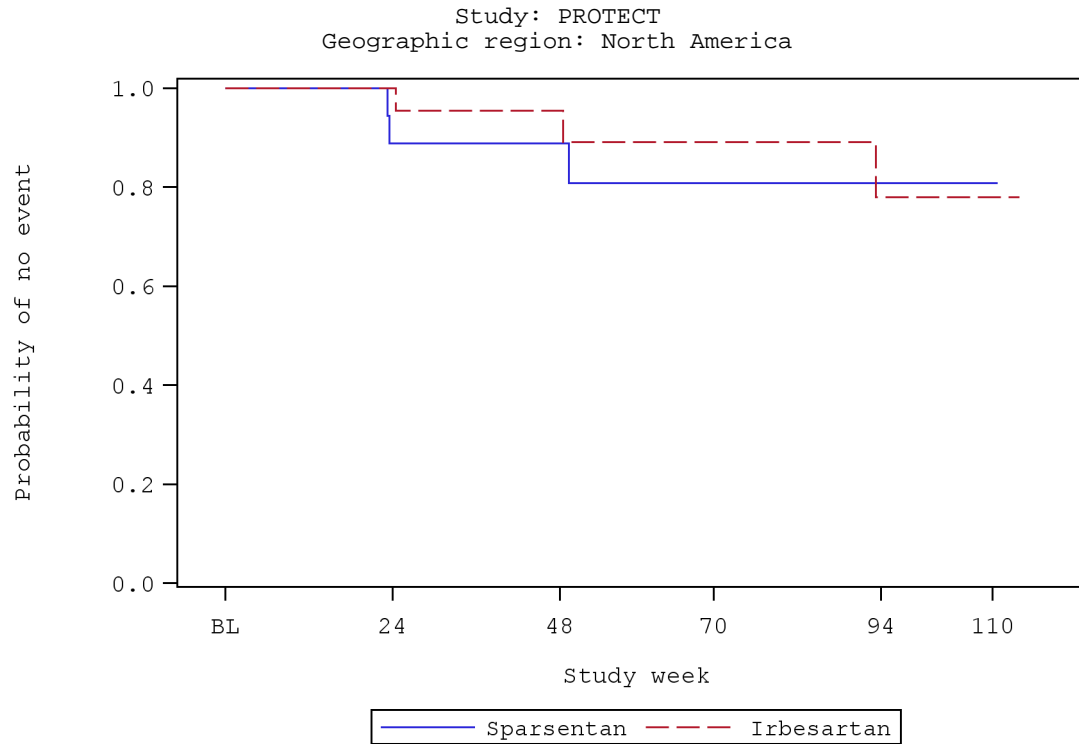
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	91	60	45	36	19	9
Irbesartan	88	45	37	24	12	4

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

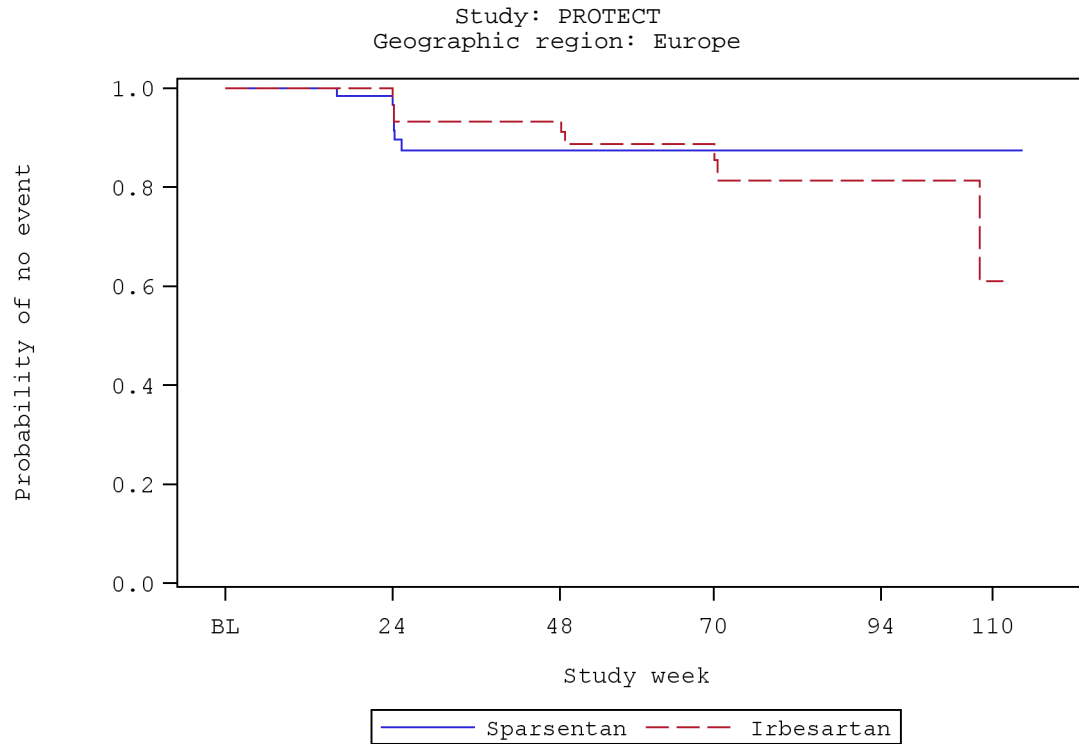


Sparsentan	35	16	11	8	5	1
Irbesartan	46	25	19	11	6	5

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM



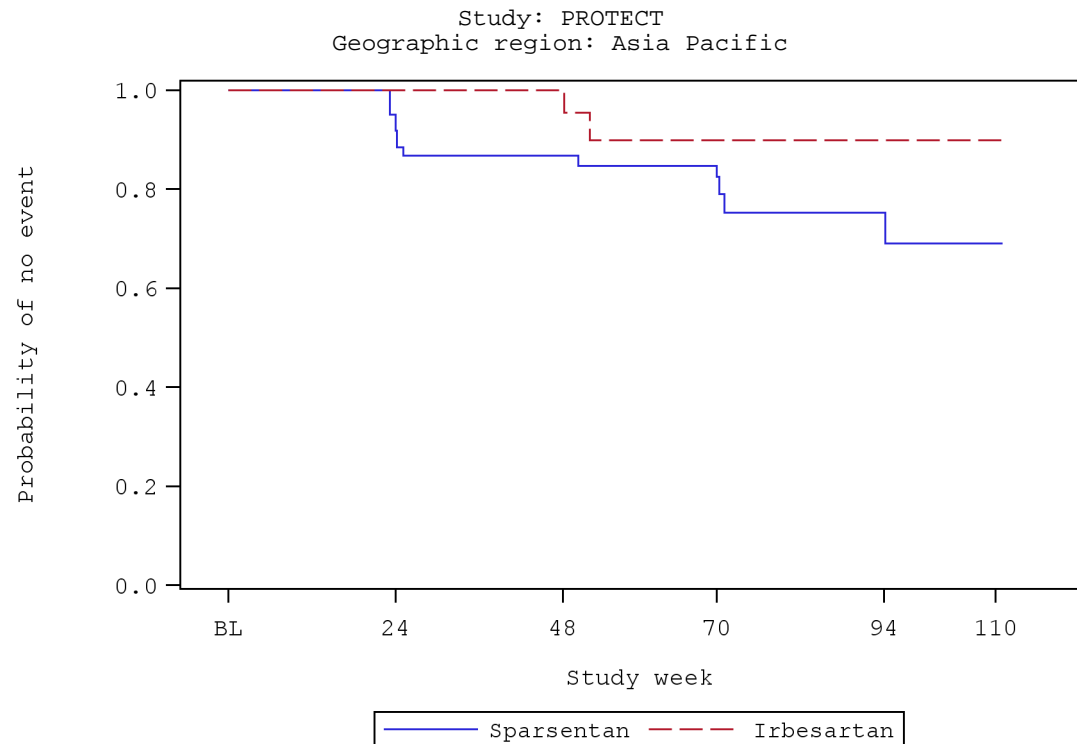
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	98	57	39	25	12	6
Irbesartan	115	60	44	28	10	2

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

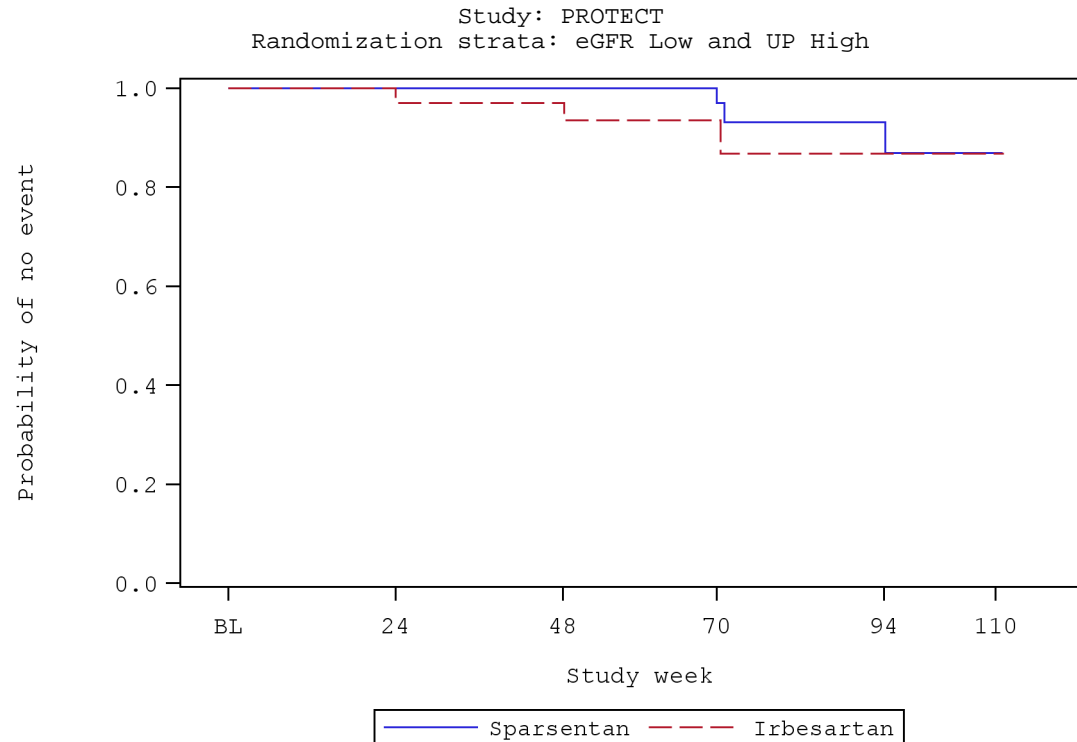
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	69	58	49	37	12	5
Irbesartan	41	30	22	12	5	2

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

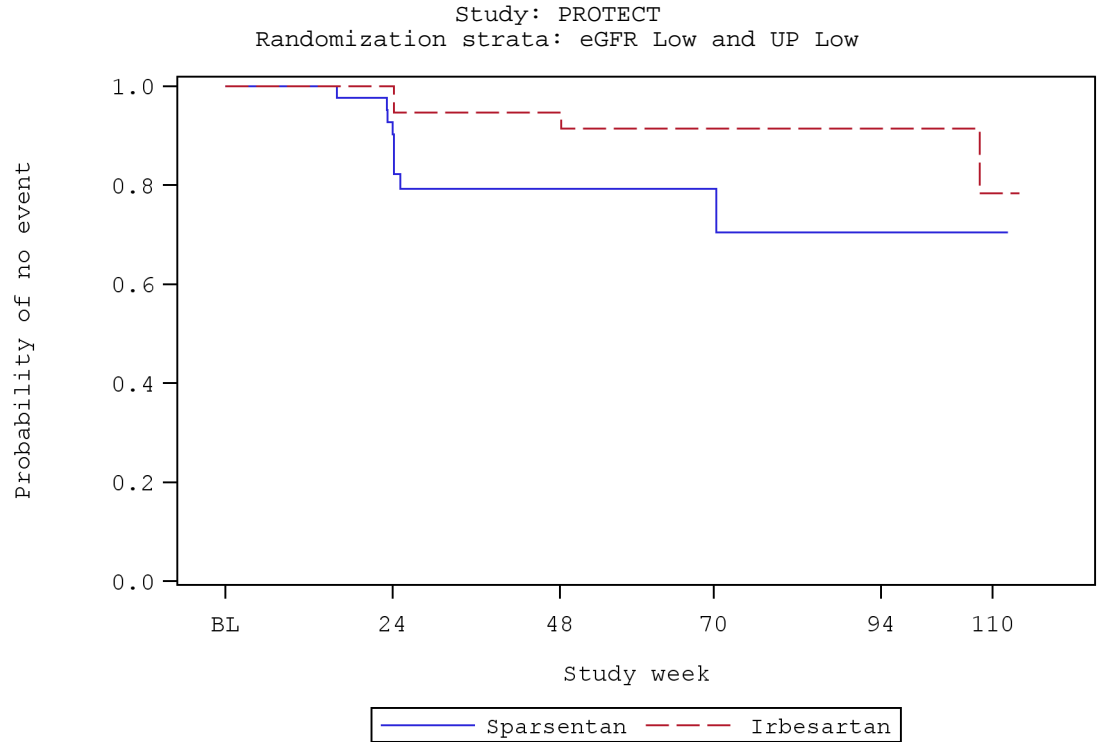
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	71	47	41	34	16	3
Irbesartan	74	34	27	14	7	2

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

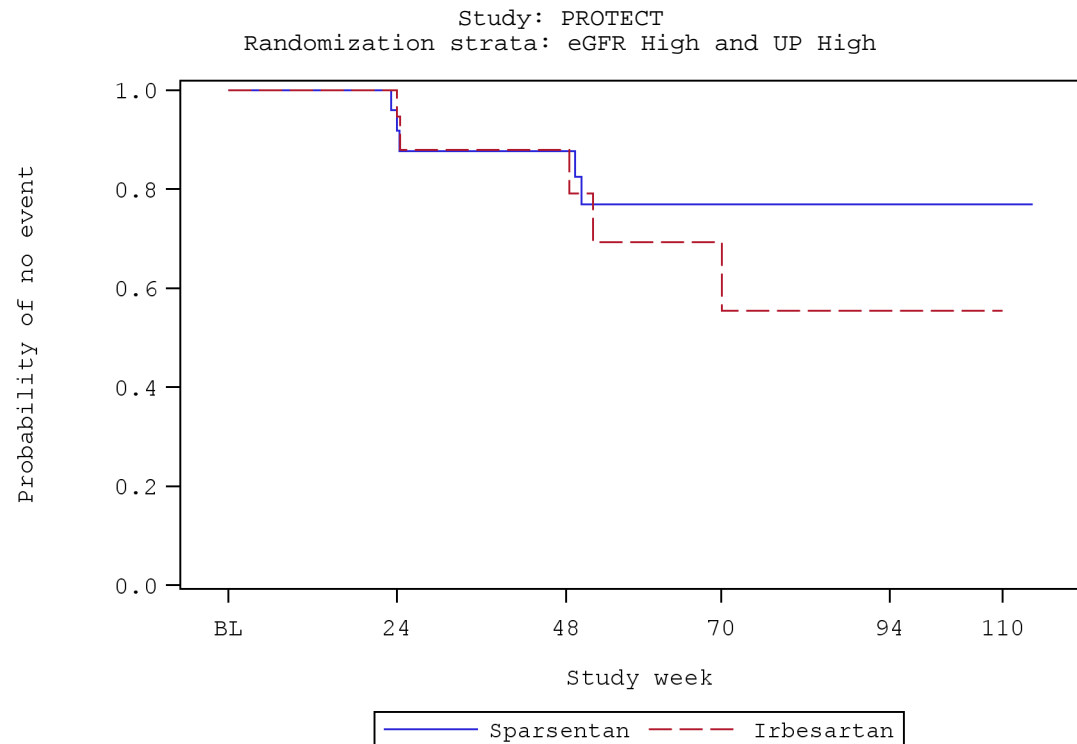
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	55	37	23	15	6	6
Irbesartan	55	38	29	20	10	5

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

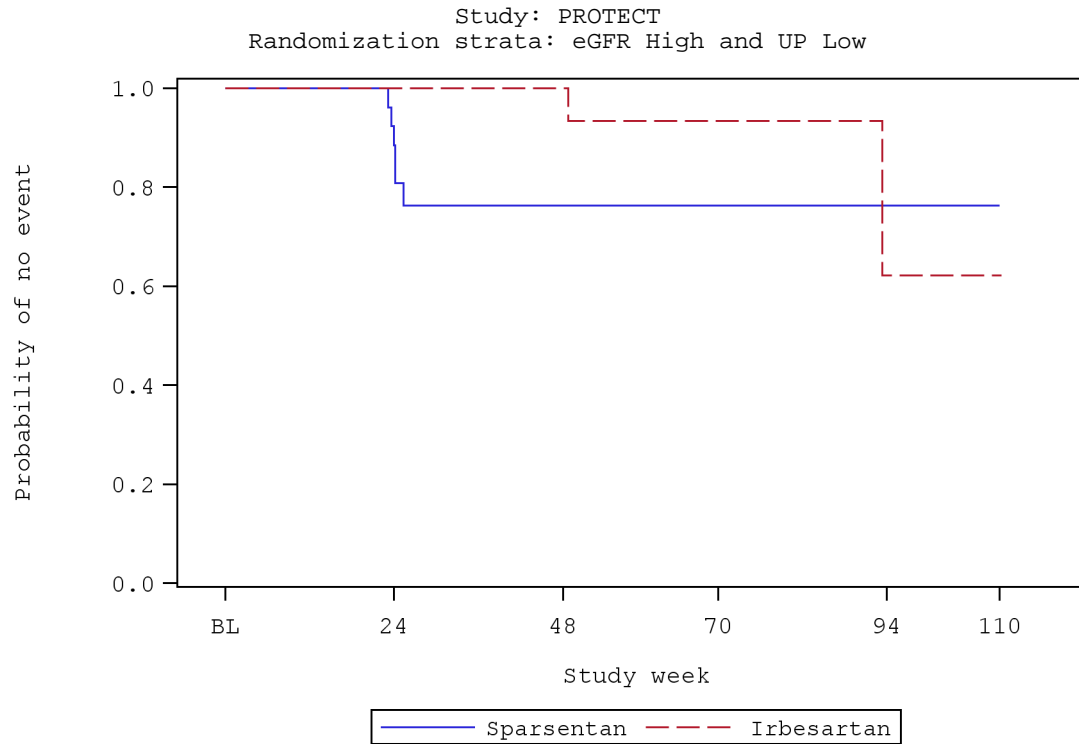
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	37	23	19	12	5	2
Irbesartan	36	19	11	5	2	1

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

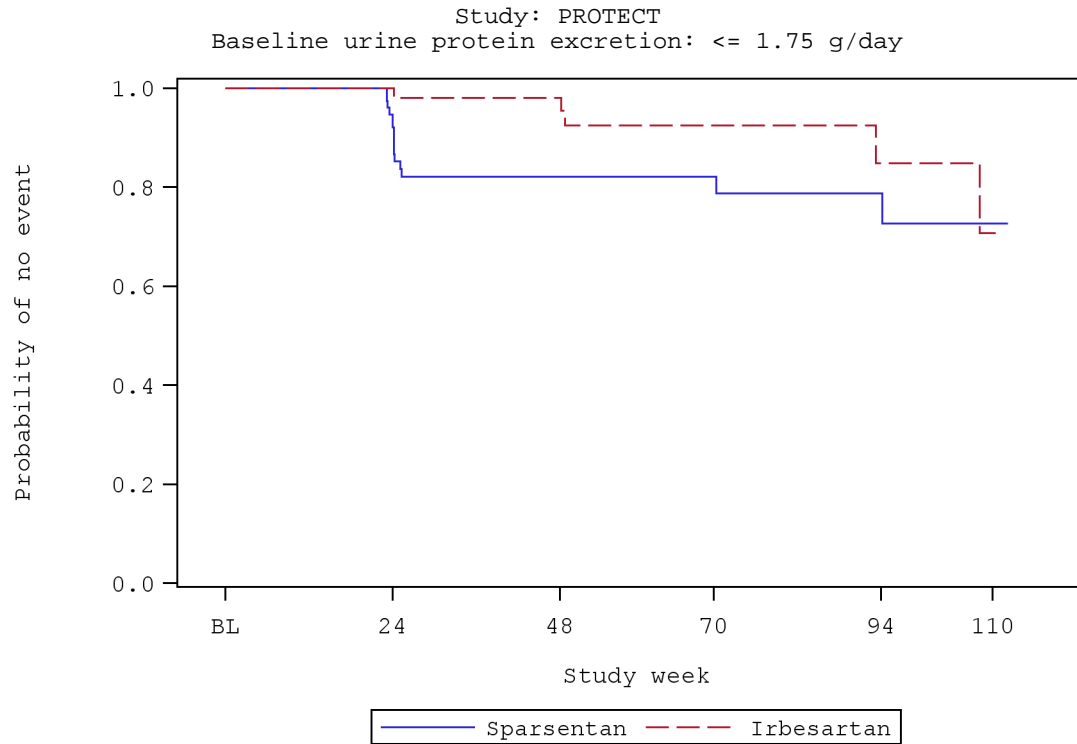
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	39	24	16	9	2	1
Irbesartan	37	24	18	12	2	1

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

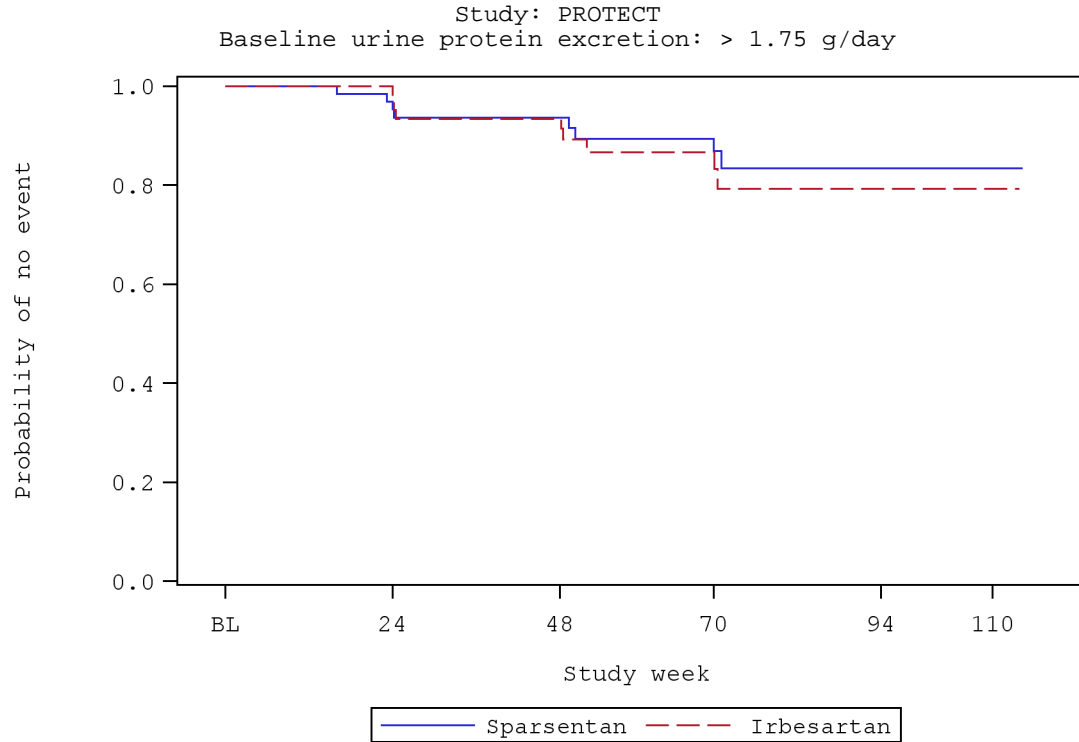
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	98	71	50	33	13	9
Irbesartan	94	52	38	25	10	4

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

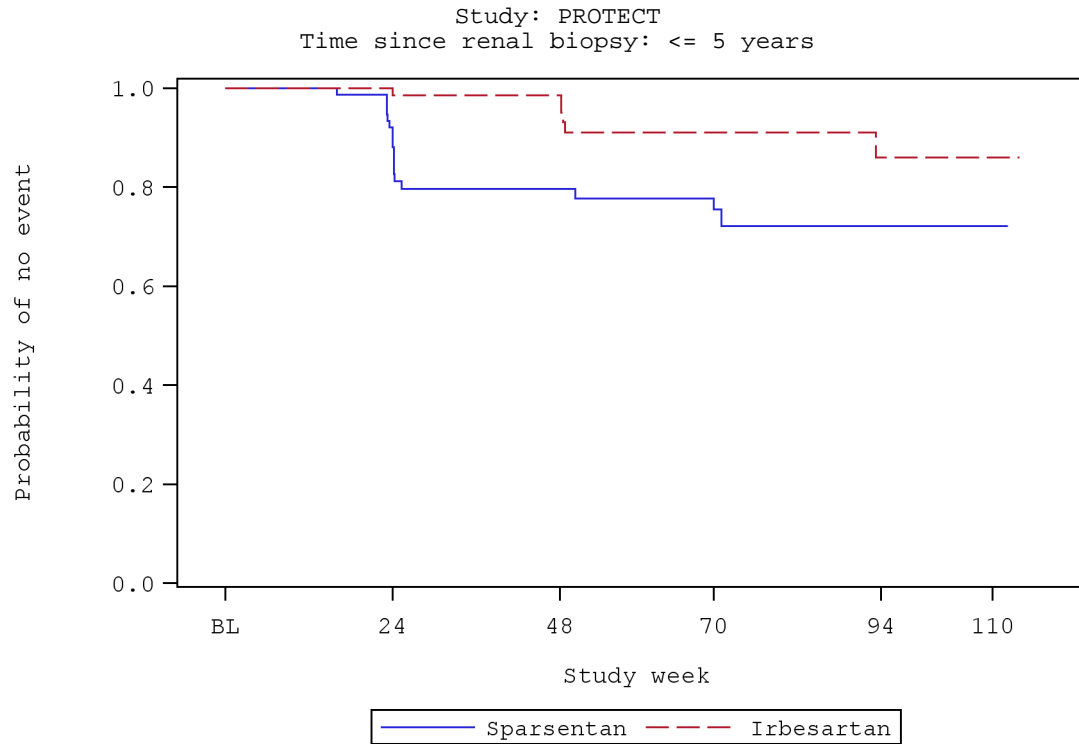


Sparsentan	104	60	49	37	16	3
Irbesartan	108	63	47	26	11	5

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM



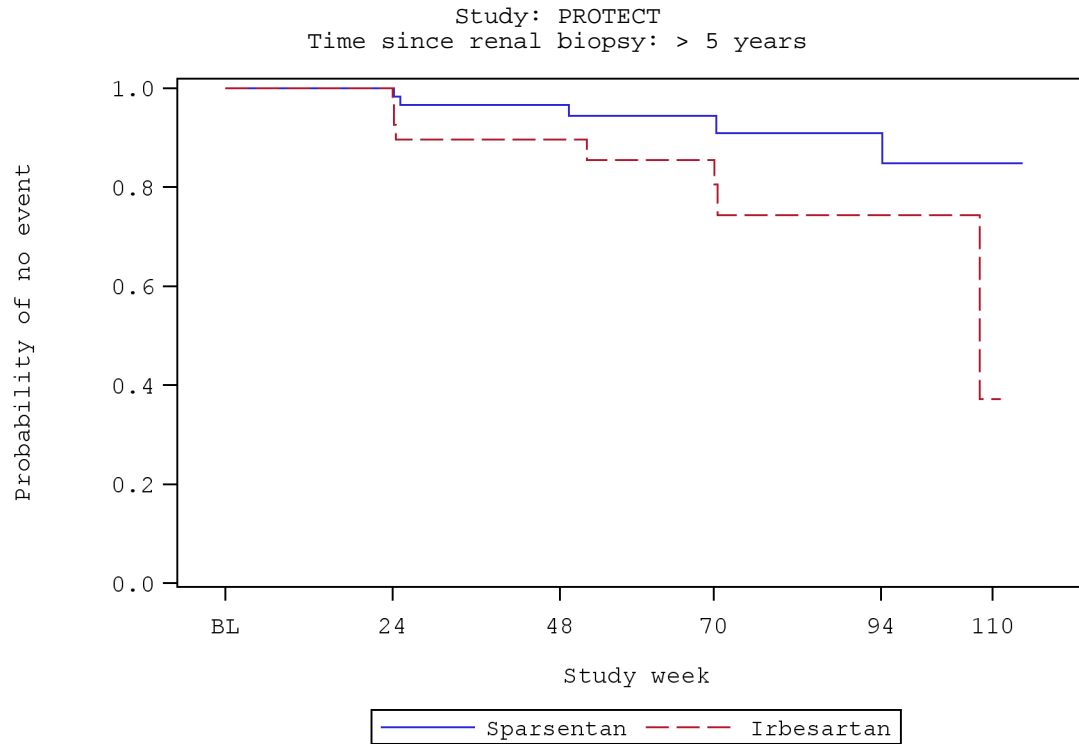
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	113	69	48	35	14	5
Irbesartan	127	73	56	33	15	8

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

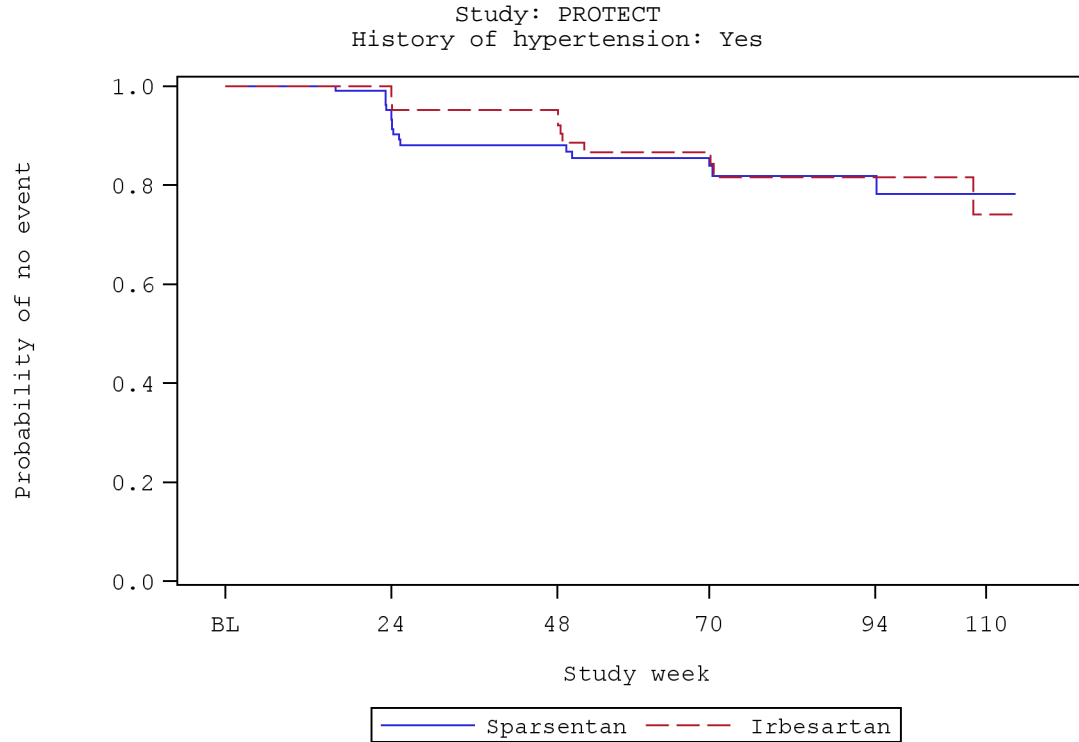
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	89	62	51	35	15	7
Irbesartan	75	42	29	18	6	1

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

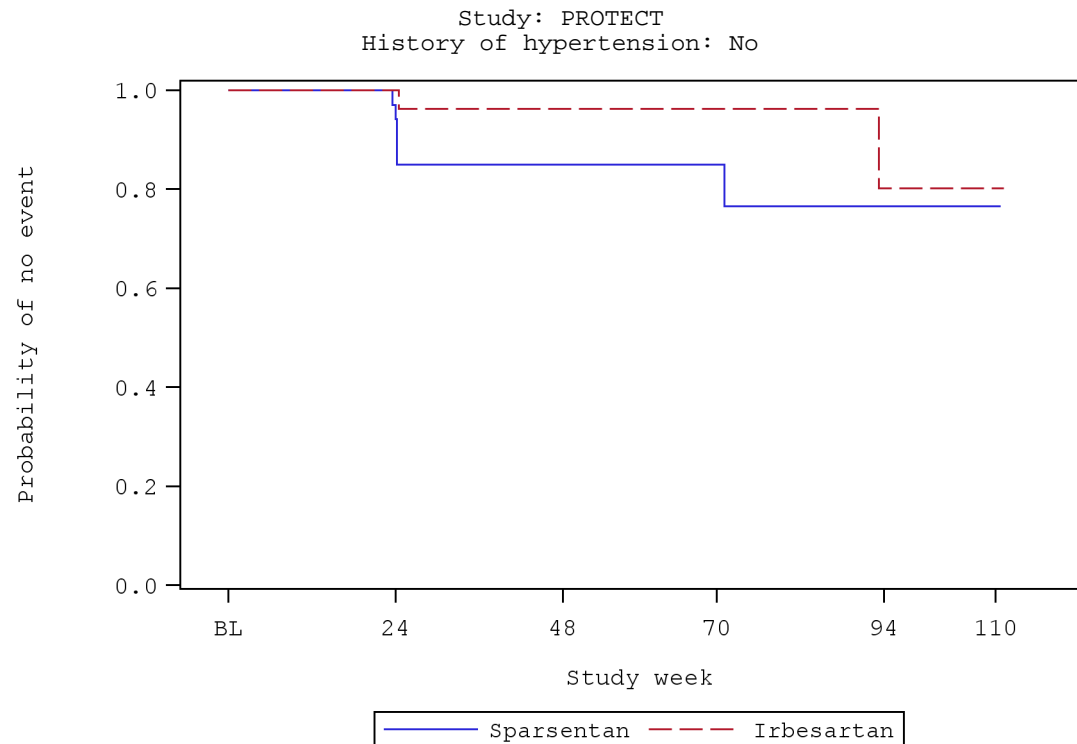
Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	153	98	75	56	23	9
Irbesartan	157	83	63	38	16	7

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

Figure PF1KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	49	33	24	14	6	3
Irbesartan	45	32	22	13	5	2

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KEFIT\_FSTM

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.047 #
Male	Sparsentan	139	17 (12.2)	NE		0.843	(0.416, 1.711)	0.637
	Irbesartan	143	15 (10.5)	NE				
Female	Sparsentan	63	1 (1.6)	NE		0.133	(0.015, 1.199)	0.072
	Irbesartan	59	5 (8.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan							
							Interaction test:	0.255
<= 45 years	Sparsentan	96	11 (11.5)	NE		0.863	(0.372, 1.998)	0.730
	Irbesartan	99	11 (11.1)	NE				
> 45 years	Sparsentan	106	7 (6.6)	NE		0.512	(0.183, 1.433)	0.202
	Irbesartan	103	9 (8.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	1 (11.1) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	11 (10.8)	NE		0.653	(0.288, 1.481)	0.308
	Irbesartan	109	13 (11.9)	NE				
> 40 years	Sparsentan	91	6 (6.6)	NE		0.606	(0.197, 1.859)	0.381
	Irbesartan	88	7 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.665
North America	Sparsentan	35	2 (5.7)	NE		0.367	(0.071, 1.900)	0.232
	Irbesartan	46	6 (13.0)	NE				
Europe	Sparsentan	98	6 (6.1)	NE		0.488	(0.174, 1.366)	0.172
	Irbesartan	115	10 (8.7)	NE				
Asia Pacific	Sparsentan	69	10 (14.5)	NE		1.132	(0.338, 3.786)	0.841
	Irbesartan	41	4 (9.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan							
							Interaction test:	0.945
< 27 kg/m**2	Sparsentan	84	7 (8.3)	NE		0.702	(0.250, 1.968)	0.501
	Irbesartan	94	8 (8.5)	NE				
≥ 27 kg/m**2	Sparsentan	118	11 (9.3)	NE		0.719	(0.305, 1.697)	0.452
	Irbesartan	107	11 (10.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan							Interaction test: 0.951
eGFR Low and UP High	Sparsentan	71	9 (12.7)	NE		0.503	(0.201, 1.256)	0.141
	Irbesartan	74	10 (13.5)	NE				
eGFR Low and UP Low	Sparsentan	55	3 (5.5)	NE		0.890	(0.198, 3.995)	0.879
	Irbesartan	55	4 (7.3)	NE				
eGFR High and UP High	Sparsentan	37	4 (10.8)	NE		0.793	(0.175, 3.591)	0.764
	Irbesartan	36	3 (8.3)	NE				
eGFR High and UP Low	Sparsentan	39	2 (5.1)	NE		0.717	(0.115, 4.458)	0.721
	Irbesartan	37	3 (8.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.762
< 60 mL/min/1.73 m**2	Sparsentan	127	13 (10.2)	NE		0.625	(0.291, 1.342)	0.228
	Irbesartan	129	15 (11.6)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	3 (6.1)	NE		1.099	(0.182, 6.634)	0.918
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	2 (7.7)	NE		0.579	(0.090, 3.717)	0.564
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: 0.090
< 45 mL/min/1.73 m**2	Sparsentan	82	7 (8.5)	NE		0.348	(0.136, 0.887)	0.027 *
	Irbesartan	80	13 (16.3)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	6 (13.3)	NE		3.229	(0.568, 18.361)	0.186
	Irbesartan	49	2 (4.1)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	3 (6.1)	NE		1.099	(0.182, 6.634)	0.918
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	2 (7.7)	NE		0.579	(0.090, 3.717)	0.564
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.863
<= 1.75 g/day	Sparsentan	98	8 (8.2)	NE		0.644	(0.231, 1.801)	0.402
	Irbesartan	94	8 (8.5)	NE				
> 1.75 g/day	Sparsentan	104	10 (9.6)	NE		0.572	(0.246, 1.333)	0.196
	Irbesartan	108	12 (11.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.824
Yes	Sparsentan	88	8 (9.1)	NE		0.562	(0.211, 1.498)	0.250
	Irbesartan	83	9 (10.8)	NE				
No	Sparsentan	114	10 (8.8)	NE		0.730	(0.308, 1.727)	0.474
	Irbesartan	119	11 (9.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.867
<= 5 years	Sparsentan	113	9 (8.0)	NE		0.565	(0.240, 1.333)	0.192
	Irbesartan	127	13 (10.2)	NE				
> 5 years	Sparsentan	89	9 (10.1)	NE		0.643	(0.232, 1.780)	0.395
	Irbesartan	75	7 (9.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

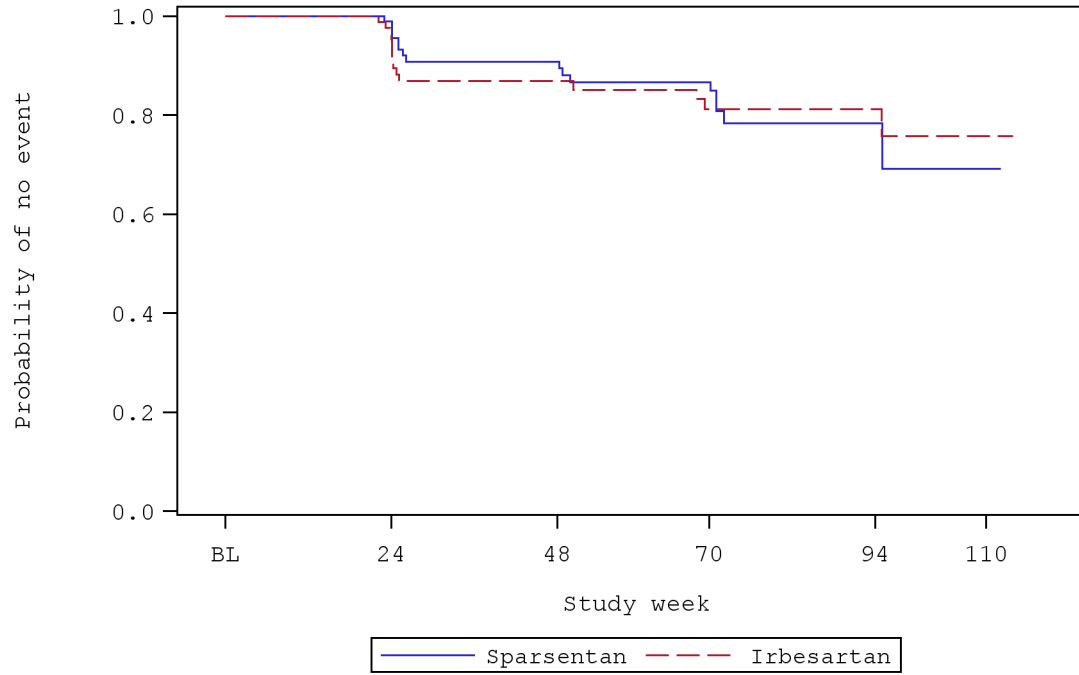
KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.949
Yes	Sparsentan	153	10 (6.5)	NE		0.569	(0.240, 1.348)	0.200
	Irbesartan	157	11 (7.0)	NE				
No	Sparsentan	49	8 (16.3)	NE		0.718	(0.268, 1.926)	0.510
	Irbesartan	45	9 (20.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Figure PF1KEFDT\_FSKM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

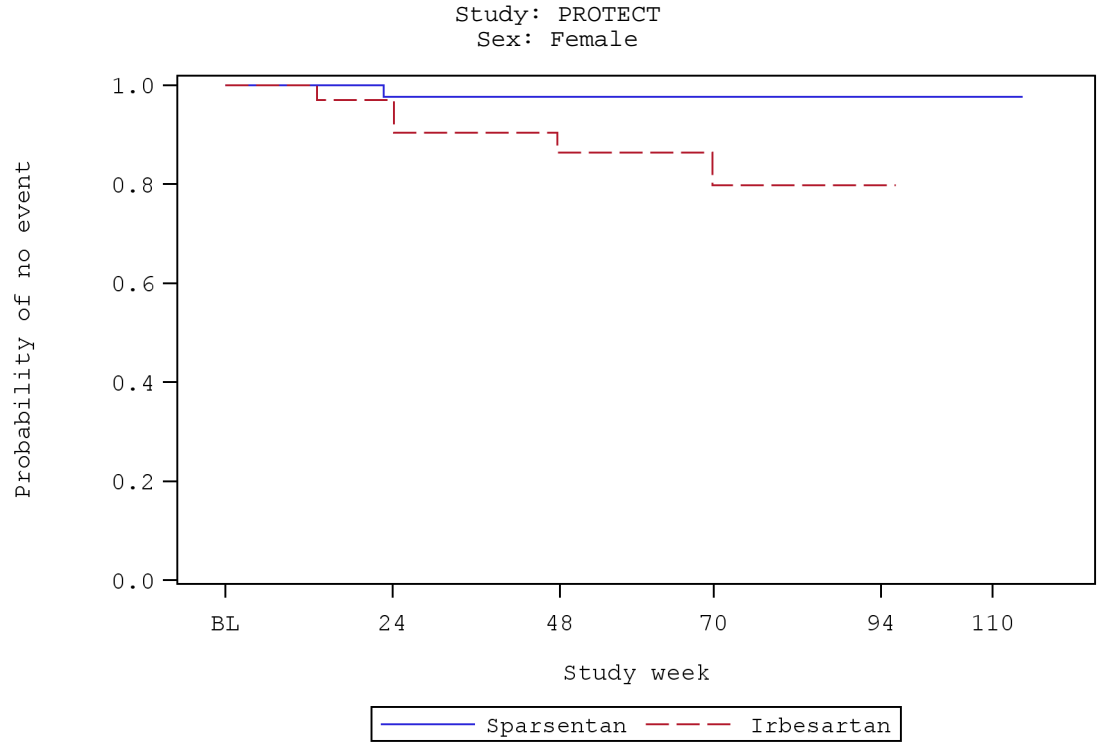
Study: PROTECT  
 Sex: Male



Sparsentan	139	91	71	52	25	8
Irbesartan	143	84	61	39	19	8

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KEFDT\_FSTM

Figure PF1KEFDT\_FSKM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	63	42	38	29	9	3
Irbesartan	59	30	21	12	2	0

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KEFDT\_FSTM

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Sex												
Male	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	139	130 (93.5)	90.82 (8.32)	61.4	86.36	93.18	97.73	100.0	
			Irbesartan	143	128 (89.5)	89.83 (11.36)	38.6	86.36	93.18	97.73	100.0	
		Week 24	Sparsentan	139	87 (62.6)	89.05 (12.90)	4.5	84.09	93.18	97.73	100.0	
			Irbesartan	143	73 (51.0)	87.33 (17.46)	4.5	86.36	93.18	97.73	100.0	
		Week 48	Sparsentan	139	72 (51.8)	87.82 (17.50)	0.0	84.09	90.91	97.73	100.0	
			Irbesartan	143	55 (38.5)	91.49 (12.04)	47.7	88.64	95.45	100.00	100.0	
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	139	87 (62.6)	-1.67 (11.01)	-72.7	-4.55	0.00	4.55	25.0
		Week 48	Irbesartan	143	73 (51.0)	-2.77 (16.40)	-90.9	-4.55	-2.27	2.27	52.3	
			Sparsentan	139	72 (51.8)	-2.81 (18.16)	-100.0	-4.55	0.00	4.55	27.3	-0.17 [-0.52, 0.18]
			Irbesartan	143	55 (38.5)	-0.25 (8.65)	-27.3	-2.27	2.27	4.55	15.9	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Female	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	63	56 (88.9)	84.25 (15.21)	31.8	77.27	88.64	95.45	100.0	
			Irbesartan	59	51 (86.4)	88.19 (10.22)	50.0	81.82	90.91	95.45	100.0	
		Week 24	Sparsentan	63	38 (60.3)	87.32 (12.90)	54.5	79.55	92.05	97.73	100.0	
			Irbesartan	59	31 (52.5)	86.95 (18.70)	4.5	81.82	95.45	97.73	100.0	
		Week 48	Sparsentan	63	38 (60.3)	88.40 (12.58)	61.4	77.27	94.32	100.00	100.0	
			Irbesartan	59	22 (37.3)	85.02 (12.79)	54.5	77.27	88.64	93.18	100.0	
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	63	38 (60.3)	4.37 (14.95)	-25.0	-2.27	2.27	11.36	63.6	0.35 [-0.12, 0.83]
			Irbesartan	59	31 (52.5)	-1.61 (19.02)	-86.4	-9.09	0.00	4.55	29.5	
		Week 48	Sparsentan	63	38 (60.3)	5.44 (15.09)	-22.7	-2.27	2.27	11.36	59.1	0.48 [-0.05, 1.01]
			Irbesartan	59	22 (37.3)	-1.24 (11.50)	-15.9	-9.09	-2.27	2.27	34.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age												
<= 45 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	96	87 (90.6)	88.53 (12.27)	31.8	84.09	90.91	97.73	100.0	
			Irbesartan	99	92 (92.9)	89.30 (11.00)	52.3	86.36	92.05	97.73	100.0	
		Week 24	Sparsentan	96	57 (59.4)	89.59 (15.00)	4.5	84.09	93.18	97.73	100.0	
			Irbesartan	99	53 (53.5)	85.76 (20.69)	4.5	86.36	90.91	97.73	100.0	
		Week 48	Sparsentan	96	53 (55.2)	87.82 (16.18)	0.0	81.82	90.91	97.73	100.0	
			Irbesartan	99	40 (40.4)	90.68 (12.89)	47.7	88.64	95.45	100.00	100.0	
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	96	57 (59.4)	1.28 (14.77)	-72.7	-2.27	2.27	4.55	63.6
		Irbesartan	99	53 (53.5)	-3.99 (18.17)	-90.9	-6.82	0.00	2.27	15.9		
		Week 48	Sparsentan	96	53 (55.2)	-0.04 (17.42)	-90.9	-4.55	0.00	4.55	59.1	0.12 [-0.29, 0.53]
		Irbesartan	99	40 (40.4)	-1.70 (8.59)	-27.3	-4.55	0.00	2.27	11.4		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 45 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	106	99 (93.4)	89.12 (10.29)	38.6	84.09	90.91	95.45	100.0	
			Irbesartan	103	87 (84.5)	89.42 (11.15)	38.6	84.09	93.18	97.73	100.0	
		Week 24	Sparsentan	106	68 (64.2)	87.63 (10.80)	50.0	82.95	90.91	95.45	100.0	
			Irbesartan	103	51 (49.5)	88.73 (14.10)	11.4	84.09	93.18	97.73	100.0	
		Week 48	Sparsentan	106	57 (53.8)	88.20 (15.81)	0.0	84.09	93.18	97.73	100.0	
			Irbesartan	103	37 (35.9)	88.51 (12.19)	47.7	84.09	93.18	95.45	100.0	
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	106	68 (64.2)	-0.77 (10.44)	-25.0	-6.82	-2.27	4.55	36.4	0.00 [-0.36, 0.37]
			Irbesartan	103	51 (49.5)	-0.80 (16.01)	-77.3	-4.55	0.00	2.27	52.3	
		Week 48	Sparsentan	106	57 (53.8)	0.12 (17.80)	-100.0	-4.55	0.00	6.82	38.6	-0.04 [-0.45, 0.37]
			Irbesartan	103	37 (35.9)	0.74 (10.32)	-27.3	-2.27	0.00	2.27	34.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Age at IgAN diagnosis													
<= 18 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	9	9 (100.0)	85.86 (9.22)	75.0	77.27	84.09	93.18	100.0		
			Irbesartan	5	5 (100.0)	87.27 (19.85)	52.3	90.91	95.45	97.73	100.0		
		Week 24	Sparsentan	9	3 (33.3)	62.12 (50.33)	4.5	4.55	84.09	97.73	97.7		
			Irbesartan	5	2 (40.0)	67.05 (30.53)	45.5	45.45	67.05	88.64	88.6		
		Week 48	Sparsentan	9	5 (55.6)	86.82 (9.29)	75.0	81.82	88.64	88.64	100.0		
			Irbesartan	5	1 (20.0)	90.91	90.9	90.91	90.91	90.91	90.9		
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	9	3 (33.3)	-25.00 (41.35)	-72.7	-72.73	-2.27	0.00	0.0	-0.54 [-2.36, 1.28]
			Irbesartan	5	2 (40.0)	-6.82 (0.00)	-6.8	-6.82	-6.82	-6.82	-6.8		
	Week 48		Sparsentan	9	5 (55.6)	-1.82 (6.31)	-11.4	-4.55	0.00	2.27	4.5	NE	
			Irbesartan	5	1 (20.0)	-4.55	-4.5	-4.55	-4.55	-4.55	-4.5		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 18 to 40 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	102	93 (91.2)	89.10 (12.16)	31.8	84.09	90.91	97.73	100.0	
			Irbesartan	109	99 (90.8)	89.69 (10.19)	52.3	86.36	93.18	97.73	100.0	
		Week 24	Sparsentan	102	62 (60.8)	90.54 (10.21)	54.5	84.09	93.18	97.73	100.0	
			Irbesartan	109	59 (54.1)	88.75 (16.06)	4.5	86.36	95.45	97.73	100.0	
		Week 48	Sparsentan	102	57 (55.9)	88.28 (16.00)	0.0	81.82	93.18	97.73	100.0	
			Irbesartan	109	44 (40.4)	89.41 (13.41)	47.7	87.50	92.05	100.00	100.0	
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	102	62 (60.8)	2.53 (10.74)	-18.2	-2.27	2.27	4.55	63.6	0.37 [0.01, 0.73]
			Irbesartan	109	59 (54.1)	-1.85 (12.88)	-86.4	-4.55	0.00	2.27	15.9	
		Week 48	Sparsentan	102	57 (55.9)	0.16 (16.97)	-90.9	-4.55	0.00	6.82	59.1	0.21 [-0.18, 0.61]
			Irbesartan	109	44 (40.4)	-2.84 (9.29)	-27.3	-7.95	0.00	2.27	11.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024



Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	91	84 (92.3)	88.88 (10.40)	38.6	85.23	90.91	95.45	100.0	
			Irbesartan	88	75 (85.2)	89.06 (11.58)	38.6	84.09	90.91	97.73	100.0	
		Week 24	Sparsentan	91	60 (65.9)	87.77 (10.69)	50.0	82.95	90.91	95.45	100.0	
			Irbesartan	88	43 (48.9)	86.05 (19.22)	4.5	81.82	90.91	95.45	100.0	
		Week 48	Sparsentan	91	48 (52.7)	87.83 (16.58)	0.0	82.95	90.91	97.73	100.0	
			Irbesartan	88	32 (36.4)	89.91 (11.60)	47.7	84.09	93.18	98.86	100.0	
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	91	60 (65.9)	-1.02 (10.79)	-25.0	-6.82	-2.27	3.41	36.4	0.12 [-0.27, 0.51]
			Irbesartan	88	43 (48.9)	-3.01 (22.12)	-90.9	-4.55	-2.27	4.55	52.3	
		Week 48	Sparsentan	91	48 (52.7)	0.09 (19.14)	-100.0	-4.55	1.14	7.95	38.6	-0.17 [-0.62, 0.28]
			Irbesartan	88	32 (36.4)	2.77 (9.01)	-11.4	-2.27	1.14	5.68	34.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Geographic region													
North America	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	35	32 (91.4)	90.98 (8.05)	75.0	85.23	93.18	97.73	100.0		
			Irbesartan	46	43 (93.5)	90.91 (10.80)	50.0	86.36	93.18	100.00	100.0		
		Week 24	Sparsentan	35	15 (42.9)	88.64 (23.96)	4.5	90.91	97.73	100.00	100.0		
			Irbesartan	46	26 (56.5)	86.63 (19.14)	4.5	81.82	90.91	97.73	100.0		
		Week 48	Sparsentan	35	13 (37.1)	93.71 (8.66)	75.0	93.18	95.45	100.00	100.0		
			Irbesartan	46	17 (37.0)	89.30 (11.58)	65.9	84.09	93.18	100.00	100.0		
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	35	15 (42.9)	-3.94 (19.60)	-72.7	-2.27	0.00	2.27	11.4	-0.07 [-0.70, 0.57]
			Irbesartan	46	26 (56.5)	-2.62 (19.01)	-86.4	-4.55	0.00	2.27	29.5		
	Week 48		Sparsentan	35	13 (37.1)	1.92 (6.62)	-4.5	-2.27	0.00	2.27	18.2	-0.02 [-0.74, 0.70]	
			Irbesartan	46	17 (37.0)	2.14 (11.21)	-15.9	-2.27	2.27	2.27	34.1		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Europe	KDQOL: symptom/ problems of kidney disease	Baseline	Sparsentan	98	87 (88.8)	88.27 (13.03)	31.8	84.09	90.91	97.73	100.0	
		Week 24	Irbesartan	115	96 (83.5)	88.68 (11.56)	38.6	84.09	90.91	96.59	100.0	
			Sparsentan	98	53 (54.1)	88.72 (10.50)	59.1	84.09	93.18	97.73	100.0	
		Week 48	Irbesartan	115	47 (40.9)	86.32 (20.10)	4.5	84.09	93.18	97.73	100.0	
			Sparsentan	98	44 (44.9)	85.64 (17.36)	0.0	79.55	88.64	97.73	100.0	
		Irbesartan	115	38 (33.0)	88.40 (14.23)	47.7	84.09	92.05	100.00	100.0		
	KDQOL: Change from baseline in symptom/ problems of kidney disease	Week 24	Sparsentan	98	53 (54.1)	1.76 (13.99)	-25.0	-4.55	0.00	4.55	63.6	0.31 [-0.08, 0.71]
			Irbesartan	115	47 (40.9)	-3.58 (20.05)	-90.9	-6.82	0.00	4.55	52.3	
		Week 48	Sparsentan	98	44 (44.9)	0.21 (21.27)	-100.0	-6.82	0.00	6.82	59.1	0.17 [-0.27, 0.60]
			Irbesartan	115	38 (33.0)	-2.63 (9.46)	-27.3	-6.82	0.00	2.27	11.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Asia Pacific	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	69	67 (97.1)	88.57 (9.98)	50.0	84.09	88.64	97.73	100.0	
			Irbesartan	41	40 (97.6)	89.32 (10.10)	52.3	85.23	93.18	95.45	100.0	
		Week 24	Sparsentan	69	57 (82.6)	88.32 (10.99)	50.0	84.09	90.91	97.73	100.0	
			Irbesartan	41	31 (75.6)	89.08 (12.30)	50.0	86.36	95.45	95.45	100.0	
		Week 48	Sparsentan	69	53 (76.8)	88.59 (15.85)	0.0	81.82	93.18	97.73	100.0	
			Irbesartan	41	22 (53.7)	92.05 (9.99)	54.5	90.91	93.18	100.00	100.0	
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	69	57 (82.6)	-0.24 (8.13)	-22.7	-4.55	0.00	4.55	13.6	0.03 [-0.41, 0.47]
			Irbesartan	41	31 (75.6)	-0.51 (9.20)	-25.0	-4.55	0.00	2.27	20.5	
		Week 48	Sparsentan	69	53 (76.8)	-0.56 (16.10)	-90.9	-4.55	0.00	9.09	27.3	-0.11 [-0.61, 0.39]
			Irbesartan	41	22 (53.7)	1.03 (7.45)	-15.9	-2.27	0.00	4.55	20.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline BMI												
< 27 kg/m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	84	76 (90.5)	89.17 (9.64)	47.7	84.09	89.77	97.73	100.0	
			Irbesartan	94	84 (89.4)	89.07 (10.98)	38.6	86.36	90.91	95.45	100.0	
		Week 24	Sparsentan	84	52 (61.9)	88.02 (15.75)	4.5	84.09	93.18	97.73	100.0	
			Irbesartan	94	50 (53.2)	86.36 (17.33)	4.5	81.82	92.05	97.73	100.0	
	Week 48	Sparsentan	84	47 (56.0)	89.22 (11.14)	65.9	79.55	93.18	100.00	100.0		
		Irbesartan	94	36 (38.3)	86.36 (15.35)	47.7	78.41	90.91	100.00	100.0		
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	84	52 (61.9)	-1.40 (13.45)	-72.7	-6.82	0.00	4.55	20.5	0.03 [-0.35, 0.42]
		Week 48	Irbesartan	94	50 (53.2)	-1.91 (16.31)	-86.4	-6.82	0.00	4.55	52.3	
		Week 48	Sparsentan	84	47 (56.0)	0.73 (10.03)	-18.2	-6.82	0.00	6.82	25.0	0.40 [-0.04, 0.84]
		Week 48	Irbesartan	94	36 (38.3)	-3.16 (9.13)	-27.3	-6.82	0.00	2.27	11.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 27 kg/m**2	KDQOL: symptom/ problems of kidney disease	Baseline	Sparsentan	118	110 (93.2)	88.62 (12.25)	31.8	84.09	90.91	97.73	100.0	
			Irbesartan	107	94 (87.9)	89.94 (10.75)	47.7	84.09	93.18	97.73	100.0	
		Week 24	Sparsentan	118	73 (61.9)	88.89 (10.45)	50.0	84.09	90.91	97.73	100.0	
			Irbesartan	107	54 (50.5)	88.01 (18.25)	4.5	86.36	93.18	97.73	100.0	
		Week 48	Sparsentan	118	63 (53.4)	87.12 (18.75)	0.0	81.82	90.91	97.73	100.0	
		Irbesartan	107	41 (38.3)	92.52 (8.58)	61.4	90.91	95.45	100.00	100.0		
	KDQOL: Change from baseline in symptom/ problems of kidney disease	Week 24	Sparsentan	118	73 (61.9)	1.28 (11.91)	-22.7	-4.55	0.00	4.55	63.6	0.28 [-0.07, 0.64]
		Irbesartan	107	54 (50.5)	-2.90 (18.01)	-90.9	-4.55	0.00	2.27	29.5		
		Week 48	Sparsentan	118	63 (53.4)	-0.47 (21.57)	-100.0	-4.55	0.00	6.82	59.1	-0.13 [-0.52, 0.27]
		Irbesartan	107	41 (38.3)	1.77 (9.28)	-15.9	-2.27	2.27	4.55	34.1		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Randomization strata													
eGFR Low and UP High	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	71	63 (88.7)	90.73 (8.57)	63.6	86.36	93.18	97.73	100.0		
			Irbesartan	74	63 (85.1)	86.51 (13.18)	38.6	81.82	88.64	95.45	100.0		
		Week 24	Sparsentan	71	44 (62.0)	89.46 (10.49)	50.0	85.23	93.18	96.59	100.0		
			Irbesartan	74	32 (43.2)	86.22 (13.19)	45.5	82.95	88.64	94.32	100.0		
		Week 48	Sparsentan	71	37 (52.1)	91.46 (9.71)	65.9	88.64	95.45	100.00	100.0		
			Irbesartan	74	22 (29.7)	84.71 (14.93)	47.7	79.55	90.91	95.45	100.0		
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	71	44 (62.0)	-2.17 (8.74)	-25.0	-4.55	-2.27	2.27	25.0	-0.03 [-0.49, 0.42]
				Irbesartan	74	32 (43.2)	-1.85 (11.84)	-22.7	-6.82	-2.27	0.00	52.3	
			Week 48	Sparsentan	71	37 (52.1)	0.31 (9.97)	-20.5	-4.55	0.00	6.82	27.3	0.35 [-0.18, 0.89]
				Irbesartan	74	22 (29.7)	-3.20 (9.86)	-27.3	-6.82	0.00	2.27	11.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR Low and UP Low	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	55	51 (92.7)	86.54 (14.68)	31.8	84.09	88.64	97.73	100.0	
		Week 24	Irbesartan	55	49 (89.1)	90.58 (7.35)	75.0	84.09	93.18	95.45	100.0	
			Sparsentan	55	38 (69.1)	88.04 (10.59)	63.6	81.82	90.91	97.73	100.0	
		Week 48	Irbesartan	55	33 (60.0)	92.22 (9.13)	68.2	88.64	95.45	97.73	100.0	
			Sparsentan	55	28 (50.9)	82.14 (25.21)	0.0	78.41	90.91	95.45	100.0	
		Irbesartan	55	25 (45.5)	90.00 (10.56)	61.4	86.36	90.91	100.00	100.0		
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	55	38 (69.1)	3.17 (14.22)	-18.2	-6.82	0.00	6.82	63.6	0.17 [-0.30, 0.64]
			Irbesartan	55	33 (60.0)	1.17 (7.85)	-18.2	-2.27	0.00	4.55	20.5	
		Week 48	Sparsentan	55	28 (50.9)	-1.95 (31.02)	-100.0	-4.55	0.00	11.36	59.1	-0.03 [-0.56, 0.51]
			Irbesartan	55	25 (45.5)	-1.36 (8.55)	-15.9	-9.09	0.00	2.27	20.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024



Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR High and UP High	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	37	35 (94.6)	88.05 (9.38)	65.9	84.09	88.64	97.73	100.0		
			Irbesartan	36	31 (86.1)	90.25 (10.63)	50.0	84.09	93.18	97.73	100.0		
		Week 24	Sparsentan	37	19 (51.4)	85.89 (21.86)	4.5	84.09	88.64	100.00	100.0		
			Irbesartan	36	18 (50.0)	86.99 (19.96)	11.4	86.36	92.05	95.45	100.0		
		Week 48	Sparsentan	37	22 (59.5)	87.09 (12.88)	61.4	77.27	89.77	97.73	100.0		
			Irbesartan	36	11 (30.6)	88.22 (14.55)	47.7	84.09	90.91	97.73	100.0		
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	37	19 (51.4)	-1.32 (19.61)	-72.7	-2.27	0.00	9.09	22.7	0.05 [-0.60, 0.69]
				Irbesartan	36	18 (50.0)	-2.27 (21.05)	-77.3	-6.82	0.00	4.55	29.5	
			Week 48	Sparsentan	37	22 (59.5)	0.93 (11.15)	-22.7	-4.55	-1.14	11.36	20.5	-0.04 [-0.77, 0.68]
				Irbesartan	36	11 (30.6)	1.45 (14.77)	-27.3	-4.55	2.27	6.82	34.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP Low	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	39	37 (94.9)	89.56 (11.19)	50.0	86.36	90.91	97.73	100.0	
			Irbesartan	37	36 (97.3)	91.92 (10.83)	52.3	89.77	95.45	97.73	100.0	
		Week 24	Sparsentan	39	24 (61.5)	89.68 (11.11)	54.5	84.09	93.18	97.73	100.0	
			Irbesartan	37	21 (56.8)	81.06 (28.23)	4.5	75.00	95.45	95.45	97.7	
		Week 48	Sparsentan	39	23 (59.0)	90.51 (9.56)	65.9	88.64	93.18	97.73	100.0	
			Irbesartan	37	19 (51.4)	95.69 (8.12)	65.9	93.18	100.00	100.00	100.00	
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	39	24 (61.5)	0.85 (7.58)	-22.7	0.00	2.27	4.55	13.6	0.51 [-0.09, 1.10]
			Irbesartan	37	21 (56.8)	-9.09 (27.41)	-90.9	-4.55	0.00	2.27	11.4	
		Week 48	Sparsentan	39	23 (59.0)	1.19 (7.34)	-11.4	-2.27	0.00	6.82	15.9	-0.21 [-0.82, 0.40]
			Irbesartan	37	19 (51.4)	2.51 (5.02)	-6.8	0.00	2.27	4.55	15.9	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 1														
< 60 mL/min/1.73 m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	127	115 (90.6)	88.99 (11.78)	31.8	84.09	90.91	97.73	100.0			
			Irbesartan	129	113 (87.6)	88.66 (11.11)	38.6	84.09	90.91	95.45	100.0			
		Week 24	Sparsentan	127	83 (65.4)	88.64 (10.34)	50.0	84.09	93.18	95.45	100.0			
			Irbesartan	129	68 (52.7)	88.03 (15.38)	4.5	84.09	93.18	97.73	100.0			
		Week 48	Sparsentan	127	65 (51.2)	87.41 (18.51)	0.0	84.09	90.91	97.73	100.0			
			Irbesartan	129	49 (38.0)	88.03 (12.82)	47.7	79.55	93.18	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	127	83 (65.4)	0.05 (11.77)	-25.0	-4.55	-2.27	4.55	63.6	0.13 [-0.19, 0.45]
					Irbesartan	129	68 (52.7)	-1.70 (14.73)	-90.9	-5.68	0.00	2.27	52.3	
				Week 48	Sparsentan	127	65 (51.2)	-0.77 (21.50)	-100.0	-4.55	0.00	6.82	59.1	0.05 [-0.32, 0.42]
					Irbesartan	129	49 (38.0)	-1.58 (9.36)	-27.3	-6.82	0.00	2.27	20.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL: symptom/ problems of kidney disease	Baseline	Sparsentan	49	47 (95.9)	88.83 (9.65)	50.0	84.09	88.64	97.73	100.0	
			Irbesartan	48	43 (89.6)	89.64 (10.62)	50.0	86.36	93.18	97.73	100.0	
		Week 24	Sparsentan	49	27 (55.1)	86.28 (19.69)	4.5	84.09	93.18	97.73	100.0	
			Irbesartan	48	24 (50.0)	87.12 (17.83)	11.4	85.23	92.05	95.45	100.0	
			Sparsentan	49	29 (59.2)	87.70 (11.14)	65.9	81.82	88.64	97.73	100.0	
	Week 48	Sparsentan	49	29 (59.2)	87.70 (11.14)	65.9	81.82	88.64	97.73	100.0		
		Irbesartan	48	19 (39.6)	90.19 (13.30)	47.7	88.64	90.91	100.00	100.0		
	KDQOL: Change from baseline in symptom/ problems of kidney disease	Week 24	Sparsentan	49	27 (55.1)	-2.27 (16.46)	-72.7	-2.27	0.00	4.55	13.6	-0.04 [-0.59, 0.51]
			Irbesartan	48	24 (50.0)	-1.61 (18.20)	-77.3	-4.55	1.14	4.55	29.5	
		Week 48	Sparsentan	49	29 (59.2)	0.08 (9.02)	-22.7	-6.82	0.00	4.55	18.2	-0.09 [-0.67, 0.49]
Irbesartan			48	19 (39.6)	0.96 (11.40)	-27.3	-2.27	0.00	2.27	34.1		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	26	24 (92.3)	88.16 (11.84)	61.4	81.82	89.77	97.73	100.0		
			Irbesartan	25	23 (92.0)	92.29 (11.48)	52.3	88.64	97.73	100.00	100.0		
		Week 24	Sparsentan	26	15 (57.7)	91.97 (9.94)	70.5	88.64	97.73	100.00	100.0		
			Irbesartan	25	12 (48.0)	82.77 (28.58)	4.5	81.82	95.45	97.73	100.0		
		Week 48	Sparsentan	26	16 (61.5)	91.05 (11.66)	61.4	81.82	96.59	100.00	100.0		
		Irbesartan	25	9 (36.0)	97.22 (4.92)	88.6	97.73	100.00	100.00	100.0			
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	26	15 (57.7)	5.15 (7.16)	-6.8	0.00	2.27	9.09	22.7	0.73 [-0.06, 1.51]
				Irbesartan	25	12 (48.0)	-8.14 (26.39)	-86.4	-5.68	-1.14	2.27	11.4	
			Week 48	Sparsentan	26	16 (61.5)	3.27 (9.82)	-18.2	-1.14	2.27	11.36	20.5	0.15 [-0.67, 0.97]
				Irbesartan	25	9 (36.0)	2.02 (4.01)	-6.8	2.27	2.27	2.27	6.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline eGFR Group 2												
< 45 mL/min/1.73 m**2	KDQOL: symptom/ problems of kidney disease	Baseline	Sparsentan	82	71 (86.6)	87.74 (11.84)	31.8	81.82	90.91	95.45	100.0	
			Irbesartan	80	69 (86.3)	87.09 (12.66)	38.6	81.82	90.91	95.45	100.0	
		Week 24	Sparsentan	82	53 (64.6)	88.68 (10.11)	50.0	84.09	90.91	95.45	100.0	
			Irbesartan	80	45 (56.3)	87.73 (12.82)	45.5	84.09	90.91	97.73	100.0	
	Week 48	Sparsentan	82	42 (51.2)	86.53 (16.91)	0.0	79.55	90.91	95.45	100.0		
		Irbesartan	80	29 (36.3)	85.03 (14.95)	47.7	72.73	90.91	95.45	100.0		
		KDQOL: Change from baseline in symptom/ problems of kidney disease	Week 24	Sparsentan	82	53 (64.6)	0.81 (12.54)	-22.7	-6.82	-2.27	4.55	63.6
		Week 48	Irbesartan	80	45 (56.3)	-0.71 (11.09)	-22.7	-6.82	-2.27	2.27	52.3	
			Sparsentan	82	42 (51.2)	-0.27 (20.08)	-90.9	-4.55	0.00	9.09	59.1	0.17 [-0.30, 0.64]
			Irbesartan	80	29 (36.3)	-3.13 (10.48)	-27.3	-9.09	0.00	2.27	20.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
45 to < 60 mL/min/1.73 m**2	KDQOL: symptom/ problems of kidney disease	Baseline	Sparsentan	45	44 (97.8)	91.01 (11.55)	38.6	86.36	95.45	98.86	100.0	
			Irbesartan	49	44 (89.8)	91.12 (7.61)	68.2	86.36	93.18	97.73	100.0	
		Week 24	Sparsentan	45	30 (66.7)	88.56 (10.90)	59.1	81.82	93.18	95.45	100.0	
			Irbesartan	49	23 (46.9)	88.64 (19.78)	4.5	88.64	95.45	97.73	100.0	
		Week 48	Sparsentan	45	23 (51.1)	89.03 (21.44)	0.0	86.36	97.73	100.00	100.0	
			Irbesartan	49	20 (40.8)	92.39 (7.24)	77.3	88.64	93.18	98.86	100.0	
	KDQOL: Change from baseline in symptom/ problems of kidney disease	Week 24	Sparsentan	45	30 (66.7)	-1.29 (10.33)	-25.0	-4.55	-2.27	4.55	36.4	0.15 [-0.39, 0.70]
			Irbesartan	49	23 (46.9)	-3.66 (20.21)	-90.9	-4.55	0.00	2.27	15.9	
		Week 48	Sparsentan	45	23 (51.1)	-1.68 (24.34)	-100.0	-2.27	0.00	6.82	38.6	-0.13 [-0.73, 0.47]
			Irbesartan	49	20 (40.8)	0.68 (7.11)	-13.6	-1.14	0.00	4.55	15.9	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	49	47 (95.9)	88.83 (9.65)	50.0	84.09	88.64	97.73	100.0	
			Irbesartan	48	43 (89.6)	89.64 (10.62)	50.0	86.36	93.18	97.73	100.0	
		Week 24	Sparsentan	49	27 (55.1)	86.28 (19.69)	4.5	84.09	93.18	97.73	100.0	
			Irbesartan	48	24 (50.0)	87.12 (17.83)	11.4	85.23	92.05	95.45	100.0	
			Week 48	Sparsentan	49	29 (59.2)	87.70 (11.14)	65.9	81.82	88.64	97.73	100.0
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	49	27 (55.1)	-2.27 (16.46)	-72.7	-2.27	0.00	4.55	13.6	-0.04 [-0.59, 0.51]
			Irbesartan	48	24 (50.0)	-1.61 (18.20)	-77.3	-4.55	1.14	4.55	29.5	
		Week 48	Sparsentan	49	29 (59.2)	0.08 (9.02)	-22.7	-6.82	0.00	4.55	18.2	-0.09 [-0.67, 0.49]
			Irbesartan	48	19 (39.6)	0.96 (11.40)	-27.3	-2.27	0.00	2.27	34.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	26	24 (92.3)	88.16 (11.84)	61.4	81.82	89.77	97.73	100.0	
			Irbesartan	25	23 (92.0)	92.29 (11.48)	52.3	88.64	97.73	100.00	100.0	
		Week 24	Sparsentan	26	15 (57.7)	91.97 (9.94)	70.5	88.64	97.73	100.00	100.0	
			Irbesartan	25	12 (48.0)	82.77 (28.58)	4.5	81.82	95.45	97.73	100.0	
		Week 48	Sparsentan	26	16 (61.5)	91.05 (11.66)	61.4	81.82	96.59	100.00	100.0	
		Irbesartan	25	9 (36.0)	97.22 (4.92)	88.6	97.73	100.00	100.00	100.0		
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	26	15 (57.7)	5.15 (7.16)	-6.8	0.00	2.27	9.09	22.7	0.73 [-0.06, 1.51]
			Irbesartan	25	12 (48.0)	-8.14 (26.39)	-86.4	-5.68	-1.14	2.27	11.4	
		Week 48	Sparsentan	26	16 (61.5)	3.27 (9.82)	-18.2	-1.14	2.27	11.36	20.5	0.15 [-0.67, 0.97]
			Irbesartan	25	9 (36.0)	2.02 (4.01)	-6.8	2.27	2.27	2.27	6.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline urine protein excretion													
<= 1.75 g/day	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	98	92 (93.9)	88.02 (13.43)	31.8	84.09	90.91	97.73	100.0		
			Irbesartan	94	82 (87.2)	90.55 (9.51)	52.3	86.36	93.18	97.73	100.0		
		Week 24	Sparsentan	98	66 (67.3)	87.78 (10.66)	54.5	81.82	89.77	97.73	100.0		
			Irbesartan	94	43 (45.7)	87.32 (21.28)	4.5	86.36	95.45	97.73	100.0		
		Week 48	Sparsentan	98	59 (60.2)	86.90 (19.26)	0.0	79.55	93.18	97.73	100.0		
			Irbesartan	94	34 (36.2)	93.65 (9.56)	65.9	90.91	98.86	100.00	100.0		
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	98	66 (67.3)	0.90 (12.19)	-22.7	-6.82	0.00	4.55	63.6	0.24 [-0.14, 0.63]
			Irbesartan	94	43 (45.7)	-3.01 (20.47)	-90.9	-2.27	0.00	4.55	20.5		
	Week 48		Sparsentan	98	59 (60.2)	-0.04 (22.18)	-100.0	-4.55	0.00	6.82	59.1	-0.08 [-0.51, 0.34]	
			Irbesartan	94	34 (36.2)	1.47 (6.50)	-15.9	0.00	2.27	4.55	20.5		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 1.75 g/day	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	104	94 (90.4)	89.65 (8.56)	65.9	84.09	90.91	97.73	100.0	
			Irbesartan	108	97 (89.8)	88.36 (12.15)	38.6	84.09	90.91	97.73	100.0	
		Week 24	Sparsentan	104	59 (56.7)	89.37 (15.01)	4.5	88.64	93.18	97.73	100.0	
			Irbesartan	108	61 (56.5)	87.15 (14.95)	11.4	84.09	90.91	95.45	100.0	
		Week 48	Sparsentan	104	51 (49.0)	89.30 (10.90)	61.4	81.82	90.91	97.73	100.0	
			Irbesartan	108	43 (39.8)	86.47 (13.73)	47.7	79.55	90.91	95.45	100.0	
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	104	59 (56.7)	-0.65 (13.08)	-72.7	-4.55	0.00	4.55	25.0	0.10 [-0.26, 0.46]
			Irbesartan	108	61 (56.5)	-2.01 (14.51)	-77.3	-6.82	-2.27	2.27	52.3	
		Week 48	Sparsentan	104	51 (49.0)	0.13 (9.99)	-20.5	-6.82	0.00	6.82	27.3	0.21 [-0.19, 0.62]
			Irbesartan	108	43 (39.8)	-2.11 (11.12)	-27.3	-9.09	0.00	2.27	34.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline use of antihypertensives													
Yes	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	88	77 (87.5)	88.34 (10.76)	38.6	84.09	90.91	95.45	100.0		
			Irbesartan	83	70 (84.3)	89.06 (10.21)	38.6	86.36	90.91	95.45	100.0		
		Week 24	Sparsentan	88	50 (56.8)	88.00 (11.39)	50.0	84.09	88.64	97.73	100.0		
			Irbesartan	83	39 (47.0)	90.09 (8.89)	56.8	86.36	93.18	95.45	100.0		
		Week 48	Sparsentan	88	41 (46.6)	90.02 (10.21)	61.4	88.64	90.91	97.73	100.0		
			Irbesartan	83	28 (33.7)	89.69 (11.99)	47.7	90.91	93.18	95.45	100.0		
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	88	50 (56.8)	0.23 (10.96)	-25.0	-4.55	0.00	4.55	36.4	-0.01 [-0.42, 0.41]
			Irbesartan	83	39 (47.0)	0.29 (10.80)	-22.7	-4.55	0.00	2.27	52.3		
	Week 48		Sparsentan	88	41 (46.6)	2.99 (11.17)	-18.2	-2.27	2.27	6.82	38.6	0.29 [-0.20, 0.77]	
			Irbesartan	83	28 (33.7)	0.00 (9.34)	-27.3	-2.27	0.00	4.55	15.9		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL: symptom/ problems of kidney disease	Baseline	Sparsentan	114	109 (95.6)	89.20 (11.60)	31.8	84.09	90.91	97.73	100.0	
		Week 24	Irbesartan	119	109 (91.6)	89.55 (11.59)	47.7	86.36	93.18	97.73	100.0	
			Sparsentan	114	75 (65.8)	88.88 (13.83)	4.5	84.09	93.18	97.73	100.0	
		Week 48	Irbesartan	119	65 (54.6)	85.49 (21.25)	4.5	81.82	93.18	97.73	100.0	
			Sparsentan	114	69 (60.5)	86.82 (18.46)	0.0	79.55	93.18	97.73	100.0	
		Irbesartan	119	49 (41.2)	89.61 (12.94)	47.7	84.09	90.91	100.00	100.0		
	KDQOL: Change from baseline in symptom/ problems of kidney disease	Week 24	Sparsentan	114	75 (65.8)	0.12 (13.64)	-72.7	-4.55	0.00	4.55	63.6	0.25 [-0.09, 0.58]
			Irbesartan	119	65 (54.6)	-4.06 (19.90)	-90.9	-6.82	0.00	2.27	29.5	
		Week 48	Sparsentan	114	69 (60.5)	-1.71 (20.28)	-100.0	-4.55	0.00	6.82	59.1	-0.05 [-0.42, 0.31]
			Irbesartan	119	49 (41.2)	-0.83 (9.64)	-27.3	-6.82	0.00	2.27	34.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Time since renal biopsy													
<= 5 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	113	104 (92.0)	88.81 (11.12)	38.6	84.09	90.91	97.73	100.0		
			Irbesartan	127	116 (91.3)	88.81 (12.03)	38.6	84.09	92.05	97.73	100.0		
		Week 24	Sparsentan	113	70 (61.9)	90.10 (10.53)	50.0	86.36	93.18	97.73	100.0		
			Irbesartan	127	67 (52.8)	85.45 (20.51)	4.5	81.82	93.18	97.73	100.0		
		Week 48	Sparsentan	113	61 (54.0)	89.79 (14.79)	0.0	86.36	93.18	100.00	100.0		
			Irbesartan	127	51 (40.2)	89.71 (13.68)	47.7	88.64	93.18	100.00	100.0		
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	113	70 (61.9)	2.31 (8.92)	-22.7	-2.27	2.27	4.55	36.4	0.35 [0.01, 0.68]
			Irbesartan	127	67 (52.8)	-3.22 (20.90)	-90.9	-4.55	0.00	2.27	52.3		
			Week 48	Sparsentan	113	61 (54.0)	2.42 (16.70)	-100.0	-2.27	2.27	11.36	38.6	0.13 [-0.24, 0.50]
				Irbesartan	127	51 (40.2)	0.62 (9.61)	-27.3	-2.27	2.27	4.55	34.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 5 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	89	82 (92.1)	88.89 (11.45)	31.8	84.09	90.91	97.73	100.0	
			Irbesartan	75	63 (84.0)	90.37 (8.95)	52.3	86.36	93.18	97.73	100.0	
		Week 24	Sparsentan	89	55 (61.8)	86.53 (15.21)	4.5	79.55	88.64	97.73	100.0	
			Irbesartan	75	37 (49.3)	90.42 (10.63)	45.5	88.64	93.18	97.73	100.0	
		Week 48	Sparsentan	89	49 (55.1)	85.81 (17.12)	0.0	79.55	90.91	97.73	100.0	
			Irbesartan	75	26 (34.7)	89.51 (10.12)	61.4	84.09	90.91	95.45	100.0	
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	89	55 (61.8)	-2.56 (15.77)	-72.7	-6.82	-2.27	2.27	63.6	-0.12 [-0.54, 0.29]
			Irbesartan	75	37 (49.3)	-0.98 (6.00)	-18.2	-4.55	-2.27	2.27	11.4	
		Week 48	Sparsentan	89	49 (55.1)	-2.92 (18.27)	-90.9	-6.82	-2.27	4.55	59.1	-0.01 [-0.48, 0.47]
			Irbesartan	75	26 (34.7)	-2.80 (8.96)	-27.3	-9.09	-1.14	2.27	11.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: History of hypertension													
Yes	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	153	138 (90.2)	88.18 (10.81)	38.6	84.09	90.91	95.45	100.0		
			Irbesartan	157	135 (86.0)	89.68 (10.49)	38.6	84.09	93.18	97.73	100.0		
		Week 24	Sparsentan	153	94 (61.4)	88.47 (11.24)	50.0	84.09	92.05	97.73	100.0		
			Irbesartan	157	76 (48.4)	89.41 (13.81)	4.5	86.36	93.18	97.73	100.0		
		Week 48	Sparsentan	153	81 (52.9)	87.88 (14.63)	0.0	81.82	90.91	97.73	100.0		
			Irbesartan	157	57 (36.3)	89.87 (12.73)	47.7	88.64	93.18	100.00	100.0		
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	153	94 (61.4)	0.85 (9.46)	-25.0	-4.55	0.00	4.55	36.4	0.13 [-0.17, 0.44]
			Irbesartan	157	76 (48.4)	-0.72 (13.96)	-90.9	-4.55	0.00	2.27	52.3		
	Week 48		Sparsentan	153	81 (52.9)	0.28 (14.76)	-90.9	-4.55	0.00	6.82	38.6	0.06 [-0.28, 0.40]	
			Irbesartan	157	57 (36.3)	-0.48 (10.08)	-27.3	-4.55	0.00	4.55	34.1		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	49	48 (98.0)	90.77 (12.30)	31.8	87.50	95.45	100.00	100.0	
		Week 24	Irbesartan	45	44 (97.8)	88.38 (12.67)	47.7	86.36	90.91	98.86	100.0	
			Sparsentan	49	31 (63.3)	88.71 (17.11)	4.5	84.09	93.18	97.73	100.0	
		Week 48	Irbesartan	45	28 (62.2)	81.25 (24.91)	4.5	71.59	92.05	97.73	100.0	
			Sparsentan	49	29 (59.2)	88.40 (19.35)	0.0	86.36	93.18	100.00	100.0	
		Irbesartan	45	20 (44.4)	88.98 (12.19)	54.5	85.23	90.91	100.00	100.0		
	KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	49	31 (63.3)	-1.91 (19.29)	-72.7	-6.82	0.00	4.55	63.6	0.24 [-0.27, 0.75]
			Irbesartan	45	28 (62.2)	-7.06 (23.43)	-86.4	-7.95	-2.27	4.55	20.5	
		Week 48	Sparsentan	49	29 (59.2)	-0.63 (23.96)	-100.0	-6.82	0.00	11.36	59.1	0.00 [-0.57, 0.57]
			Irbesartan	45	20 (44.4)	-0.68 (7.74)	-13.6	-4.55	0.00	2.27	20.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Sex	Overall		Sparsentan							Interaction: 0.117
Male	Week 24	Sparsentan	139	87 (62.6)	-1.63 (1.50)	(-4.58, 1.31)	1.32 (2.22)	(-3.05, 5.68)	0.553	
		Irbesartan	143	73 (51.0)	-2.95 (1.63)	(-6.16, 0.26)				
	Week 48	Sparsentan	139	72 (51.8)	-2.87 (1.64)	(-6.10, 0.36)	-2.94 (2.51)	(-7.87, 2.00)	0.243	
		Irbesartan	143	55 (38.5)	0.06 (1.89)	(-3.66, 3.79)				
Female	Week 24	Sparsentan	63	38 (60.3)	3.06 (2.15)	(-1.21, 7.33)	3.42 (3.33)	(-3.20, 10.04)	0.307	
		Irbesartan	59	31 (52.5)	-0.36 (2.51)	(-5.35, 4.62)				
	Week 48	Sparsentan	63	38 (60.3)	4.21 (2.14)	(-0.03, 8.46)	7.07 (3.51)	(0.10, 14.03)	0.047 *	
		Irbesartan	59	22 (37.3)	-2.85 (2.75)	(-8.30, 2.59)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.355
<= 45 years	Week 24	Sparsentan	96	57 (59.4)	0.77 (1.99)	(-3.15, 4.69)	4.36 (2.87)	(-1.29, 10.02)	0.130
		Irbesartan	99	53 (53.5)	-3.60 (2.05)	(-7.65, 0.46)			
	Week 48	Sparsentan	96	53 (55.2)	-0.90 (2.06)	(-4.96, 3.16)	0.20 (3.15)	(-6.02, 6.42)	0.950
		Irbesartan	99	40 (40.4)	-1.10 (2.38)	(-5.79, 3.59)			
> 45 years	Week 24	Sparsentan	106	68 (64.2)	-1.56 (1.55)	(-4.62, 1.49)	-1.62 (2.39)	(-6.34, 3.09)	0.497
		Irbesartan	103	51 (49.5)	0.06 (1.81)	(-3.52, 3.64)			
	Week 48	Sparsentan	106	57 (53.8)	0.11 (1.67)	(-3.18, 3.39)	0.59 (2.66)	(-4.66, 5.84)	0.824
		Irbesartan	103	37 (35.9)	-0.49 (2.07)	(-4.57, 3.59)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Age at IgAN diagnosis	Overall		Sparsentan						Interaction:	0.677
<= 18 years	Week 24		Sparsentan	9	3 (33.3)	-11.63 (14.66)	(-50.27, 27.00)	-9.73 (28.35)	(-88.56, 69.10)	0.749
			Irbesartan	5	2 (40.0)	-1.90 (24.05)	(-70.32, 66.51)			
	Week 48		Sparsentan	9	5 (55.6)	-11.52 (13.46)	(-49.14, 26.10)	-11.72 (28.63)	(-90.30, 66.85)	0.703
			Irbesartan	5	1 (20.0)	0.21 (25.12)	(-68.03, 68.44)			
> 18 to 40 years	Week 24		Sparsentan	102	62 (60.8)	2.18 (1.55)	(-0.88, 5.24)	3.55 (2.23)	(-0.84, 7.95)	0.113
			Irbesartan	109	59 (54.1)	-1.37 (1.59)	(-4.52, 1.77)			
	Week 48		Sparsentan	102	57 (55.9)	-0.39 (1.62)	(-3.58, 2.80)	2.49 (2.46)	(-2.37, 7.35)	0.314
			Irbesartan	109	44 (40.4)	-2.88 (1.84)	(-6.51, 0.75)			
> 40 years	Week 24		Sparsentan	91	60 (65.9)	-1.03 (1.89)	(-4.75, 2.70)	1.49 (2.89)	(-4.22, 7.20)	0.608
			Irbesartan	88	43 (48.9)	-2.51 (2.21)	(-6.87, 1.84)			
	Week 48		Sparsentan	91	48 (52.7)	-0.21 (2.10)	(-4.35, 3.93)	-2.09 (3.34)	(-8.68, 4.49)	0.532
			Irbesartan	88	32 (36.4)	1.88 (2.60)	(-3.25, 7.01)			

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Geographic region	Overall		Sparsentan							Interaction: 0.633
North America	Week 24		Sparsentan	35	15 (42.9)	-2.45 (4.53)	(-11.60, 6.70)	0.31 (5.86)	(-11.51, 12.14)	0.957
			Irbesartan	46	26 (56.5)	-2.76 (3.65)	(-10.13, 4.61)			
	Week 48		Sparsentan	35	13 (37.1)	-0.99 (4.61)	(-10.29, 8.31)	2.07 (6.03)	(-10.07, 14.21)	0.733
			Irbesartan	46	17 (37.0)	-3.06 (3.81)	(-10.72, 4.60)			
Europe	Week 24		Sparsentan	98	53 (54.1)	0.78 (2.06)	(-3.29, 4.86)	3.31 (3.01)	(-2.64, 9.26)	0.273
			Irbesartan	115	47 (40.9)	-2.53 (2.20)	(-6.87, 1.81)			
	Week 48		Sparsentan	98	44 (44.9)	-0.98 (2.28)	(-5.48, 3.51)	0.57 (3.38)	(-6.10, 7.23)	0.867
			Irbesartan	115	38 (33.0)	-1.55 (2.46)	(-6.40, 3.30)			
Asia Pacific	Week 24		Sparsentan	69	57 (82.6)	-0.27 (1.46)	(-3.16, 2.61)	0.38 (2.47)	(-4.51, 5.27)	0.879
			Irbesartan	41	31 (75.6)	-0.65 (1.99)	(-4.57, 3.27)			
	Week 48		Sparsentan	69	53 (76.8)	-0.53 (1.52)	(-3.53, 2.47)	-1.88 (2.82)	(-7.45, 3.70)	0.507
			Irbesartan	41	22 (53.7)	1.34 (2.37)	(-3.33, 6.02)			

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline BMI	Overall		Sparsentan							Interaction: 0.462
< 27 kg/m**2	Week 24		Sparsentan	84	52 (61.9)	-1.51 (1.80)	(-5.07, 2.06)	0.41 (2.59)	(-4.72, 5.53)	0.876
			Irbesartan	94	50 (53.2)	-1.91 (1.85)	(-5.58, 1.75)			
	Week 48		Sparsentan	84	47 (56.0)	-0.11 (1.86)	(-3.79, 3.57)	5.15 (2.79)	(-0.36, 10.66)	0.067
			Irbesartan	94	36 (38.3)	-5.26 (2.07)	(-9.34, -1.17)			
≥ 27 kg/m**2	Week 24		Sparsentan	118	73 (61.9)	0.40 (1.68)	(-2.92, 3.71)	1.87 (2.59)	(-3.24, 6.98)	0.471
			Irbesartan	107	54 (50.5)	-1.47 (1.95)	(-5.32, 2.38)			
	Week 48		Sparsentan	118	63 (53.4)	-1.31 (1.80)	(-4.86, 2.24)	-4.40 (2.88)	(-10.08, 1.29)	0.129
			Irbesartan	107	41 (38.3)	3.09 (2.25)	(-1.34, 7.52)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Randomization strata	Overall		Sparsentan							Interaction: 0.365
eGFR Low and UP High	Week 24	Sparsentan	71	44 (62.0)	-1.55 (1.42)	(-4.36, 1.26)	0.93 (2.20)	(-3.41, 5.28)	0.671	
		Irbesartan	74	32 (43.2)	-2.48 (1.66)	(-5.78, 0.81)				
	Week 48	Sparsentan	71	37 (52.1)	0.75 (1.54)	(-2.30, 3.81)	4.97 (2.54)	(-0.05, 9.99)	0.052	
		Irbesartan	74	22 (29.7)	-4.22 (2.01)	(-8.19, -0.25)				
eGFR Low and UP Low	Week 24	Sparsentan	55	38 (69.1)	0.73 (2.39)	(-4.00, 5.46)	-3.02 (3.54)	(-10.02, 3.99)	0.395	
		Irbesartan	55	33 (60.0)	3.75 (2.57)	(-1.34, 8.84)				
	Week 48	Sparsentan	55	28 (50.9)	-4.59 (2.79)	(-10.12, 0.93)	-5.82 (4.09)	(-13.93, 2.29)	0.158	
		Irbesartan	55	25 (45.5)	1.23 (2.95)	(-4.62, 7.08)				
eGFR High and UP High	Week 24	Sparsentan	37	19 (51.4)	-0.74 (3.93)	(-8.63, 7.15)	0.57 (5.85)	(-11.21, 12.35)	0.923	
		Irbesartan	36	18 (50.0)	-1.31 (4.33)	(-10.03, 7.41)				
	Week 48	Sparsentan	37	22 (59.5)	-1.90 (3.81)	(-9.57, 5.77)	2.20 (6.16)	(-10.15, 14.54)	0.723	
		Irbesartan	36	11 (30.6)	-4.09 (4.83)	(-13.75, 5.57)				
eGFR High and UP Low	Week 24	Sparsentan	39	24 (61.5)	0.42 (2.96)	(-5.47, 6.30)	9.53 (4.34)	(0.90, 18.16)	0.031 *	
		Irbesartan	37	21 (56.8)	-9.11 (3.17)	(-15.42, -2.80)				
	Week 48	Sparsentan	39	23 (59.0)	0.91 (3.03)	(-5.11, 6.93)	-2.48 (4.55)	(-11.53, 6.57)	0.587	
		Irbesartan	37	19 (51.4)	3.39 (3.38)	(-3.33, 10.12)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline eGFR Group 1		Overall	Sparsentan							Interaction: 0.380
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	83 (65.4)	-0.29 (1.48)	(-3.21, 2.63)	0.94 (2.22)	(-3.43, 5.30)	0.673	
		Irbesartan	129	68 (52.7)	-1.23 (1.65)	(-4.47, 2.01)				
	Week 48	Sparsentan	127	65 (51.2)	-1.27 (1.68)	(-4.58, 2.03)	-0.15 (2.57)	(-5.22, 4.92)	0.954	
		Irbesartan	129	49 (38.0)	-1.13 (1.94)	(-4.95, 2.70)				
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	27 (55.1)	-2.09 (2.89)	(-7.87, 3.69)	-1.22 (4.26)	(-9.73, 7.29)	0.776	
		Irbesartan	48	24 (50.0)	-0.87 (3.08)	(-7.03, 5.29)				
	Week 48	Sparsentan	49	29 (59.2)	-1.77 (2.86)	(-7.49, 3.95)	0.75 (4.37)	(-7.97, 9.46)	0.865	
		Irbesartan	48	19 (39.6)	-2.52 (3.24)	(-8.98, 3.94)				
≥ 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	15 (57.7)	4.35 (3.96)	(-3.72, 12.43)	11.29 (5.93)	(-0.78, 23.36)	0.066	
		Irbesartan	25	12 (48.0)	-6.94 (4.40)	(-15.90, 2.02)				
	Week 48	Sparsentan	26	16 (61.5)	2.42 (3.90)	(-5.55, 10.39)	-7.22 (6.26)	(-19.89, 5.45)	0.256	
		Irbesartan	25	9 (36.0)	9.64 (4.87)	(-0.21, 19.48)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024



Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
	Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.465
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	53 (64.6)	0.84 (1.69)	(-2.51, 4.19)	1.23 (2.49)	(-3.70, 6.15)	0.623	
		Irbesartan	80	45 (56.3)	-0.39 (1.84)	(-4.01, 3.24)				
	Week 48	Sparsentan	82	42 (51.2)	-0.76 (1.90)	(-4.51, 3.00)	2.36 (2.98)	(-3.52, 8.24)	0.430	
		Irbesartan	80	29 (36.3)	-3.11 (2.29)	(-7.64, 1.41)				
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	30 (66.7)	-2.53 (2.88)	(-8.26, 3.21)	-0.15 (4.49)	(-9.07, 8.78)	0.974	
		Irbesartan	49	23 (46.9)	-2.38 (3.38)	(-9.09, 4.33)				
	Week 48	Sparsentan	45	23 (51.1)	-2.18 (3.32)	(-8.77, 4.42)	-4.08 (4.95)	(-13.92, 5.76)	0.412	
		Irbesartan	49	20 (40.8)	1.91 (3.63)	(-5.31, 9.12)				
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	27 (55.1)	-2.09 (2.89)	(-7.87, 3.69)	-1.22 (4.26)	(-9.73, 7.29)	0.776	
		Irbesartan	48	24 (50.0)	-0.87 (3.08)	(-7.03, 5.29)				
	Week 48	Sparsentan	49	29 (59.2)	-1.77 (2.86)	(-7.49, 3.95)	0.75 (4.37)	(-7.97, 9.46)	0.865	
		Irbesartan	48	19 (39.6)	-2.52 (3.24)	(-8.98, 3.94)				
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	15 (57.7)	4.35 (3.96)	(-3.72, 12.43)	11.29 (5.93)	(-0.78, 23.36)	0.066	
		Irbesartan	25	12 (48.0)	-6.94 (4.40)	(-15.90, 2.02)				
	Week 48	Sparsentan	26	16 (61.5)	2.42 (3.90)	(-5.55, 10.39)	-7.22 (6.26)	(-19.89, 5.45)	0.256	
		Irbesartan	25	9 (36.0)	9.64 (4.87)	(-0.21, 19.48)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline urine protein excretion	Overall		Sparsentan							Interaction: 0.335
<= 1.75 g/day	Week 24		Sparsentan	98	66 (67.3)	-0.28 (1.91)	(-4.04, 3.47)	1.60 (3.06)	(-4.43, 7.62)	0.602
			Irbesartan	94	43 (45.7)	-1.88 (2.38)	(-6.58, 2.82)			
	Week 48		Sparsentan	98	59 (60.2)	-1.09 (2.02)	(-5.06, 2.89)	-4.56 (3.38)	(-11.22, 2.10)	0.179
			Irbesartan	94	34 (36.2)	3.47 (2.68)	(-1.81, 8.76)			
> 1.75 g/day	Week 24		Sparsentan	104	59 (56.7)	-0.32 (1.59)	(-3.47, 2.82)	1.83 (2.25)	(-2.61, 6.27)	0.418
			Irbesartan	108	61 (56.5)	-2.15 (1.58)	(-5.26, 0.96)			
	Week 48		Sparsentan	104	51 (49.0)	-0.03 (1.70)	(-3.39, 3.33)	3.78 (2.54)	(-1.23, 8.79)	0.138
			Irbesartan	108	43 (39.8)	-3.81 (1.85)	(-7.46, -0.16)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline use of antihypertensives		Overall	Sparsentan							Interaction: 0.699
Yes	Week 24	Sparsentan	88	50 (56.8)	-0.11 (1.25)	(-2.57, 2.36)	-1.06 (1.89)	(-4.79, 2.66)	0.574	
		Irbesartan	83	39 (47.0)	0.96 (1.41)	(-1.84, 3.75)				
	Week 48	Sparsentan	88	41 (46.6)	2.36 (1.39)	(-0.38, 5.10)	1.68 (2.19)	(-2.64, 6.01)	0.443	
		Irbesartan	83	28 (33.7)	0.68 (1.68)	(-2.65, 4.00)				
No	Week 24	Sparsentan	114	75 (65.8)	-0.48 (1.87)	(-4.16, 3.20)	3.34 (2.74)	(-2.06, 8.75)	0.224	
		Irbesartan	119	65 (54.6)	-3.83 (2.01)	(-7.79, 0.14)				
	Week 48	Sparsentan	114	69 (60.5)	-2.29 (1.94)	(-6.10, 1.53)	-0.85 (3.00)	(-6.77, 5.07)	0.777	
		Irbesartan	119	49 (41.2)	-1.43 (2.30)	(-5.96, 3.09)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.062
<= 5 years	Week 24	Sparsentan	113	70 (61.9)	1.92 (1.68)	(-1.38, 5.23)	4.86 (2.40)	(0.13, 9.58)	0.044 *
		Irbesartan	127	67 (52.8)	-2.93 (1.71)	(-6.31, 0.44)			
	Week 48	Sparsentan	113	61 (54.0)	2.10 (1.80)	(-1.45, 5.65)	1.45 (2.67)	(-3.81, 6.71)	0.587
		Irbesartan	127	51 (40.2)	0.64 (1.98)	(-3.25, 4.54)			
> 5 years	Week 24	Sparsentan	89	55 (61.8)	-2.81 (1.79)	(-6.34, 0.73)	-2.94 (2.85)	(-8.56, 2.69)	0.304
		Irbesartan	75	37 (49.3)	0.13 (2.22)	(-4.26, 4.52)			
	Week 48	Sparsentan	89	49 (55.1)	-3.71 (1.91)	(-7.48, 0.06)	-1.50 (3.26)	(-7.93, 4.94)	0.647
		Irbesartan	75	26 (34.7)	-2.21 (2.61)	(-7.37, 2.95)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Table PT1KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.302
Yes	Week 24	Sparsentan	153	94 (61.4)	0.41 (1.17)	(-1.89, 2.71)	0.56 (1.75)	(-2.88, 4.00)	0.748
		Irbesartan	157	76 (48.4)	-0.15 (1.30)	(-2.71, 2.41)			
	Week 48	Sparsentan	153	81 (52.9)	-0.22 (1.26)	(-2.70, 2.25)	-0.30 (1.97)	(-4.17, 3.57)	0.878
		Irbesartan	157	57 (36.3)	0.08 (1.51)	(-2.89, 3.04)			
No	Week 24	Sparsentan	49	31 (63.3)	-3.57 (3.88)	(-11.32, 4.18)	3.14 (5.69)	(-8.24, 14.52)	0.584
		Irbesartan	45	28 (62.2)	-6.71 (4.14)	(-14.98, 1.57)			
	Week 48	Sparsentan	49	29 (59.2)	-2.40 (3.90)	(-10.20, 5.39)	3.57 (5.84)	(-8.07, 15.22)	0.542
		Irbesartan	45	20 (44.4)	-5.98 (4.31)	(-14.57, 2.62)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aqs, created on: 04APR2024

Figure PF1KSYC\_FSGM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

Not done: No significant subgroup found.

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KSYC\_FSCM.

Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.245
Male	Sparsentan	139	5 (3.6)	NE		5.238	(0.729, 37.628)	0.100
	Irbesartan	143	4 (2.8)	NE				
Female	Sparsentan	63	10 (15.9)	NE		0.907	(0.275, 2.988)	0.872
	Irbesartan	59	5 (8.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.075
<= 45 years	Sparsentan	96	8 (8.3)	NE		4.343	(0.627, 30.094)	0.137
	Irbesartan	99	2 (2.0)	NE				
> 45 years	Sparsentan	106	7 (6.6)	NE		0.549	(0.175, 1.717)	0.303
	Irbesartan	103	7 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	8 (7.8)	NE		2.163	(0.481, 9.728)	0.315
	Irbesartan	109	3 (2.8)	NE				
> 40 years	Sparsentan	91	7 (7.7)	NE		0.646	(0.187, 2.232)	0.490
	Irbesartan	88	6 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.246
North America	Sparsentan	35	1 (2.9)	NE		0.666	(0.053, 8.425)	0.754
	Irbesartan	46	4 (8.7)	NE				
Europe	Sparsentan	98	8 (8.2)	NE		4.379	(0.705, 27.201)	0.113
	Irbesartan	115	2 (1.7)	NE				
Asia Pacific	Sparsentan	69	6 (8.7)	NE		0.355	(0.070, 1.810)	0.213
	Irbesartan	41	3 (7.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.414
< 27 kg/m**2	Sparsentan	84	3 (3.6)	NE		0.699	(0.052, 9.333)	0.786
	Irbesartan	94	2 (2.1)	NE				
≥ 27 kg/m**2	Sparsentan	118	12 (10.2)	NE		0.988	(0.357, 2.738)	0.982
	Irbesartan	107	7 (6.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.176
eGFR Low and UP High	Sparsentan	71	3 (4.2)	NE		51.733	(0.982, NE)	NE
	Irbesartan	74	2 (2.7)	NE				
eGFR Low and UP Low	Sparsentan	55	7 (12.7)	NE		0.715	(0.149, 3.417)	0.674
	Irbesartan	55	4 (7.3)	NE				
eGFR High and UP High	Sparsentan	37	4 (10.8)	NE		3.984	(0.339, 46.873)	0.272
	Irbesartan	36	2 (5.6)	94.1	(94.1, NE)			
eGFR High and UP Low	Sparsentan	39	1 (2.6)	NE		0.540	(0.021, 13.995)	0.711
	Irbesartan	37	1 (2.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

Subgroup	Treatment	N	nev (%)	Kaplan-Meier analysis		Cox regression		
				Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	NE
< 60 mL/min/1.73 m**2	Sparsentan	127	10 (7.9)	NE		1.029	(0.344, 3.080)	0.959
	Irbesartan	129	7 (5.4)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	2 (4.1)	NE		0.996	(0.072, 13.717)	0.998
	Irbesartan	48	2 (4.2)	94.1	(94.1, NE)			
≥ 90 mL/min/1.73 m**2	Sparsentan	26	3 (11.5) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	NE
< 45 mL/min/1.73 m**2	Sparsentan	82	7 (8.5)	NE		1.767	(0.453, 6.889)	0.413
	Irbesartan	80	4 (5.0)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	3 (6.7)	NE		0.214	(0.014, 3.383)	0.274
	Irbesartan	49	3 (6.1)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	2 (4.1)	NE		0.996	(0.072, 13.717)	0.998
	Irbesartan	48	2 (4.2)	94.1	(94.1, NE)			
>= 90 mL/min/1.73 m**2	Sparsentan	26	3 (11.5) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.748
<= 1.75 g/day	Sparsentan	98	10 (10.2)	NE		1.791	(0.331, 9.694)	0.499
	Irbesartan	94	2 (2.1)	NE				
> 1.75 g/day	Sparsentan	104	5 (4.8)	NE		2.848	(0.544, 14.912)	0.215
	Irbesartan	108	7 (6.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment  
 was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan						Interaction test:	0.895
Yes	Sparsentan	88	6 (6.8)	NE		3.275	(0.477, 22.488)	0.227
	Irbesartan	83	4 (4.8)	NE				
No	Sparsentan	114	9 (7.9)	NE		1.144	(0.358, 3.654)	0.820
	Irbesartan	119	5 (4.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.897
<= 5 years	Sparsentan	113	10 (8.8)	NE		1.385	(0.501, 3.828)	0.530
	Irbesartan	127	8 (6.3)	NE				
> 5 years	Sparsentan	89	5 (5.6)	NE		1.548	(0.109, 21.980)	0.747
	Irbesartan	75	1 (1.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							
							Interaction test:	0.461
Yes	Sparsentan	153	12 (7.8)	NE		1.363	(0.459, 4.044)	0.577
	Irbesartan	157	7 (4.5)	NE				
No	Sparsentan	49	3 (6.1)	NE		0.707	(0.081, 6.185)	0.754
	Irbesartan	45	2 (4.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KSYIT\_FSKM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_ An increase reflects an improvement of the status of the patient.  
Reference table: PT1KSYIT\_FSTM

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.079
Male	Sparsentan	139	18 (12.9)	NE		1.488	(0.690, 3.210)	0.310
	Irbesartan	143	11 (7.7)	NE				
Female	Sparsentan	63	5 (7.9)	NE		0.353	(0.099, 1.257)	0.108
	Irbesartan	59	7 (11.9)	96.1	(70.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.201
<= 45 years	Sparsentan	96	9 (9.4)	NE		0.671	(0.278, 1.621)	0.375
	Irbesartan	99	12 (12.1)	NE				
> 45 years	Sparsentan	106	14 (13.2)	NE		1.575	(0.596, 4.160)	0.359
	Irbesartan	103	6 (5.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: 0.508
<= 18 years	Sparsentan	9	1 (11.1)	NE		52.163	(NE, NE)	NE
	Irbesartan	5	1 (20.0)	70.0	(NE, NE)			
> 18 to 40 years	Sparsentan	102	10 (9.8)	NE		0.807	(0.332, 1.966)	0.637
	Irbesartan	109	11 (10.1)	NE				
> 40 years	Sparsentan	91	12 (13.2)	NE		1.362	(0.505, 3.672)	0.541
	Irbesartan	88	6 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.429
North America	Sparsentan	35	1 (2.9)	NE		0.429	(0.038, 4.795)	0.492
	Irbesartan	46	3 (6.5)	NE				
Europe	Sparsentan	98	9 (9.2)	NE		0.726	(0.301, 1.753)	0.477
	Irbesartan	115	12 (10.4)	NE				
Asia Pacific	Sparsentan	69	13 (18.8)	NE		1.591	(0.444, 5.698)	0.476
	Irbesartan	41	3 (7.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.265
< 27 kg/m**2	Sparsentan	84	9 (10.7)	NE		0.623	(0.246, 1.573)	0.316
	Irbesartan	94	10 (10.6)	NE				
≥ 27 kg/m**2	Sparsentan	118	14 (11.9)	NE		1.537	(0.608, 3.884)	0.364
	Irbesartan	107	7 (6.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.896
eGFR Low and UP High	Sparsentan	71	9 (12.7)	NE		0.940	(0.330, 2.676)	0.908
	Irbesartan	74	7 (9.5)	NE				
eGFR Low and UP Low	Sparsentan	55	5 (9.1)	NE		1.127	(0.332, 3.822)	0.848
	Irbesartan	55	6 (10.9)	NE				
eGFR High and UP High	Sparsentan	37	6 (16.2)	NE		1.702	(0.338, 8.564)	0.519
	Irbesartan	36	2 (5.6)	NE				
eGFR High and UP Low	Sparsentan	39	3 (7.7)	NE		0.994	(0.199, 4.978)	0.995
	Irbesartan	37	3 (8.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.920
< 60 mL/min/1.73 m**2	Sparsentan	127	15 (11.8)	NE		0.895	(0.426, 1.882)	0.770
	Irbesartan	129	14 (10.9)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	4 (8.2)	NE		1.609	(0.290, 8.936)	0.587
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	4 (15.4)	NE		1.296	(0.215, 7.794)	0.777
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.328
< 45 mL/min/1.73 m**2	Sparsentan	82	11 (13.4)	NE		0.702	(0.312, 1.577)	0.391
	Irbesartan	80	13 (16.3)	96.1	(94.9, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	4 (8.9)	NE		3.492	(0.351, 34.729)	0.286
	Irbesartan	49	1 (2.0)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	4 (8.2)	NE		1.609	(0.290, 8.936)	0.587
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	4 (15.4)	NE		1.296	(0.215, 7.794)	0.777
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.937
<= 1.75 g/day	Sparsentan	98	11 (11.2)	NE		1.102	(0.411, 2.960)	0.846
	Irbesartan	94	7 (7.4)	NE				
> 1.75 g/day	Sparsentan	104	12 (11.5)	NE		1.092	(0.448, 2.659)	0.847
	Irbesartan	108	11 (10.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan						Interaction test:	0.248
Yes	Sparsentan	88	10 (11.4)	NE		1.623	(0.548, 4.805)	0.382
	Irbesartan	83	5 (6.0)	NE				
No	Sparsentan	114	13 (11.4)	NE		0.766	(0.352, 1.666)	0.501
	Irbesartan	119	13 (10.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment  
 was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.296
<= 5 years	Sparsentan	113	9 (8.0)	NE		0.656	(0.275, 1.568)	0.343
	Irbesartan	127	12 (9.4)	NE				
> 5 years	Sparsentan	89	14 (15.7)	NE		1.480	(0.553, 3.962)	0.435
	Irbesartan	75	6 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan						Interaction test:	0.974
Yes	Sparsentan	153	15 (9.8)	NE		0.932	(0.436, 1.995)	0.857
	Irbesartan	157	12 (7.6)	NE				
No	Sparsentan	49	8 (16.3)	NE		0.826	(0.261, 2.618)	0.745
	Irbesartan	45	6 (13.3)	96.1	(95.3, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KSYDT\_FSKM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_ reflects a worsening of the status of the patient.  
Reference table: PT1KSYDT\_FSTM



Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Sex												
Male	KDQOL-SF12: PCS	Baseline	Sparsentan	139	130 (93.5)	51.62 (7.25)	25.5	47.38	53.46	56.74	62.3	
			Irbesartan	143	132 (92.3)	52.00 (7.33)	21.6	49.77	54.43	56.42	64.3	
		Week 24	Sparsentan	139	88 (63.3)	51.56 (7.11)	16.3	47.65	52.97	56.49	62.0	
			Irbesartan	143	81 (56.6)	51.54 (7.63)	20.8	47.89	53.82	56.21	64.3	
		Week 48	Sparsentan	139	74 (53.2)	51.54 (6.74)	33.2	46.91	53.22	56.15	66.1	
			Irbesartan	143	58 (40.6)	52.74 (6.86)	32.8	50.22	55.09	56.71	64.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	139	88 (63.3)	0.71 (5.56)	-13.4	-3.23	0.29	3.73	15.5	0.13 [-0.18, 0.43]
			Irbesartan	143	81 (56.6)	-0.01 (6.02)	-20.0	-1.86	0.00	2.01	16.9	
		Week 48	Sparsentan	139	74 (53.2)	0.51 (6.74)	-18.7	-3.24	-0.14	4.31	20.6	0.01 [-0.34, 0.35]
			Irbesartan	143	58 (40.6)	0.46 (7.82)	-24.7	-2.98	0.42	3.64	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Female	KDQOL-SF12: PCS	Baseline	Sparsentan	63	57 (90.5)	49.75 (9.72)	12.4	44.71	52.33	56.15	63.5	
			Irbesartan	59	52 (88.1)	50.94 (6.16)	29.6	48.38	51.16	55.60	61.1	
	Week 24	Sparsentan	63	40 (63.5)	49.90 (8.20)	27.3	42.75	52.26	56.19	62.9		
		Irbesartan	59	31 (52.5)	50.73 (5.99)	39.0	45.08	53.18	55.09	59.1		
	Week 48	Sparsentan	63	38 (60.3)	49.98 (7.34)	29.8	46.56	50.05	55.09	61.7		
		Irbesartan	59	22 (37.3)	48.53 (7.30)	33.7	44.25	48.44	54.51	60.0		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	63	40 (63.5)	-0.36 (8.94)	-24.8	-6.06	-0.22	4.08	20.4	0.04 [-0.43, 0.51]
			Irbesartan	59	31 (52.5)	-0.67 (4.83)	-12.8	-3.24	-0.78	3.24	7.8	
Week 48		Sparsentan	63	38 (60.3)	0.58 (8.40)	-22.3	-3.11	1.07	5.05	24.7	0.30 [-0.23, 0.83]	
		Irbesartan	59	22 (37.3)	-1.84 (7.33)	-18.8	-5.29	-0.55	1.62	16.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age												
<= 45 years	KDQOL-SF12: PCS	Baseline	Sparsentan	96	88 (91.7)	52.37 (7.89)	12.4	49.42	54.38	57.08	63.5	
			Irbesartan	99	92 (92.9)	52.20 (7.28)	21.6	50.06	54.38	56.39	64.3	
		Week 24	Sparsentan	96	59 (61.5)	52.61 (6.88)	27.3	50.37	53.26	56.95	61.8	
			Irbesartan	99	56 (56.6)	52.10 (7.44)	20.8	51.03	54.51	56.18	64.3	
		Week 48	Sparsentan	96	53 (55.2)	52.42 (6.97)	29.8	48.13	54.23	56.45	66.1	
			Irbesartan	99	40 (40.4)	52.64 (7.02)	32.8	49.19	55.20	56.71	63.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	96	59 (61.5)	0.43 (6.24)	-24.8	-3.37	0.82	4.08	14.7	0.15 [-0.21, 0.52]
			Irbesartan	99	56 (56.6)	-0.44 (5.02)	-15.6	-2.69	-0.77	1.54	13.3	
		Week 48	Sparsentan	96	53 (55.2)	0.57 (7.48)	-22.3	-2.38	0.27	3.77	24.7	0.20 [-0.21, 0.61]
			Irbesartan	99	40 (40.4)	-0.91 (7.05)	-24.7	-2.15	0.00	2.30	16.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 45 years	KDQOL-SF12: PCS	Baseline	Sparsentan	106	99 (93.4)	49.88 (8.14)	25.5	45.62	52.30	56.15	61.8	
			Irbesartan	103	92 (89.3)	51.21 (6.75)	27.9	49.01	53.04	56.15	59.7	
	Week 24	Sparsentan	106	69 (65.1)	49.70 (7.75)	16.3	43.91	51.81	54.80	62.9		
		Irbesartan	103	56 (54.4)	50.53 (6.92)	31.2	44.62	52.77	56.15	61.9		
	Week 48	Sparsentan	106	59 (55.7)	49.74 (6.75)	33.2	45.63	51.57	54.51	60.2		
		Irbesartan	103	40 (38.8)	50.52 (7.29)	33.7	44.78	52.51	56.31	64.5		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	106	69 (65.1)	0.33 (7.25)	-14.9	-4.03	-0.27	3.53	20.4	0.04 [-0.31, 0.39]
			Irbesartan	103	56 (54.4)	0.04 (6.35)	-20.0	-1.72	0.40	3.61	16.9	
		Week 48	Sparsentan	106	59 (55.7)	0.49 (7.21)	-15.4	-3.76	-0.29	5.05	20.6	-0.01 [-0.41, 0.39]
			Irbesartan	103	40 (38.8)	0.57 (8.35)	-18.8	-4.31	0.55	4.96	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age at IgAN diagnosis												
<= 18 years	KDQOL-SF12: PCS	Baseline	Sparsentan	9	9 (100.0)	49.16 (14.77)	12.4	53.62	54.80	55.47	58.8	
			Irbesartan	5	5 (100.0)	50.57 (8.50)	39.0	44.78	53.18	56.15	59.7	
		Week 24	Sparsentan	9	3 (33.3)	59.89 (1.04)	58.9	58.85	59.89	60.93	60.9	
			Irbesartan	5	2 (40.0)	48.12 (4.57)	44.9	44.90	48.12	51.35	51.4	
			Irbesartan	5	1 (20.0)	55.59	55.6	55.59	55.59	55.59	55.6	
	Week 48	Sparsentan	9	5 (55.6)	51.76 (8.53)	37.1	52.50	54.23	56.15	58.8		
		Irbesartan	5	1 (20.0)	55.59	55.6	55.59	55.59	55.59	55.6		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	9	3 (33.3)	5.39 (1.41)	3.8	3.76	6.13	6.27	6.3	4.46 [1.17, 7.76]
			Irbesartan	5	2 (40.0)	-0.86 (1.38)	-1.8	-1.83	-0.86	0.12	0.1	
		Week 48	Sparsentan	9	5 (55.6)	4.69 (11.54)	-3.0	-2.69	0.60	3.75	24.7	NE
Irbesartan			5	1 (20.0)	2.40	2.4	2.40	2.40	2.40	2.4		
Irbesartan			5	1 (20.0)	2.40	2.4	2.40	2.40	2.40	2.4		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 18 to 40 years	KDQOL-SF12: PCS	Baseline	Sparsentan	102	94 (92.2)	52.86 (7.28)	25.5	49.93	54.52	57.30	63.5	
			Irbesartan	109	100 (91.7)	52.36 (6.99)	21.6	50.06	54.33	56.34	64.3	
		Week 24	Sparsentan	102	64 (62.7)	52.00 (6.83)	27.3	49.91	52.97	56.49	61.8	
			Irbesartan	109	61 (56.0)	52.57 (6.86)	20.8	51.90	54.51	56.22	64.3	
		Week 48	Sparsentan	102	57 (55.9)	52.89 (6.84)	29.8	49.81	54.32	57.76	66.1	
			Irbesartan	109	45 (41.3)	52.31 (6.80)	32.8	49.48	54.80	56.71	63.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	102	64 (62.7)	0.34 (6.61)	-24.8	-3.53	0.31	4.08	14.7	0.07 [-0.28, 0.42]
			Irbesartan	109	61 (56.0)	-0.05 (4.71)	-15.0	-2.02	0.00	1.83	13.3	
		Week 48	Sparsentan	102	57 (55.9)	0.37 (6.67)	-22.3	-2.43	0.02	3.69	15.6	0.15 [-0.24, 0.54]
			Irbesartan	109	45 (41.3)	-0.65 (7.39)	-24.7	-2.15	0.00	1.62	16.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	KDQOL-SF12: PCS	Baseline	Sparsentan	91	84 (92.3)	49.23 (7.69)	25.8	44.61	50.82	55.33	60.2	
			Irbesartan	88	79 (89.8)	50.95 (6.98)	27.9	48.47	52.45	56.15	59.7	
		Week 24	Sparsentan	91	61 (67.0)	49.59 (7.89)	16.3	43.35	52.11	54.80	62.9	
			Irbesartan	88	49 (55.7)	49.90 (7.48)	31.2	43.80	52.91	56.15	61.9	
		Week 48	Sparsentan	91	50 (54.9)	48.80 (6.41)	33.2	45.02	49.07	53.83	59.4	
			Irbesartan	88	34 (38.6)	50.50 (7.73)	33.7	44.25	52.51	56.15	64.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	91	61 (67.0)	0.17 (7.08)	-14.9	-3.94	-0.98	3.49	20.4	0.07 [-0.30, 0.45]
			Irbesartan	88	49 (55.7)	-0.35 (6.88)	-20.0	-3.49	0.00	3.78	16.9	
		Week 48	Sparsentan	91	50 (54.9)	0.30 (7.57)	-15.4	-4.33	-0.25	5.28	20.6	-0.01 [-0.45, 0.42]
			Irbesartan	88	34 (38.6)	0.39 (8.29)	-18.8	-4.24	0.42	4.53	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Geographic region												
North America	KDQOL-SF12: PCS	Baseline	Sparsentan	35	32 (91.4)	52.20 (9.84)	12.4	50.86	54.80	57.76	62.3	
			Irbesartan	46	44 (95.7)	50.87 (8.24)	27.9	47.98	54.38	56.15	61.1	
		Week 24	Sparsentan	35	15 (42.9)	54.73 (4.24)	45.7	52.64	55.66	57.49	61.3	
			Irbesartan	46	28 (60.9)	50.77 (7.36)	31.2	45.48	53.32	56.03	59.2	
			Sparsentan	35	13 (37.1)	53.19 (7.55)	37.1	53.17	55.59	57.76	61.7	
	Week 48	Sparsentan	35	13 (37.1)	53.19 (7.55)	37.1	53.17	55.59	57.76	61.7		
		Irbesartan	46	17 (37.0)	51.93 (7.14)	39.8	46.17	55.32	56.71	64.5		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	35	15 (42.9)	-0.30 (4.48)	-9.7	-3.22	0.00	3.24	6.1	-0.25 [-0.87, 0.38]
			Irbesartan	46	28 (60.9)	1.16 (6.55)	-20.0	-1.60	1.17	4.70	13.3	
		Week 48	Sparsentan	35	13 (37.1)	1.86 (8.93)	-15.4	-0.52	0.02	3.45	24.7	-0.07 [-0.79, 0.65]
Irbesartan			46	17 (37.0)	2.43 (7.84)	-8.9	-1.11	1.06	7.14	20.4		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Europe	KDQOL-SF12: PCS	Baseline	Sparsentan	98	88 (89.8)	51.17 (7.68)	25.5	47.76	53.32	56.66	62.3	
			Irbesartan	115	99 (86.1)	52.29 (6.36)	29.2	49.79	53.95	56.71	62.0	
		Week 24	Sparsentan	98	56 (57.1)	51.18 (8.01)	16.3	48.69	52.59	56.83	62.9	
			Irbesartan	115	50 (43.5)	51.64 (6.42)	36.8	46.58	52.89	56.15	64.3	
		Week 48	Sparsentan	98	44 (44.9)	51.09 (6.33)	36.5	46.97	53.15	54.95	61.9	
			Irbesartan	115	40 (34.8)	52.41 (7.24)	32.8	49.86	54.80	57.35	63.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	98	56 (57.1)	0.61 (6.34)	-13.4	-3.47	0.22	3.92	14.7	0.39 [0.01, 0.78]
			Irbesartan	115	50 (43.5)	-1.60 (4.80)	-15.0	-3.49	-0.94	0.56	10.0	
		Week 48	Sparsentan	98	44 (44.9)	0.56 (6.19)	-18.7	-2.83	1.09	4.51	11.4	0.21 [-0.22, 0.64]
			Irbesartan	115	40 (34.8)	-0.91 (7.90)	-24.7	-3.84	0.14	2.29	16.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Asia Pacific	KDQOL-SF12: PCS	Baseline	Sparsentan	69	67 (97.1)	50.34 (7.79)	32.4	44.69	52.08	56.15	63.5	
			Irbesartan	41	41 (100.0)	51.18 (7.18)	21.6	49.48	51.57	55.86	64.3	
		Week 24	Sparsentan	69	57 (82.6)	49.94 (7.38)	27.3	44.79	52.31	56.15	62.0	
			Irbesartan	41	34 (82.9)	51.30 (8.27)	20.8	47.89	54.44	56.15	59.4	
		Week 48	Sparsentan	69	55 (79.7)	50.44 (7.31)	29.8	46.11	52.35	56.15	66.1	
			Irbesartan	41	23 (56.1)	49.88 (7.16)	33.7	43.48	53.17	55.09	57.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	69	57 (82.6)	0.32 (7.72)	-24.8	-3.94	0.00	4.90	20.4	-0.06 [-0.49, 0.36]
			Irbesartan	41	34 (82.9)	0.75 (5.88)	-15.6	-1.62	0.28	3.78	16.9	
		Week 48	Sparsentan	69	55 (79.7)	0.20 (7.81)	-22.3	-3.58	-0.58	4.37	20.6	0.13 [-0.36, 0.62]
			Irbesartan	41	23 (56.1)	-0.80 (7.20)	-18.8	-4.39	0.00	3.24	15.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline BMI												
< 27 kg/m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	84	76 (90.5)	51.57 (8.78)	12.4	47.39	53.48	57.49	63.5	
			Irbesartan	94	87 (92.6)	53.21 (5.20)	39.0	49.84	54.53	56.71	61.5	
		Week 24	Sparsentan	84	52 (61.9)	51.83 (6.49)	37.6	47.99	53.46	57.36	62.0	
			Irbesartan	94	54 (57.4)	52.06 (5.63)	36.8	49.56	53.52	55.88	60.8	
		Week 48	Sparsentan	84	48 (57.1)	51.25 (6.89)	33.2	46.74	53.02	56.15	66.1	
			Irbesartan	94	37 (39.4)	51.57 (7.19)	32.8	46.35	54.51	56.71	63.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	84	52 (61.9)	0.63 (6.66)	-11.9	-4.39	-0.22	6.02	20.4	0.26 [-0.12, 0.64]
			Irbesartan	94	54 (57.4)	-0.80 (4.14)	-15.0	-2.38	-0.64	1.35	10.4	
		Week 48	Sparsentan	84	48 (57.1)	0.57 (7.27)	-13.9	-4.35	-0.53	5.16	24.7	0.34 [-0.09, 0.78]
			Irbesartan	94	37 (39.4)	-1.81 (6.46)	-24.7	-4.22	0.00	1.61	11.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 27 kg/m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	118	111 (94.1)	50.70 (7.63)	25.5	46.31	52.40	56.15	62.3	
			Irbesartan	107	96 (89.7)	50.47 (8.07)	21.6	47.36	52.90	56.10	64.3	
		Week 24	Sparsentan	118	76 (64.4)	50.50 (8.08)	16.3	46.38	52.45	56.19	62.9	
			Irbesartan	107	58 (54.2)	50.62 (8.39)	20.8	44.55	53.46	56.22	64.3	
		Week 48	Sparsentan	118	64 (54.2)	50.83 (7.05)	29.8	46.34	53.18	55.73	61.9	
			Irbesartan	107	43 (40.2)	51.59 (7.28)	33.7	48.01	54.80	56.40	64.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	118	76 (64.4)	0.20 (6.90)	-24.8	-3.41	0.31	3.51	15.5	-0.02 [-0.36, 0.32]
			Irbesartan	107	58 (54.2)	0.36 (6.84)	-20.0	-2.16	0.78	3.77	16.9	
		Week 48	Sparsentan	118	64 (54.2)	0.50 (7.39)	-22.3	-2.83	0.04	3.72	20.6	-0.09 [-0.48, 0.29]
			Irbesartan	107	43 (40.2)	1.24 (8.47)	-18.8	-3.45	1.07	5.40	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Randomization strata												
eGFR Low and UP High	KDQOL-SF12: PCS	Baseline	Sparsentan	71	64 (90.1)	51.45 (7.23)	29.7	46.69	53.18	56.41	63.5	
			Irbesartan	74	64 (86.5)	50.34 (7.86)	29.2	44.67	52.50	56.15	60.1	
		Week 24	Sparsentan	71	45 (63.4)	51.07 (7.77)	16.3	47.80	52.91	56.15	60.9	
			Irbesartan	74	36 (48.6)	49.69 (7.24)	34.9	43.14	52.16	56.01	61.9	
			Week 48	Sparsentan	71	39 (54.9)	52.28 (6.40)	33.2	47.92	53.46	57.49	61.9
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	71	45 (63.4)	-0.62 (6.81)	-13.4	-4.04	-0.64	3.10	20.4	0.03 [-0.41, 0.47]
			Irbesartan	74	36 (48.6)	-0.82 (5.84)	-15.6	-2.69	0.06	2.60	12.3	
		Week 48	Sparsentan	71	39 (54.9)	0.70 (7.94)	-18.7	-3.58	0.00	4.71	20.6	0.18 [-0.34, 0.70]
			Irbesartan	74	23 (31.1)	-0.84 (9.49)	-24.7	-5.82	0.00	2.96	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR Low and UP Low	KDQOL-SF12: PCS	Baseline	Sparsentan	55	51 (92.7)	49.09 (8.20)	25.8	45.40	50.55	55.14	61.1		
			Irbesartan	55	51 (92.7)	52.43 (4.53)	39.6	50.07	52.91	56.15	62.0		
		Week 24	Sparsentan	55	39 (70.9)	50.04 (7.26)	31.2	43.35	52.40	54.26	62.0		
			Irbesartan	55	35 (63.6)	51.82 (6.61)	32.9	47.25	53.46	56.22	64.3		
		Week 48	Sparsentan	55	28 (50.9)	50.24 (7.05)	36.0	45.48	53.18	55.33	59.4		
			Irbesartan	55	25 (45.5)	51.60 (6.88)	37.7	47.21	53.46	56.97	63.5		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	55	39 (70.9)	1.54 (6.42)	-14.9	-3.25	0.69	4.90	14.7	0.34 [-0.12, 0.80]	
			Irbesartan	55	35 (63.6)	-0.61 (6.28)	-20.0	-2.55	0.00	2.22	16.9		
			Week 48	Sparsentan	55	28 (50.9)	2.16 (4.61)	-7.4	-0.84	1.58	6.57	10.7	0.42 [-0.12, 0.97]
				Irbesartan	55	25 (45.5)	-0.33 (7.08)	-11.9	-2.86	0.56	1.61	16.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP High	KDQOL-SF12: PCS	Baseline	Sparsentan	37	35 (94.6)	52.49 (9.30)	12.4	50.75	54.80	56.71	62.9	
			Irbesartan	36	33 (91.7)	51.69 (7.00)	32.5	48.47	53.46	56.15	64.3	
		Week 24	Sparsentan	37	20 (54.1)	53.09 (8.47)	27.3	50.71	56.72	57.74	61.8	
			Irbesartan	36	19 (52.8)	52.59 (4.28)	43.0	51.35	53.46	56.15	58.9	
		Week 48	Sparsentan	37	22 (59.5)	50.41 (8.00)	29.8	45.68	52.98	55.09	61.7	
			Irbesartan	36	13 (36.1)	50.40 (4.98)	40.3	46.17	51.57	54.80	55.6	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	37	20 (54.1)	-0.02 (7.50)	-24.8	-1.45	1.59	3.64	9.9	-0.27 [-0.90, 0.36]
			Irbesartan	36	19 (52.8)	1.75 (5.21)	-5.9	-1.83	1.35	3.64	13.3	
		Week 48	Sparsentan	37	22 (59.5)	-0.78 (9.31)	-22.3	-3.76	-2.02	3.39	24.7	-0.09 [-0.77, 0.60]
			Irbesartan	36	13 (36.1)	-0.02 (7.58)	-15.8	-3.90	-1.06	4.31	13.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP Low	KDQOL-SF12: PCS	Baseline	Sparsentan	39	37 (94.9)	51.70 (8.03)	25.5	46.73	53.46	57.30	62.3	
			Irbesartan	37	36 (97.3)	53.11 (8.14)	21.6	51.72	55.73	56.96	60.8	
	Week 24	Sparsentan	39	24 (61.5)	50.89 (6.44)	39.4	46.74	51.57	55.24	62.9		
		Irbesartan	37	22 (59.5)	52.08 (9.65)	20.8	53.18	54.95	56.48	60.8		
	Week 48	Sparsentan	39	23 (59.0)	50.37 (6.87)	35.7	46.11	49.35	55.87	66.1		
		Irbesartan	37	19 (51.4)	54.01 (7.51)	33.7	54.80	56.40	57.49	62.5		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	39	24 (61.5)	0.68 (6.77)	-8.4	-3.98	-0.98	5.03	14.1	0.15 [-0.43, 0.73]
			Irbesartan	37	22 (59.5)	-0.20 (4.86)	-12.8	-2.16	-0.77	2.42	10.4	
		Week 48	Sparsentan	39	23 (59.0)	-0.48 (6.74)	-13.6	-5.01	0.27	3.77	11.4	-0.18 [-0.79, 0.43]
			Irbesartan	37	19 (51.4)	0.75 (6.67)	-18.8	-1.04	1.32	3.24	12.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline eGFR Group 1												
< 60 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	127	116 (91.3)	50.33 (7.67)	25.8	46.15	52.34	56.15	63.5	
			Irbesartan	129	116 (89.9)	51.42 (6.60)	27.9	48.61	52.90	56.15	62.0	
	Week 24	Sparsentan	127	85 (66.9)	50.39 (7.44)	16.3	45.43	52.40	55.66	62.0		
		Irbesartan	129	73 (56.6)	50.64 (7.05)	31.2	44.69	52.50	56.15	64.3		
	Week 48	Sparsentan	127	67 (52.8)	51.58 (6.60)	33.2	46.96	53.39	56.15	61.9		
		Irbesartan	129	50 (38.8)	51.43 (7.51)	32.8	46.35	53.58	56.71	64.5		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	127	85 (66.9)	0.27 (6.73)	-14.9	-3.69	0.00	3.69	20.4	0.10 [-0.21, 0.42]
			Irbesartan	129	73 (56.6)	-0.38 (5.87)	-20.0	-2.69	0.00	2.31	16.9	
		Week 48	Sparsentan	127	67 (52.8)	1.70 (6.49)	-18.7	-2.43	0.69	6.54	20.6	0.17 [-0.20, 0.54]
			Irbesartan	129	50 (38.8)	0.46 (8.29)	-24.7	-4.24	1.06	4.27	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	49	47 (95.9)	53.00 (6.69)	37.4	48.78	53.79	57.76	61.8	
		Week 24	Irbesartan	48	44 (91.7)	52.03 (7.49)	29.6	49.15	54.95	56.30	64.3	
			Sparsentan	49	28 (57.1)	52.61 (7.74)	27.3	48.92	53.07	58.39	62.9	
	Week 48	Irbesartan	48	26 (54.2)	53.08 (6.00)	34.9	52.17	54.52	56.15	60.8		
		Sparsentan	49	29 (59.2)	49.94 (8.14)	29.8	43.15	51.29	55.59	66.1		
		Irbesartan	48	20 (41.7)	50.46 (7.10)	33.7	45.94	53.18	55.87	58.6		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	49	28 (57.1)	0.18 (7.43)	-24.8	-3.63	0.25	5.14	11.8	0.03 [-0.50, 0.57]
		Week 48	Irbesartan	48	26 (54.2)	-0.06 (6.31)	-15.6	-2.16	-0.04	3.24	13.3	
			Sparsentan	49	29 (59.2)	-2.24 (7.65)	-22.3	-5.01	-2.13	3.45	11.4	0.05 [-0.52, 0.62]
			Week 48	Irbesartan	48	20 (41.7)	-2.58 (7.17)	-18.8	-5.97	-1.20	0.55	9.5

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	26	24 (92.3)	50.73 (11.70)	12.4	49.48	54.33	56.81	62.9		
			Irbesartan	25	24 (96.0)	52.47 (8.27)	21.6	51.67	55.35	56.15	61.1		
		Week 24	Sparsentan	26	15 (57.7)	51.81 (7.16)	36.9	48.47	54.33	57.49	58.0		
			Irbesartan	25	13 (52.0)	51.62 (9.82)	20.8	52.50	54.36	56.40	58.6		
		Week 48	Sparsentan	26	16 (61.5)	50.56 (6.23)	37.1	46.13	51.27	55.48	58.6		
			Irbesartan	25	10 (40.0)	54.58 (5.26)	40.3	54.51	56.28	57.49	58.0		
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	26	15 (57.7)	1.35 (6.15)	-6.4	-3.22	1.57	2.98	14.1	0.15 [-0.59, 0.90]
				Irbesartan	25	13 (52.0)	0.58 (3.20)	-3.2	-1.06	0.00	1.36	10.0	
			Week 48	Sparsentan	26	16 (61.5)	0.64 (8.93)	-11.7	-3.62	0.04	2.08	24.7	-0.11 [-0.90, 0.68]
				Irbesartan	25	10 (40.0)	1.48 (4.68)	-3.9	-1.04	1.33	1.63	13.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 2													
< 45 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	82	72 (87.8)	49.73 (7.85)	29.2	44.55	51.65	56.05	60.8		
			Irbesartan	80	70 (87.5)	50.48 (6.75)	29.2	46.31	52.11	56.06	60.1		
		Week 24	Sparsentan	82	54 (65.9)	50.20 (7.46)	16.3	45.37	52.40	54.80	60.9		
			Irbesartan	80	48 (60.0)	50.24 (6.36)	36.8	44.17	52.11	56.01	61.9		
		Week 48	Sparsentan	82	44 (53.7)	51.72 (6.68)	36.0	46.74	53.42	56.82	61.9		
			Irbesartan	80	30 (37.5)	50.35 (7.92)	32.8	45.30	51.22	56.48	64.5		
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	82	54 (65.9)	0.47 (6.22)	-13.4	-3.37	-0.09	4.07	15.5	0.12 [-0.27, 0.51]
			Irbesartan	80	48 (60.0)	-0.25 (5.71)	-15.0	-2.25	0.00	2.84	16.9		
			Week 48	Sparsentan	82	44 (53.7)	2.15 (6.33)	-18.7	-1.25	1.46	6.32	20.6	0.32 [-0.15, 0.79]
				Irbesartan	80	30 (37.5)	-0.33 (9.48)	-24.7	-4.39	-0.55	2.19	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
45 to < 60 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	45	44 (97.8)	51.30 (7.34)	25.8	46.86	52.52	56.15	63.5	
		Week 24	Irbesartan	49	46 (93.9)	52.86 (6.16)	27.9	50.34	53.66	57.20	62.0	
			Sparsentan	45	31 (68.9)	50.71 (7.52)	31.2	45.43	53.03	56.15	62.0	
	Week 48	Irbesartan	49	25 (51.0)	51.40 (8.30)	31.2	47.89	54.78	56.15	64.3		
		Sparsentan	45	23 (51.1)	51.33 (6.58)	33.2	47.92	52.91	55.57	60.7		
		Irbesartan	49	20 (40.8)	53.05 (6.71)	39.0	51.02	54.67	57.76	62.5		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	45	31 (68.9)	-0.09 (7.62)	-14.9	-4.25	0.00	2.80	20.4	0.08 [-0.45, 0.61]
		Week 48	Irbesartan	49	25 (51.0)	-0.64 (6.29)	-20.0	-3.84	0.00	2.22	10.1	
			Sparsentan	45	23 (51.1)	0.84 (6.85)	-13.9	-3.11	-0.58	6.61	17.0	-0.13 [-0.72, 0.47]
		Irbesartan	49	20 (40.8)	1.65 (6.12)	-11.3	0.81	2.15	4.29	12.5		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	49	47 (95.9)	53.00 (6.69)	37.4	48.78	53.79	57.76	61.8	
			Irbesartan	48	44 (91.7)	52.03 (7.49)	29.6	49.15	54.95	56.30	64.3	
	Week 24	Sparsentan	49	28 (57.1)	52.61 (7.74)	27.3	48.92	53.07	58.39	62.9		
		Irbesartan	48	26 (54.2)	53.08 (6.00)	34.9	52.17	54.52	56.15	60.8		
	Week 48	Sparsentan	49	29 (59.2)	49.94 (8.14)	29.8	43.15	51.29	55.59	66.1		
		Irbesartan	48	20 (41.7)	50.46 (7.10)	33.7	45.94	53.18	55.87	58.6		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	49	28 (57.1)	0.18 (7.43)	-24.8	-3.63	0.25	5.14	11.8	0.03 [-0.50, 0.57]
			Irbesartan	48	26 (54.2)	-0.06 (6.31)	-15.6	-2.16	-0.04	3.24	13.3	
		Week 48	Sparsentan	49	29 (59.2)	-2.24 (7.65)	-22.3	-5.01	-2.13	3.45	11.4	0.05 [-0.52, 0.62]
			Irbesartan	48	20 (41.7)	-2.58 (7.17)	-18.8	-5.97	-1.20	0.55	9.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	26	24 (92.3)	50.73 (11.70)	12.4	49.48	54.33	56.81	62.9		
			Irbesartan	25	24 (96.0)	52.47 (8.27)	21.6	51.67	55.35	56.15	61.1		
		Week 24	Sparsentan	26	15 (57.7)	51.81 (7.16)	36.9	48.47	54.33	57.49	58.0		
			Irbesartan	25	13 (52.0)	51.62 (9.82)	20.8	52.50	54.36	56.40	58.6		
		Week 48	Sparsentan	26	16 (61.5)	50.56 (6.23)	37.1	46.13	51.27	55.48	58.6		
			Irbesartan	25	10 (40.0)	54.58 (5.26)	40.3	54.51	56.28	57.49	58.0		
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	26	15 (57.7)	1.35 (6.15)	-6.4	-3.22	1.57	2.98	14.1	0.15 [-0.59, 0.90]
				Irbesartan	25	13 (52.0)	0.58 (3.20)	-3.2	-1.06	0.00	1.36	10.0	
			Week 48	Sparsentan	26	16 (61.5)	0.64 (8.93)	-11.7	-3.62	0.04	2.08	24.7	-0.11 [-0.90, 0.68]
				Irbesartan	25	10 (40.0)	1.48 (4.68)	-3.9	-1.04	1.33	1.63	13.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline urine protein excretion												
<= 1.75 g/day	KDQOL-SF12: PCS	Baseline	Sparsentan	98	92 (93.9)	50.30 (8.31)	25.5	45.89	52.36	56.84	62.3	
			Irbesartan	94	84 (89.4)	52.02 (6.72)	21.6	49.93	53.69	56.15	60.8	
		Week 24	Sparsentan	98	67 (68.4)	50.05 (7.46)	27.3	43.91	52.35	54.80	62.9	
			Irbesartan	94	47 (50.0)	51.97 (7.60)	20.8	50.61	54.51	56.44	60.8	
		Week 48	Sparsentan	98	60 (61.2)	50.56 (7.32)	29.8	46.34	51.96	55.73	66.1	
			Irbesartan	94	34 (36.2)	52.56 (6.57)	33.7	49.68	54.95	57.20	62.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	98	67 (68.4)	0.73 (7.76)	-24.8	-3.82	0.00	5.07	20.4	0.15 [-0.23, 0.52]
			Irbesartan	94	47 (50.0)	-0.33 (6.10)	-20.0	-2.16	0.00	2.31	16.9	
		Week 48	Sparsentan	98	60 (61.2)	0.74 (6.74)	-22.3	-2.53	1.07	4.74	17.0	0.18 [-0.24, 0.61]
			Irbesartan	94	34 (36.2)	-0.46 (6.06)	-18.8	-2.15	0.27	1.61	15.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 1.75 g/day	KDQOL-SF12: PCS	Baseline	Sparsentan	104	95 (91.3)	51.77 (7.87)	12.4	48.84	53.50	56.68	63.5	
			Irbesartan	108	100 (92.6)	51.44 (7.29)	27.9	48.39	53.46	56.34	64.3	
		Week 24	Sparsentan	104	61 (58.7)	52.12 (7.40)	16.3	50.30	53.46	57.49	61.8	
			Irbesartan	108	65 (60.2)	50.85 (6.91)	31.2	45.73	52.91	56.15	64.3	
		Week 48	Sparsentan	104	52 (50.0)	51.53 (6.55)	33.2	46.97	53.36	56.82	60.7	
			Irbesartan	108	46 (42.6)	50.86 (7.61)	32.8	45.72	53.04	56.48	64.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	104	61 (58.7)	-0.01 (5.54)	-13.4	-3.08	0.29	2.98	15.5	0.02 [-0.33, 0.37]
			Irbesartan	108	65 (60.2)	-0.10 (5.44)	-15.6	-2.38	0.29	2.69	13.3	
		Week 48	Sparsentan	104	52 (50.0)	0.29 (7.97)	-18.7	-3.36	-0.52	4.12	24.7	0.03 [-0.37, 0.43]
			Irbesartan	108	46 (42.6)	0.04 (8.80)	-24.7	-4.66	0.14	5.13	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline use of antihypertensives												
Yes	KDQOL-SF12: PCS	Baseline	Sparsentan	88	78 (88.6)	50.39 (8.19)	25.8	46.31	52.90	56.15	61.2	
			Irbesartan	83	73 (88.0)	50.33 (7.47)	27.9	45.74	51.28	55.88	64.3	
		Week 24	Sparsentan	88	53 (60.2)	51.16 (7.01)	36.9	45.67	52.70	56.95	62.9	
			Irbesartan	83	43 (51.8)	49.70 (6.56)	31.2	44.55	52.11	55.86	58.9	
		Week 48	Sparsentan	88	43 (48.9)	50.79 (7.39)	35.7	44.69	53.24	57.49	61.9	
			Irbesartan	83	30 (36.1)	50.18 (8.06)	32.8	43.48	51.22	56.15	64.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	88	53 (60.2)	1.04 (7.12)	-14.9	-3.45	-0.64	4.30	20.4	0.13 [-0.27, 0.54]
			Irbesartan	83	43 (51.8)	0.11 (6.75)	-15.0	-3.49	0.00	3.64	16.9	
		Week 48	Sparsentan	88	43 (48.9)	1.78 (7.50)	-15.4	-3.10	0.60	6.61	20.6	0.07 [-0.40, 0.53]
			Irbesartan	83	30 (36.1)	1.20 (10.41)	-24.7	-4.24	1.48	9.50	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL-SF12: PCS	Baseline	Sparsentan	114	109 (95.6)	51.52 (8.04)	12.4	47.38	53.44	56.98	63.5	
			Irbesartan	119	111 (93.3)	52.61 (6.59)	21.6	50.34	54.51	56.30	62.0	
		Week 24	Sparsentan	114	75 (65.8)	50.95 (7.84)	16.3	48.65	52.64	56.24	62.0	
			Irbesartan	119	69 (58.0)	52.32 (7.44)	20.8	51.90	54.53	56.22	64.3	
		Week 48	Sparsentan	114	69 (60.5)	51.15 (6.73)	29.8	46.91	53.17	55.59	66.1	
			Irbesartan	119	50 (42.0)	52.42 (6.56)	37.7	48.88	54.80	56.71	63.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	114	75 (65.8)	-0.09 (6.54)	-24.8	-3.70	0.29	3.49	14.1	0.05 [-0.28, 0.38]
			Irbesartan	119	69 (58.0)	-0.38 (4.99)	-20.0	-2.13	0.00	1.83	10.4	
		Week 48	Sparsentan	114	69 (60.5)	-0.24 (7.12)	-22.3	-3.28	-0.52	3.24	24.7	0.12 [-0.25, 0.48]
			Irbesartan	119	50 (42.0)	-0.99 (5.48)	-15.8	-2.98	0.00	1.62	13.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Time since renal biopsy												
<= 5 years	KDQOL-SF12: PCS	Baseline	Sparsentan	113	105 (92.9)	49.79 (7.95)	25.8	45.62	51.57	56.15	63.5	
			Irbesartan	127	119 (93.7)	51.63 (7.63)	21.6	48.83	54.13	56.42	64.3	
		Week 24	Sparsentan	113	73 (64.6)	51.09 (7.47)	16.3	47.50	52.50	56.24	62.0	
			Irbesartan	127	73 (57.5)	50.60 (7.79)	20.8	45.08	53.82	56.15	59.4	
		Week 48	Sparsentan	113	61 (54.0)	50.08 (6.72)	33.2	45.70	51.29	55.57	60.7	
			Irbesartan	127	53 (41.7)	51.92 (7.67)	32.8	46.35	54.80	56.71	64.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	113	73 (64.6)	1.65 (6.57)	-13.4	-3.08	1.35	4.84	20.4	0.33 [0.00, 0.66]
			Irbesartan	127	73 (57.5)	-0.47 (6.33)	-20.0	-3.24	-1.06	2.42	16.9	
		Week 48	Sparsentan	113	61 (54.0)	1.05 (7.30)	-18.7	-3.15	0.02	5.28	20.6	0.05 [-0.32, 0.42]
			Irbesartan	127	53 (41.7)	0.64 (8.17)	-24.7	-2.98	1.09	4.27	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 5 years	KDQOL-SF12: PCS	Baseline	Sparsentan	89	82 (92.1)	52.67 (8.06)	12.4	49.93	54.80	57.25	62.3	
			Irbesartan	75	65 (86.7)	51.85 (5.81)	34.9	49.51	53.43	56.11	62.0	
	Week 24	Sparsentan	89	55 (61.8)	50.97 (7.56)	27.3	44.79	52.77	56.95	62.9		
		Irbesartan	75	39 (52.0)	52.66 (5.80)	39.0	51.35	53.46	56.40	64.3		
	Week 48	Sparsentan	89	51 (57.3)	52.13 (7.13)	29.8	48.86	53.39	56.15	66.1		
		Irbesartan	75	27 (36.0)	50.90 (6.23)	37.7	48.01	52.11	56.42	58.6		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	89	55 (61.8)	-1.32 (6.74)	-24.8	-5.08	-0.27	2.69	14.1	-0.28 [-0.69, 0.13]
			Irbesartan	75	39 (52.0)	0.31 (4.33)	-12.7	-0.83	0.56	2.31	10.8	
		Week 48	Sparsentan	89	51 (57.3)	-0.09 (7.33)	-22.3	-3.47	-0.04	3.69	24.7	0.24 [-0.23, 0.70]
			Irbesartan	75	27 (36.0)	-1.76 (6.58)	-12.5	-7.08	-1.81	1.33	13.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: History of hypertension												
Yes	KDQOL-SF12: PCS	Baseline	Sparsentan	153	139 (90.8)	50.02 (8.60)	12.4	45.69	52.30	56.19	62.3	
			Irbesartan	157	140 (89.2)	51.65 (6.66)	27.9	49.14	53.21	56.15	64.3	
		Week 24	Sparsentan	153	97 (63.4)	50.59 (7.61)	16.3	45.84	52.35	56.15	62.9	
			Irbesartan	157	83 (52.9)	50.64 (6.92)	31.2	44.90	52.91	55.88	61.9	
		Week 48	Sparsentan	153	83 (54.2)	50.39 (6.72)	33.2	45.70	52.50	55.59	61.9	
			Irbesartan	157	59 (37.6)	51.42 (7.48)	32.8	45.72	54.80	56.71	64.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	153	97 (63.4)	0.77 (6.64)	-14.9	-3.37	0.21	4.08	20.4	0.19 [-0.10, 0.48]
			Irbesartan	157	83 (52.9)	-0.45 (6.25)	-20.0	-3.24	0.00	2.42	16.9	
		Week 48	Sparsentan	153	83 (54.2)	0.87 (7.45)	-18.7	-3.47	0.00	5.11	24.7	0.07 [-0.26, 0.41]
			Irbesartan	157	59 (37.6)	0.30 (8.50)	-24.7	-3.90	1.06	3.64	20.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL-SF12: PCS	Baseline	Sparsentan	49	48 (98.0)	54.03 (5.53)	37.4	51.94	54.80	57.39	63.5	
			Irbesartan	45	44 (97.8)	51.89 (8.15)	21.6	49.37	55.07	56.42	62.0	
		Week 24	Sparsentan	49	31 (63.3)	52.45 (6.97)	27.3	49.67	54.26	57.49	60.9	
			Irbesartan	45	29 (64.4)	53.25 (7.74)	20.8	52.39	54.80	56.48	64.3	
		Week 48	Sparsentan	49	29 (59.2)	52.78 (7.43)	29.8	50.03	53.71	56.45	66.1	
			Irbesartan	45	21 (46.7)	52.02 (6.47)	39.0	49.48	53.17	56.42	63.5	
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	49	31 (63.3)	-0.87 (7.15)	-24.8	-4.04	0.00	2.70	12.0	-0.24 [-0.75, 0.27]
			Irbesartan	45	29 (64.4)	0.52 (3.71)	-8.9	-1.49	0.29	2.22	10.4	
		Week 48	Sparsentan	49	29 (59.2)	-0.45 (6.92)	-22.3	-2.88	-0.04	1.93	15.6	0.17 [-0.39, 0.73]
			Irbesartan	45	21 (46.7)	-1.49 (4.78)	-11.3	-4.66	0.00	1.32	6.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.980
Male	Week 24	Sparsentan	139	88 (63.3)	0.64 (0.61)	(-0.56, 1.85)	0.57 (0.88)	(-1.17, 2.31)	0.517
		Irbesartan	143	81 (56.6)	0.07 (0.64)	(-1.18, 1.32)			
	Week 48	Sparsentan	139	74 (53.2)	0.45 (0.65)	(-0.83, 1.74)	0.10 (0.98)	(-1.84, 2.03)	0.921
		Irbesartan	143	58 (40.6)	0.36 (0.73)	(-1.08, 1.80)			
Female	Week 24	Sparsentan	63	40 (63.5)	-0.29 (1.01)	(-2.30, 1.72)	-0.24 (1.55)	(-3.31, 2.84)	0.879
		Irbesartan	59	31 (52.5)	-0.05 (1.17)	(-2.37, 2.27)			
	Week 48	Sparsentan	63	38 (60.3)	-0.44 (1.03)	(-2.49, 1.60)	0.95 (1.71)	(-2.43, 4.33)	0.578
		Irbesartan	59	22 (37.3)	-1.40 (1.35)	(-4.06, 1.27)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024



Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.395
<= 45 years	Week 24	Sparsentan	96	59 (61.5)	0.33 (0.75)	(-1.14, 1.81)	0.63 (1.07)	(-1.48, 2.74)	0.558
		Irbesartan	99	56 (56.6)	-0.29 (0.77)	(-1.81, 1.22)			
	Week 48	Sparsentan	96	53 (55.2)	0.25 (0.78)	(-1.30, 1.80)	1.08 (1.19)	(-1.27, 3.43)	0.365
		Irbesartan	99	40 (40.4)	-0.83 (0.90)	(-2.60, 0.94)			
> 45 years	Week 24	Sparsentan	106	69 (65.1)	0.25 (0.73)	(-1.20, 1.70)	-0.22 (1.11)	(-2.41, 1.96)	0.841
		Irbesartan	103	56 (54.4)	0.47 (0.83)	(-1.16, 2.11)			
	Week 48	Sparsentan	106	59 (55.7)	-0.05 (0.77)	(-1.58, 1.47)	-0.86 (1.21)	(-3.25, 1.53)	0.477
		Irbesartan	103	40 (38.8)	0.81 (0.93)	(-1.02, 2.64)			

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.510
<= 18 years	Week 24	Sparsentan	9	3 (33.3)	NE		NE		NE
		Irbesartan	5	2 (40.0)	NE				
	Week 48	Sparsentan	9	5 (55.6)	NE		NE		NE
		Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Week 24	Sparsentan	102	64 (62.7)	0.09 (0.70)	(-1.29, 1.47)	-0.00 (1.00)	(-1.98, 1.98)	0.999
		Irbesartan	109	61 (56.0)	0.09 (0.72)	(-1.33, 1.51)			
	Week 48	Sparsentan	102	57 (55.9)	0.44 (0.74)	(-1.02, 1.90)	1.15 (1.12)	(-1.06, 3.35)	0.306
		Irbesartan	109	45 (41.3)	-0.71 (0.83)	(-2.35, 0.93)			
> 40 years	Week 24	Sparsentan	91	61 (67.0)	0.30 (0.82)	(-1.33, 1.93)	0.31 (1.23)	(-2.13, 2.75)	0.801
		Irbesartan	88	49 (55.7)	-0.01 (0.92)	(-1.83, 1.80)			
	Week 48	Sparsentan	91	50 (54.9)	-0.56 (0.87)	(-2.28, 1.17)	-1.33 (1.35)	(-4.01, 1.34)	0.326
		Irbesartan	88	34 (38.6)	0.78 (1.03)	(-1.26, 2.82)			

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For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.854
North America	Week 24	Sparsentan	35	15 (42.9)	1.77 (1.45)	(-1.13, 4.68)	1.43 (1.82)	(-2.21, 5.07)	0.436
		Irbesartan	46	28 (60.9)	0.34 (1.06)	(-1.78, 2.47)			
	Week 48	Sparsentan	35	13 (37.1)	2.25 (1.55)	(-0.85, 5.36)	0.89 (2.07)	(-3.25, 5.02)	0.670
		Irbesartan	46	17 (37.0)	1.36 (1.35)	(-1.33, 4.06)			
Europe	Week 24	Sparsentan	98	56 (57.1)	0.25 (0.76)	(-1.24, 1.75)	1.28 (1.11)	(-0.92, 3.47)	0.253
		Irbesartan	115	50 (43.5)	-1.02 (0.81)	(-2.62, 0.57)			
	Week 48	Sparsentan	98	44 (44.9)	-0.04 (0.84)	(-1.71, 1.62)	0.43 (1.23)	(-2.00, 2.86)	0.728
		Irbesartan	115	40 (34.8)	-0.47 (0.89)	(-2.22, 1.28)			
Asia Pacific	Week 24	Sparsentan	69	57 (82.6)	0.21 (0.86)	(-1.49, 1.91)	-0.72 (1.42)	(-3.54, 2.10)	0.615
		Irbesartan	41	34 (82.9)	0.93 (1.13)	(-1.30, 3.16)			
	Week 48	Sparsentan	69	55 (79.7)	-0.07 (0.87)	(-1.79, 1.65)	0.93 (1.57)	(-2.17, 4.02)	0.554
		Irbesartan	41	23 (56.1)	-1.00 (1.29)	(-3.55, 1.55)			

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For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.412
< 27 kg/m**2	Week 24	Sparsentan	84	52 (61.9)	0.36 (0.73)	(-1.07, 1.79)	0.57 (1.03)	(-1.45, 2.60)	0.577
		Irbesartan	94	54 (57.4)	-0.21 (0.72)	(-1.63, 1.20)			
	Week 48	Sparsentan	84	48 (57.1)	0.01 (0.75)	(-1.48, 1.49)	1.23 (1.14)	(-1.02, 3.48)	0.281
		Irbesartan	94	37 (39.4)	-1.22 (0.85)	(-2.89, 0.44)			
≥ 27 kg/m**2	Week 24	Sparsentan	118	76 (64.4)	0.31 (0.74)	(-1.15, 1.77)	0.01 (1.13)	(-2.22, 2.24)	0.991
		Irbesartan	107	58 (54.2)	0.30 (0.85)	(-1.38, 1.98)			
	Week 48	Sparsentan	118	64 (54.2)	0.23 (0.79)	(-1.32, 1.79)	-0.68 (1.24)	(-3.13, 1.76)	0.582
		Irbesartan	107	43 (40.2)	0.92 (0.96)	(-0.97, 2.81)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan						Interaction:	0.722
eGFR Low and UP High	Week 24	Sparsentan	71	45 (63.4)	-0.37 (0.96)	(-2.28, 1.54)	0.79 (1.46)	(-2.09, 3.68)	0.586
		Irbesartan	74	36 (48.6)	-1.17 (1.09)	(-3.33, 0.99)			
	Week 48	Sparsentan	71	39 (54.9)	0.77 (1.01)	(-1.23, 2.78)	1.32 (1.63)	(-1.91, 4.54)	0.421
		Irbesartan	74	23 (31.1)	-0.54 (1.28)	(-3.07, 1.98)			
eGFR Low and UP Low	Week 24	Sparsentan	55	39 (70.9)	0.90 (0.90)	(-0.88, 2.67)	0.53 (1.33)	(-2.10, 3.16)	0.690
		Irbesartan	55	35 (63.6)	0.37 (0.95)	(-1.52, 2.26)			
	Week 48	Sparsentan	55	28 (50.9)	0.82 (1.04)	(-1.24, 2.89)	0.85 (1.53)	(-2.18, 3.89)	0.579
		Irbesartan	55	25 (45.5)	-0.03 (1.10)	(-2.22, 2.16)			
eGFR High and UP High	Week 24	Sparsentan	37	20 (54.1)	0.99 (1.26)	(-1.52, 3.51)	-0.36 (1.82)	(-3.99, 3.26)	0.843
		Irbesartan	36	19 (52.8)	1.36 (1.31)	(-1.25, 3.97)			
	Week 48	Sparsentan	37	22 (59.5)	-0.73 (1.20)	(-3.14, 1.67)	-0.06 (1.96)	(-3.96, 3.84)	0.975
		Irbesartan	36	13 (36.1)	-0.67 (1.54)	(-3.75, 2.41)			
eGFR High and UP Low	Week 24	Sparsentan	39	24 (61.5)	0.50 (1.16)	(-1.82, 2.81)	0.40 (1.68)	(-2.96, 3.75)	0.814
		Irbesartan	37	22 (59.5)	0.10 (1.21)	(-2.31, 2.51)			
	Week 48	Sparsentan	39	23 (59.0)	-0.66 (1.18)	(-3.02, 1.70)	-1.71 (1.76)	(-5.22, 1.81)	0.337
		Irbesartan	37	19 (51.4)	1.05 (1.30)	(-1.54, 3.63)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.841
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	85 (66.9)	0.17 (0.65)	(-1.10, 1.45)	0.34 (0.96)	(-1.55, 2.23)	0.725
		Irbesartan	129	73 (56.6)	-0.17 (0.71)	(-1.56, 1.22)			
	Week 48	Sparsentan	127	67 (52.8)	1.18 (0.71)	(-0.23, 2.58)	0.76 (1.09)	(-1.39, 2.91)	0.486
		Irbesartan	129	50 (38.8)	0.42 (0.82)	(-1.20, 2.04)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	28 (57.1)	0.50 (1.18)	(-1.85, 2.85)	0.23 (1.73)	(-3.21, 3.66)	0.895
		Irbesartan	48	26 (54.2)	0.28 (1.24)	(-2.19, 2.74)			
	Week 48	Sparsentan	49	29 (59.2)	-2.16 (1.17)	(-4.48, 0.17)	0.21 (1.81)	(-3.39, 3.82)	0.906
		Irbesartan	48	20 (41.7)	-2.37 (1.36)	(-5.08, 0.34)			
≥ 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	15 (57.7)	1.11 (1.23)	(-1.36, 3.58)	0.49 (1.80)	(-3.14, 4.11)	0.789
		Irbesartan	25	13 (52.0)	0.62 (1.32)	(-2.03, 3.28)			
	Week 48	Sparsentan	26	16 (61.5)	0.15 (1.20)	(-2.25, 2.56)	-2.39 (1.96)	(-6.33, 1.55)	0.229
		Irbesartan	25	10 (40.0)	2.54 (1.54)	(-0.54, 5.63)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.912
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	54 (65.9)	0.34 (0.81)	(-1.27, 1.95)	0.32 (1.19)	(-2.03, 2.68)	0.787
		Irbesartan	80	48 (60.0)	0.02 (0.87)	(-1.70, 1.74)			
	Week 48	Sparsentan	82	44 (53.7)	1.55 (0.88)	(-0.19, 3.29)	1.52 (1.37)	(-1.19, 4.22)	0.270
		Irbesartan	80	30 (37.5)	0.03 (1.05)	(-2.03, 2.10)			
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	31 (68.9)	-0.15 (1.10)	(-2.35, 2.04)	0.36 (1.68)	(-2.99, 3.72)	0.829
		Irbesartan	49	25 (51.0)	-0.52 (1.25)	(-3.01, 1.98)			
	Week 48	Sparsentan	45	23 (51.1)	0.45 (1.26)	(-2.05, 2.94)	-0.58 (1.88)	(-4.32, 3.15)	0.757
		Irbesartan	49	20 (40.8)	1.03 (1.38)	(-1.70, 3.77)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	28 (57.1)	0.50 (1.18)	(-1.85, 2.85)	0.23 (1.73)	(-3.21, 3.66)	0.895
		Irbesartan	48	26 (54.2)	0.28 (1.24)	(-2.19, 2.74)			
	Week 48	Sparsentan	49	29 (59.2)	-2.16 (1.17)	(-4.48, 0.17)	0.21 (1.81)	(-3.39, 3.82)	0.906
		Irbesartan	48	20 (41.7)	-2.37 (1.36)	(-5.08, 0.34)			
≥ 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	15 (57.7)	1.11 (1.23)	(-1.36, 3.58)	0.49 (1.80)	(-3.14, 4.11)	0.789
		Irbesartan	25	13 (52.0)	0.62 (1.32)	(-2.03, 3.28)			
	Week 48	Sparsentan	26	16 (61.5)	0.15 (1.20)	(-2.25, 2.56)	-2.39 (1.96)	(-6.33, 1.55)	0.229
		Irbesartan	25	10 (40.0)	2.54 (1.54)	(-0.54, 5.63)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.629
<= 1.75 g/day	Week 24	Sparsentan	98	67 (68.4)	0.37 (0.73)	(-1.08, 1.82)	0.12 (1.16)	(-2.17, 2.41)	0.916
		Irbesartan	94	47 (50.0)	0.25 (0.89)	(-1.50, 2.00)			
	Week 48	Sparsentan	98	60 (61.2)	0.11 (0.76)	(-1.40, 1.62)	0.35 (1.27)	(-2.17, 2.86)	0.786
		Irbesartan	94	34 (36.2)	-0.23 (1.00)	(-2.21, 1.74)			
> 1.75 g/day	Week 24	Sparsentan	104	61 (58.7)	0.52 (0.76)	(-0.98, 2.02)	0.81 (1.07)	(-1.29, 2.92)	0.447
		Irbesartan	108	65 (60.2)	-0.30 (0.74)	(-1.76, 1.17)			
	Week 48	Sparsentan	104	52 (50.0)	0.36 (0.82)	(-1.25, 1.97)	0.56 (1.20)	(-1.80, 2.93)	0.640
		Irbesartan	108	46 (42.6)	-0.20 (0.86)	(-1.91, 1.50)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024



Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.478
Yes	Week 24	Sparsentan	88	53 (60.2)	1.39 (0.86)	(-0.31, 3.10)	1.19 (1.29)	(-1.35, 3.73)	0.357
		Irbesartan	83	43 (51.8)	0.21 (0.95)	(-1.68, 2.09)			
	Week 48	Sparsentan	88	43 (48.9)	1.25 (0.94)	(-0.61, 3.10)	0.22 (1.46)	(-2.67, 3.10)	0.883
		Irbesartan	83	30 (36.1)	1.03 (1.11)	(-1.17, 3.23)			
No	Week 24	Sparsentan	114	75 (65.8)	-0.20 (0.65)	(-1.49, 1.09)	-0.11 (0.95)	(-1.98, 1.76)	0.909
		Irbesartan	119	69 (58.0)	-0.09 (0.69)	(-1.45, 1.26)			
	Week 48	Sparsentan	114	69 (60.5)	-0.47 (0.67)	(-1.80, 0.85)	0.36 (1.04)	(-1.68, 2.40)	0.730
		Irbesartan	119	50 (42.0)	-0.83 (0.78)	(-2.38, 0.71)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.466
<= 5 years	Week 24	Sparsentan	113	73 (64.6)	1.42 (0.71)	(0.02, 2.83)	1.51 (1.01)	(-0.47, 3.50)	0.135
		Irbesartan	127	73 (57.5)	-0.09 (0.71)	(-1.49, 1.32)			
	Week 48	Sparsentan	113	61 (54.0)	0.34 (0.76)	(-1.16, 1.85)	-0.41 (1.11)	(-2.60, 1.79)	0.715
		Irbesartan	127	53 (41.7)	0.75 (0.81)	(-0.85, 2.35)			
> 5 years	Week 24	Sparsentan	89	55 (61.8)	-1.13 (0.77)	(-2.65, 0.38)	-1.66 (1.20)	(-4.04, 0.71)	0.168
		Irbesartan	75	39 (52.0)	0.53 (0.93)	(-1.31, 2.36)			
	Week 48	Sparsentan	89	51 (57.3)	-0.24 (0.79)	(-1.80, 1.33)	1.29 (1.34)	(-1.35, 3.93)	0.335
		Irbesartan	75	27 (36.0)	-1.53 (1.07)	(-3.64, 0.59)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.772
Yes	Week 24	Sparsentan	153	97 (63.4)	0.70 (0.61)	(-0.49, 1.89)	0.77 (0.89)	(-0.99, 2.53)	0.389
		Irbesartan	157	83 (52.9)	-0.07 (0.66)	(-1.37, 1.23)			
	Week 48	Sparsentan	153	83 (54.2)	0.24 (0.64)	(-1.03, 1.50)	-0.31 (0.99)	(-2.27, 1.64)	0.754
		Irbesartan	157	59 (37.6)	0.55 (0.76)	(-0.94, 2.04)			
No	Week 24	Sparsentan	49	31 (63.3)	-0.71 (1.02)	(-2.73, 1.31)	-1.33 (1.48)	(-4.26, 1.60)	0.371
		Irbesartan	45	29 (64.4)	0.62 (1.06)	(-1.49, 2.73)			
	Week 48	Sparsentan	49	29 (59.2)	-0.22 (1.05)	(-2.30, 1.85)	1.65 (1.61)	(-1.55, 4.85)	0.309
		Irbesartan	45	21 (46.7)	-1.87 (1.22)	(-4.30, 0.55)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 04APR2024

Figure PF1KPSC\_FSGM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

Not done: No significant subgroup found.

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KPSC\_FSCM.

Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.439
Male	Sparsentan	139	14 (10.1)	NE		1.072	(0.489, 2.351)	0.862
	Irbesartan	143	14 (9.8)	NE				
Female	Sparsentan	63	8 (12.7)	NE		1.109	(0.075, 16.413)	0.940
	Irbesartan	59	1 (1.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.488
<= 45 years	Sparsentan	96	8 (8.3)	NE		0.768	(0.243, 2.426)	0.653
	Irbesartan	99	6 (6.1)	NE				
> 45 years	Sparsentan	106	14 (13.2)	NE		1.109	(0.441, 2.784)	0.826
	Irbesartan	103	9 (8.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	1 (11.1) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	10 (9.8)	NE		1.282	(0.485, 3.384)	0.617
	Irbesartan	109	8 (7.3)	NE				
> 40 years	Sparsentan	91	11 (12.1)	NE		1.126	(0.376, 3.371)	0.832
	Irbesartan	88	7 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan							Interaction test: <0.001 #
North America	Sparsentan	35	1 (2.9)	NE		0.093	(0.007, 1.237)	0.072
	Irbesartan	46	6 (13.0)	NE				
Europe	Sparsentan	98	7 (7.1)	NE		0.436	(0.080, 2.362)	0.336
	Irbesartan	115	5 (4.3)	NE				
Asia Pacific	Sparsentan	69	14 (20.3)	NE		1.939	(0.607, 6.199)	0.264
	Irbesartan	41	4 (9.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.956
< 27 kg/m**2	Sparsentan	84	10 (11.9)	NE		1.202	(0.357, 4.054)	0.767
	Irbesartan	94	5 (5.3)	NE				
≥ 27 kg/m**2	Sparsentan	118	12 (10.2)	NE		1.150	(0.466, 2.839)	0.761
	Irbesartan	107	10 (9.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.012 #
eGFR Low and UP High	Sparsentan	71	6 (8.5)	NE		1.140	(0.355, 3.662)	0.826
	Irbesartan	74	6 (8.1)	NE				
eGFR Low and UP Low	Sparsentan	55	7 (12.7)	NE		0.875	(0.175, 4.389)	0.871
	Irbesartan	55	3 (5.5)	NE				
eGFR High and UP High	Sparsentan	37	3 (8.1)	NE		0.261	(0.031, 2.183)	0.215
	Irbesartan	36	3 (8.3)	NE				
eGFR High and UP Low	Sparsentan	39	6 (15.4)	NE		3.957	(0.642, 24.379)	0.138
	Irbesartan	37	3 (8.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.897
< 60 mL/min/1.73 m**2	Sparsentan	127	14 (11.0)	NE		1.126	(0.466, 2.723)	0.792
	Irbesartan	129	11 (8.5)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	4 (8.2)	NE		0.972	(0.101, 9.341)	0.981
	Irbesartan	48	3 (6.3)	NE				
>= 90 mL/min/1.73 m**2	Sparsentan	26	4 (15.4)	NE		1.447	(0.129, 16.254)	0.765
	Irbesartan	25	1 (4.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.971
< 45 mL/min/1.73 m**2	Sparsentan	82	8 (9.8)	NE		0.865	(0.254, 2.947)	0.816
	Irbesartan	80	6 (7.5)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	6 (13.3)	NE		1.250	(0.319, 4.896)	0.748
	Irbesartan	49	5 (10.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	4 (8.2)	NE		0.972	(0.101, 9.341)	0.981
	Irbesartan	48	3 (6.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	4 (15.4)	NE		1.447	(0.129, 16.254)	0.765
	Irbesartan	25	1 (4.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.063
<= 1.75 g/day	Sparsentan	98	14 (14.3)	NE		2.448	(0.769, 7.790)	0.130
	Irbesartan	94	4 (4.3)	NE				
> 1.75 g/day	Sparsentan	104	8 (7.7)	NE		0.453	(0.158, 1.302)	0.142
	Irbesartan	108	11 (10.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.828
Yes	Sparsentan	88	10 (11.4)	NE		0.792	(0.298, 2.106)	0.640
	Irbesartan	83	10 (12.0)	NE				
No	Sparsentan	114	12 (10.5)	NE		1.809	(0.612, 5.351)	0.284
	Irbesartan	119	5 (4.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.242
<= 5 years	Sparsentan	113	15 (13.3)	NE		1.484	(0.659, 3.341)	0.340
	Irbesartan	127	11 (8.7)	NE				
> 5 years	Sparsentan	89	7 (7.9)	NE		0.939	(0.225, 3.917)	0.931
	Irbesartan	75	4 (5.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

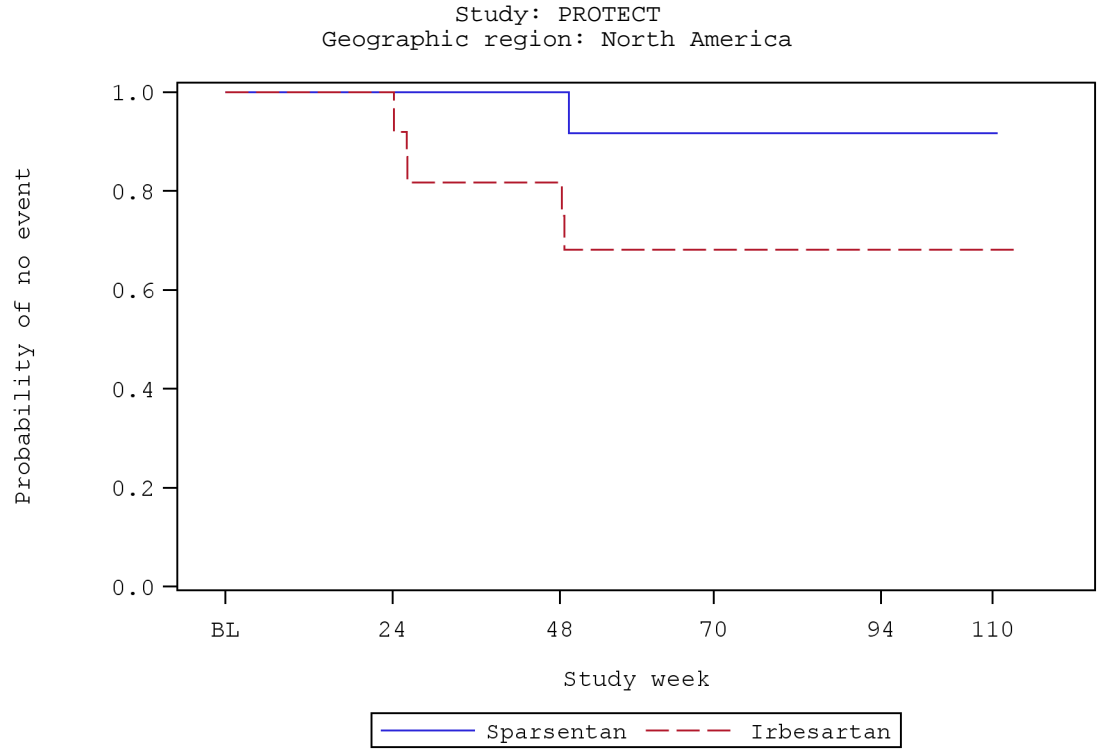
Table PT1KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.040 #
Yes	Sparsentan	153	16 (10.5)	NE		0.705	(0.312, 1.595)	0.402
	Irbesartan	157	12 (7.6)	NE				
No	Sparsentan	49	6 (12.2)	NE		4.395	(0.738, 26.182)	0.104
	Irbesartan	45	3 (6.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



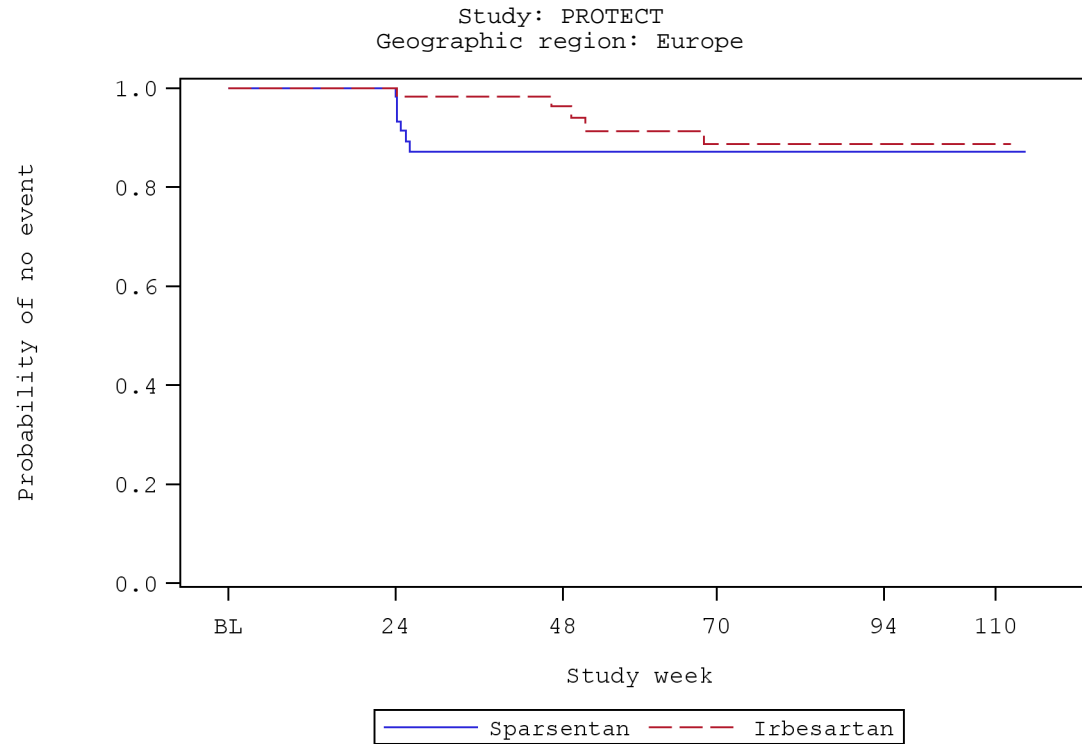
Figure PF1KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	35	18	13	9	6	1
Irbesartan	46	26	16	8	3	2

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KPSIT\_FSTM

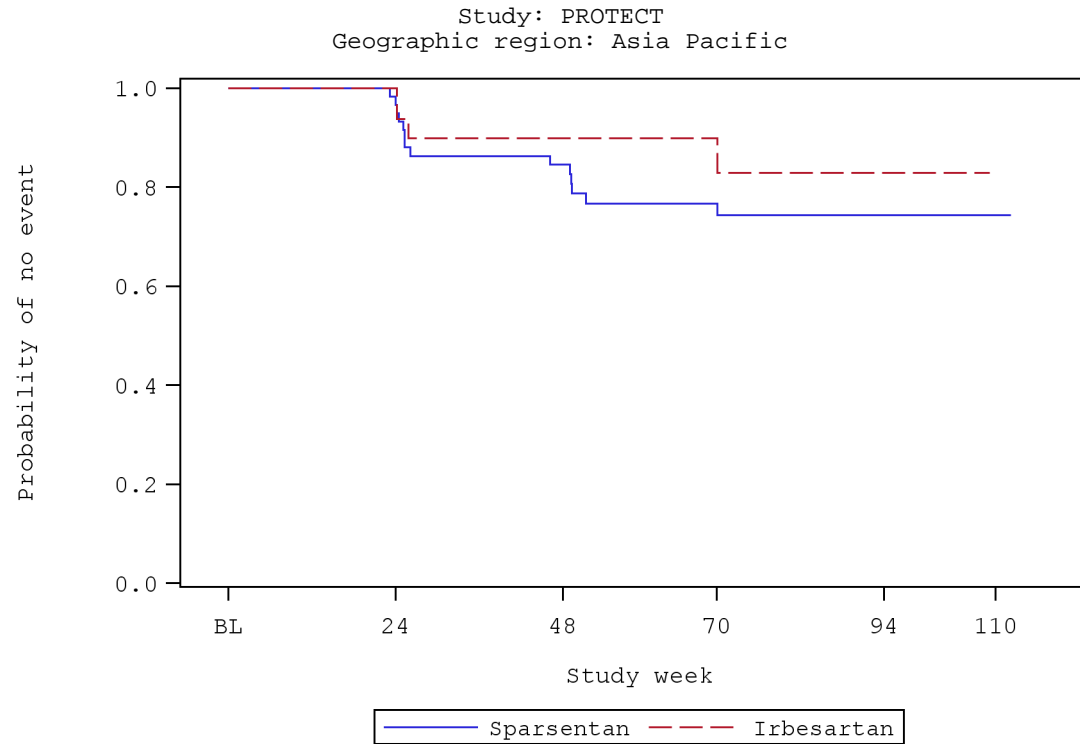
Figure PF1KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	98	59	40	27	13	6
Irbesartan	115	61	46	30	12	3

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KPSIT\_FSTM

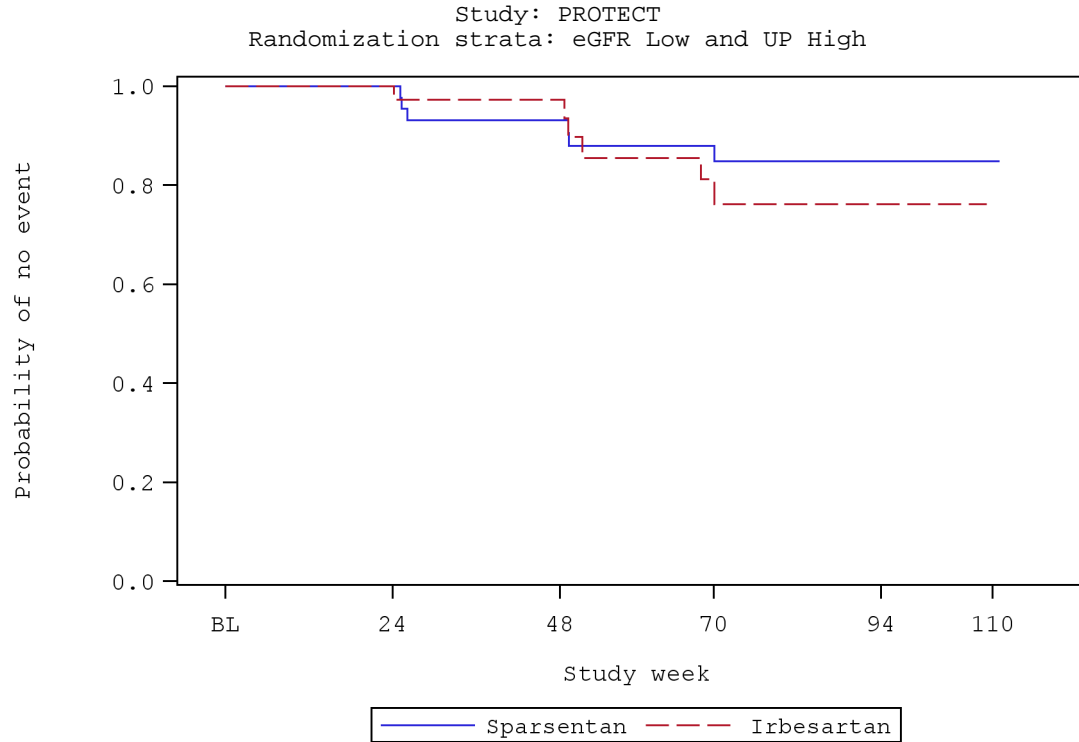
Figure PF1KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	69	59	48	36	13	8
Irbesartan	41	32	21	13	2	0

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KPSIT\_FSTM

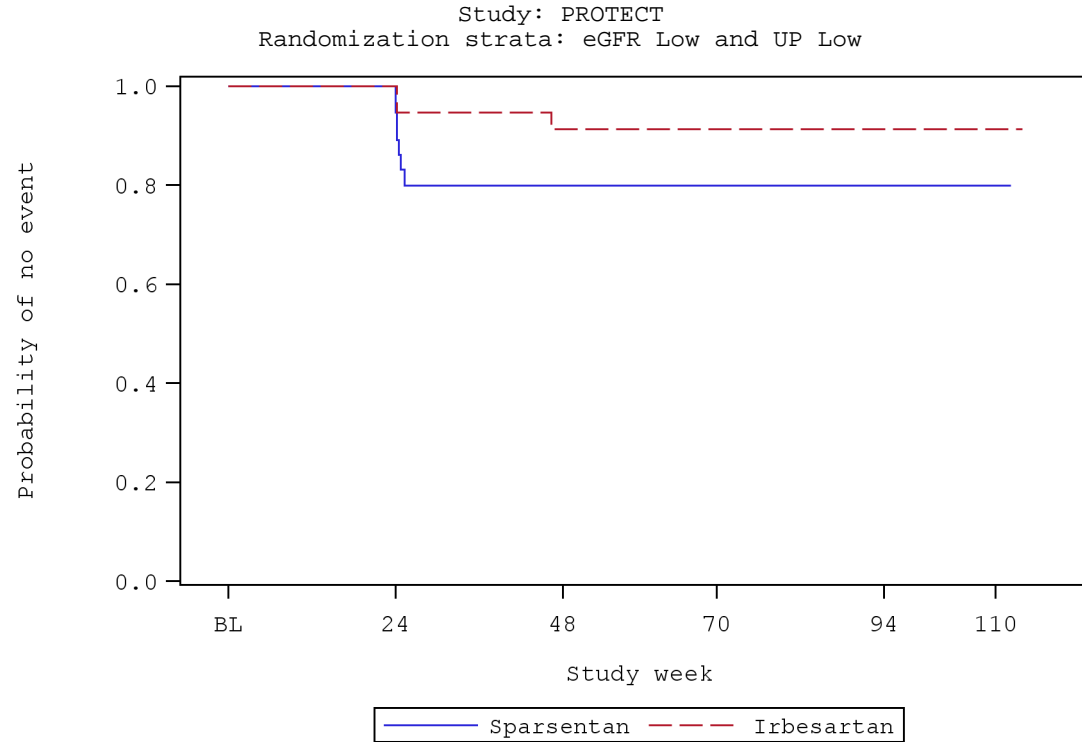
Figure PF1KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	71	48	39	30	15	5
Irbesartan	74	37	29	17	6	0

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KPSIT\_FSTM

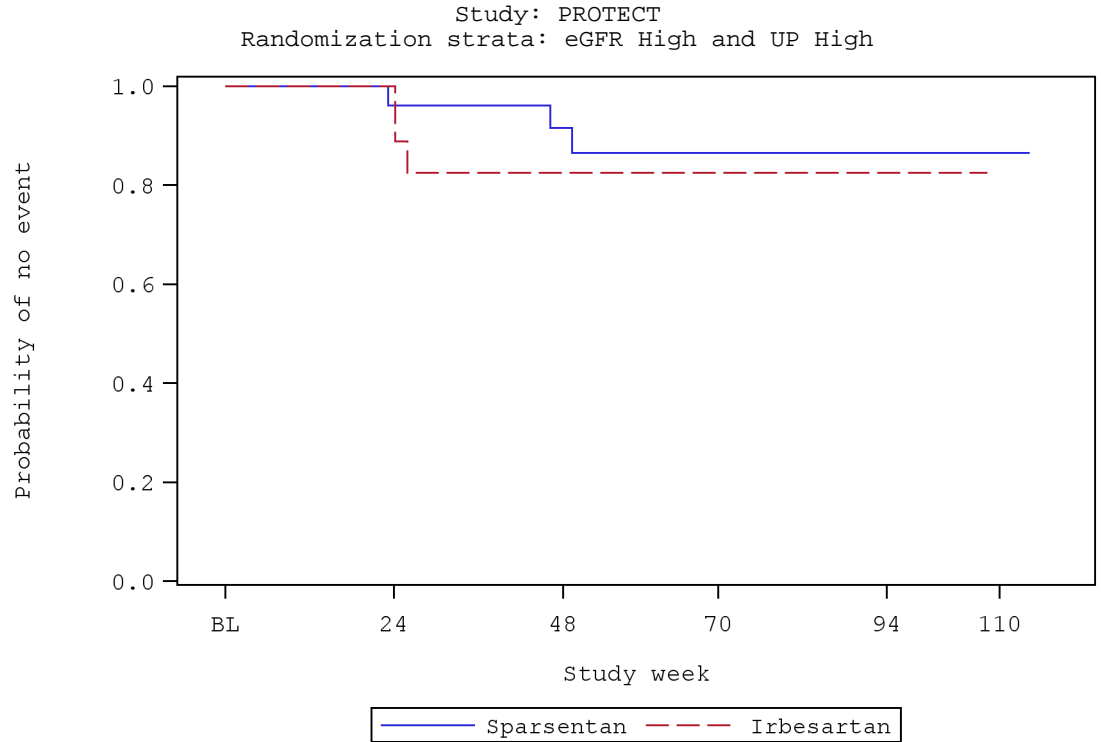
Figure PF1KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	55	38	23	15	8	6
Irbesartan	55	38	27	18	8	4

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KPSIT\_FSTM

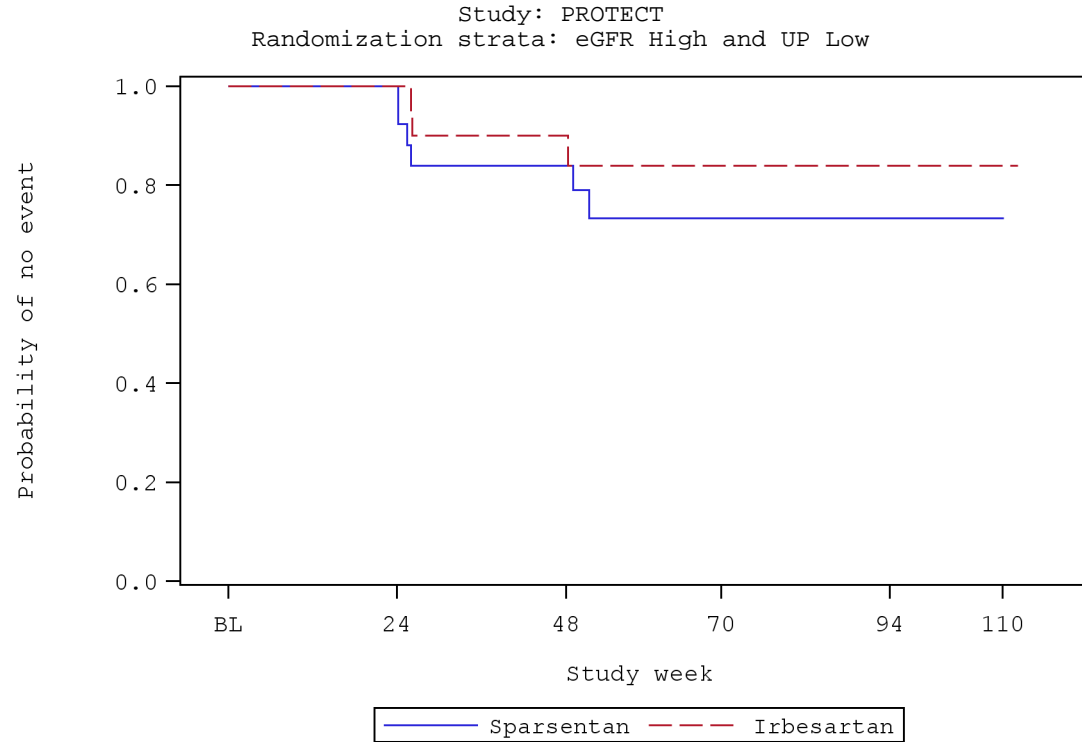
Figure PF1KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	37	24	20	14	6	3
Irbesartan	36	20	11	5	1	0

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KPSIT\_FSTM

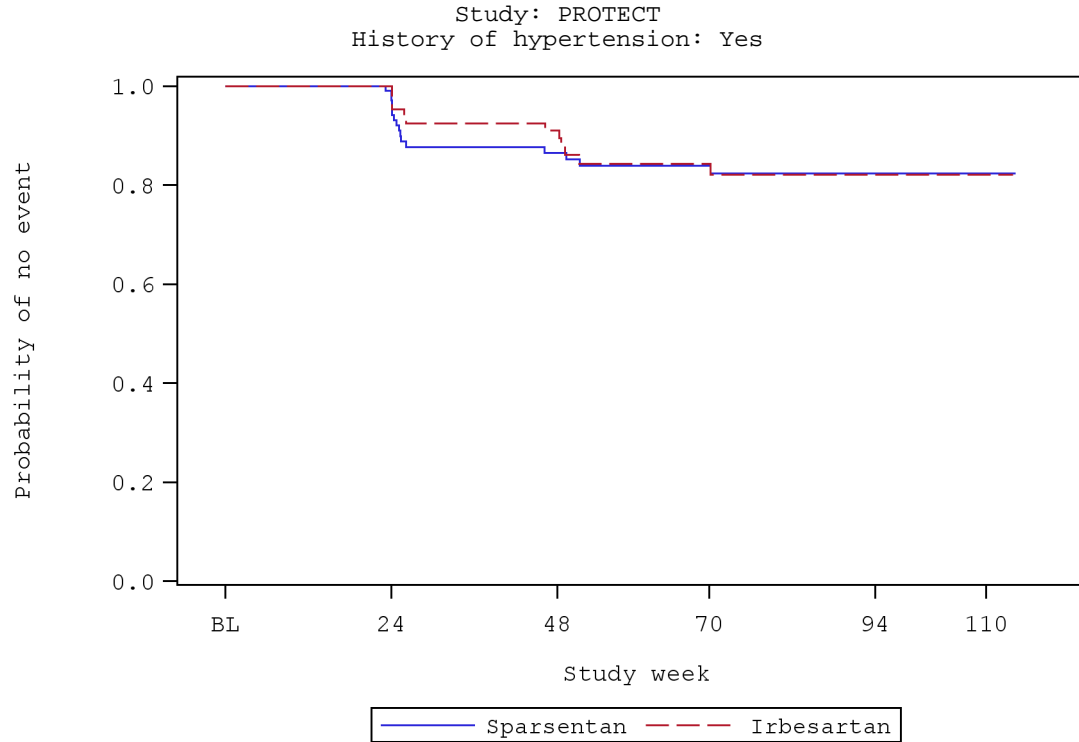
Figure PF1KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	39	26	19	13	3	1
Irbesartan	37	24	16	11	2	1

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KPSIT\_FSTM

Figure PF1KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

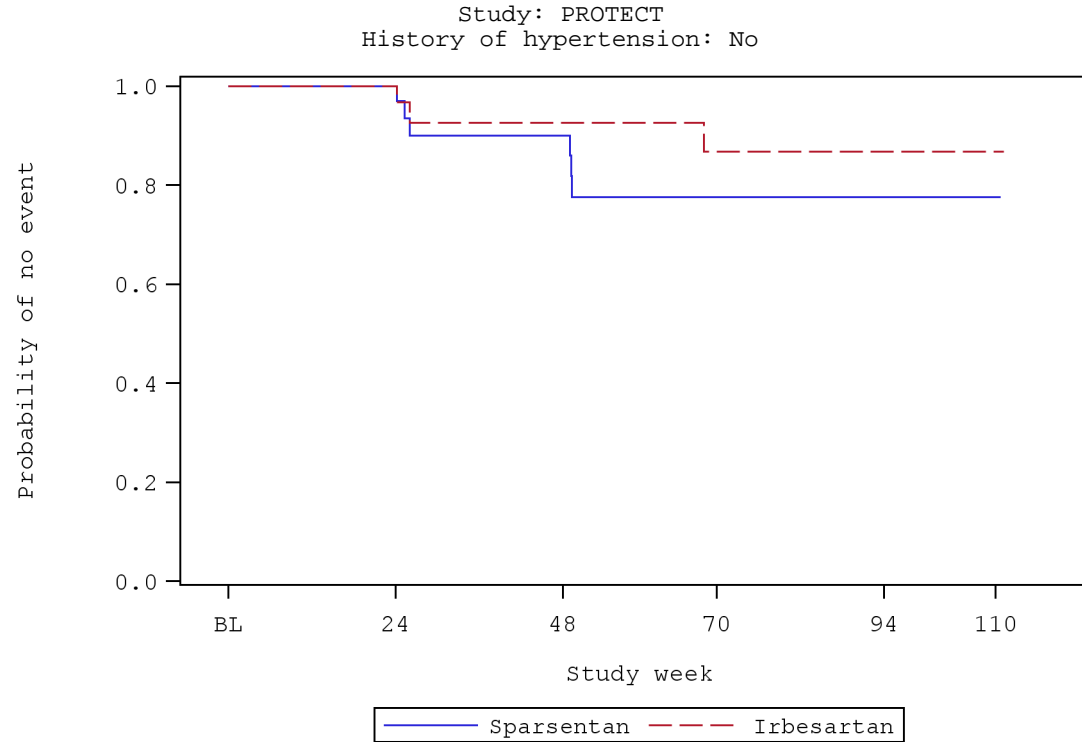


Sparsentan	153	102	75	56	26	12
Irbesartan	157	87	62	39	13	4

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KPSIT\_FSTM



Figure PF1KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	49	34	26	16	6	3
Irbesartan	45	32	21	12	4	1

An increase reflects an improvement of the status of the patient.  
 Reference table: PT1KPSIT\_FSTM

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.795
Male	Sparsentan	139	13 (9.4)	NE		0.870	(0.391, 1.939)	0.734
	Irbesartan	143	12 (8.4)	NE				
Female	Sparsentan	63	12 (19.0)	NE		0.948	(0.323, 2.776)	0.922
	Irbesartan	59	6 (10.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.325
<= 45 years	Sparsentan	96	12 (12.5)	NE		1.424	(0.549, 3.689)	0.467
	Irbesartan	99	7 (7.1)	NE				
> 45 years	Sparsentan	106	13 (12.3)	NE		0.823	(0.365, 1.856)	0.638
	Irbesartan	103	11 (10.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	15 (14.7)	NE		1.502	(0.635, 3.553)	0.354
	Irbesartan	109	9 (8.3)	NE				
> 40 years	Sparsentan	91	10 (11.0)	NE		0.832	(0.335, 2.064)	0.692
	Irbesartan	88	9 (10.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.517
North America	Sparsentan	35	3 (8.6)	NE		0.629	(0.130, 3.034)	0.563
	Irbesartan	46	5 (10.9)	NE				
Europe	Sparsentan	98	9 (9.2)	NE		0.817	(0.325, 2.050)	0.666
	Irbesartan	115	10 (8.7)	NE				
Asia Pacific	Sparsentan	69	13 (18.8)	NE		1.817	(0.507, 6.510)	0.359
	Irbesartan	41	3 (7.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.863
< 27 kg/m**2	Sparsentan	84	8 (9.5)	NE		0.849	(0.290, 2.485)	0.766
	Irbesartan	94	7 (7.4)	NE				
≥ 27 kg/m**2	Sparsentan	118	17 (14.4)	NE		1.130	(0.525, 2.433)	0.754
	Irbesartan	107	11 (10.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.035 #
eGFR Low and UP High	Sparsentan	71	10 (14.1)	NE		1.004	(0.358, 2.817)	0.994
	Irbesartan	74	6 (8.1)	NE				
eGFR Low and UP Low	Sparsentan	55	1 (1.8)	NE		0.137	(0.017, 1.109)	0.062
	Irbesartan	55	8 (14.5)	NE				
eGFR High and UP High	Sparsentan	37	7 (18.9)	NE		1.585	(0.314, 8.017)	0.577
	Irbesartan	36	2 (5.6)	NE				
eGFR High and UP Low	Sparsentan	39	7 (17.9)	NE		3.027	(0.602, 15.224)	0.179
	Irbesartan	37	2 (5.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: NE
< 60 mL/min/1.73 m**2	Sparsentan	127	12 (9.4)	NE		0.784	(0.351, 1.751)	0.553
	Irbesartan	129	13 (10.1)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	8 (16.3)	NE		0.873	(0.271, 2.812)	0.820
	Irbesartan	48	5 (10.4)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	5 (19.2) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	NE
< 45 mL/min/1.73 m**2	Sparsentan	82	6 (7.3)	NE		0.445	(0.158, 1.255)	0.126
	Irbesartan	80	10 (12.5)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	6 (13.3)	NE		2.363	(0.553, 10.107)	0.246
	Irbesartan	49	3 (6.1)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	8 (16.3)	NE		0.873	(0.271, 2.812)	0.820
	Irbesartan	48	5 (10.4)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	5 (19.2)	NE		NE		NE
	Irbesartan	25	0 (0.0) No events in 1 group	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.615
<= 1.75 g/day	Sparsentan	98	12 (12.2)	NE		1.157	(0.435, 3.080)	0.770
	Irbesartan	94	7 (7.4)	NE				
> 1.75 g/day	Sparsentan	104	13 (12.5)	NE		0.998	(0.418, 2.382)	0.996
	Irbesartan	108	11 (10.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.649
Yes	Sparsentan	88	10 (11.4)	NE		0.878	(0.337, 2.284)	0.790
	Irbesartan	83	8 (9.6)	NE				
No	Sparsentan	114	15 (13.2)	NE		1.139	(0.508, 2.552)	0.753
	Irbesartan	119	10 (8.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.795
<= 5 years	Sparsentan	113	12 (10.6)	NE		1.075	(0.462, 2.503)	0.867
	Irbesartan	127	10 (7.9)	NE				
> 5 years	Sparsentan	89	13 (14.6)	NE		0.857	(0.335, 2.192)	0.748
	Irbesartan	75	8 (10.7)	NE				

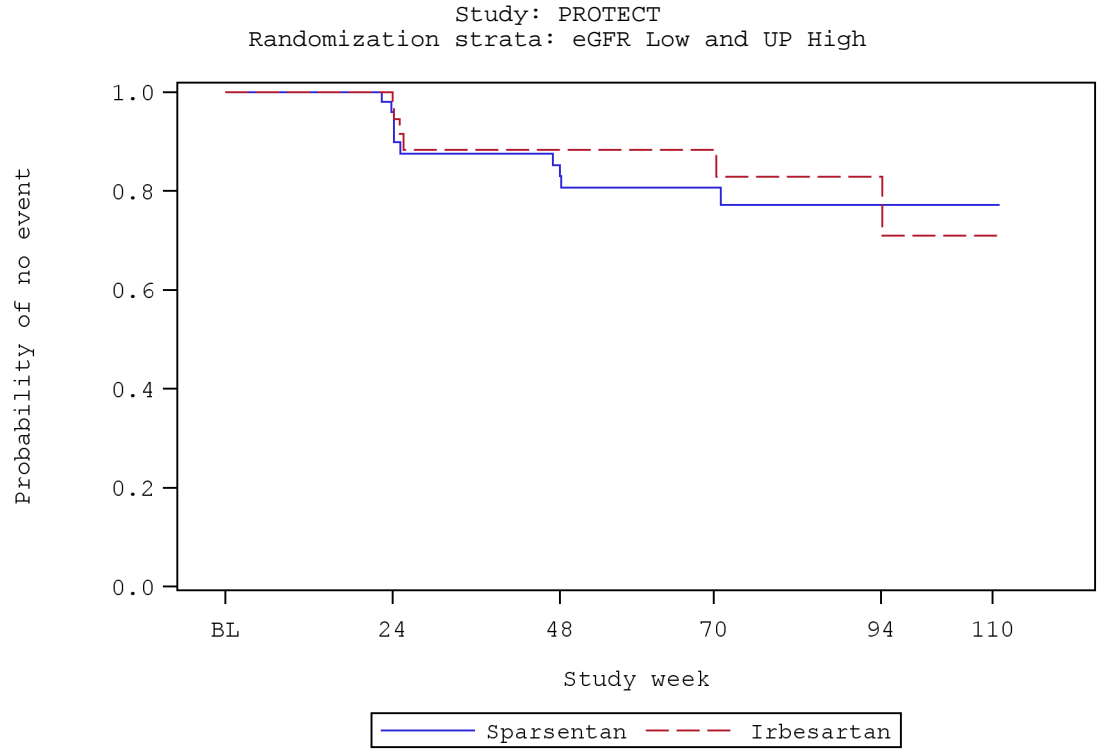
N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan						Interaction test:	0.583
Yes	Sparsentan	153	19 (12.4)	NE		0.968	(0.487, 1.921)	0.925
	Irbesartan	157	15 (9.6)	NE				
No	Sparsentan	49	6 (12.2)	NE		1.584	(0.373, 6.733)	0.533
	Irbesartan	45	3 (6.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

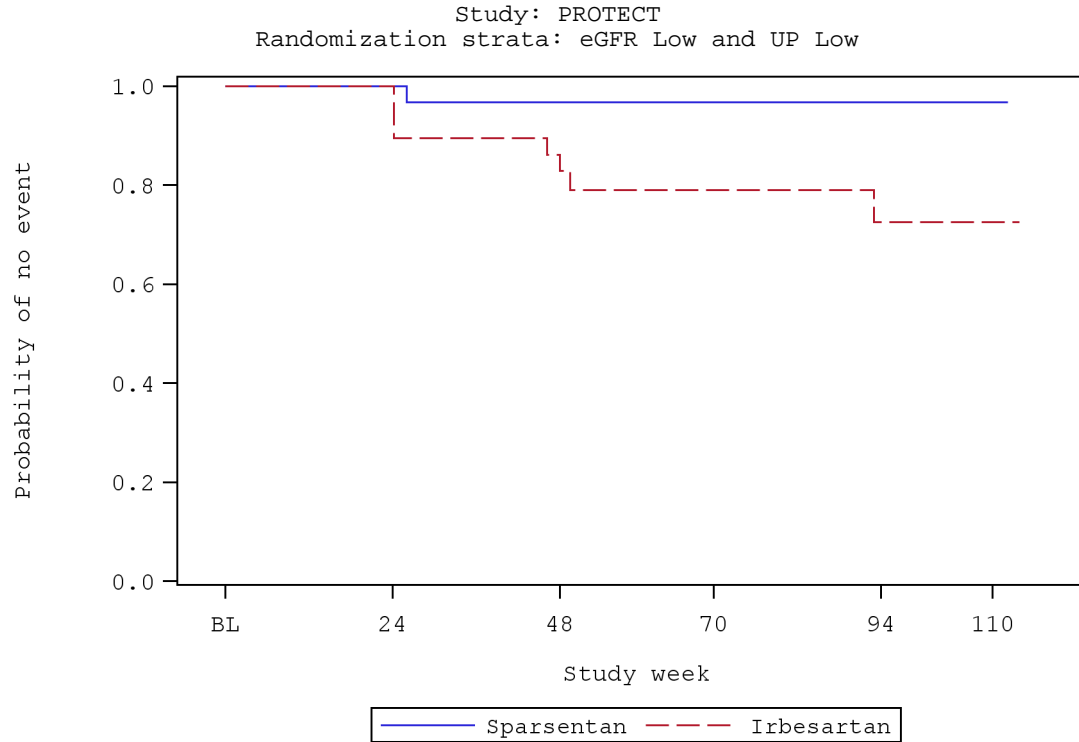
Figure PF1KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	71	47	37	29	15	2
Irbesartan	74	37	26	18	7	2

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KPSDT\_FSTM

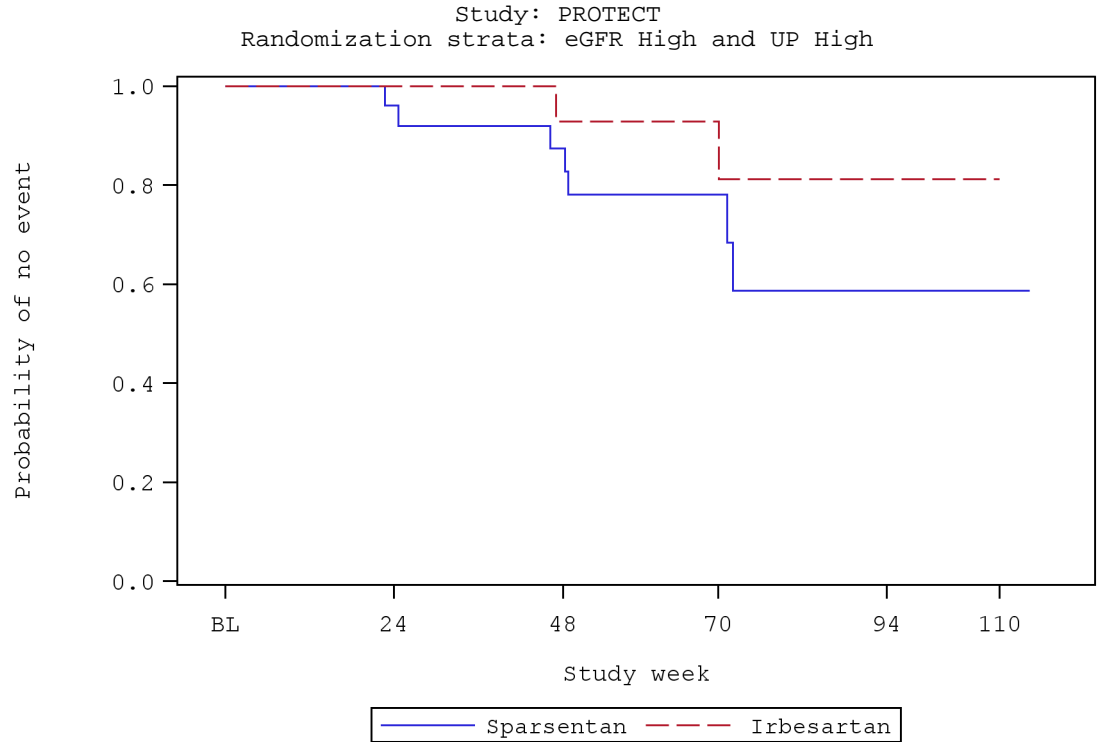
Figure PF1KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	55	38	29	20	11	6
Irbesartan	55	38	26	17	9	4

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KPSDT\_FSTM

Figure PF1KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

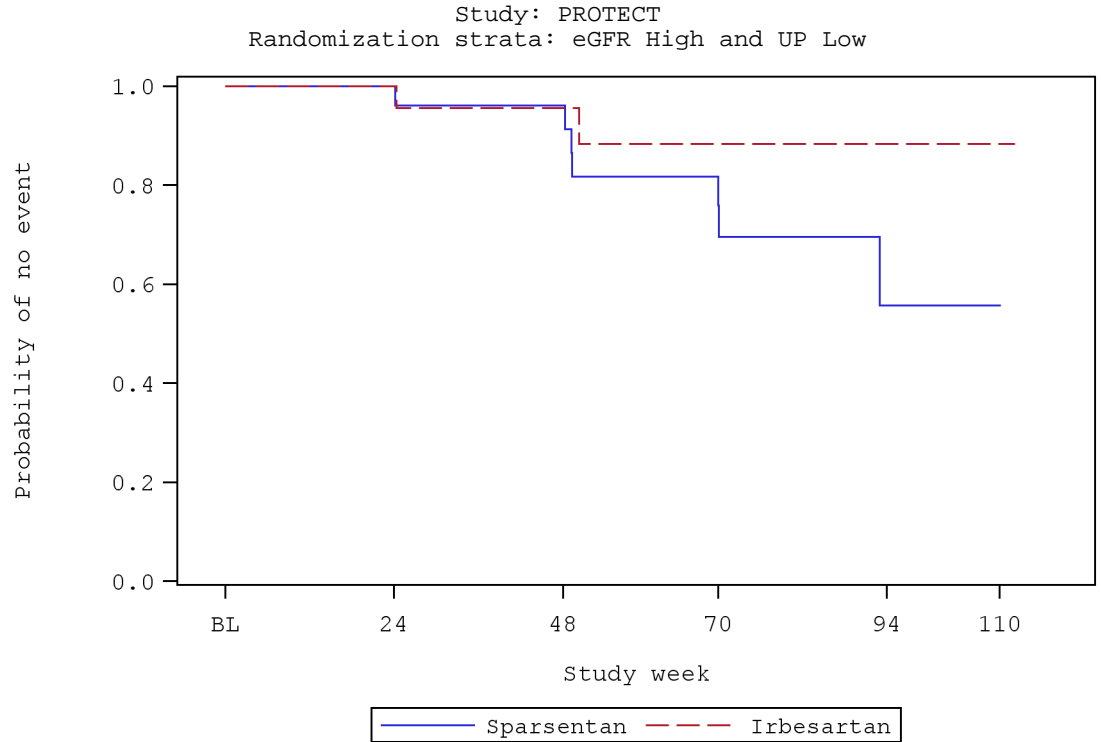


Sparsentan	37	24	19	13	3	1
Irbesartan	36	20	13	8	3	1

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KPSDT\_FSTM



Figure PF1KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	39	26	22	14	3	1
Irbesartan	37	24	17	11	3	2

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT1KPSDT\_FSTM

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Sex												
Male	KDQOL-SF12: MCS	Baseline	Sparsentan	139	130 (93.5)	50.45 (7.97)	30.1	45.01	51.54	57.06	66.5	
			Irbesartan	143	132 (92.3)	51.61 (8.91)	24.3	48.28	53.36	57.16	66.6	
		Week 24	Sparsentan	139	88 (63.3)	51.32 (7.78)	20.8	47.69	51.92	57.16	66.8	
			Irbesartan	143	81 (56.6)	52.38 (8.56)	24.2	48.43	55.77	58.15	64.2	
		Week 48	Sparsentan	139	74 (53.2)	50.78 (7.92)	27.5	46.89	52.13	57.16	62.5	
			Irbesartan	143	58 (40.6)	52.97 (8.50)	28.5	49.58	54.88	58.58	64.6	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	139	88 (63.3)	0.17 (7.74)	-19.1	-4.42	-0.05	4.50	25.3	-0.01 [-0.31, 0.29]
			Irbesartan	143	81 (56.6)	0.25 (7.03)	-15.4	-4.36	0.00	2.98	28.7	
		Week 48	Sparsentan	139	74 (53.2)	0.86 (7.78)	-18.6	-3.85	0.69	6.32	18.7	0.06 [-0.28, 0.41]
			Irbesartan	143	58 (40.6)	0.39 (6.92)	-17.3	-2.88	0.00	4.92	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Female	KDQOL-SF12: MCS	Baseline	Sparsentan	63	57 (90.5)	50.24 (9.11)	16.2	44.79	51.77	56.05	65.2	
			Irbesartan	59	52 (88.1)	50.78 (7.91)	31.5	45.73	52.81	57.16	61.3	
		Week 24	Sparsentan	63	40 (63.5)	52.17 (7.51)	30.7	47.38	53.24	58.30	63.5	
			Irbesartan	59	31 (52.5)	53.55 (7.70)	32.5	51.19	54.45	57.76	62.6	
		Week 48	Sparsentan	63	38 (60.3)	50.61 (9.24)	27.5	42.67	51.71	59.46	63.4	
			Irbesartan	59	22 (37.3)	46.84 (12.07)	19.6	39.39	50.08	56.11	64.5	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	63	40 (63.5)	1.96 (9.36)	-23.6	-1.63	0.99	7.80	26.4	0.08 [-0.38, 0.55]
			Irbesartan	59	31 (52.5)	1.27 (6.09)	-9.4	-2.96	1.41	3.95	20.2	
		Week 48	Sparsentan	63	38 (60.3)	0.19 (11.16)	-23.7	-4.74	-0.10	4.70	28.1	0.51 [-0.02, 1.04]
			Irbesartan	59	22 (37.3)	-5.38 (10.48)	-33.9	-13.37	-2.99	2.79	6.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age												
<= 45 years	KDQOL-SF12: MCS	Baseline	Sparsentan	96	88 (91.7)	48.94 (8.49)	27.8	43.44	50.17	55.34	65.2	
			Irbesartan	99	92 (92.9)	50.34 (9.59)	24.3	44.08	52.45	57.16	64.2	
		Week 24	Sparsentan	96	59 (61.5)	50.89 (8.33)	20.8	46.59	52.22	57.24	63.5	
			Irbesartan	99	56 (56.6)	51.80 (9.33)	24.2	48.37	54.41	58.02	62.7	
		Week 48	Sparsentan	96	53 (55.2)	49.15 (8.58)	27.5	42.67	50.84	56.29	62.5	
			Irbesartan	99	40 (40.4)	51.51 (10.15)	21.5	47.22	54.50	58.35	64.6	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	96	59 (61.5)	0.14 (8.74)	-23.6	-5.04	0.00	5.32	26.4	-0.07 [-0.44, 0.30]
			Irbesartan	99	56 (56.6)	0.69 (6.96)	-14.1	-3.74	0.68	5.01	21.7	
		Week 48	Sparsentan	96	53 (55.2)	-0.27 (9.72)	-23.7	-4.61	0.00	4.70	21.1	0.10 [-0.31, 0.51]
			Irbesartan	99	40 (40.4)	-1.12 (7.66)	-23.5	-4.02	0.00	3.28	13.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 45 years	KDQOL-SF12: MCS	Baseline	Sparsentan	106	99 (93.4)	51.67 (7.97)	16.2	48.28	53.37	57.16	66.5	
			Irbesartan	103	92 (89.3)	52.41 (7.45)	26.4	49.05	53.29	57.16	66.6	
	Week 24	Sparsentan	106	69 (65.1)	52.18 (7.09)	34.3	48.18	53.96	57.33	66.8		
		Irbesartan	103	56 (54.4)	53.61 (7.13)	30.8	50.56	56.34	57.87	64.2		
	Week 48	Sparsentan	106	59 (55.7)	52.14 (7.95)	29.5	48.76	53.51	58.32	63.4		
		Irbesartan	103	40 (38.8)	51.06 (9.81)	19.6	47.43	52.82	57.37	64.5		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	106	69 (65.1)	1.23 (7.90)	-18.1	-2.86	0.61	4.20	25.9	0.12 [-0.24, 0.47]
			Irbesartan	103	56 (54.4)	0.38 (6.64)	-15.4	-4.23	0.00	3.27	28.7	
		Week 48	Sparsentan	106	59 (55.7)	1.45 (8.35)	-16.9	-3.83	1.89	6.43	28.1	0.31 [-0.09, 0.72]
			Irbesartan	103	40 (38.8)	-1.28 (9.17)	-33.9	-3.52	-0.62	4.90	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age at IgAN diagnosis												
<= 18 years	KDQOL-SF12: MCS	Baseline	Sparsentan	9	9 (100.0)	44.94 (11.76)	31.1	35.44	42.40	51.24	62.5	
			Irbesartan	5	5 (100.0)	50.09 (9.56)	34.8	46.63	54.94	56.91	57.2	
		Week 24	Sparsentan	9	3 (33.3)	37.69 (15.49)	20.8	20.79	41.07	51.21	51.2	
			Irbesartan	5	2 (40.0)	40.06 (14.65)	29.7	29.71	40.06	50.42	50.4	
		Week 48	Sparsentan	9	5 (55.6)	47.97 (7.75)	37.2	43.46	50.84	51.24	57.2	
	Irbesartan		5	1 (20.0)	49.58	49.6	49.58	49.58	49.58	49.6		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	9	3 (33.3)	-13.54 (4.80)	-19.1	-19.05	-11.28	-10.30	-10.3	-2.23 [-4.49, 0.03]
			Irbesartan	5	2 (40.0)	-4.81 (0.41)	-5.1	-5.10	-4.81	-4.51	-4.5	
		Week 48	Sparsentan	9	5 (55.6)	1.85 (8.68)	-11.3	0.45	1.73	5.92	12.4	NE
			Irbesartan	5	1 (20.0)	-5.36	-5.4	-5.36	-5.36	-5.36	-5.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 18 to 40 years	KDQOL-SF12: MCS	Baseline	Sparsentan	102	94 (92.2)	50.15 (7.60)	27.8	44.79	51.32	55.70	65.2	
			Irbesartan	109	100 (91.7)	50.61 (9.27)	24.3	45.26	52.05	57.16	64.2	
		Week 24	Sparsentan	102	64 (62.7)	51.50 (7.21)	30.7	46.85	52.83	57.20	63.5	
			Irbesartan	109	61 (56.0)	52.35 (8.62)	24.2	48.87	55.93	57.76	62.7	
		Week 48	Sparsentan	102	57 (55.9)	49.84 (8.46)	27.5	42.75	51.44	56.52	62.5	
			Irbesartan	109	45 (41.3)	51.59 (9.74)	21.5	48.28	54.79	58.12	62.5	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	102	64 (62.7)	0.57 (7.78)	-23.6	-3.61	0.00	4.82	26.4	-0.05 [-0.40, 0.30]
			Irbesartan	109	61 (56.0)	0.96 (7.20)	-14.1	-2.96	0.17	5.23	21.7	
		Week 48	Sparsentan	102	57 (55.9)	-0.15 (9.51)	-23.7	-4.61	-0.17	4.70	21.1	0.10 [-0.29, 0.49]
			Irbesartan	109	45 (41.3)	-1.00 (7.60)	-23.5	-2.96	0.00	2.79	13.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	KDQOL-SF12: MCS	Baseline	Sparsentan	91	84 (92.3)	51.24 (8.52)	16.2	46.28	53.18	57.16	66.5	
			Irbesartan	88	79 (89.8)	52.42 (7.67)	26.4	48.29	54.18	57.57	66.6	
	Week 24	Sparsentan	91	61 (67.0)	52.35 (7.21)	34.3	48.87	53.96	57.33	66.8		
		Irbesartan	88	49 (55.7)	53.67 (7.40)	30.8	51.06	55.77	58.15	64.2		
	Week 48	Sparsentan	91	50 (54.9)	52.01 (8.25)	29.5	48.76	53.85	58.32	63.4		
		Irbesartan	88	34 (38.6)	50.93 (10.43)	19.6	47.27	52.82	57.33	64.6		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	91	61 (67.0)	1.59 (8.36)	-18.1	-2.79	0.99	5.08	25.9	0.18 [-0.20, 0.56]
			Irbesartan	88	49 (55.7)	0.23 (6.31)	-15.4	-3.28	0.00	2.95	28.7	
		Week 48	Sparsentan	91	50 (54.9)	1.41 (8.57)	-16.9	-3.83	1.97	6.43	28.1	0.31 [-0.13, 0.74]
			Irbesartan	88	34 (38.6)	-1.34 (9.54)	-33.9	-3.95	-1.56	4.87	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Geographic region												
North America	KDQOL-SF12: MCS	Baseline	Sparsentan	35	32 (91.4)	49.84 (8.24)	32.9	44.11	51.11	56.51	62.6	
			Irbesartan	46	44 (95.7)	53.62 (7.01)	35.7	47.55	56.83	58.01	64.2	
		Week 24	Sparsentan	35	15 (42.9)	51.91 (4.62)	41.1	49.25	53.07	54.96	57.7	
			Irbesartan	46	28 (60.9)	54.59 (7.99)	24.2	54.26	57.20	58.43	63.2	
		Week 48	Sparsentan	35	13 (37.1)	50.34 (10.27)	30.9	41.17	50.84	57.06	63.4	
			Irbesartan	46	17 (37.0)	50.31 (12.30)	19.6	49.41	54.79	57.76	62.4	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	35	15 (42.9)	-0.36 (7.57)	-19.1	-2.86	-0.10	3.27	14.0	0.02 [-0.61, 0.64]
			Irbesartan	46	28 (60.9)	-0.46 (5.67)	-14.1	-3.12	0.28	2.41	13.0	
		Week 48	Sparsentan	35	13 (37.1)	1.87 (8.89)	-23.4	-0.03	3.95	6.32	14.7	0.46 [-0.27, 1.19]
			Irbesartan	46	17 (37.0)	-2.82 (11.20)	-33.9	-6.62	0.00	3.95	12.1	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Europe	KDQOL-SF12: MCS	Baseline	Sparsentan	98	88 (89.8)	50.56 (9.27)	16.2	45.11	51.83	57.16	66.5	
			Irbesartan	115	99 (86.1)	50.45 (9.15)	26.4	46.56	52.31	57.16	64.6	
		Week 24	Sparsentan	98	56 (57.1)	50.56 (8.23)	20.8	46.84	49.75	56.39	63.5	
			Irbesartan	115	50 (43.5)	52.32 (8.17)	29.7	47.84	54.29	58.29	62.7	
		Week 48	Sparsentan	98	44 (44.9)	49.65 (8.10)	29.5	42.65	51.24	56.43	62.5	
			Irbesartan	115	40 (34.8)	51.29 (9.06)	31.4	46.87	53.91	57.37	64.5	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	98	56 (57.1)	-0.39 (7.43)	-17.0	-4.93	0.00	3.25	25.9	-0.21 [-0.59, 0.17]
			Irbesartan	115	50 (43.5)	1.26 (8.15)	-15.4	-4.54	0.00	5.33	28.7	
		Week 48	Sparsentan	98	44 (44.9)	-0.79 (9.42)	-20.3	-5.13	-1.52	4.09	28.1	0.02 [-0.41, 0.44]
			Irbesartan	115	40 (34.8)	-0.92 (7.50)	-23.5	-3.23	-0.42	3.26	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Asia Pacific	KDQOL-SF12: MCS	Baseline	Sparsentan	69	67 (97.1)	50.42 (7.02)	27.8	45.01	51.25	55.70	64.2	
			Irbesartan	41	41 (100.0)	51.19 (8.65)	24.3	49.04	51.77	57.16	66.6	
		Week 24	Sparsentan	69	57 (82.6)	52.50 (7.74)	30.7	47.94	54.20	57.34	66.8	
			Irbesartan	41	34 (82.9)	51.73 (8.76)	30.8	48.43	53.11	57.31	64.2	
		Week 48	Sparsentan	69	55 (79.7)	51.67 (8.11)	27.5	48.28	52.63	57.33	62.1	
			Irbesartan	41	23 (56.1)	51.98 (9.83)	21.5	47.27	54.20	59.29	64.6	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	69	57 (82.6)	2.11 (9.14)	-23.6	-2.29	1.28	7.00	26.4	0.23 [-0.20, 0.66]
			Irbesartan	41	34 (82.9)	0.29 (5.25)	-9.4	-2.88	0.18	3.16	15.5	
		Week 48	Sparsentan	69	55 (79.7)	1.49 (8.75)	-23.7	-4.61	1.97	6.67	21.1	0.23 [-0.26, 0.72]
			Irbesartan	41	23 (56.1)	-0.47 (7.70)	-18.5	-3.09	0.00	4.92	12.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline BMI												
< 27 kg/m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	84	76 (90.5)	48.43 (8.99)	16.2	42.67	50.05	54.47	64.6	
			Irbesartan	94	87 (92.6)	51.23 (8.04)	27.9	46.93	52.48	57.16	62.7	
		Week 24	Sparsentan	84	52 (61.9)	50.35 (9.44)	20.8	43.84	51.31	57.33	66.8	
			Irbesartan	94	54 (57.4)	52.27 (9.37)	24.2	49.52	55.45	57.76	62.7	
		Week 48	Sparsentan	84	48 (57.1)	49.42 (9.00)	27.5	42.61	51.33	57.08	61.1	
			Irbesartan	94	37 (39.4)	50.08 (10.71)	19.6	46.16	52.00	57.33	64.5	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	84	52 (61.9)	1.37 (10.39)	-23.6	-5.45	-0.08	8.20	26.4	0.05 [-0.33, 0.43]
			Irbesartan	94	54 (57.4)	0.93 (7.20)	-14.1	-4.36	0.00	4.51	28.7	
		Week 48	Sparsentan	84	48 (57.1)	1.22 (9.80)	-23.4	-3.80	0.24	6.45	28.1	0.26 [-0.17, 0.69]
			Irbesartan	94	37 (39.4)	-1.27 (9.22)	-33.9	-3.50	-0.60	3.74	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 27 kg/m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	118	111 (94.1)	51.73 (7.56)	31.4	47.07	53.19	57.16	66.5	
			Irbesartan	107	96 (89.7)	51.76 (8.86)	24.3	47.57	54.14	57.45	66.6	
	Week 24	Sparsentan	118	76 (64.4)	52.43 (6.12)	37.4	48.23	53.20	57.20	62.7		
		Irbesartan	107	58 (54.2)	53.11 (7.26)	33.5	48.43	55.28	58.15	64.2		
	Week 48	Sparsentan	118	64 (54.2)	51.71 (7.76)	27.5	48.52	52.34	58.01	63.4		
		Irbesartan	107	43 (40.2)	52.32 (9.19)	28.5	50.06	54.20	60.12	64.6		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	118	76 (64.4)	0.29 (6.50)	-18.1	-2.59	0.93	4.23	16.4	0.02 [-0.32, 0.36]
			Irbesartan	107	58 (54.2)	0.16 (6.39)	-15.4	-4.10	0.13	3.16	20.2	
Week 48		Sparsentan	118	64 (54.2)	0.20 (8.45)	-23.7	-4.53	0.19	5.77	18.7	0.16 [-0.22, 0.55]	
		Irbesartan	107	43 (40.2)	-1.13 (7.72)	-23.5	-3.95	0.25	4.87	12.1		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Randomization strata												
eGFR Low and UP High	KDQOL-SF12: MCS	Baseline	Sparsentan	71	64 (90.1)	51.84 (7.80)	31.4	45.00	54.20	57.25	64.6	
			Irbesartan	74	64 (86.5)	50.93 (8.84)	26.4	47.48	53.01	56.83	64.6	
		Week 24	Sparsentan	71	45 (63.4)	52.41 (6.90)	36.1	47.94	54.20	57.33	63.4	
			Irbesartan	74	36 (48.6)	53.48 (8.43)	29.7	49.86	57.11	59.42	62.6	
		Week 48	Sparsentan	71	39 (54.9)	51.85 (6.94)	37.3	47.66	52.63	57.33	63.4	
			Irbesartan	74	23 (31.1)	52.67 (12.98)	19.6	49.23	57.16	61.43	64.6	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	71	45 (63.4)	-0.75 (6.72)	-14.5	-5.09	-0.45	2.43	20.8	-0.18 [-0.62, 0.26]
			Irbesartan	74	36 (48.6)	0.51 (7.20)	-15.4	-3.12	0.34	3.25	28.7	
		Week 48	Sparsentan	71	39 (54.9)	-1.03 (7.22)	-16.9	-5.18	-1.04	4.45	11.8	-0.06 [-0.58, 0.45]
			Irbesartan	74	23 (31.1)	-0.46 (11.03)	-33.9	-2.96	0.69	4.87	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR Low and UP Low	KDQOL-SF12: MCS	Baseline	Sparsentan	55	51 (92.7)	50.29 (9.67)	16.2	44.79	52.99	57.16	65.2	
			Irbesartan	55	51 (92.7)	52.28 (7.70)	31.9	48.74	52.09	57.73	66.6	
		Week 24	Sparsentan	55	39 (70.9)	52.22 (7.78)	34.3	46.86	53.61	57.90	63.5	
			Irbesartan	55	35 (63.6)	52.65 (8.40)	30.8	47.84	54.45	57.66	64.2	
		Week 48	Sparsentan	55	28 (50.9)	50.24 (8.69)	29.5	43.33	52.24	56.27	62.5	
			Irbesartan	55	25 (45.5)	49.40 (9.71)	31.4	42.63	50.63	57.06	62.5	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	55	39 (70.9)	1.75 (7.58)	-17.0	-2.66	1.16	5.84	25.9	0.27 [-0.19, 0.73]
			Irbesartan	55	35 (63.6)	-0.20 (6.56)	-9.4	-6.30	0.00	4.66	20.2	
		Week 48	Sparsentan	55	28 (50.9)	0.41 (8.80)	-20.3	-4.74	-1.16	5.77	28.1	0.48 [-0.07, 1.02]
			Irbesartan	55	25 (45.5)	-3.68 (8.33)	-23.5	-6.62	-2.96	2.37	12.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR High and UP High	KDQOL-SF12: MCS	Baseline	Sparsentan	37	35 (94.6)	48.30 (7.97)	27.8	43.44	49.67	54.20	62.4		
			Irbesartan	36	33 (91.7)	52.02 (8.60)	27.9	46.56	54.45	57.33	64.2		
		Week 24	Sparsentan	37	20 (54.1)	48.78 (8.27)	20.8	46.84	49.62	54.20	57.3		
			Irbesartan	36	19 (52.8)	52.72 (7.11)	36.7	48.43	54.37	57.76	62.4		
		Week 48	Sparsentan	37	22 (59.5)	48.68 (10.07)	27.5	41.17	50.09	58.30	62.5		
			Irbesartan	36	13 (36.1)	52.04 (8.14)	32.8	50.11	51.36	60.12	61.3		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	37	20 (54.1)	1.72 (10.11)	-19.1	-3.71	2.30	7.52	26.4	0.06 [-0.57, 0.68]	
			Irbesartan	36	19 (52.8)	1.23 (7.34)	-13.4	-2.96	1.02	3.16	21.7		
			Week 48	Sparsentan	37	22 (59.5)	2.90 (11.90)	-23.7	-1.47	5.25	12.38	21.1	0.16 [-0.52, 0.85]
				Irbesartan	36	13 (36.1)	1.29 (4.83)	-8.1	-2.20	2.43	5.33	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024



Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP Low	KDQOL-SF12: MCS	Baseline	Sparsentan	39	37 (94.9)	49.98 (7.21)	30.1	46.33	50.94	54.63	66.5	
			Irbesartan	37	36 (97.3)	50.30 (9.63)	24.3	45.44	51.94	57.16	62.5	
		Week 24	Sparsentan	39	24 (61.5)	51.34 (8.32)	30.7	48.13	50.43	57.63	66.8	
			Irbesartan	37	22 (59.5)	51.52 (9.31)	24.2	47.77	54.51	57.16	62.5	
		Week 48	Sparsentan	39	23 (59.0)	51.35 (8.50)	27.5	49.02	52.25	57.10	60.3	
	Irbesartan		37	19 (51.4)	51.56 (6.94)	37.5	46.16	54.20	57.16	62.5		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	39	24 (61.5)	1.02 (10.31)	-23.6	-3.88	0.62	7.08	25.3	-0.02 [-0.59, 0.56]
			Irbesartan	37	22 (59.5)	1.15 (6.19)	-13.9	-1.32	0.00	4.80	15.5	
		Week 48	Sparsentan	39	23 (59.0)	1.56 (8.93)	-18.6	-2.06	1.15	3.95	19.4	0.27 [-0.34, 0.88]
			Irbesartan	37	19 (51.4)	-0.53 (6.19)	-17.3	-2.85	0.00	2.79	13.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 1													
< 60 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	127	116 (91.3)	51.36 (8.61)	16.2	46.28	53.55	57.19	65.2		
			Irbesartan	129	116 (89.9)	51.58 (8.12)	26.4	48.07	52.60	57.13	66.6		
		Week 24	Sparsentan	127	85 (66.9)	52.32 (7.26)	34.3	47.77	54.17	57.34	63.5		
			Irbesartan	129	73 (56.6)	52.84 (8.15)	29.7	49.28	55.77	58.15	64.2		
		Week 48	Sparsentan	127	67 (52.8)	50.90 (7.75)	29.5	44.50	52.05	57.33	62.5		
			Irbesartan	129	50 (38.8)	50.08 (11.06)	19.6	42.63	53.32	57.56	64.5		
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	127	85 (66.9)	0.24 (7.23)	-17.0	-3.78	0.00	3.27	25.9	0.00 [-0.31, 0.31]
				Irbesartan	129	73 (56.6)	0.23 (6.84)	-15.4	-4.70	0.00	3.95	28.7	
			Week 48	Sparsentan	127	67 (52.8)	-0.92 (8.03)	-20.3	-5.08	-1.34	4.38	28.1	0.20 [-0.16, 0.57]
				Irbesartan	129	50 (38.8)	-2.72 (9.77)	-33.9	-6.62	-1.68	2.79	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	49	47 (95.9)	48.70 (7.66)	30.1	43.44	48.53	54.37	66.5	
			Irbesartan	48	44 (91.7)	51.20 (9.16)	27.9	44.31	54.27	57.45	64.2	
	Week 24	Sparsentan	49	28 (57.1)	48.72 (8.99)	20.8	45.38	49.41	54.20	66.8		
		Irbesartan	48	26 (54.2)	52.83 (8.42)	24.2	49.52	55.97	57.73	62.7		
	Week 48	Sparsentan	49	29 (59.2)	48.31 (10.22)	27.5	41.17	49.58	56.52	63.4		
		Irbesartan	48	20 (41.7)	53.59 (8.27)	32.8	50.19	53.97	60.62	64.6		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	49	28 (57.1)	0.05 (9.96)	-23.6	-3.95	0.19	5.31	25.3	-0.08 [-0.61, 0.46]
			Irbesartan	48	26 (54.2)	0.72 (6.66)	-13.9	-2.29	0.00	2.96	21.7	
		Week 48	Sparsentan	49	29 (59.2)	1.05 (9.54)	-23.7	-1.47	1.89	7.36	15.7	-0.12 [-0.69, 0.45]
			Irbesartan	48	20 (41.7)	1.97 (3.60)	-5.4	0.00	2.45	4.95	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	26	24 (92.3)	49.00 (7.58)	27.8	47.06	50.17	53.71	61.5	
			Irbesartan	25	24 (96.0)	50.68 (10.24)	24.3	46.60	53.61	57.51	62.5	
	Week 24	Sparsentan	26	15 (57.7)	52.76 (6.46)	40.3	47.09	52.62	58.72	62.4		
		Irbesartan	25	13 (52.0)	51.73 (9.58)	36.1	46.16	51.77	60.12	62.5		
	Week 48	Sparsentan	26	16 (61.5)	54.37 (5.61)	42.5	50.59	53.40	59.79	62.5		
		Irbesartan	25	10 (40.0)	52.71 (5.58)	41.8	52.00	54.20	54.96	60.0		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	26	15 (57.7)	4.76 (9.90)	-11.1	-0.57	4.25	10.63	26.4	0.33 [-0.41, 1.08]
			Irbesartan	25	13 (52.0)	1.86 (6.97)	-13.4	0.00	2.77	5.23	15.5	
		Week 48	Sparsentan	26	16 (61.5)	6.41 (10.11)	-14.6	0.00	5.45	15.07	21.1	0.72 [-0.09, 1.54]
			Irbesartan	25	10 (40.0)	0.06 (5.90)	-8.1	-2.47	-2.09	2.79	13.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 2													
< 45 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	82	72 (87.8)	50.09 (9.01)	16.2	44.31	51.92	57.13	64.2		
			Irbesartan	80	70 (87.5)	50.90 (8.98)	26.4	47.81	52.12	57.16	66.6		
		Week 24	Sparsentan	82	54 (65.9)	52.52 (7.65)	34.3	47.15	54.29	57.65	63.5		
			Irbesartan	80	48 (60.0)	52.53 (8.47)	29.7	49.07	55.19	58.23	64.2		
		Week 48	Sparsentan	82	44 (53.7)	50.76 (7.77)	29.5	47.22	51.45	57.11	62.1		
			Irbesartan	80	30 (37.5)	48.83 (12.46)	19.6	42.02	51.14	58.58	64.5		
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	82	54 (65.9)	1.50 (7.30)	-13.0	-2.79	1.20	4.59	25.9	0.15 [-0.24, 0.54]
			Irbesartan	80	48 (60.0)	0.36 (7.69)	-15.4	-4.81	0.00	4.06	28.7		
			Week 48	Sparsentan	82	44 (53.7)	0.18 (7.92)	-20.3	-4.17	-0.08	4.58	28.1	0.35 [-0.12, 0.82]
				Irbesartan	80	30 (37.5)	-3.16 (11.51)	-33.9	-13.91	-1.48	4.19	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
45 to < 60 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	45	44 (97.8)	53.44 (7.56)	33.8	49.54	55.60	57.82	65.2	
		Week 24	Irbesartan	49	46 (93.9)	52.63 (6.54)	31.9	49.35	54.14	57.09	64.6	
			Sparsentan	45	31 (68.9)	51.98 (6.62)	41.0	47.77	50.91	57.16	63.5	
	Week 48	Irbesartan	49	25 (51.0)	53.41 (7.64)	33.5	51.37	55.77	57.16	63.2		
		Sparsentan	45	23 (51.1)	51.16 (7.88)	39.1	42.35	52.60	58.32	62.5		
	Week 24	KDQOL-SF12: change from baseline in MCS	Sparsentan	45	31 (68.9)	-1.97 (6.65)	-17.0	-6.80	-1.05	1.38	13.1	-0.33 [-0.86, 0.20]
			Irbesartan	49	25 (51.0)	-0.00 (4.95)	-8.6	-4.10	1.02	3.95	7.1	
	Week 48	MCS	Sparsentan	45	23 (51.1)	-3.04 (7.99)	-18.6	-5.18	-2.98	4.34	11.8	-0.13 [-0.73, 0.47]
			Irbesartan	49	20 (40.8)	-2.06 (6.57)	-17.3	-4.15	-1.68	2.30	8.2	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	49	47 (95.9)	48.70 (7.66)	30.1	43.44	48.53	54.37	66.5	
			Irbesartan	48	44 (91.7)	51.20 (9.16)	27.9	44.31	54.27	57.45	64.2	
	Week 24	Sparsentan	49	28 (57.1)	48.72 (8.99)	20.8	45.38	49.41	54.20	66.8		
		Irbesartan	48	26 (54.2)	52.83 (8.42)	24.2	49.52	55.97	57.73	62.7		
	Week 48	Sparsentan	49	29 (59.2)	48.31 (10.22)	27.5	41.17	49.58	56.52	63.4		
		Irbesartan	48	20 (41.7)	53.59 (8.27)	32.8	50.19	53.97	60.62	64.6		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	49	28 (57.1)	0.05 (9.96)	-23.6	-3.95	0.19	5.31	25.3	-0.08 [-0.61, 0.46]
			Irbesartan	48	26 (54.2)	0.72 (6.66)	-13.9	-2.29	0.00	2.96	21.7	
		Week 48	Sparsentan	49	29 (59.2)	1.05 (9.54)	-23.7	-1.47	1.89	7.36	15.7	-0.12 [-0.69, 0.45]
			Irbesartan	48	20 (41.7)	1.97 (3.60)	-5.4	0.00	2.45	4.95	6.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	26	24 (92.3)	49.00 (7.58)	27.8	47.06	50.17	53.71	61.5	
			Irbesartan	25	24 (96.0)	50.68 (10.24)	24.3	46.60	53.61	57.51	62.5	
	Week 24	Sparsentan	26	15 (57.7)	52.76 (6.46)	40.3	47.09	52.62	58.72	62.4		
		Irbesartan	25	13 (52.0)	51.73 (9.58)	36.1	46.16	51.77	60.12	62.5		
	Week 48	Sparsentan	26	16 (61.5)	54.37 (5.61)	42.5	50.59	53.40	59.79	62.5		
		Irbesartan	25	10 (40.0)	52.71 (5.58)	41.8	52.00	54.20	54.96	60.0		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	26	15 (57.7)	4.76 (9.90)	-11.1	-0.57	4.25	10.63	26.4	0.33 [-0.41, 1.08]
		Week 48	Irbesartan	25	13 (52.0)	1.86 (6.97)	-13.4	0.00	2.77	5.23	15.5	
			Sparsentan	26	16 (61.5)	6.41 (10.11)	-14.6	0.00	5.45	15.07	21.1	0.72 [-0.09, 1.54]
			Irbesartan	25	10 (40.0)	0.06 (5.90)	-8.1	-2.47	-2.09	2.79	13.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline urine protein excretion												
<= 1.75 g/day	KDQOL-SF12: MCS	Baseline	Sparsentan	98	92 (93.9)	50.12 (8.77)	16.2	45.77	51.52	56.26	66.5	
			Irbesartan	94	84 (89.4)	50.87 (8.76)	24.3	47.39	52.05	57.16	66.6	
		Week 24	Sparsentan	98	67 (68.4)	50.91 (8.72)	20.8	46.85	50.71	57.35	66.8	
			Irbesartan	94	47 (50.0)	52.11 (8.83)	24.2	47.77	54.37	57.59	64.2	
		Week 48	Sparsentan	98	60 (61.2)	49.51 (9.09)	27.5	42.55	50.99	56.71	62.5	
			Irbesartan	94	34 (36.2)	51.24 (8.89)	19.6	47.58	53.61	57.16	62.5	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	98	67 (68.4)	0.93 (8.80)	-23.6	-3.44	0.61	5.84	25.9	0.11 [-0.26, 0.49]
			Irbesartan	94	47 (50.0)	0.07 (5.79)	-13.9	-4.70	0.00	3.95	15.5	
		Week 48	Sparsentan	98	60 (61.2)	-0.01 (9.61)	-23.7	-4.66	-0.20	4.52	28.1	0.12 [-0.30, 0.54]
			Irbesartan	94	34 (36.2)	-1.14 (8.48)	-33.9	-2.96	-0.05	2.47	13.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 1.75 g/day	KDQOL-SF12: MCS	Baseline	Sparsentan	104	95 (91.3)	50.65 (7.88)	27.8	44.79	51.83	57.16	64.6	
			Irbesartan	108	100 (92.6)	51.80 (8.54)	26.4	47.17	54.16	57.33	64.6	
		Week 24	Sparsentan	104	61 (58.7)	52.32 (6.33)	36.1	47.94	54.20	57.16	63.4	
			Irbesartan	108	65 (60.2)	53.13 (7.96)	29.7	49.52	55.93	58.17	62.6	
		Week 48	Sparsentan	104	52 (50.0)	52.13 (7.25)	36.0	48.53	53.07	57.33	63.4	
			Irbesartan	108	46 (42.6)	51.32 (10.71)	21.5	47.27	54.20	60.12	64.6	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	104	61 (58.7)	0.50 (7.74)	-19.1	-4.09	0.00	4.25	26.4	-0.05 [-0.40, 0.30]
			Irbesartan	108	65 (60.2)	0.87 (7.43)	-15.4	-3.28	0.72	3.55	28.7	
		Week 48	Sparsentan	104	52 (50.0)	1.38 (8.33)	-16.9	-3.43	1.36	6.94	21.1	0.31 [-0.09, 0.71]
			Irbesartan	108	46 (42.6)	-1.24 (8.42)	-23.5	-5.33	0.00	4.87	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline use of antihypertensives												
Yes	KDQOL-SF12: MCS	Baseline	Sparsentan	88	78 (88.6)	49.63 (9.22)	16.2	44.21	51.69	57.16	65.2	
			Irbesartan	83	73 (88.0)	52.01 (8.02)	27.2	48.29	54.13	57.16	66.6	
		Week 24	Sparsentan	88	53 (60.2)	51.10 (7.85)	20.8	48.18	51.21	57.06	62.4	
			Irbesartan	83	43 (51.8)	53.69 (7.69)	30.8	51.94	56.99	58.17	64.2	
		Week 48	Sparsentan	88	43 (48.9)	51.21 (7.57)	29.5	47.66	52.25	57.33	63.4	
			Irbesartan	83	30 (36.1)	51.88 (9.11)	28.5	49.23	53.32	57.76	64.5	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	88	53 (60.2)	0.59 (8.06)	-18.1	-3.78	0.61	4.80	25.9	0.04 [-0.37, 0.44]
			Irbesartan	83	43 (51.8)	0.33 (6.18)	-7.2	-4.54	0.00	3.16	28.7	
		Week 48	Sparsentan	88	43 (48.9)	0.96 (8.62)	-16.9	-4.49	1.48	6.04	28.1	0.23 [-0.24, 0.70]
			Irbesartan	83	30 (36.1)	-1.07 (9.21)	-23.5	-4.01	-0.62	4.87	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL-SF12: MCS	Baseline	Sparsentan	114	109 (95.6)	50.93 (7.59)	27.8	45.63	51.53	56.44	66.5	
			Irbesartan	119	111 (93.3)	50.96 (9.01)	24.3	46.56	52.48	57.16	64.6	
		Week 24	Sparsentan	114	75 (65.8)	51.92 (7.59)	30.7	47.15	53.41	57.35	66.8	
			Irbesartan	119	69 (58.0)	52.10 (8.68)	24.2	47.84	54.37	57.76	63.2	
		Week 48	Sparsentan	114	69 (60.5)	50.42 (8.85)	27.5	44.50	51.44	57.16	62.5	
			Irbesartan	119	50 (42.0)	50.93 (10.45)	19.6	46.16	54.20	57.56	64.6	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	114	75 (65.8)	0.83 (8.49)	-23.6	-3.44	0.00	5.08	26.4	0.02 [-0.31, 0.35]
			Irbesartan	119	69 (58.0)	0.67 (7.16)	-15.4	-2.96	0.00	4.66	21.7	
		Week 48	Sparsentan	114	69 (60.5)	0.43 (9.32)	-23.7	-4.61	0.03	6.32	21.1	0.19 [-0.17, 0.56]
			Irbesartan	119	50 (42.0)	-1.27 (7.96)	-33.9	-3.09	0.00	3.76	13.4	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Time since renal biopsy												
<= 5 years	KDQOL-SF12: MCS	Baseline	Sparsentan	113	105 (92.9)	50.34 (8.81)	16.2	45.01	52.02	57.16	66.5	
			Irbesartan	127	119 (93.7)	51.23 (9.06)	24.3	46.63	53.45	57.32	66.6	
		Week 24	Sparsentan	113	73 (64.6)	51.96 (8.13)	20.8	47.61	53.04	57.35	66.8	
			Irbesartan	127	73 (57.5)	53.20 (8.34)	24.2	49.28	56.47	58.29	64.2	
		Week 48	Sparsentan	113	61 (54.0)	51.59 (7.35)	36.8	46.92	52.43	57.73	62.5	
			Irbesartan	127	53 (41.7)	50.69 (10.65)	19.6	47.27	54.20	57.16	64.6	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	113	73 (64.6)	1.63 (9.05)	-23.6	-4.29	1.38	6.42	26.4	0.05 [-0.28, 0.37]
			Irbesartan	127	73 (57.5)	1.25 (7.19)	-14.1	-2.88	0.72	4.80	28.7	
		Week 48	Sparsentan	113	61 (54.0)	2.11 (8.99)	-18.6	-4.49	1.97	7.21	28.1	0.41 [0.04, 0.78]
			Irbesartan	127	53 (41.7)	-1.55 (8.89)	-33.9	-3.50	0.00	3.74	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 5 years	KDQOL-SF12: MCS	Baseline	Sparsentan	89	82 (92.1)	50.44 (7.66)	30.1	44.79	51.32	56.08	64.2	
			Irbesartan	75	65 (86.7)	51.63 (7.83)	26.4	47.72	52.31	57.16	63.0	
	Week 24	Sparsentan	89	55 (61.8)	51.09 (7.07)	34.3	47.83	52.01	57.16	62.7		
		Irbesartan	75	39 (52.0)	51.78 (8.30)	29.7	48.43	53.00	57.33	62.7		
	Week 48	Sparsentan	89	51 (57.3)	49.69 (9.38)	27.5	42.75	51.62	57.16	63.4		
		Irbesartan	75	27 (36.0)	52.44 (8.36)	31.4	49.58	54.20	59.29	61.4		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	89	55 (61.8)	-0.47 (7.04)	-19.1	-2.86	0.00	3.02	20.8	0.05 [-0.36, 0.46]
			Irbesartan	75	39 (52.0)	-0.80 (5.76)	-15.4	-4.54	0.00	2.77	20.2	
		Week 48	Sparsentan	89	51 (57.3)	-1.12 (8.83)	-23.7	-5.16	-0.99	3.61	19.4	-0.07 [-0.54, 0.39]
			Irbesartan	75	27 (36.0)	-0.51 (7.43)	-18.5	-3.95	0.00	4.85	12.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: History of hypertension												
Yes	KDQOL-SF12: MCS	Baseline	Sparsentan	153	139 (90.8)	50.20 (8.31)	16.2	44.79	51.55	56.44	65.2	
			Irbesartan	157	140 (89.2)	51.81 (8.50)	26.4	48.07	54.10	57.33	66.6	
		Week 24	Sparsentan	153	97 (63.4)	51.99 (7.70)	20.8	48.18	53.04	57.34	66.8	
			Irbesartan	157	83 (52.9)	53.53 (7.44)	29.7	50.06	56.75	58.29	64.2	
		Week 48	Sparsentan	153	83 (54.2)	50.91 (8.03)	29.5	44.50	52.00	57.33	63.4	
			Irbesartan	157	59 (37.6)	51.61 (9.25)	19.6	48.28	54.20	57.41	64.6	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	153	97 (63.4)	1.15 (7.78)	-17.0	-3.78	0.99	4.84	25.9	0.07 [-0.22, 0.37]
			Irbesartan	157	83 (52.9)	0.61 (6.77)	-15.4	-4.36	0.00	3.89	28.7	
		Week 48	Sparsentan	153	83 (54.2)	1.08 (8.44)	-23.4	-4.49	1.15	6.43	28.1	0.30 [-0.03, 0.64]
			Irbesartan	157	59 (37.6)	-1.49 (8.67)	-33.9	-4.01	-0.11	2.79	17.3	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL-SF12: MCS	Baseline	Sparsentan	49	48 (98.0)	50.93 (8.36)	27.8	47.06	51.68	57.11	66.5	
			Irbesartan	45	44 (97.8)	49.98 (8.96)	24.3	44.78	51.25	57.16	62.5	
		Week 24	Sparsentan	49	31 (63.3)	50.31 (7.60)	30.7	44.08	52.01	57.16	62.7	
			Irbesartan	45	29 (64.4)	50.35 (10.21)	24.2	46.16	54.37	57.16	62.6	
		Week 48	Sparsentan	49	29 (59.2)	50.19 (9.36)	27.5	46.89	51.44	57.16	62.5	
			Irbesartan	45	21 (46.7)	50.36 (11.81)	21.5	42.63	54.20	60.02	64.5	
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	49	31 (63.3)	-0.60 (9.71)	-23.6	-2.86	-0.10	3.86	26.4	-0.11 [-0.62, 0.40]
			Irbesartan	45	29 (64.4)	0.33 (6.90)	-14.1	-2.96	0.00	3.55	15.5	
		Week 48	Sparsentan	49	29 (59.2)	-0.64 (10.59)	-23.7	-4.61	-0.24	4.38	21.1	-0.03 [-0.59, 0.53]
			Irbesartan	45	21 (46.7)	-0.37 (7.72)	-17.6	-2.47	0.00	4.85	12.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 04APR2024



Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.210
Male	Week 24	Sparsentan	139	88 (63.3)	0.00 (0.70)	(-1.38, 1.39)	-0.51 (1.02)	(-2.53, 1.50)	0.615
		Irbesartan	143	81 (56.6)	0.52 (0.73)	(-0.93, 1.96)			
	Week 48	Sparsentan	139	74 (53.2)	0.20 (0.76)	(-1.30, 1.69)	-0.69 (1.15)	(-2.95, 1.57)	0.546
		Irbesartan	143	58 (40.6)	0.89 (0.85)	(-0.79, 2.57)			
Female	Week 24	Sparsentan	63	40 (63.5)	1.33 (1.31)	(-1.26, 3.93)	-0.69 (1.98)	(-4.62, 3.23)	0.727
		Irbesartan	59	31 (52.5)	2.03 (1.48)	(-0.91, 4.97)			
	Week 48	Sparsentan	63	38 (60.3)	-0.27 (1.34)	(-2.91, 2.38)	4.37 (2.22)	(-0.03, 8.77)	0.052
		Irbesartan	59	22 (37.3)	-4.64 (1.76)	(-8.13, -1.14)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

MCS = mental component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.371
<= 45 years	Week 24	Sparsentan	96	59 (61.5)	0.12 (0.98)	(-1.81, 2.06)	-0.66 (1.40)	(-3.43, 2.11)	0.641
		Irbesartan	99	56 (56.6)	0.78 (1.00)	(-1.20, 2.76)			
	Week 48	Sparsentan	96	53 (55.2)	-0.96 (1.03)	(-2.99, 1.08)	-0.59 (1.57)	(-3.69, 2.50)	0.705
		Irbesartan	99	40 (40.4)	-0.36 (1.18)	(-2.69, 1.97)			
> 45 years	Week 24	Sparsentan	106	69 (65.1)	0.83 (0.83)	(-0.82, 2.47)	-0.48 (1.26)	(-2.96, 2.00)	0.701
		Irbesartan	103	56 (54.4)	1.31 (0.93)	(-0.53, 3.15)			
	Week 48	Sparsentan	106	59 (55.7)	0.88 (0.90)	(-0.89, 2.65)	1.77 (1.42)	(-1.02, 4.56)	0.213
		Irbesartan	103	40 (38.8)	-0.89 (1.09)	(-3.03, 1.26)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

MCS = mental component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Age at IgAN diagnosis	Overall		Sparsentan							Interaction: 0.612
<= 18 years	Week 24		Sparsentan	9	3 (33.3)	NE		NE		NE
			Irbesartan	5	2 (40.0)	NE				
	Week 48		Sparsentan	9	5 (55.6)	NE		NE		NE
			Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Week 24		Sparsentan	102	64 (62.7)	0.55 (0.89)	(-1.21, 2.30)	-0.50 (1.28)	(-3.02, 2.02)	0.695
			Irbesartan	109	61 (56.0)	1.05 (0.92)	(-0.76, 2.85)			
	Week 48		Sparsentan	102	57 (55.9)	-0.70 (0.95)	(-2.56, 1.16)	-0.39 (1.43)	(-3.21, 2.43)	0.785
			Irbesartan	109	45 (41.3)	-0.31 (1.06)	(-2.40, 1.79)			
> 40 years	Week 24		Sparsentan	91	61 (67.0)	1.18 (0.92)	(-0.63, 2.99)	-0.03 (1.38)	(-2.75, 2.69)	0.982
			Irbesartan	88	49 (55.7)	1.21 (1.02)	(-0.81, 3.23)			
	Week 48		Sparsentan	91	50 (54.9)	0.93 (1.00)	(-1.05, 2.90)	1.95 (1.58)	(-1.16, 5.06)	0.218
			Irbesartan	88	34 (38.6)	-1.02 (1.21)	(-3.41, 1.37)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

MCS = mental component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Geographic region	Overall		Sparsentan							Interaction: 0.365
North America	Week 24		Sparsentan	35	15 (42.9)	-0.45 (2.06)	(-4.57, 3.67)	-0.82 (2.58)	(-5.96, 4.33)	0.752
			Irbesartan	46	28 (60.9)	0.37 (1.53)	(-2.69, 3.43)			
	Week 48		Sparsentan	35	13 (37.1)	0.60 (2.31)	(-4.02, 5.22)	3.31 (3.04)	(-2.76, 9.37)	0.280
			Irbesartan	46	17 (37.0)	-2.71 (1.95)	(-6.61, 1.19)			
Europe	Week 24		Sparsentan	98	56 (57.1)	-0.37 (0.91)	(-2.16, 1.42)	-1.67 (1.32)	(-4.27, 0.94)	0.209
			Irbesartan	115	50 (43.5)	1.30 (0.96)	(-0.60, 3.20)			
	Week 48		Sparsentan	98	44 (44.9)	-0.96 (1.02)	(-2.97, 1.05)	-0.32 (1.48)	(-3.24, 2.61)	0.831
			Irbesartan	115	40 (34.8)	-0.64 (1.07)	(-2.75, 1.47)			
Asia Pacific	Week 24		Sparsentan	69	57 (82.6)	1.90 (0.98)	(-0.03, 3.84)	1.31 (1.62)	(-1.88, 4.51)	0.418
			Irbesartan	41	34 (82.9)	0.59 (1.27)	(-1.93, 3.11)			
	Week 48		Sparsentan	69	55 (79.7)	1.41 (1.00)	(-0.56, 3.38)	1.14 (1.83)	(-2.47, 4.75)	0.533
			Irbesartan	41	23 (56.1)	0.27 (1.52)	(-2.72, 3.27)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

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For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.997
< 27 kg/m**2	Week 24	Sparsentan	84	52 (61.9)	0.92 (1.14)	(-1.34, 3.18)	-0.54 (1.61)	(-3.73, 2.64)	0.738
		Irbesartan	94	54 (57.4)	1.46 (1.13)	(-0.77, 3.69)			
	Week 48	Sparsentan	84	48 (57.1)	0.51 (1.19)	(-1.84, 2.86)	1.23 (1.81)	(-2.34, 4.80)	0.498
		Irbesartan	94	37 (39.4)	-0.72 (1.35)	(-3.38, 1.94)			
≥ 27 kg/m**2	Week 24	Sparsentan	118	76 (64.4)	0.31 (0.72)	(-1.12, 1.73)	-0.13 (1.10)	(-2.30, 2.04)	0.906
		Irbesartan	107	58 (54.2)	0.44 (0.83)	(-1.19, 2.07)			
	Week 48	Sparsentan	118	64 (54.2)	-0.27 (0.78)	(-1.81, 1.27)	0.28 (1.24)	(-2.16, 2.71)	0.824
		Irbesartan	107	43 (40.2)	-0.55 (0.96)	(-2.43, 1.33)			

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LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan						Interaction:	0.539
eGFR Low and UP High	Week 24	Sparsentan	71	45 (63.4)	-0.70 (1.09)	(-2.85, 1.45)	-1.24 (1.63)	(-4.47, 1.98)	0.448
		Irbesartan	74	36 (48.6)	0.54 (1.22)	(-1.86, 2.95)			
	Week 48	Sparsentan	71	39 (54.9)	-1.17 (1.16)	(-3.46, 1.13)	-0.51 (1.90)	(-4.27, 3.25)	0.790
		Irbesartan	74	23 (31.1)	-0.66 (1.51)	(-3.64, 2.32)			
eGFR Low and UP Low	Week 24	Sparsentan	55	39 (70.9)	1.34 (1.07)	(-0.78, 3.46)	1.01 (1.56)	(-2.08, 4.10)	0.518
		Irbesartan	55	35 (63.6)	0.33 (1.13)	(-1.91, 2.57)			
	Week 48	Sparsentan	55	28 (50.9)	-0.20 (1.27)	(-2.70, 2.31)	2.76 (1.85)	(-0.90, 6.42)	0.138
		Irbesartan	55	25 (45.5)	-2.96 (1.34)	(-5.61, -0.31)			
eGFR High and UP High	Week 24	Sparsentan	37	20 (54.1)	1.04 (1.77)	(-2.49, 4.58)	-1.97 (2.58)	(-7.11, 3.17)	0.448
		Irbesartan	36	19 (52.8)	3.01 (1.84)	(-0.66, 6.68)			
	Week 48	Sparsentan	37	22 (59.5)	1.23 (1.71)	(-2.17, 4.64)	-1.37 (2.82)	(-7.00, 4.26)	0.628
		Irbesartan	36	13 (36.1)	2.61 (2.21)	(-1.80, 7.01)			
eGFR High and UP Low	Week 24	Sparsentan	39	24 (61.5)	0.51 (1.49)	(-2.45, 3.46)	-0.60 (2.14)	(-4.87, 3.67)	0.780
		Irbesartan	37	22 (59.5)	1.11 (1.55)	(-1.97, 4.19)			
	Week 48	Sparsentan	39	23 (59.0)	1.03 (1.52)	(-1.98, 4.05)	0.52 (2.25)	(-3.96, 5.01)	0.817
		Irbesartan	37	19 (51.4)	0.51 (1.66)	(-2.79, 3.81)			

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.123
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	85 (66.9)	0.16 (0.76)	(-1.34, 1.65)	-0.27 (1.12)	(-2.48, 1.93)	0.808
		Irbesartan	129	73 (56.6)	0.43 (0.82)	(-1.19, 2.05)			
	Week 48	Sparsentan	127	67 (52.8)	-1.15 (0.85)	(-2.82, 0.53)	1.27 (1.31)	(-1.30, 3.85)	0.332
		Irbesartan	129	50 (38.8)	-2.42 (0.99)	(-4.36, -0.47)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	28 (57.1)	-0.35 (1.47)	(-3.27, 2.56)	-1.91 (2.14)	(-6.16, 2.34)	0.375
		Irbesartan	48	26 (54.2)	1.56 (1.53)	(-1.48, 4.59)			
	Week 48	Sparsentan	49	29 (59.2)	0.15 (1.46)	(-2.75, 3.05)	-2.56 (2.30)	(-7.13, 2.02)	0.270
		Irbesartan	48	20 (41.7)	2.71 (1.74)	(-0.75, 6.16)			
≥ 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	15 (57.7)	4.15 (1.54)	(1.04, 7.27)	1.92 (2.27)	(-2.66, 6.49)	0.403
		Irbesartan	25	13 (52.0)	2.24 (1.66)	(-1.10, 5.58)			
	Week 48	Sparsentan	26	16 (61.5)	5.59 (1.51)	(2.54, 8.64)	3.09 (2.45)	(-1.84, 8.02)	0.213
		Irbesartan	25	10 (40.0)	2.50 (1.89)	(-1.32, 6.31)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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A decrease reflects a worsening of the status of the patient.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

MCS = mental component summary.

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Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.070
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	54 (65.9)	1.37 (1.02)	(-0.64, 3.38)	0.72 (1.49)	(-2.21, 3.65)	0.629
		Irbesartan	80	48 (60.0)	0.65 (1.08)	(-1.48, 2.78)			
	Week 48	Sparsentan	82	44 (53.7)	-0.16 (1.13)	(-2.38, 2.06)	2.83 (1.77)	(-0.66, 6.32)	0.111
		Irbesartan	80	30 (37.5)	-3.00 (1.36)	(-5.68, -0.31)			
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	31 (68.9)	-2.02 (1.09)	(-4.19, 0.15)	-2.32 (1.66)	(-5.62, 0.98)	0.166
		Irbesartan	49	25 (51.0)	0.30 (1.23)	(-2.15, 2.75)			
	Week 48	Sparsentan	45	23 (51.1)	-2.92 (1.25)	(-5.40, -0.44)	-1.44 (1.86)	(-5.14, 2.26)	0.442
		Irbesartan	49	20 (40.8)	-1.49 (1.36)	(-4.20, 1.23)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	28 (57.1)	-0.35 (1.47)	(-3.27, 2.56)	-1.91 (2.14)	(-6.16, 2.34)	0.375
		Irbesartan	48	26 (54.2)	1.56 (1.53)	(-1.48, 4.59)			
	Week 48	Sparsentan	49	29 (59.2)	0.15 (1.46)	(-2.75, 3.05)	-2.56 (2.30)	(-7.13, 2.02)	0.270
		Irbesartan	48	20 (41.7)	2.71 (1.74)	(-0.75, 6.16)			
≥ 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	15 (57.7)	4.15 (1.54)	(1.04, 7.27)	1.92 (2.27)	(-2.66, 6.49)	0.403
		Irbesartan	25	13 (52.0)	2.24 (1.66)	(-1.10, 5.58)			
	Week 48	Sparsentan	26	16 (61.5)	5.59 (1.51)	(2.54, 8.64)	3.09 (2.45)	(-1.84, 8.02)	0.213
		Irbesartan	25	10 (40.0)	2.50 (1.89)	(-1.32, 6.31)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

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Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.666
<= 1.75 g/day	Week 24	Sparsentan	98	67 (68.4)	0.63 (0.91)	(-1.16, 2.42)	0.25 (1.42)	(-2.55, 3.05)	0.860
		Irbesartan	94	47 (50.0)	0.38 (1.09)	(-1.77, 2.53)			
	Week 48	Sparsentan	98	60 (61.2)	-0.36 (0.96)	(-2.25, 1.54)	0.11 (1.61)	(-3.07, 3.28)	0.948
		Irbesartan	94	34 (36.2)	-0.46 (1.28)	(-2.98, 2.06)			
> 1.75 g/day	Week 24	Sparsentan	104	61 (58.7)	0.27 (0.89)	(-1.48, 2.02)	-0.86 (1.24)	(-3.31, 1.59)	0.491
		Irbesartan	108	65 (60.2)	1.13 (0.86)	(-0.58, 2.83)			
	Week 48	Sparsentan	104	52 (50.0)	0.47 (0.96)	(-1.42, 2.36)	1.39 (1.41)	(-1.40, 4.17)	0.327
		Irbesartan	108	46 (42.6)	-0.92 (1.02)	(-2.92, 1.09)			

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Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.695
Yes	Week 24	Sparsentan	88	53 (60.2)	-0.02 (0.92)	(-1.84, 1.80)	-1.28 (1.38)	(-4.01, 1.46)	0.358
		Irbesartan	83	43 (51.8)	1.26 (1.03)	(-0.77, 3.28)			
	Week 48	Sparsentan	88	43 (48.9)	0.24 (1.02)	(-1.78, 2.26)	0.58 (1.60)	(-2.58, 3.74)	0.716
		Irbesartan	83	30 (36.1)	-0.34 (1.22)	(-2.75, 2.07)			
No	Week 24	Sparsentan	114	75 (65.8)	0.89 (0.87)	(-0.83, 2.61)	0.10 (1.26)	(-2.39, 2.59)	0.937
		Irbesartan	119	69 (58.0)	0.79 (0.91)	(-1.01, 2.59)			
	Week 48	Sparsentan	114	69 (60.5)	-0.10 (0.91)	(-1.88, 1.69)	0.73 (1.40)	(-2.03, 3.49)	0.603
		Irbesartan	119	50 (42.0)	-0.83 (1.07)	(-2.92, 1.27)			

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Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.326
<= 5 years	Week 24	Sparsentan	113	73 (64.6)	1.40 (0.85)	(-0.28, 3.07)	-0.34 (1.21)	(-2.72, 2.03)	0.776
		Irbesartan	127	73 (57.5)	1.74 (0.85)	(0.06, 3.42)			
	Week 48	Sparsentan	113	61 (54.0)	1.39 (0.93)	(-0.44, 3.22)	2.29 (1.36)	(-0.39, 4.96)	0.094
		Irbesartan	127	53 (41.7)	-0.90 (0.99)	(-2.85, 1.05)			
> 5 years	Week 24	Sparsentan	89	55 (61.8)	-0.58 (0.94)	(-2.43, 1.27)	-0.04 (1.46)	(-2.93, 2.85)	0.980
		Irbesartan	75	39 (52.0)	-0.55 (1.13)	(-2.78, 1.68)			
	Week 48	Sparsentan	89	51 (57.3)	-1.68 (0.98)	(-3.62, 0.26)	-1.62 (1.67)	(-4.93, 1.68)	0.333
		Irbesartan	75	27 (36.0)	-0.05 (1.34)	(-2.70, 2.60)			

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LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

MCS = mental component summary.

Source Data: aqs, created on: 04APR2024

Table PT1KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.692
Yes	Week 24	Sparsentan	153	97 (63.4)	0.78 (0.69)	(-0.57, 2.13)	-0.54 (1.01)	(-2.53, 1.46)	0.597
		Irbesartan	157	83 (52.9)	1.32 (0.74)	(-0.15, 2.78)			
	Week 48	Sparsentan	153	83 (54.2)	0.28 (0.74)	(-1.18, 1.73)	1.22 (1.15)	(-1.05, 3.49)	0.291
		Irbesartan	157	59 (37.6)	-0.94 (0.88)	(-2.66, 0.79)			
No	Week 24	Sparsentan	49	31 (63.3)	-0.47 (1.51)	(-3.46, 2.53)	-0.47 (2.18)	(-4.79, 3.86)	0.831
		Irbesartan	45	29 (64.4)	0.00 (1.57)	(-3.11, 3.11)			
	Week 48	Sparsentan	49	29 (59.2)	-0.71 (1.56)	(-3.79, 2.38)	-0.83 (2.42)	(-5.64, 3.98)	0.732
		Irbesartan	45	21 (46.7)	0.13 (1.85)	(-3.53, 3.79)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. NE= Not evaluable.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. A first order regressive covariance structure was used.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 58 are included in the model and also presented.

A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

MCS = mental component summary.

Source Data: aqs, created on: 04APR2024

Figure PF1KMSC\_FSGM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

Not done: No significant subgroup found.

Number of available records are presented for key visits up to Week 58 only. LS-Mean = Least square mean.  
95% CI = 95% confidence interval. eGFR = estimated glomerular filtration rate. UP = urine protein. BMI = body mass index.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
IgAN = immunoglobulin A nephropathy. A decrease reflects a worsening of the status of the patient. Reference table: PT1KMSC\_FSCM.

Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.563
Male	Sparsentan	139	20 (14.4)	NE		1.567	(0.742, 3.311)	0.239
	Irbesartan	143	13 (9.1)	NE				
Female	Sparsentan	63	13 (20.6)	NE		2.406	(0.476, 12.166)	0.288
	Irbesartan	59	2 (3.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.621
<= 45 years	Sparsentan	96	15 (15.6)	NE		1.845	(0.733, 4.646)	0.193
	Irbesartan	99	9 (9.1)	NE				
> 45 years	Sparsentan	106	18 (17.0)	NE		1.502	(0.577, 3.907)	0.404
	Irbesartan	103	6 (5.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	1 (11.1) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	16 (15.7)	NE		1.903	(0.773, 4.689)	0.162
	Irbesartan	109	11 (10.1)	NE				
> 40 years	Sparsentan	91	16 (17.6)	NE		2.666	(0.860, 8.271)	0.090
	Irbesartan	88	4 (4.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.117
North America	Sparsentan	35	4 (11.4)	NE		2.796	(0.336, 23.267)	0.342
	Irbesartan	46	3 (6.5)	NE				
Europe	Sparsentan	98	9 (9.2)	NE		0.959	(0.308, 2.980)	0.942
	Irbesartan	115	8 (7.0)	NE				
Asia Pacific	Sparsentan	69	20 (29.0)	NE		3.804	(1.249, 11.587)	0.019 *
	Irbesartan	41	4 (9.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.814
< 27 kg/m**2	Sparsentan	84	17 (20.2)	NE		1.613	(0.634, 4.102)	0.315
	Irbesartan	94	7 (7.4)	NE				
≥ 27 kg/m**2	Sparsentan	118	16 (13.6)	NE		1.992	(0.774, 5.126)	0.153
	Irbesartan	107	8 (7.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.240
eGFR Low and UP High	Sparsentan	71	7 (9.9)	NE		2.396	(0.539, 10.645)	0.251
	Irbesartan	74	4 (5.4)	NE				
eGFR Low and UP Low	Sparsentan	55	8 (14.5)	NE		0.739	(0.230, 2.379)	0.612
	Irbesartan	55	6 (10.9)	NE				
eGFR High and UP High	Sparsentan	37	11 (29.7)	94.4	(46.1, NE)	2.070	(0.440, 9.739)	0.357
	Irbesartan	36	2 (5.6)	NE				
eGFR High and UP Low	Sparsentan	39	7 (17.9)	NE		4.492	(0.907, 22.247)	0.066
	Irbesartan	37	3 (8.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.113
< 60 mL/min/1.73 m**2	Sparsentan	127	14 (11.0)	NE		1.163	(0.510, 2.653)	0.720
	Irbesartan	129	10 (7.8)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	10 (20.4)	94.4	(69.7, NE)	1.649	(0.445, 6.112)	0.455
	Irbesartan	48	3 (6.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	9 (34.6)	72.1	(24.1, NE)	36.556	(2.758, 484.571)	0.006 *
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: 0.200
< 45 mL/min/1.73 m**2	Sparsentan	82	11 (13.4)	NE		1.314	(0.493, 3.499)	0.585
	Irbesartan	80	7 (8.8)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	3 (6.7)	NE		0.563	(0.104, 3.054)	0.506
	Irbesartan	49	3 (6.1)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	10 (20.4)	94.4	(69.7, NE)	1.649	(0.445, 6.112)	0.455
	Irbesartan	48	3 (6.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	9 (34.6)	72.1	(24.1, NE)	36.556	(2.758, 484.571)	0.006 *
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.184
<= 1.75 g/day	Sparsentan	98	18 (18.4)	NE		2.497	(0.875, 7.125)	0.087
	Irbesartan	94	5 (5.3)	NE				
> 1.75 g/day	Sparsentan	104	15 (14.4)	NE		1.610	(0.636, 4.079)	0.315
	Irbesartan	108	10 (9.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.295
Yes	Sparsentan	88	14 (15.9)	NE		3.405	(1.023, 11.329)	0.046 *
	Irbesartan	83	4 (4.8)	NE				
No	Sparsentan	114	19 (16.7)	NE		1.480	(0.661, 3.310)	0.340
	Irbesartan	119	11 (9.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.058
<= 5 years	Sparsentan	113	24 (21.2)	NE		2.645	(1.237, 5.656)	0.012 *
	Irbesartan	127	10 (7.9)	NE				
> 5 years	Sparsentan	89	9 (10.1)	NE		0.729	(0.221, 2.410)	0.605
	Irbesartan	75	5 (6.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension		Sparsentan				Interaction test: 0.214		
Yes	Sparsentan	153	26 (17.0)	NE		3.123	(1.329, 7.341)	0.009 *
	Irbesartan	157	8 (5.1)	NE				
No	Sparsentan	49	7 (14.3)	NE		1.733	(0.446, 6.737)	0.428
	Irbesartan	45	7 (15.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KMSIT\_FSKM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_   
An increase reflects an improvement of the status of the patient.  
Reference table: PT1KMSIT\_FSTM

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.642
Male	Sparsentan	139	21 (15.1)	111.1	(111.1, NE)	1.478	(0.738, 2.961)	0.270
	Irbesartan	143	15 (10.5)	NE				
Female	Sparsentan	63	13 (20.6)	108.7	(87.7, 114.3)	1.166	(0.449, 3.026)	0.752
	Irbesartan	59	7 (11.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.226
<= 45 years	Sparsentan	96	18 (18.8)	108.7	(87.7, NE)	1.973	(0.889, 4.380)	0.095
	Irbesartan	99	10 (10.1)	NE				
> 45 years	Sparsentan	106	16 (15.1)	111.1	(96.1, 114.3)	0.944	(0.421, 2.115)	0.888
	Irbesartan	103	12 (11.7)	108.1	(95.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	3 (33.3) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	19 (18.6)	108.7	(87.7, NE)	1.777	(0.836, 3.778)	0.135
	Irbesartan	109	12 (11.0)	NE				
> 40 years	Sparsentan	91	12 (13.2)	111.1	(111.1, 114.3)	0.901	(0.368, 2.203)	0.819
	Irbesartan	88	10 (11.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.785
North America	Sparsentan	35	5 (14.3)	NE		0.736	(0.203, 2.660)	0.640
	Irbesartan	46	7 (15.2)	NE				
Europe	Sparsentan	98	18 (18.4)	111.1	(95.1, 114.3)	2.032	(0.922, 4.478)	0.078
	Irbesartan	115	11 (9.6)	NE				
Asia Pacific	Sparsentan	69	11 (15.9)	NE		1.526	(0.472, 4.928)	0.480
	Irbesartan	41	4 (9.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.877
< 27 kg/m**2	Sparsentan	84	16 (19.0)	111.1	(111.1, 114.3)	1.350	(0.600, 3.036)	0.469
	Irbesartan	94	11 (11.7)	NE				
≥ 27 kg/m**2	Sparsentan	118	18 (15.3)	NE		1.419	(0.650, 3.095)	0.380
	Irbesartan	107	11 (10.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.675
eGFR Low and UP High	Sparsentan	71	12 (16.9)	NE		1.066	(0.431, 2.636)	0.890
	Irbesartan	74	8 (10.8)	NE				
eGFR Low and UP Low	Sparsentan	55	10 (18.2)	111.1	(95.1, 111.1)	1.328	(0.522, 3.379)	0.552
	Irbesartan	55	9 (16.4)	NE				
eGFR High and UP High	Sparsentan	37	8 (21.6)	114.3	(87.7, 114.3)	3.699	(0.648, 21.114)	0.141
	Irbesartan	36	2 (5.6)	NE				
eGFR High and UP Low	Sparsentan	39	4 (10.3)	NE		1.372	(0.297, 6.344)	0.686
	Irbesartan	37	3 (8.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024



Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

Subgroup	Treatment	KDQOL-SF12: time to decrease in MCS by at least 9.0		Kaplan-Meier analysis		Cox regression		
		N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.220
< 60 mL/min/1.73 m**2	Sparsentan	127	23 (18.1)	111.1	(96.1, 111.1)	1.271	(0.665, 2.427)	0.468
	Irbesartan	129	19 (14.7)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	8 (16.3)	114.3	(NE, NE)	5.740	(0.669, 49.217)	0.111
	Irbesartan	48	1 (2.1)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	3 (11.5)	NE		1.397	(0.170, 11.486)	0.756
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.101
< 45 mL/min/1.73 m**2	Sparsentan	82	12 (14.6)	111.1	(96.1, 111.1)	0.734	(0.329, 1.641)	0.452
	Irbesartan	80	13 (16.3)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	11 (24.4)	108.7	(69.4, NE)	3.294	(1.036, 10.479)	0.043 *
	Irbesartan	49	6 (12.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	8 (16.3)	114.3	(NE, NE)	5.740	(0.669, 49.217)	0.111
	Irbesartan	48	1 (2.1)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	3 (11.5)	NE		1.397	(0.170, 11.486)	0.756
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.200
<= 1.75 g/day	Sparsentan	98	19 (19.4)	111.1	(95.1, 111.1)	2.037	(0.809, 5.129)	0.131
	Irbesartan	94	7 (7.4)	NE				
> 1.75 g/day	Sparsentan	104	15 (14.4)	114.3	(NE, NE)	1.260	(0.569, 2.793)	0.568
	Irbesartan	108	15 (13.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.154
Yes	Sparsentan	88	17 (19.3)	111.1	(72.1, 111.1)	2.198	(0.894, 5.405)	0.086
	Irbesartan	83	8 (9.6)	NE				
No	Sparsentan	114	17 (14.9)	114.3	(108.7, 114.3)	1.050	(0.500, 2.205)	0.898
	Irbesartan	119	14 (11.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.672
<= 5 years	Sparsentan	113	15 (13.3)	NE		1.156	(0.544, 2.456)	0.705
	Irbesartan	127	14 (11.0)	NE				
> 5 years	Sparsentan	89	19 (21.3)	111.1	(96.1, 114.3)	1.712	(0.715, 4.099)	0.228
	Irbesartan	75	8 (10.7)	108.1	(108.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Table PT1KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan						Interaction test:	0.987
Yes	Sparsentan	153	22 (14.4)	111.1	(108.7, 114.3)	1.444	(0.710, 2.937)	0.310
	Irbesartan	157	14 (8.9)	NE				
No	Sparsentan	49	12 (24.5)	NE		1.292	(0.518, 3.223)	0.583
	Irbesartan	45	8 (17.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 04APR2024

Figure PF1KMSDT\_FSKM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_ reflects a worsening of the status of the patient.  
Reference table: PT1KMSDT\_FSTM

Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.871
Male	Double-blind period	Sparsentan	139	114 (82.0)	1.128 [0.993, 1.280]	1.710 [0.969, 3.018]	9.3 [-1.1, 19.7]	0.066
		Irbesartan	143	104 (72.7)				
Female	Double-blind period	Sparsentan	63	52 (82.5)	1.107 [0.918, 1.335]	1.612 [0.671, 3.868]	8.0 [-8.2, 24.1]	0.377
		Irbesartan	59	44 (74.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.579
<= 45 years	Double-blind period	Sparsentan	96	83 (86.5)	1.157 [1.006, 1.329]	2.157 [1.029, 4.520]	11.7 [-0.3, 23.7]	0.047 *
		Irbesartan	99	74 (74.7)				
> 45 years	Double-blind period	Sparsentan	106	83 (78.3)	1.090 [0.931, 1.275]	1.414 [0.753, 2.657]	6.5 [-6.2, 19.1]	0.337
		Irbesartan	103	74 (71.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.761
<= 18 years	Double-blind period	Sparsentan	9	8 (88.9)	1.481 [0.698, 3.143]	5.333 [0.343, 82.827]	28.9 [-34.3, 92.0]	0.505
		Irbesartan	5	3 (60.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	87 (85.3)	1.120 [0.981, 1.279]	1.817 [0.899, 3.670]	9.1 [-2.3, 20.6]	0.117
		Irbesartan	109	83 (76.1)				
> 40 years	Double-blind period	Sparsentan	91	71 (78.0)	1.107 [0.931, 1.318]	1.489 [0.758, 2.924]	7.6 [-6.3, 21.5]	0.305
		Irbesartan	88	62 (70.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.383
North America	Double-blind period	Sparsentan	35	28 (80.0)	1.082 [0.853, 1.374]	1.412 [0.490, 4.066]	6.1 [-14.8, 26.9]	0.603
		Irbesartan	46	34 (73.9)				
Europe	Double-blind period	Sparsentan	98	79 (80.6)	1.173 [1.003, 1.373]	1.895 [1.002, 3.584]	11.9 [-0.6, 24.4]	0.059
		Irbesartan	115	79 (68.7)				
Asia Pacific	Double-blind period	Sparsentan	69	59 (85.5)	1.002 [0.854, 1.175]	1.011 [0.338, 3.024]	0.1 [-15.4, 15.7]	1.000
		Irbesartan	41	35 (85.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.962
< 27 kg/m**2	Double-blind period	Sparsentan	84	68 (81.0)	1.119 [0.951, 1.316]	1.625 [0.801, 3.298]	8.6 [-4.9, 22.1]	0.217
		Irbesartan	94	68 (72.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	98 (83.1)	1.125 [0.979, 1.293]	1.737 [0.910, 3.313]	9.2 [-2.4, 20.8]	0.105
		Irbesartan	107	79 (73.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
Safety Set

TEAEs					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.250
eGFR Low and UP High	Double-blind period	Sparsentan	71	61 (85.9)	1.177	2.259	12.9	0.066
		Irbesartan	74	54 (73.0)	[0.996, 1.392]	[0.973, 5.248]	[-1.4, 27.3]	
eGFR Low and UP Low	Double-blind period	Sparsentan	55	44 (80.0)	1.023	1.116	1.8	1.000
		Irbesartan	55	43 (78.2)	[0.844, 1.240]	[0.445, 2.800]	[-15.2, 18.8]	
eGFR High and UP High	Double-blind period	Sparsentan	37	31 (83.8)	1.371	3.288	22.7	0.038 *
		Irbesartan	36	22 (61.1)	[1.019, 1.844]	[1.093, 9.892]	[0.1, 45.3]	
eGFR High and UP Low	Double-blind period	Sparsentan	39	30 (76.9)	0.981	0.920	-1.5	1.000
		Irbesartan	37	29 (78.4)	[0.771, 1.249]	[0.312, 2.709]	[-22.8, 19.9]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.861
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	106 (83.5)	1.099 [0.970, 1.244]	1.597 [0.860, 2.963]	7.5 [-3.1, 18.1]	0.163
		Irbesartan	129	98 (76.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	40 (81.6)	1.152 [0.920, 1.443]	1.830 [0.705, 4.751]	10.8 [-8.1, 29.7]	0.240
		Irbesartan	48	34 (70.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	20 (76.9)	1.202 [0.837, 1.726]	1.875 [0.551, 6.379]	12.9 [-15.8, 41.7]	0.368
		Irbesartan	25	16 (64.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.896
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	68 (82.9)	1.070	1.410	5.4	0.433
		Irbesartan	80	62 (77.5)	[0.918, 1.248]	[0.647, 3.072]	[-8.1, 18.9]	
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	38 (84.4)	1.149	1.960	11.0	0.217
		Irbesartan	49	36 (73.5)	[0.932, 1.418]	[0.703, 5.467]	[-7.4, 29.4]	
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	40 (81.6)	1.152	1.830	10.8	0.240
		Irbesartan	48	34 (70.8)	[0.920, 1.443]	[0.705, 4.751]	[-8.1, 29.7]	
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	20 (76.9)	1.202	1.875	12.9	0.368
		Irbesartan	25	16 (64.0)	[0.837, 1.726]	[0.551, 6.379]	[-15.8, 41.7]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.128
<= 1.75 g/day	Double-blind period	Sparsentan	98	75 (76.5)	1.028 [0.875, 1.208]	1.118 [0.579, 2.159]	2.1 [-11.1, 15.3]	0.867
		Irbesartan	94	70 (74.5)				
> 1.75 g/day	Double-blind period	Sparsentan	104	91 (87.5)	1.212 [1.056, 1.390]	2.692 [1.314, 5.518]	15.3 [3.8, 26.8]	0.006 *
		Irbesartan	108	78 (72.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
Safety Set

TEAEs					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.178
Yes	Double-blind period	Sparsentan	88	70 (79.5)	1.032	1.155	2.4	0.714
		Irbesartan	83	64 (77.1)	[0.881, 1.208]	[0.557, 2.392]	[-11.1, 16.0]	
No	Double-blind period	Sparsentan	114	96 (84.2)	1.193	2.222	13.6	0.019 *
		Irbesartan	119	84 (70.6)	[1.037, 1.373]	[1.172, 4.212]	[2.2, 25.1]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.486
<= 5 years	Double-blind period	Sparsentan	113	92 (81.4)	1.088 [0.952, 1.244]	1.476 [0.793, 2.745]	6.6 [-4.6, 17.9]	0.275
		Irbesartan	127	95 (74.8)				
> 5 years	Double-blind period	Sparsentan	89	74 (83.1)	1.177 [0.989, 1.399]	2.048 [0.972, 4.314]	12.5 [-1.7, 26.6]	0.063
		Irbesartan	75	53 (70.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.897
Yes	Double-blind period	Sparsentan	153	125 (81.7)	1.125 [0.996, 1.271]	1.684 [0.982, 2.888]	9.1 [-0.8, 19.0]	0.060
		Irbesartan	157	114 (72.6)				
No	Double-blind period	Sparsentan	49	41 (83.7)	1.107 [0.900, 1.362]	1.658 [0.599, 4.588]	8.1 [-10.3, 26.5]	0.442
		Irbesartan	45	34 (75.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.607
Male	Double-blind period	Sparsentan	139	34 (24.5)	1.128 [0.736, 1.729]	1.170 [0.672, 2.037]	2.8 [-7.8, 13.3]	0.672
		Irbesartan	143	31 (21.7)				
Female	Double-blind period	Sparsentan	63	15 (23.8)	1.405 [0.686, 2.877]	1.531 [0.627, 3.743]	6.9 [-9.0, 22.7]	0.378
		Irbesartan	59	10 (16.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.204
<= 45 years	Double-blind period	Sparsentan	96	29 (30.2)	1.495 [0.911, 2.455]	1.710 [0.887, 3.295]	10.0 [-3.1, 23.2]	0.137
		Irbesartan	99	20 (20.2)				
> 45 years	Double-blind period	Sparsentan	106	20 (18.9)	0.925 [0.534, 1.602]	0.908 [0.459, 1.798]	-1.5 [-13.2, 10.2]	0.862
		Irbesartan	103	21 (20.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.617
<= 18 years	Double-blind period	Sparsentan	9	5 (55.6)	2.778 [0.438, 17.629]	5.000 [0.388, 64.387]	35.6 [-27.8, 98.9]	0.301
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	23 (22.5)	1.069 [0.641, 1.781]	1.089 [0.566, 2.093]	1.4 [-10.7, 13.6]	0.868
		Irbesartan	109	23 (21.1)				
> 40 years	Double-blind period	Sparsentan	91	21 (23.1)	1.195 [0.677, 2.109]	1.253 [0.610, 2.573]	3.8 [-9.3, 16.8]	0.586
		Irbesartan	88	17 (19.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.571
North America	Double-blind period	Sparsentan	35	7 (20.0)	1.533 [0.565, 4.159]	1.667 [0.506, 5.493]	7.0 [-12.0, 25.9]	0.543
		Irbesartan	46	6 (13.0)				
Europe	Double-blind period	Sparsentan	98	24 (24.5)	1.006 [0.626, 1.616]	1.008 [0.538, 1.887]	0.1 [-12.4, 12.7]	1.000
		Irbesartan	115	28 (24.3)				
Asia Pacific	Double-blind period	Sparsentan	69	18 (26.1)	1.528 [0.698, 3.343]	1.714 [0.647, 4.545]	9.0 [-8.4, 26.4]	0.350
		Irbesartan	41	7 (17.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.028 #
< 27 kg/m**2	Double-blind period	Sparsentan	84	16 (19.0)	0.746 [0.426, 1.306]	0.686 [0.336, 1.403]	-6.5 [-19.8, 6.8]	0.369
		Irbesartan	94	24 (25.5)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	33 (28.0)	1.760 [1.043, 2.971]	2.055 [1.067, 3.961]	12.1 [0.5, 23.6]	0.037 *
		Irbesartan	107	17 (15.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.987
eGFR Low and UP High	Double-blind period	Sparsentan	71	21 (29.6)	1.152 [0.679, 1.954]	1.216 [0.586, 2.521]	3.9 [-12.0, 19.8]	0.711
		Irbesartan	74	19 (25.7)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	16 (29.1)	1.333 [0.697, 2.550]	1.470 [0.619, 3.491]	7.3 [-10.8, 25.3]	0.512
		Irbesartan	55	12 (21.8)				
eGFR High and UP High	Double-blind period	Sparsentan	37	6 (16.2)	1.168 [0.391, 3.488]	1.200 [0.331, 4.346]	2.3 [-16.8, 21.5]	1.000
		Irbesartan	36	5 (13.9)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	6 (15.4)	1.138 [0.380, 3.414]	1.164 [0.323, 4.196]	1.9 [-16.6, 20.3]	1.000
		Irbesartan	37	5 (13.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.699
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	36 (28.3)	1.306 [0.851, 2.005]	1.427 [0.808, 2.522]	6.6 [-4.7, 18.0]	0.249
		Irbesartan	129	28 (21.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	9 (18.4)	0.882 [0.393, 1.977]	0.855 [0.313, 2.334]	-2.5 [-20.3, 15.4]	0.803
		Irbesartan	48	10 (20.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	4 (15.4)	1.282 [0.318, 5.161]	1.333 [0.267, 6.666]	3.4 [-19.4, 26.1]	1.000
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.825
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	26 (31.7)	1.208 [0.743, 1.963]	1.304 [0.660, 2.579]	5.5 [-9.7, 20.6]	0.491
		Irbesartan	80	21 (26.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	10 (22.2)	1.556 [0.647, 3.739]	1.714 [0.591, 4.973]	7.9 [-9.8, 25.7]	0.423
		Irbesartan	49	7 (14.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	9 (18.4)	0.882 [0.393, 1.977]	0.855 [0.313, 2.334]	-2.5 [-20.3, 15.4]	0.803
		Irbesartan	48	10 (20.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	4 (15.4)	1.282 [0.318, 5.161]	1.333 [0.267, 6.666]	3.4 [-19.4, 26.1]	1.000
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.200
<= 1.75 g/day	Double-blind period	Sparsentan	98	21 (21.4)	1.679 [0.876, 3.217]	1.864 [0.859, 4.043]	8.7 [-2.9, 20.3]	0.128
		Irbesartan	94	12 (12.8)				
> 1.75 g/day	Double-blind period	Sparsentan	104	28 (26.9)	1.003 [0.643, 1.563]	1.004 [0.547, 1.842]	0.1 [-12.8, 13.0]	1.000
		Irbesartan	108	29 (26.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.080
Yes	Double-blind period	Sparsentan	88	26 (29.5)	1.752 [0.984, 3.117]	2.067 [0.991, 4.309]	12.7 [-1.0, 26.3]	0.070
		Irbesartan	83	14 (16.9)				
No	Double-blind period	Sparsentan	114	23 (20.2)	0.889 [0.543, 1.456]	0.861 [0.460, 1.612]	-2.5 [-13.9, 8.9]	0.750
		Irbesartan	119	27 (22.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.072
<= 5 years	Double-blind period	Sparsentan	113	31 (27.4)	1.584 [0.976, 2.570]	1.804 [0.973, 3.348]	10.1 [-1.3, 21.5]	0.063
		Irbesartan	127	22 (17.3)				
> 5 years	Double-blind period	Sparsentan	89	18 (20.2)	0.798 [0.453, 1.407]	0.747 [0.359, 1.556]	-5.1 [-19.2, 9.0]	0.459
		Irbesartan	75	19 (25.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.009 #
Yes	Double-blind period	Sparsentan	153	44 (28.8)	1.557 [1.031, 2.352]	1.782 [1.045, 3.039]	10.3 [0.2, 20.3]	0.044 *
		Irbesartan	157	29 (18.5)				
No	Double-blind period	Sparsentan	49	5 (10.2)	0.383 [0.146, 1.001]	0.313 [0.100, 0.974]	-16.5 [-34.0, 1.1]	0.059
		Irbesartan	45	12 (26.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
 Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.683
Male	Double-blind period	Sparsentan	139	10 (7.2)	1.029 [0.442, 2.395]	1.031 [0.415, 2.560]	0.2 [-6.5, 6.9]	1.000
		Irbesartan	143	10 (7.0)				
Female	Double-blind period	Sparsentan	63	4 (6.3)	0.749 [0.211, 2.657]	0.732 [0.187, 2.869]	-2.1 [-13.1, 8.8]	0.738
		Irbesartan	59	5 (8.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
 Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.718
<= 45 years	Double-blind period	Sparsentan	96	10 (10.4)	1.031 [0.450, 2.366]	1.035 [0.410, 2.611]	0.3 [-9.2, 9.9]	1.000
		Irbesartan	99	10 (10.1)				
> 45 years	Double-blind period	Sparsentan	106	4 (3.8)	0.777 [0.215, 2.814]	0.769 [0.201, 2.946]	-1.1 [-7.6, 5.4]	0.746
		Irbesartan	103	5 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.690
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	3.000 + [0.171, 52.527]	3.667 + [0.145, 92.653]	22.2 [-20.5, 64.9]	0.505
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	7 (6.9)	0.831 [0.321, 2.149]	0.819 [0.293, 2.286]	-1.4 [-9.5, 6.7]	0.798
		Irbesartan	109	9 (8.3)				
> 40 years	Double-blind period	Sparsentan	91	5 (5.5)	0.806 [0.255, 2.545]	0.795 [0.233, 2.704]	-1.3 [-9.5, 6.8]	0.764
		Irbesartan	88	6 (6.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
 Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.303
North America	Double-blind period	Sparsentan	35	5 (14.3)	2.190 [0.561, 8.553]	2.389 [0.530, 10.764]	7.8 [-8.4, 23.9]	0.282
		Irbesartan	46	3 (6.5)				
Europe	Double-blind period	Sparsentan	98	7 (7.1)	0.913 [0.353, 2.361]	0.906 [0.325, 2.529]	-0.7 [-8.7, 7.3]	1.000
		Irbesartan	115	9 (7.8)				
Asia Pacific	Double-blind period	Sparsentan	69	2 (2.9)	0.396 [0.069, 2.273]	0.378 [0.060, 2.364]	-4.4 [-15.3, 6.4]	0.359
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.740
< 27 kg/m**2	Double-blind period	Sparsentan	84	5 (6.0)	0.799 [0.264, 2.424]	0.787 [0.240, 2.579]	-1.5 [-10.0, 7.0]	0.771
		Irbesartan	94	7 (7.4)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	9 (7.6)	1.020 [0.408, 2.549]	1.022 [0.379, 2.751]	0.2 [-7.7, 8.0]	1.000
		Irbesartan	107	8 (7.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
 Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.726
eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	1.390 [0.508, 3.805]	1.439 [0.473, 4.378]	3.2 [-7.9, 14.2]	0.582
		Irbesartan	74	6 (8.1)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)	0.500 [0.095, 2.618]	0.481 [0.084, 2.742]	-3.6 [-13.9, 6.6]	0.679
		Irbesartan	55	4 (7.3)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4)	0.649 [0.115, 3.656]	0.629 [0.099, 4.003]	-2.9 [-17.3, 11.4]	0.674
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)	0.949 [0.141, 6.392]	0.946 [0.126, 7.086]	-0.3 [-13.0, 12.4]	1.000
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.506
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	10 (7.9)	0.846 [0.379, 1.889]	0.833 [0.347, 2.004]	-1.4 [-9.1, 6.2]	0.824
		Irbesartan	129	12 (9.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.653 [0.114, 3.737]	0.638 [0.102, 4.000]	-2.2 [-13.0, 8.7]	0.678
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)	4.815 + [0.243, 95.576]	5.204 + [0.238, 113.979]	7.7 [-6.5, 21.9]	0.490
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.715
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	6 (7.3)	0.836 [0.294, 2.380]	0.823 [0.264, 2.566]	-1.4 [-11.0, 8.2]	0.780
		Irbesartan	80	7 (8.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	0.871 [0.249, 3.043]	0.859 [0.216, 3.419]	-1.3 [-15.3, 12.7]	1.000
		Irbesartan	49	5 (10.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.653 [0.114, 3.737]	0.638 [0.102, 4.000]	-2.2 [-13.0, 8.7]	0.678
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)	4.815 + [0.243, 95.576]	5.204 + [0.238, 113.979]	7.7 [-6.5, 21.9]	0.490
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.386
<= 1.75 g/day	Double-blind period	Sparsentan	98	5 (5.1)	1.599 [0.393, 6.503]	1.631 [0.379, 7.024]	1.9 [-4.8, 8.6]	0.721
		Irbesartan	94	3 (3.2)				
> 1.75 g/day	Double-blind period	Sparsentan	104	9 (8.7)	0.779 [0.343, 1.771]	0.758 [0.305, 1.882]	-2.5 [-11.4, 6.5]	0.648
		Irbesartan	108	12 (11.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.723
Yes	Double-blind period	Sparsentan	88	6 (6.8)	0.808	0.794	-1.6	0.777
		Irbesartan	83	7 (8.4)	[0.283, 2.306]	[0.256, 2.470]	[-10.8, 7.5]	
No	Double-blind period	Sparsentan	114	8 (7.0)	1.044	1.047	0.3	1.000
		Irbesartan	119	8 (6.7)	[0.405, 2.688]	[0.379, 2.891]	[-7.1, 7.7]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
 Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.538
<= 5 years	Double-blind period	Sparsentan	113	9 (8.0)	0.843 [0.369, 1.926]	0.829 [0.336, 2.048]	-1.5 [-9.4, 6.5]	0.820
		Irbesartan	127	12 (9.4)				
> 5 years	Double-blind period	Sparsentan	89	5 (5.6)	1.404 [0.347, 5.684]	1.429 [0.330, 6.186]	1.6 [-6.1, 9.4]	0.728
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
 Safety Set

Severe TEAEs					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.282
Yes	Double-blind period	Sparsentan	153	13 (8.5)	1.112 [0.524, 2.359]	1.122 [0.495, 2.543]	0.9 [-5.9, 7.6]	0.837
		Irbesartan	157	12 (7.6)				
No	Double-blind period	Sparsentan	49	1 (2.0)	0.306 [0.033, 2.838]	0.292 [0.029, 2.911]	-4.6 [-15.1, 5.8]	0.346
		Irbesartan	45	3 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.847
Male	Double-blind period	Sparsentan	139	19 (13.7)	0.931 [0.524, 1.654]	0.920 [0.471, 1.797]	-1.0 [-9.9, 7.8]	0.865
		Irbesartan	143	21 (14.7)				
Female	Double-blind period	Sparsentan	63	9 (14.3)	0.843 [0.368, 1.928]	0.817 [0.306, 2.176]	-2.7 [-17.2, 11.9]	0.804
		Irbesartan	59	10 (16.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.801
<= 45 years	Double-blind period	Sparsentan	96	15 (15.6)	0.859 [0.460, 1.606]	0.833 [0.393, 1.766]	-2.6 [-14.1, 9.0]	0.704
		Irbesartan	99	18 (18.2)				
> 45 years	Double-blind period	Sparsentan	106	13 (12.3)	0.972 [0.473, 1.995]	0.968 [0.426, 2.201]	-0.4 [-10.3, 9.6]	1.000
		Irbesartan	103	13 (12.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.461
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	3.000 + [0.171, 52.527]	3.667 + [0.145, 92.653]	22.2 [-20.5, 64.9]	0.505
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	14 (13.7)	1.069 [0.536, 2.130]	1.080 [0.487, 2.392]	0.9 [-9.2, 11.0]	1.000
		Irbesartan	109	14 (12.8)				
> 40 years	Double-blind period	Sparsentan	91	12 (13.2)	0.683 [0.346, 1.345]	0.634 [0.283, 1.420]	-6.1 [-18.0, 5.8]	0.313
		Irbesartan	88	17 (19.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.680
North America	Double-blind period	Sparsentan	35	6 (17.1)	1.127 [0.415, 3.056]	1.153 [0.350, 3.795]	1.9 [-16.8, 20.7]	1.000
		Irbesartan	46	7 (15.2)				
Europe	Double-blind period	Sparsentan	98	10 (10.2)	0.690 [0.332, 1.437]	0.655 [0.285, 1.506]	-4.6 [-14.4, 5.2]	0.409
		Irbesartan	115	17 (14.8)				
Asia Pacific	Double-blind period	Sparsentan	69	12 (17.4)	1.019 [0.436, 2.379]	1.023 [0.367, 2.848]	0.3 [-16.2, 16.8]	1.000
		Irbesartan	41	7 (17.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.996
< 27 kg/m**2	Double-blind period	Sparsentan	84	13 (15.5)	0.909 [0.465, 1.777]	0.893 [0.401, 1.985]	-1.5 [-13.5, 10.4]	0.841
		Irbesartan	94	16 (17.0)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	15 (12.7)	0.907 [0.466, 1.765]	0.893 [0.414, 1.927]	-1.3 [-11.1, 8.5]	0.845
		Irbesartan	107	15 (14.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.382
eGFR Low and UP High	Double-blind period	Sparsentan	71	12 (16.9)	0.834 [0.420, 1.655]	0.800 [0.345, 1.854]	-3.4 [-17.4, 10.7]	0.672
		Irbesartan	74	15 (20.3)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	8 (14.5)	1.333 [0.495, 3.589]	1.390 [0.448, 4.310]	3.6 [-10.6, 17.9]	0.776
		Irbesartan	55	6 (10.9)				
eGFR High and UP High	Double-blind period	Sparsentan	37	6 (16.2)	1.459 [0.449, 4.745]	1.548 [0.398, 6.022]	5.1 [-13.3, 23.5]	0.736
		Irbesartan	36	4 (11.1)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)	0.316 [0.068, 1.469]	0.279 [0.053, 1.483]	-11.1 [-27.5, 5.3]	0.148
		Irbesartan	37	6 (16.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.694
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	20 (15.7)	0.813 [0.476, 1.387]	0.778 [0.407, 1.485]	-3.6 [-13.7, 6.5]	0.512
		Irbesartan	129	25 (19.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	7 (14.3)	1.371 [0.467, 4.024]	1.433 [0.422, 4.874]	3.9 [-11.3, 19.0]	0.759
		Irbesartan	48	5 (10.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.694
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	14 (17.1)	0.976 [0.497, 1.914]	0.971 [0.430, 2.192]	-0.4 [-13.3, 12.5]	1.000
		Irbesartan	80	14 (17.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	6 (13.3)	0.594 [0.239, 1.474]	0.531 [0.179, 1.581]	-9.1 [-26.6, 8.3]	0.293
		Irbesartan	49	11 (22.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	7 (14.3)	1.371 [0.467, 4.024]	1.433 [0.422, 4.874]	3.9 [-11.3, 19.0]	0.759
		Irbesartan	48	5 (10.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.186
<= 1.75 g/day	Double-blind period	Sparsentan	98	14 (14.3)	1.343 [0.628, 2.873]	1.400 [0.589, 3.328]	3.6 [-6.7, 14.0]	0.516
		Irbesartan	94	10 (10.6)				
> 1.75 g/day	Double-blind period	Sparsentan	104	14 (13.5)	0.692 [0.372, 1.287]	0.644 [0.308, 1.348]	-6.0 [-16.9, 4.9]	0.270
		Irbesartan	108	21 (19.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.484
Yes	Double-blind period	Sparsentan	88	11 (12.5)	0.741 [0.357, 1.539]	0.704 [0.300, 1.654]	-4.4 [-16.2, 7.4]	0.517
		Irbesartan	83	14 (16.9)				
No	Double-blind period	Sparsentan	114	17 (14.9)	1.044 [0.561, 1.943]	1.052 [0.508, 2.177]	0.6 [-9.3, 10.6]	1.000
		Irbesartan	119	17 (14.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.040 #
<= 5 years	Double-blind period	Sparsentan	113	14 (12.4)	0.629 [0.344, 1.150]	0.577 [0.284, 1.174]	-7.3 [-17.3, 2.7]	0.161
		Irbesartan	127	25 (19.7)				
> 5 years	Double-blind period	Sparsentan	89	14 (15.7)	1.966 [0.795, 4.864]	2.147 [0.781, 5.898]	7.7 [-3.2, 18.7]	0.156
		Irbesartan	75	6 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.704
Yes	Double-blind period	Sparsentan	153	22 (14.4)	0.868 [0.515, 1.464]	0.846 [0.456, 1.569]	-2.2 [-10.9, 6.5]	0.640
		Irbesartan	157	26 (16.6)				
No	Double-blind period	Sparsentan	49	6 (12.2)	1.102 [0.361, 3.363]	1.116 [0.316, 3.945]	1.1 [-14.0, 16.2]	1.000
		Irbesartan	45	5 (11.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.312
Male	Double-blind period	Sparsentan	139	10 (7.2)	2.572 [0.826, 8.008]	2.694 [0.824, 8.802]	4.4 [-1.4, 10.2]	0.105
		Irbesartan	143	4 (2.8)				
Female	Double-blind period	Sparsentan	63	6 (9.5)	1.124 [0.362, 3.487]	1.137 [0.328, 3.943]	1.0 [-10.7, 12.8]	1.000
		Irbesartan	59	5 (8.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.196
<= 45 years	Double-blind period	Sparsentan	96	8 (8.3)	1.179 [0.445, 3.124]	1.195 [0.416, 3.434]	1.3 [-7.3, 9.8]	0.793
		Irbesartan	99	7 (7.1)				
> 45 years	Double-blind period	Sparsentan	106	8 (7.5)	3.887 [0.845, 17.871]	4.122 [0.854, 19.899]	5.6 [-1.0, 12.3]	0.101
		Irbesartan	103	2 (1.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
 Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.976
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	9 (8.8)	1.603 [0.591, 4.345]	1.661 [0.570, 4.845]	3.3 [-4.6, 11.2]	0.426
		Irbesartan	109	6 (5.5)				
> 40 years	Double-blind period	Sparsentan	91	6 (6.6)	1.934 [0.499, 7.495]	2.000 [0.484, 8.259]	3.2 [-4.3, 10.7]	0.497
		Irbesartan	88	3 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
 Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.675
North America	Double-blind period	Sparsentan	35	5 (14.3)	3.286 [0.677, 15.949]	3.667 [0.667, 20.156]	9.9 [-5.6, 25.5]	0.230
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	6 (6.1)	1.408 [0.443, 4.473]	1.435 [0.424, 4.854]	1.8 [-5.2, 8.8]	0.758
		Irbesartan	115	5 (4.3)				
Asia Pacific	Double-blind period	Sparsentan	69	5 (7.2)	1.486 [0.302, 7.311]	1.523 [0.282, 8.235]	2.4 [-8.6, 13.3]	1.000
		Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.823
< 27 kg/m**2	Double-blind period	Sparsentan	84	7 (8.3)	1.958 [0.594, 6.455]	2.045 [0.577, 7.252]	4.1 [-4.2, 12.4]	0.353
		Irbesartan	94	4 (4.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	9 (7.6)	1.632 [0.565, 4.718]	1.684 [0.546, 5.194]	3.0 [-4.2, 10.1]	0.417
		Irbesartan	107	5 (4.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
 Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.305
eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	2.085 [0.657, 6.618]	2.222 [0.638, 7.737]	5.9 [-4.5, 16.2]	0.238
		Irbesartan	74	4 (5.4)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	4 (7.3)	9.000 + [0.496, 163.268]	9.699 + [0.510, 184.615]	7.3 [-1.4, 16.0]	0.118
		Irbesartan	55	0 (0.0)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4)	0.486 [0.095, 2.493]	0.457 [0.078, 2.667]	-5.7 [-21.0, 9.6]	0.430
		Irbesartan	36	4 (11.1)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)	1.897 [0.180, 20.054]	1.946 [0.169, 22.413]	2.4 [-8.9, 13.7]	1.000
		Irbesartan	37	1 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
 Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.281
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	12 (9.4)	3.047 [1.010, 9.198]	3.261 [1.023, 10.398]	6.3 [-0.3, 13.0]	0.041 *
		Irbesartan	129	4 (3.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	3 (6.1)	0.735 [0.174, 3.110]	0.717 [0.152, 3.390]	-2.2 [-14.6, 10.2]	0.715
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.420
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	7 (8.5)	2.276 [0.610, 8.496]	2.396 [0.597, 9.612]	4.8 [-3.8, 13.4]	0.328
		Irbesartan	80	3 (3.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	5 (11.1)	5.444 [0.661, 44.842]	6.000 [0.673, 53.486]	9.1 [-3.1, 21.2]	0.101
		Irbesartan	49	1 (2.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	3 (6.1)	0.735 [0.174, 3.110]	0.717 [0.152, 3.390]	-2.2 [-14.6, 10.2]	0.715
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.200
<= 1.75 g/day	Double-blind period	Sparsentan	98	6 (6.1)	5.755 [0.706, 46.904]	6.065 [0.716, 51.372]	5.1 [-1.2, 11.3]	0.119
		Irbesartan	94	1 (1.1)				
> 1.75 g/day	Double-blind period	Sparsentan	104	10 (9.6)	1.298 [0.533, 3.160]	1.330 [0.503, 3.513]	2.2 [-6.3, 10.7]	0.628
		Irbesartan	108	8 (7.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.647
Yes	Double-blind period	Sparsentan	88	9 (10.2)	2.122 [0.679, 6.629]	2.250 [0.665, 7.609]	5.4 [-3.6, 14.4]	0.251
		Irbesartan	83	4 (4.8)				
No	Double-blind period	Sparsentan	114	7 (6.1)	1.461 [0.478, 4.472]	1.492 [0.459, 4.842]	1.9 [-4.6, 8.5]	0.564
		Irbesartan	119	5 (4.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.901
<= 5 years	Double-blind period	Sparsentan	113	10 (8.8)	1.873 [0.703, 4.990]	1.958 [0.688, 5.571]	4.1 [-3.1, 11.4]	0.300
		Irbesartan	127	6 (4.7)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	1.685 [0.436, 6.511]	1.735 [0.419, 7.188]	2.7 [-5.3, 10.8]	0.510
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
 Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.248
Yes	Double-blind period	Sparsentan	153	12 (7.8)	2.463 [0.889, 6.824]	2.587 [0.889, 7.529]	4.7 [-1.1, 10.4]	0.084
		Irbesartan	157	5 (3.2)				
No	Double-blind period	Sparsentan	49	4 (8.2)	0.918 [0.244, 3.457]	0.911 [0.214, 3.881]	-0.7 [-14.2, 12.7]	1.000
		Irbesartan	45	4 (8.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AD\_SSIM: Incidence of fatal TEAEs during double-blind period by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.764
Male	Double-blind period	Sparsentan	139	112 (80.6)	1.130 [0.990, 1.289]	1.667 [0.957, 2.904]	9.2 [-1.4, 19.9]	0.072
		Irbesartan	143	102 (71.3)				
Female	Double-blind period	Sparsentan	63	50 (79.4)	1.089 [0.891, 1.330]	1.431 [0.619, 3.307]	6.5 [-10.3, 23.2]	0.524
		Irbesartan	59	43 (72.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.872
<= 45 years	Double-blind period	Sparsentan	96	81 (84.4)	1.129 [0.978, 1.303]	1.824 [0.894, 3.724]	9.6 [-2.6, 21.9]	0.112
		Irbesartan	99	74 (74.7)				
> 45 years	Double-blind period	Sparsentan	106	81 (76.4)	1.109 [0.938, 1.310]	1.460 [0.791, 2.694]	7.5 [-5.5, 20.5]	0.277
		Irbesartan	103	71 (68.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.920
<= 18 years	Double-blind period	Sparsentan	9	7 (77.8)	1.296 [0.585, 2.874]	2.333 [0.216, 25.245]	17.8 [-48.6, 84.1]	0.580
		Irbesartan	5	3 (60.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	86 (84.3)	1.107 [0.968, 1.266]	1.684 [0.843, 3.363]	8.2 [-3.5, 19.8]	0.168
		Irbesartan	109	83 (76.1)				
> 40 years	Double-blind period	Sparsentan	91	69 (75.8)	1.131 [0.938, 1.363]	1.542 [0.801, 2.965]	8.8 [-5.5, 23.1]	0.246
		Irbesartan	88	59 (67.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.182
North America	Double-blind period	Sparsentan	35	27 (77.1)	1.044 [0.814, 1.339]	1.191 [0.426, 3.328]	3.2 [-18.1, 24.6]	0.799
		Irbesartan	46	34 (73.9)				
Europe	Double-blind period	Sparsentan	98	78 (79.6)	1.204 [1.021, 1.420]	2.001 [1.071, 3.738]	13.5 [0.8, 26.2]	0.032 *
		Irbesartan	115	76 (66.1)				
Asia Pacific	Double-blind period	Sparsentan	69	57 (82.6)	0.968 [0.819, 1.143]	0.814 [0.280, 2.366]	-2.8 [-18.7, 13.2]	0.794
		Irbesartan	41	35 (85.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.998
< 27 kg/m**2	Double-blind period	Sparsentan	84	67 (79.8)	1.119 [0.946, 1.323]	1.588 [0.793, 3.182]	8.5 [-5.2, 22.2]	0.225
		Irbesartan	94	67 (71.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	95 (80.5)	1.119 [0.965, 1.297]	1.609 [0.865, 2.994]	8.5 [-3.5, 20.6]	0.157
		Irbesartan	107	77 (72.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.233
eGFR Low and UP High	Double-blind period	Sparsentan	71	60 (84.5)	1.180 [0.991, 1.405]	2.161 [0.954, 4.896]	12.9 [-1.8, 27.5]	0.073
		Irbesartan	74	53 (71.6)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	41 (74.5)	0.976 [0.789, 1.208]	0.906 [0.380, 2.161]	-1.8 [-19.7, 16.1]	1.000
		Irbesartan	55	42 (76.4)				
eGFR High and UP High	Double-blind period	Sparsentan	37	31 (83.8)	1.371 [1.019, 1.844]	3.288 [1.093, 9.892]	22.7 [0.1, 45.3]	0.038 *
		Irbesartan	36	22 (61.1)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	30 (76.9)	1.016 [0.791, 1.306]	1.071 [0.372, 3.086]	1.2 [-20.5, 23.0]	1.000
		Irbesartan	37	28 (75.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.828
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	103 (81.1)	1.090 [0.956, 1.243]	1.475 [0.814, 2.674]	6.7 [-4.2, 17.6]	0.230
		Irbesartan	129	96 (74.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	39 (79.6)	1.158 [0.913, 1.468]	1.773 [0.703, 4.469]	10.8 [-8.5, 30.2]	0.252
		Irbesartan	48	33 (68.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	20 (76.9)	1.202 [0.837, 1.726]	1.875 [0.551, 6.379]	12.9 [-15.8, 41.7]	0.368
		Irbesartan	25	16 (64.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.866
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	65 (79.3)	1.057 [0.893, 1.250]	1.275 [0.611, 2.660]	4.3 [-9.9, 18.4]	0.577
		Irbesartan	80	60 (75.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	38 (84.4)	1.149 [0.932, 1.418]	1.960 [0.703, 5.467]	11.0 [-7.4, 29.4]	0.217
		Irbesartan	49	36 (73.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	39 (79.6)	1.158 [0.913, 1.468]	1.773 [0.703, 4.469]	10.8 [-8.5, 30.2]	0.252
		Irbesartan	48	33 (68.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	20 (76.9)	1.202 [0.837, 1.726]	1.875 [0.551, 6.379]	12.9 [-15.8, 41.7]	0.368
		Irbesartan	25	16 (64.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.360
<= 1.75 g/day	Double-blind period	Sparsentan	98	75 (76.5)	1.058 [0.896, 1.249]	1.247 [0.651, 2.388]	4.2 [-9.2, 17.6]	0.513
		Irbesartan	94	68 (72.3)				
> 1.75 g/day	Double-blind period	Sparsentan	104	87 (83.7)	1.173 [1.013, 1.359]	2.060 [1.058, 4.011]	12.4 [0.3, 24.4]	0.034 *
		Irbesartan	108	77 (71.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.100
Yes	Double-blind period	Sparsentan	88	67 (76.1)	1.003 [0.848, 1.187]	1.013 [0.502, 2.044]	0.2 [-13.7, 14.2]	1.000
		Irbesartan	83	63 (75.9)				
No	Double-blind period	Sparsentan	114	95 (83.3)	1.209 [1.045, 1.399]	2.256 [1.205, 4.224]	14.4 [2.8, 26.1]	0.014 *
		Irbesartan	119	82 (68.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.395
<= 5 years	Double-blind period	Sparsentan	113	90 (79.6)	1.076 [0.936, 1.236]	1.374 [0.750, 2.518]	5.6 [-5.8, 17.1]	0.360
		Irbesartan	127	94 (74.0)				
> 5 years	Double-blind period	Sparsentan	89	72 (80.9)	1.190 [0.989, 1.432]	1.993 [0.973, 4.084]	12.9 [-1.7, 27.5]	0.071
		Irbesartan	75	51 (68.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.822
Yes	Double-blind period	Sparsentan	153	121 (79.1)	1.109 [0.975, 1.260]	1.519 [0.902, 2.558]	7.7 [-2.5, 18.0]	0.117
		Irbesartan	157	112 (71.3)				
No	Double-blind period	Sparsentan	49	41 (83.7)	1.141 [0.920, 1.415]	1.864 [0.682, 5.092]	10.3 [-8.3, 29.0]	0.313
		Irbesartan	45	33 (73.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.705
Male	Double-blind period	Sparsentan	139	112 (80.6)	1.141 [0.998, 1.304]	1.725 [0.992, 3.000]	9.9 [-0.7, 20.6]	0.054
		Irbesartan	143	101 (70.6)				
Female	Double-blind period	Sparsentan	63	50 (79.4)	1.089 [0.891, 1.330]	1.431 [0.619, 3.307]	6.5 [-10.3, 23.2]	0.524
		Irbesartan	59	43 (72.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.972
<= 45 years	Double-blind period	Sparsentan	96	81 (84.4)	1.129 [0.978, 1.303]	1.824 [0.894, 3.724]	9.6 [-2.6, 21.9]	0.112
		Irbesartan	99	74 (74.7)				
> 45 years	Double-blind period	Sparsentan	106	81 (76.4)	1.124 [0.949, 1.332]	1.527 [0.830, 2.812]	8.5 [-4.6, 21.5]	0.216
		Irbesartan	103	70 (68.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.892
<= 18 years	Double-blind period	Sparsentan	9	7 (77.8)	1.296 [0.585, 2.874]	2.333 [0.216, 25.245]	17.8 [-48.6, 84.1]	0.580
		Irbesartan	5	3 (60.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	86 (84.3)	1.107 [0.968, 1.266]	1.684 [0.843, 3.363]	8.2 [-3.5, 19.8]	0.168
		Irbesartan	109	83 (76.1)				
> 40 years	Double-blind period	Sparsentan	91	69 (75.8)	1.150 [0.952, 1.391]	1.622 [0.846, 3.112]	9.9 [-4.4, 24.3]	0.188
		Irbesartan	88	58 (65.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.150
North America	Double-blind period	Sparsentan	35	27 (77.1)	1.044 [0.814, 1.339]	1.191 [0.426, 3.328]	3.2 [-18.1, 24.6]	0.799
		Irbesartan	46	34 (73.9)				
Europe	Double-blind period	Sparsentan	98	78 (79.6)	1.220 [1.033, 1.442]	2.080 [1.115, 3.880]	14.4 [1.6, 27.1]	0.022 *
		Irbesartan	115	75 (65.2)				
Asia Pacific	Double-blind period	Sparsentan	69	57 (82.6)	0.968 [0.819, 1.143]	0.814 [0.280, 2.366]	-2.8 [-18.7, 13.2]	0.794
		Irbesartan	41	35 (85.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.911
< 27 kg/m**2	Double-blind period	Sparsentan	84	67 (79.8)	1.119 [0.946, 1.323]	1.588 [0.793, 3.182]	8.5 [-5.2, 22.2]	0.225
		Irbesartan	94	67 (71.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	95 (80.5)	1.133 [0.976, 1.317]	1.685 [0.908, 3.126]	9.5 [-2.6, 21.6]	0.118
		Irbesartan	107	76 (71.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.269
eGFR Low and UP High	Double-blind period	Sparsentan	71	60 (84.5)	1.180 [0.991, 1.405]	2.161 [0.954, 4.896]	12.9 [-1.8, 27.5]	0.073
		Irbesartan	74	53 (71.6)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	41 (74.5)	0.976 [0.789, 1.208]	0.906 [0.380, 2.161]	-1.8 [-19.7, 16.1]	1.000
		Irbesartan	55	42 (76.4)				
eGFR High and UP High	Double-blind period	Sparsentan	37	31 (83.8)	1.371 [1.019, 1.844]	3.288 [1.093, 9.892]	22.7 [0.1, 45.3]	0.038 *
		Irbesartan	36	22 (61.1)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	30 (76.9)	1.054 [0.812, 1.368]	1.235 [0.436, 3.492]	4.0 [-18.2, 26.1]	0.793
		Irbesartan	37	27 (73.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.869
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	103 (81.1)	1.101 [0.964, 1.258]	1.536 [0.850, 2.777]	7.5 [-3.5, 18.4]	0.180
		Irbesartan	129	95 (73.6)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	39 (79.6)	1.158 [0.913, 1.468]	1.773 [0.703, 4.469]	10.8 [-8.5, 30.2]	0.252
		Irbesartan	48	33 (68.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	20 (76.9)	1.202 [0.837, 1.726]	1.875 [0.551, 6.379]	12.9 [-15.8, 41.7]	0.368
		Irbesartan	25	16 (64.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.822
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	65 (79.3)	1.057 [0.893, 1.250]	1.275 [0.611, 2.660]	4.3 [-9.9, 18.4]	0.577
		Irbesartan	80	60 (75.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	38 (84.4)	1.182 [0.952, 1.469]	2.171 [0.785, 6.003]	13.0 [-5.6, 31.6]	0.146
		Irbesartan	49	35 (71.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	39 (79.6)	1.158 [0.913, 1.468]	1.773 [0.703, 4.469]	10.8 [-8.5, 30.2]	0.252
		Irbesartan	48	33 (68.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	20 (76.9)	1.202 [0.837, 1.726]	1.875 [0.551, 6.379]	12.9 [-15.8, 41.7]	0.368
		Irbesartan	25	16 (64.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.437
<= 1.75 g/day	Double-blind period	Sparsentan	98	75 (76.5)	1.074 [0.907, 1.271]	1.314 [0.688, 2.508]	5.3 [-8.2, 18.7]	0.417
		Irbesartan	94	67 (71.3)				
> 1.75 g/day	Double-blind period	Sparsentan	104	87 (83.7)	1.173 [1.013, 1.359]	2.060 [1.058, 4.011]	12.4 [0.3, 24.4]	0.034 *
		Irbesartan	108	77 (71.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.136
Yes	Double-blind period	Sparsentan	88	67 (76.1)	1.019 [0.859, 1.210]	1.081 [0.539, 2.169]	1.4 [-12.6, 15.5]	0.860
		Irbesartan	83	62 (74.7)				
No	Double-blind period	Sparsentan	114	95 (83.3)	1.209 [1.045, 1.399]	2.256 [1.205, 4.224]	14.4 [2.8, 26.1]	0.014 *
		Irbesartan	119	82 (68.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.316
<= 5 years	Double-blind period	Sparsentan	113	90 (79.6)	1.076 [0.936, 1.236]	1.374 [0.750, 2.518]	5.6 [-5.8, 17.1]	0.360
		Irbesartan	127	94 (74.0)				
> 5 years	Double-blind period	Sparsentan	89	72 (80.9)	1.213 [1.004, 1.466]	2.118 [1.037, 4.325]	14.2 [-0.4, 28.9]	0.048 *
		Irbesartan	75	50 (66.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.877
Yes	Double-blind period	Sparsentan	153	121 (79.1)	1.119 [0.983, 1.273]	1.567 [0.932, 2.635]	8.4 [-1.9, 18.6]	0.116
		Irbesartan	157	111 (70.7)				
No	Double-blind period	Sparsentan	49	41 (83.7)	1.141 [0.920, 1.415]	1.864 [0.682, 5.092]	10.3 [-8.3, 29.0]	0.313
		Irbesartan	45	33 (73.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.566
Male	Double-blind period	Sparsentan	139	8 (5.8)	1.176 [0.438, 3.155]	1.186 [0.418, 3.365]	0.9 [-5.1, 6.8]	0.796
		Irbesartan	143	7 (4.9)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	0.702 [0.164, 3.007]	0.688 [0.147, 3.210]	-2.0 [-12.0, 7.9]	0.711
		Irbesartan	59	4 (6.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.945
<= 45 years	Double-blind period	Sparsentan	96	7 (7.3)	1.031 [0.376, 2.829]	1.034 [0.348, 3.067]	0.2 [-8.1, 8.5]	1.000
		Irbesartan	99	7 (7.1)				
> 45 years	Double-blind period	Sparsentan	106	4 (3.8)	0.972 [0.250, 3.783]	0.971 [0.236, 3.988]	-0.1 [-6.3, 6.1]	1.000
		Irbesartan	103	4 (3.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.868
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	5 (4.9)	1.069 [0.319, 3.583]	1.072 [0.301, 3.818]	0.3 [-6.4, 7.0]	1.000
		Irbesartan	109	5 (4.6)				
> 40 years	Double-blind period	Sparsentan	91	5 (5.5)	0.806 [0.255, 2.545]	0.795 [0.233, 2.704]	-1.3 [-9.5, 6.8]	0.764
		Irbesartan	88	6 (6.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.434
North America	Double-blind period	Sparsentan	35	5 (14.3)	2.190 [0.561, 8.553]	2.389 [0.530, 10.764]	7.8 [-8.4, 23.9]	0.282
		Irbesartan	46	3 (6.5)				
Europe	Double-blind period	Sparsentan	98	4 (4.1)	0.782 [0.227, 2.693]	0.773 [0.212, 2.822]	-1.1 [-7.7, 5.5]	0.756
		Irbesartan	115	6 (5.2)				
Asia Pacific	Double-blind period	Sparsentan	69	2 (2.9)	0.594 [0.087, 4.059]	0.582 [0.079, 4.298]	-2.0 [-11.6, 7.7]	0.628
		Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.808
< 27 kg/m**2	Double-blind period	Sparsentan	84	4 (4.8)	1.119 [0.289, 4.335]	1.125 [0.272, 4.646]	0.5 [-6.7, 7.7]	1.000
		Irbesartan	94	4 (4.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	7 (5.9)	0.907 [0.329, 2.501]	0.901 [0.305, 2.658]	-0.6 [-7.8, 6.6]	1.000
		Irbesartan	107	7 (6.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	5 (7.0)				NE
		Irbesartan	74	4 (5.4)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)				NE
		Irbesartan	55	3 (5.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4)				NE
		Irbesartan	36	2 (5.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)				NE
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.528
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	7 (5.5)	0.790 [0.303, 2.057]	0.778 [0.281, 2.156]	-1.5 [-8.2, 5.2]	0.797
		Irbesartan	129	9 (7.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.980 [0.144, 6.676]	0.979 [0.132, 7.244]	-0.1 [-10.1, 9.9]	1.000
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)	4.815 + [0.243, 95.576]	5.204 + [0.238, 113.979]	7.7 [-6.5, 21.9]	0.490
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	3 (3.7)				NE
		Irbesartan	80	4 (5.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)				NE
		Irbesartan	49	5 (10.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)				NE
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)				NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.417
<= 1.75 g/day	Double-blind period	Sparsentan	98	5 (5.1)	1.599 [0.393, 6.503]	1.631 [0.379, 7.024]	1.9 [-4.8, 8.6]	0.721
		Irbesartan	94	3 (3.2)				
> 1.75 g/day	Double-blind period	Sparsentan	104	6 (5.8)	0.779 [0.280, 2.168]	0.765 [0.256, 2.287]	-1.6 [-9.3, 6.0]	0.784
		Irbesartan	108	8 (7.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.310
Yes	Double-blind period	Sparsentan	88	3 (3.4)	0.566 [0.140, 2.294]	0.551 [0.127, 2.380]	-2.6 [-10.2, 4.9]	0.487
		Irbesartan	83	5 (6.0)				
No	Double-blind period	Sparsentan	114	8 (7.0)	1.392 [0.498, 3.886]	1.421 [0.477, 4.233]	2.0 [-5.0, 9.0]	0.589
		Irbesartan	119	6 (5.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.227
<= 5 years	Double-blind period	Sparsentan	113	7 (6.2)	0.787 [0.310, 1.998]	0.773 [0.284, 2.102]	-1.7 [-9.0, 5.6]	0.802
		Irbesartan	127	10 (7.9)				
> 5 years	Double-blind period	Sparsentan	89	4 (4.5)	3.371 [0.385, 29.512]	3.482 [0.381, 31.851]	3.2 [-3.1, 9.4]	0.377
		Irbesartan	75	1 (1.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.229
Yes	Double-blind period	Sparsentan	153	11 (7.2)	1.254 [0.535, 2.941]	1.274 [0.512, 3.166]	1.5 [-4.7, 7.6]	0.649
		Irbesartan	157	9 (5.7)				
No	Double-blind period	Sparsentan	49	0 (0.0)	0.184 + [0.009, 3.732]	0.176 + [0.008, 3.762]	-4.4 [-12.6, 3.7]	0.226
		Irbesartan	45	2 (4.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.878
Male	Double-blind period	Sparsentan	139	16 (11.5)	0.914 [0.486, 1.720]	0.903 [0.441, 1.852]	-1.1 [-9.4, 7.2]	0.856
		Irbesartan	143	18 (12.6)				
Female	Double-blind period	Sparsentan	63	9 (14.3)	0.843 [0.368, 1.928]	0.817 [0.306, 2.176]	-2.7 [-17.2, 11.9]	0.804
		Irbesartan	59	10 (16.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.550
<= 45 years	Double-blind period	Sparsentan	96	12 (12.5)	0.773 [0.386, 1.548]	0.741 [0.330, 1.662]	-3.7 [-14.5, 7.2]	0.542
		Irbesartan	99	16 (16.2)				
> 45 years	Double-blind period	Sparsentan	106	13 (12.3)	1.053 [0.504, 2.198]	1.060 [0.459, 2.446]	0.6 [-9.1, 10.4]	1.000
		Irbesartan	103	12 (11.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.680
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	12 (11.8)	1.069 [0.503, 2.270]	1.078 [0.461, 2.522]	0.8 [-8.8, 10.3]	1.000
		Irbesartan	109	12 (11.0)				
> 40 years	Double-blind period	Sparsentan	91	12 (13.2)	0.725 [0.364, 1.444]	0.684 [0.303, 1.542]	-5.0 [-16.8, 6.8]	0.413
		Irbesartan	88	16 (18.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.572
North America	Double-blind period	Sparsentan	35	6 (17.1)	1.127 [0.415, 3.056]	1.153 [0.350, 3.795]	1.9 [-16.8, 20.7]	1.000
		Irbesartan	46	7 (15.2)				
Europe	Double-blind period	Sparsentan	98	8 (8.2)	0.626 [0.277, 1.413]	0.593 [0.240, 1.464]	-4.9 [-14.0, 4.3]	0.277
		Irbesartan	115	15 (13.0)				
Asia Pacific	Double-blind period	Sparsentan	69	11 (15.9)	1.089 [0.436, 2.724]	1.106 [0.376, 3.256]	1.3 [-14.5, 17.1]	1.000
		Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.736
< 27 kg/m**2	Double-blind period	Sparsentan	84	11 (13.1)	0.821 [0.399, 1.686]	0.794 [0.342, 1.839]	-2.9 [-14.3, 8.6]	0.673
		Irbesartan	94	15 (16.0)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	14 (11.9)	0.977 [0.481, 1.983]	0.973 [0.435, 2.177]	-0.3 [-9.7, 9.1]	1.000
		Irbesartan	107	13 (12.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.363
eGFR Low and UP High	Double-blind period	Sparsentan	71	10 (14.1)	0.802 [0.376, 1.710]	0.769 [0.314, 1.887]	-3.5 [-16.7, 9.8]	0.652
		Irbesartan	74	13 (17.6)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	8 (14.5)	1.333 [0.495, 3.589]	1.390 [0.448, 4.310]	3.6 [-10.6, 17.9]	0.776
		Irbesartan	55	6 (10.9)				
eGFR High and UP High	Double-blind period	Sparsentan	37	5 (13.5)	1.622 [0.418, 6.292]	1.719 [0.379, 7.794]	5.2 [-11.8, 22.2]	0.711
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)	0.316 [0.068, 1.469]	0.279 [0.053, 1.483]	-11.1 [-27.5, 5.3]	0.148
		Irbesartan	37	6 (16.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.662
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	18 (14.2)	0.795 [0.451, 1.400]	0.761 [0.389, 1.491]	-3.7 [-13.4, 6.1]	0.496
		Irbesartan	129	23 (17.8)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	1.469 [0.442, 4.883]	1.535 [0.405, 5.822]	3.9 [-10.2, 18.0]	0.740
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.684
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	12 (14.6)	0.976 [0.466, 2.042]	0.971 [0.408, 2.312]	-0.4 [-12.5, 11.8]	1.000
		Irbesartan	80	12 (15.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	6 (13.3)	0.594 [0.239, 1.474]	0.531 [0.179, 1.581]	-9.1 [-26.6, 8.3]	0.293
		Irbesartan	49	11 (22.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	1.469 [0.442, 4.883]	1.535 [0.405, 5.822]	3.9 [-10.2, 18.0]	0.740
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.263
<= 1.75 g/day	Double-blind period	Sparsentan	98	13 (13.3)	1.247 [0.575, 2.705]	1.285 [0.534, 3.091]	2.6 [-7.6, 12.8]	0.659
		Irbesartan	94	10 (10.6)				
> 1.75 g/day	Double-blind period	Sparsentan	104	12 (11.5)	0.692 [0.351, 1.365]	0.652 [0.297, 1.431]	-5.1 [-15.4, 5.1]	0.328
		Irbesartan	108	18 (16.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.461
Yes	Double-blind period	Sparsentan	88	9 (10.2)	0.707 [0.315, 1.591]	0.674 [0.268, 1.694]	-4.2 [-15.3, 6.8]	0.487
		Irbesartan	83	12 (14.5)				
No	Double-blind period	Sparsentan	114	16 (14.0)	1.044 [0.548, 1.987]	1.051 [0.498, 2.216]	0.6 [-9.1, 10.3]	1.000
		Irbesartan	119	16 (13.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.014 #
<= 5 years	Double-blind period	Sparsentan	113	11 (9.7)	0.538 [0.274, 1.053]	0.488 [0.226, 1.052]	-8.4 [-17.9, 1.1]	0.067
		Irbesartan	127	23 (18.1)				
> 5 years	Double-blind period	Sparsentan	89	14 (15.7)	2.360 [0.891, 6.248]	2.613 [0.895, 7.633]	9.1 [-1.6, 19.7]	0.088
		Irbesartan	75	5 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.681
Yes	Double-blind period	Sparsentan	153	19 (12.4)	0.848 [0.482, 1.492]	0.826 [0.430, 1.587]	-2.2 [-10.5, 6.0]	0.620
		Irbesartan	157	23 (14.6)				
No	Double-blind period	Sparsentan	49	6 (12.2)	1.102 [0.361, 3.363]	1.116 [0.316, 3.945]	1.1 [-14.0, 16.2]	1.000
		Irbesartan	45	5 (11.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Sex	Double-blind period	Sparsentan					Interaction test:	0.732
	Male	Double-blind period	Sparsentan	139	12 (8.6)	2.469 [0.893, 6.825]	2.608 [0.894, 7.609]	5.1 [-1.1, 11.4]	0.083
			Irbesartan	143	5 (3.5)				
	Female	Double-blind period	Sparsentan	63	4 (6.3)	3.746 [0.431, 32.559]	3.932 [0.427, 36.244]	4.7 [-3.9, 13.2]	0.366
			Irbesartan	59	1 (1.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Age	Double-blind period	Sparsentan					Interaction test:	0.697
	<= 45 years	Double-blind period	Sparsentan	96	9 (9.4)	2.320 [0.739, 7.283]	2.457 [0.730, 8.265]	5.3 [-2.7, 13.4]	0.160
			Irbesartan	99	4 (4.0)				
	> 45 years	Double-blind period	Sparsentan	106	7 (6.6)	3.401 [0.723, 15.990]	3.571 [0.724, 17.610]	4.7 [-1.7, 11.0]	0.171
			Irbesartan	103	2 (1.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Age at IgAN diagnosis	Double-blind period	Sparsentan					Interaction test:	0.733
	<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	1.111 [0.131, 9.416]	1.143 [0.077, 16.947]	2.2 [-57.7, 62.1]	1.000
			Irbesartan	5	1 (20.0)				
	> 18 to 40 years	Double-blind period	Sparsentan	102	8 (7.8)	2.850 [0.777, 10.447]	3.007 [0.775, 11.665]	5.1 [-1.9, 12.1]	0.125
			Irbesartan	109	3 (2.8)				
	> 40 years	Double-blind period	Sparsentan	91	6 (6.6)	2.901 [0.602, 13.990]	3.035 [0.596, 15.463]	4.3 [-2.8, 11.4]	0.278
			Irbesartan	88	2 (2.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Geographic region	Double-blind period	Sparsentan					Interaction test:	0.985
	North America	Double-blind period	Sparsentan	35	2 (5.7)	2.629 [0.248, 27.835]	2.727 [0.237, 31.357]	3.5 [-7.7, 14.8]	0.575
			Irbesartan	46	1 (2.2)				
	Europe	Double-blind period	Sparsentan	98	10 (10.2)	2.934 [0.950, 9.062]	3.153 [0.957, 10.395]	6.7 [-1.1, 14.5]	0.056
			Irbesartan	115	4 (3.5)				
	Asia Pacific	Double-blind period	Sparsentan	69	4 (5.8)	2.377 [0.275, 20.547]	2.462 [0.266, 22.810]	3.4 [-5.8, 12.6]	0.649
			Irbesartan	41	1 (2.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.110
	< 27 kg/m**2	Double-blind period	Sparsentan	84	6 (7.1)	1.343 [0.425, 4.240]	1.369 [0.402, 4.662]	1.8 [-6.4, 10.1]	0.758
			Irbesartan	94	5 (5.3)				
	>= 27 kg/m**2	Double-blind period	Sparsentan	118	10 (8.5)	9.068 [1.180, 69.661]	9.815 [1.235, 78.016]	7.5 [1.3, 13.8]	0.011 *
			Irbesartan	107	1 (0.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Randomization strata	Double-blind period	Sparsentan					Interaction test:	0.454
	eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	2.779 [0.768, 10.060]	3.005 [0.764, 11.822]	7.2 [-2.8, 17.2]	0.124
			Irbesartan	74	3 (4.1)				
	eGFR Low and UP Low	Double-blind period	Sparsentan	55	6 (10.9)	6.000 [0.747, 48.207]	6.612 [0.769, 56.881]	9.1 [-1.7, 19.9]	0.113
			Irbesartan	55	1 (1.8)				
	eGFR High and UP High	Double-blind period	Sparsentan	37	1 (2.7)	2.921 + [0.123, 69.434]	3.000 + [0.118, 76.091]	2.7 [-5.3, 10.7]	1.000
			Irbesartan	36	0 (0.0)				
	eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)	0.474 [0.045, 5.014]	0.461 [0.040, 5.305]	-2.8 [-14.3, 8.6]	0.610
			Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Baseline eGFR Group 1	Double-blind period	Sparsentan					Interaction test:	0.516
	< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	14 (11.0)	3.555 [1.203, 10.509]	3.872 [1.238, 12.105]	7.9 [0.9, 14.9]	0.015 *
			Irbesartan	129	4 (3.1)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	1 (2.0)	0.980 [0.063, 15.218]	0.979 [0.059, 16.116]	-0.0 [-7.8, 7.7]	1.000
			Irbesartan	48	1 (2.1)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
			Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Baseline eGFR Group 2	Double-blind period	Sparsentan					Interaction test:	0.681
	< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	9 (11.0)	2.927 [0.822, 10.420]	3.164 [0.824, 12.149]	7.2 [-2.0, 16.4]	0.131
			Irbesartan	80	3 (3.8)				
	45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	5 (11.1)	5.444 [0.661, 44.842]	6.000 [0.673, 53.486]	9.1 [-3.1, 21.2]	0.101
			Irbesartan	49	1 (2.0)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	1 (2.0)	0.980 [0.063, 15.218]	0.979 [0.059, 16.116]	-0.0 [-7.8, 7.7]	1.000
			Irbesartan	48	1 (2.1)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
			Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
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p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.134
	<= 1.75 g/day	Double-blind period	Sparsentan	98	9 (9.2)	8.633 [1.115, 66.824]	9.404 [1.167, 75.756]	8.1 [1.0, 15.2]	0.019 *
			Irbesartan	94	1 (1.1)				
	> 1.75 g/day	Double-blind period	Sparsentan	104	7 (6.7)	1.454 [0.476, 4.436]	1.487 [0.456, 4.841]	2.1 [-5.1, 9.3]	0.564
			Irbesartan	108	5 (4.6)				

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Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.204
	Yes	Double-blind period	Sparsentan	88	8 (9.1)	7.545 [0.964, 59.031]	8.200 [1.003, 67.066]	7.9 [0.3, 15.5]	0.035 *
			Irbesartan	83	1 (1.2)				
	No	Double-blind period	Sparsentan	114	8 (7.0)	1.670 [0.563, 4.955]	1.721 [0.546, 5.425]	2.8 [-4.0, 9.6]	0.402
			Irbesartan	119	5 (4.2)				

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 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Fatigue	Time since renal biopsy	Double-blind period	Sparsentan					Interaction test:	0.368
	<= 5 years	Double-blind period	Sparsentan	113	9 (8.0)	2.023 [0.698, 5.859]	2.112 [0.686, 6.498]	4.0 [-2.8, 10.9]	0.270
			Irbesartan	127	5 (3.9)				
	> 5 years	Double-blind period	Sparsentan	89	7 (7.9)	5.899 [0.742, 46.871]	6.317 [0.759, 52.561]	6.5 [-0.9, 13.9]	0.072
			Irbesartan	75	1 (1.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs										
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
PT: Fatigue	History of hypertension	Double-blind period	Sparsentan						Interaction test:	0.021 #
			Irbesartan							
	Yes	Double-blind period	Sparsentan	153	16 (10.5)	5.473 [1.627, 18.405]	5.995 [1.710, 21.018]	8.5 [2.6, 14.5]	0.002 *	
			Irbesartan	157	3 (1.9)					
			Sparsentan	49	0 (0.0)	0.131 + [0.007, 2.476]	0.123 + [0.006, 2.442]	-6.7 [-16.1, 2.8]		0.106
			Irbesartan	45	3 (6.7)					
No	Double-blind period	Sparsentan	49	0 (0.0)	0.131 + [0.007, 2.476]	0.123 + [0.006, 2.442]	-6.7 [-16.1, 2.8]	0.106		
		Irbesartan	45	3 (6.7)						

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Sex	Double-blind period	Sparsentan					Interaction test:	0.568
	Male	Double-blind period	Sparsentan	139	14 (10.1)	2.400 [0.950, 6.068]	2.557 [0.953, 6.859]	5.9 [-0.8, 12.6]	0.065
			Irbesartan	143	6 (4.2)				
	Female	Double-blind period	Sparsentan	63	12 (19.0)	3.746 [1.112, 12.616]	4.392 [1.172, 16.455]	14.0 [1.1, 26.8]	0.026 *
			Irbesartan	59	3 (5.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Age	Double-blind period	Sparsentan					Interaction test:	0.541
	<= 45 years	Double-blind period	Sparsentan	96	14 (14.6)	2.406 [0.964, 6.003]	2.646 [0.972, 7.204]	8.5 [-1.0, 18.0]	0.060
			Irbesartan	99	6 (6.1)				
	> 45 years	Double-blind period	Sparsentan	106	12 (11.3)	3.887 [1.130, 13.374]	4.255 [1.164, 15.553]	8.4 [0.6, 16.2]	0.029 *
			Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Age at IgAN diagnosis	Double-blind period	Sparsentan					Interaction test:	0.806
	<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
			Irbesartan	5	0 (0.0)				
	> 18 to 40 years	Double-blind period	Sparsentan	102	16 (15.7)	3.420 [1.300, 8.996]	3.870 [1.362, 10.993]	11.1 [2.1, 20.1]	0.010 *
			Irbesartan	109	5 (4.6)				
	> 40 years	Double-blind period	Sparsentan	91	9 (9.9)	2.176 [0.695, 6.808]	2.305 [0.683, 7.779]	5.3 [-3.3, 14.0]	0.250
			Irbesartan	88	4 (4.5)				

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Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Geographic region	Double-blind period	Sparsentan					Interaction test:	0.305
	North America	Double-blind period	Sparsentan	35	3 (8.6)	3.943 [0.428, 36.305]	4.219 [0.420, 42.421]	6.4 [-6.3, 19.1]	0.311
			Irbesartan	46	1 (2.2)				
	Europe	Double-blind period	Sparsentan	98	6 (6.1)	7.041 [0.862, 57.484]	7.435 [0.879, 62.861]	5.3 [-0.7, 11.2]	0.050
			Irbesartan	115	1 (0.9)				
	Asia Pacific	Double-blind period	Sparsentan	69	17 (24.6)	1.443 [0.654, 3.182]	1.588 [0.596, 4.234]	7.6 [-9.7, 24.9]	0.475
			Irbesartan	41	7 (17.1)				

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N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.851
	< 27 kg/m**2	Double-blind period	Sparsentan	84	14 (16.7)	3.133 [1.179, 8.330]	3.560 [1.223, 10.358]	11.3 [1.0, 21.6]	0.016 *
			Irbesartan	94	5 (5.3)				
	>= 27 kg/m**2	Double-blind period	Sparsentan	118	12 (10.2)	2.720 [0.905, 8.180]	2.915 [0.911, 9.333]	6.4 [-1.0, 13.9]	0.072
			Irbesartan	107	4 (3.7)				

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Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Randomization strata	Double-blind period	Sparsentan					Interaction test:	0.585
	eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	1.668 [0.573, 4.856]	1.752 [0.545, 5.637]	4.5 [-6.2, 15.2]	0.394
			Irbesartan	74	5 (6.8)				
	eGFR Low and UP Low	Double-blind period	Sparsentan	55	8 (14.5)	2.667 [0.747, 9.526]	2.950 [0.739, 11.778]	9.1 [-3.8, 22.0]	0.202
			Irbesartan	55	3 (5.5)				
	eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	3.892 [0.457, 33.169]	4.242 [0.451, 39.943]	8.0 [-6.1, 22.1]	0.358
			Irbesartan	36	1 (2.8)				
	eGFR High and UP Low	Double-blind period	Sparsentan	39	6 (15.4)	12.350 + [0.720, 211.796]	14.552 + [0.790, 268.186]	15.4 [1.4, 29.3]	0.026 *
			Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.541
	< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	17 (13.4)	2.158 [0.966, 4.822]	2.338 [0.970, 5.630]	7.2 [-0.8, 15.2]	0.060
			Irbesartan	129	8 (6.2)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	4.898 [0.594, 40.392]	5.341 [0.600, 47.534]	8.1 [-3.3, 19.6]	0.204
			Irbesartan	48	1 (2.1)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	4 (15.4)	8.667 + [0.491, 153.111]	10.200 + [0.520, 200.060]	15.4 [-2.4, 33.2]	0.110
			Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Baseline eGFR Group 2	Double-blind period	Sparsentan					Interaction test:	0.737
	< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	8 (9.8)	1.951 [0.612, 6.224]	2.054 [0.593, 7.113]	4.8 [-4.5, 14.0]	0.370
			Irbesartan	80	4 (5.0)				
	45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	9 (20.0)	2.450 [0.811, 7.405]	2.813 [0.800, 9.882]	11.8 [-4.3, 27.9]	0.136
			Irbesartan	49	4 (8.2)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	4.898 [0.594, 40.392]	5.341 [0.600, 47.534]	8.1 [-3.3, 19.6]	0.204
			Irbesartan	48	1 (2.1)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	4 (15.4)	8.667 + [0.491, 153.111]	10.200 + [0.520, 200.060]	15.4 [-2.4, 33.2]	0.110
			Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.089
	<= 1.75 g/day	Double-blind period	Sparsentan	98	15 (15.3)	7.194 [1.691, 30.608]	8.313 [1.846, 37.442]	13.2 [4.4, 21.9]	0.002 *
			Irbesartan	94	2 (2.1)				
	> 1.75 g/day	Double-blind period	Sparsentan	104	11 (10.6)	1.632 [0.658, 4.048]	1.707 [0.635, 4.586]	4.1 [-4.4, 12.6]	0.331
			Irbesartan	108	7 (6.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.784
	Yes	Double-blind period	Sparsentan	88	8 (9.1)	2.515 [0.691, 9.161]	2.667 [0.683, 10.417]	5.5 [-2.9, 13.9]	0.213
			Irbesartan	83	3 (3.6)				
	No	Double-blind period	Sparsentan	114	18 (15.8)	3.132 [1.289, 7.607]	3.531 [1.348, 9.252]	10.7 [2.1, 19.4]	0.009 *
			Irbesartan	119	6 (5.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.571
	<= 5 years	Double-blind period	Sparsentan	113	16 (14.2)	2.569 [1.097, 6.018]	2.828 [1.118, 7.150]	8.6 [0.3, 17.0]	0.028 *
			Irbesartan	127	7 (5.5)				
	> 5 years	Double-blind period	Sparsentan	89	10 (11.2)	4.213 [0.953, 18.635]	4.620 [0.979, 21.794]	8.6 [-0.2, 17.3]	0.040 *
			Irbesartan	75	2 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs										
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
PT: Dizziness	History of hypertension	Double-blind period	Interaction test:							0.561
			Yes	Double-blind period	Sparsentan	153	17 (11.1)	2.492 [1.063, 5.840]	2.679 [1.078, 6.656]	6.7 [0.1, 13.2]
				Irbesartan	157	7 (4.5)				
	No	Double-blind period	Sparsentan	49	9 (18.4)	4.133 [0.943, 18.114]	4.838 [0.985, 23.758]	13.9 [-0.6, 28.5]	0.053	
			Irbesartan	45	2 (4.4)					

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Sex	Double-blind period	Sparsentan					Interaction test:	0.434
	Male	Double-blind period	Sparsentan	139	27 (19.4)	1.462 [0.853, 2.505]	1.573 [0.830, 2.984]	6.1 [-3.2, 15.5]	0.198
			Irbesartan	143	19 (13.3)				
	Female	Double-blind period	Sparsentan	63	20 (31.7)	2.081 [1.031, 4.199]	2.584 [1.065, 6.267]	16.5 [0.1, 32.8]	0.036 *
			Irbesartan	59	9 (15.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Age	Double-blind period	Sparsentan					Interaction test:	0.525
	<= 45 years	Double-blind period	Sparsentan	96	21 (21.9)	1.969 [1.004, 3.860]	2.240 [1.015, 4.945]	10.8 [-0.6, 22.1]	0.053
			Irbesartan	99	11 (11.1)				
	> 45 years	Double-blind period	Sparsentan	106	26 (24.5)	1.486 [0.859, 2.570]	1.644 [0.830, 3.255]	8.0 [-3.8, 19.9]	0.173
			Irbesartan	103	17 (16.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.663
	<= 18 years	Double-blind period	Sparsentan	9	3 (33.3)	4.200 + [0.259, 68.038]	5.923 + [0.248, 141.482]	33.3 [-13.0, 79.7]	0.258
			Irbesartan	5	0 (0.0)				
	> 18 to 40 years	Double-blind period	Sparsentan	102	18 (17.6)	1.374 [0.721, 2.616]	1.454 [0.682, 3.102]	4.8 [-5.9, 15.5]	0.345
			Irbesartan	109	14 (12.8)				
	> 40 years	Double-blind period	Sparsentan	91	26 (28.6)	1.796 [1.006, 3.207]	2.114 [1.019, 4.388]	12.7 [-0.5, 25.8]	0.049 *
			Irbesartan	88	14 (15.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Geographic region	Double-blind period	Sparsentan					Interaction test:	0.929
	North America	Double-blind period	Sparsentan	35	4 (11.4)	1.314 [0.353, 4.892]	1.355 [0.314, 5.843]	2.7 [-13.1, 18.6]	0.721
			Irbesartan	46	4 (8.7)				
	Europe	Double-blind period	Sparsentan	98	25 (25.5)	1.630 [0.947, 2.804]	1.846 [0.937, 3.634]	9.9 [-2.0, 21.7]	0.088
			Irbesartan	115	18 (15.7)				
	Asia Pacific	Double-blind period	Sparsentan	69	18 (26.1)	1.783 [0.770, 4.126]	2.059 [0.743, 5.705]	11.5 [-5.5, 28.4]	0.232
			Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Baseline BMI	Double-blind period	Sparsentan					Interaction test:	0.418
	< 27 kg/m**2	Double-blind period	Sparsentan	84	17 (20.2)	1.359 [0.714, 2.585]	1.450 [0.666, 3.157]	5.3 [-7.0, 17.7]	0.429
			Irbesartan	94	14 (14.9)				
	>= 27 kg/m**2	Double-blind period	Sparsentan	118	30 (25.4)	1.943 [1.090, 3.463]	2.265 [1.127, 4.552]	12.3 [1.3, 23.4]	0.028 *
			Irbesartan	107	14 (13.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Randomization strata	Double-blind period	Sparsentan					Interaction test:	0.802
	eGFR Low and UP High	Double-blind period	Sparsentan	71	16 (22.5)	1.516 [0.756, 3.038]	1.666 [0.713, 3.893]	7.7 [-6.4, 21.7]	0.288
			Irbesartan	74	11 (14.9)				
	eGFR Low and UP Low	Double-blind period	Sparsentan	55	16 (29.1)	2.000 [0.934, 4.285]	2.410 [0.933, 6.226]	14.5 [-2.5, 31.6]	0.105
			Irbesartan	55	8 (14.5)				
	eGFR High and UP High	Double-blind period	Sparsentan	37	9 (24.3)	2.189 [0.740, 6.477]	2.571 [0.713, 9.270]	13.2 [-6.7, 33.2]	0.221
			Irbesartan	36	4 (11.1)				
	eGFR High and UP Low	Double-blind period	Sparsentan	39	6 (15.4)	1.138 [0.380, 3.414]	1.164 [0.323, 4.196]	1.9 [-16.6, 20.3]	1.000
			Irbesartan	37	5 (13.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
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RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.892
	< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	33 (26.0)	1.764 [1.061, 2.933]	2.032 [1.085, 3.809]	11.3 [0.7, 21.8]	0.030 *
			Irbesartan	129	19 (14.7)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	10 (20.4)	1.399 [0.580, 3.374]	1.502 [0.520, 4.338]	5.8 [-11.3, 23.0]	0.595
			Irbesartan	48	7 (14.6)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	4 (15.4)	1.923 [0.386, 9.584]	2.091 [0.347, 12.589]	7.4 [-14.0, 28.8]	0.668
			Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
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p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Baseline eGFR Group 2	Double-blind period	Sparsentan					Interaction test:	0.906
	< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	21 (25.6)	1.576 [0.848, 2.928]	1.774 [0.818, 3.847]	9.4 [-4.3, 23.0]	0.178
			Irbesartan	80	13 (16.3)				
	45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	12 (26.7)	2.178 [0.892, 5.317]	2.606 [0.885, 7.673]	14.4 [-3.6, 32.4]	0.114
			Irbesartan	49	6 (12.2)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	10 (20.4)	1.399 [0.580, 3.374]	1.502 [0.520, 4.338]	5.8 [-11.3, 23.0]	0.595
			Irbesartan	48	7 (14.6)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	4 (15.4)	1.923 [0.386, 9.584]	2.091 [0.347, 12.589]	7.4 [-14.0, 28.8]	0.668
			Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.040 #
	<= 1.75 g/day	Double-blind period	Sparsentan	98	25 (25.5)	2.997 [1.424, 6.309]	3.682 [1.566, 8.657]	17.0 [5.6, 28.4]	0.002 *
			Irbesartan	94	8 (8.5)				
	> 1.75 g/day	Double-blind period	Sparsentan	104	22 (21.2)	1.142 [0.664, 1.965]	1.180 [0.600, 2.321]	2.6 [-9.0, 14.3]	0.731
			Irbesartan	108	20 (18.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.543
	Yes	Double-blind period	Sparsentan	88	21 (23.9)	1.981 [0.993, 3.952]	2.288 [1.005, 5.210]	11.8 [-0.7, 24.3]	0.049 *
			Irbesartan	83	10 (12.0)				
	No	Double-blind period	Sparsentan	114	26 (22.8)	1.508 [0.876, 2.596]	1.658 [0.852, 3.225]	7.7 [-3.2, 18.6]	0.180
			Irbesartan	119	18 (15.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
SOC: Vascular disorders	Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.307
	<= 5 years	Double-blind period	Sparsentan	113	27 (23.9)	2.023 [1.135, 3.606]	2.344 [1.175, 4.678]	12.1 [1.6, 22.6]	0.017 *
			Irbesartan	127	15 (11.8)				
	> 5 years	Double-blind period	Sparsentan	89	20 (22.5)	1.296 [0.693, 2.427]	1.382 [0.635, 3.009]	5.1 [-8.3, 18.6]	0.441
			Irbesartan	75	13 (17.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
<b>SOC:</b> Vascular disorders	History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.102
	Yes	Double-blind period	Sparsentan	153	41 (26.8)	2.003 [1.244, 3.226]	2.371 [1.324, 4.244]	13.4 [4.0, 22.9]	0.004 *
			Irbesartan	157	21 (13.4)				
	No	Double-blind period	Sparsentan	49	6 (12.2)	0.787 [0.286, 2.167]	0.757 [0.234, 2.452]	-3.3 [-19.5, 12.8]	0.768
			Irbesartan	45	7 (15.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Sex	Double-blind period	Sparsentan					Interaction test:	0.538
	Male	Double-blind period	Sparsentan	139	9 (6.5)	4.629 [1.018, 21.046]	4.881 [1.035, 23.009]	5.1 [-0.2, 10.3]	0.033 *
			Irbesartan	143	2 (1.4)				
	Female	Double-blind period	Sparsentan	63	11 (17.5)	2.575 [0.868, 7.644]	2.909 [0.871, 9.711]	10.7 [-2.3, 23.7]	0.099
			Irbesartan	59	4 (6.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Age	Double-blind period	Sparsentan					Interaction test:	0.877
	<= 45 years	Double-blind period	Sparsentan	96	9 (9.4)	3.094 [0.864, 11.084]	3.310 [0.868, 12.623]	6.3 [-1.4, 14.1]	0.079
			Irbesartan	99	3 (3.0)				
	> 45 years	Double-blind period	Sparsentan	106	11 (10.4)	3.563 [1.023, 12.404]	3.860 [1.044, 14.263]	7.5 [-0.1, 15.1]	0.050 *
			Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Age at IgAN diagnosis	Double-blind period	Sparsentan					Interaction test:	0.914
	<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
			Irbesartan	5	0 (0.0)				
	> 18 to 40 years	Double-blind period	Sparsentan	102	10 (9.8)	3.562 [1.009, 12.578]	3.841 [1.026, 14.378]	7.1 [-0.4, 14.5]	0.044 *
			Irbesartan	109	3 (2.8)				
	> 40 years	Double-blind period	Sparsentan	91	9 (9.9)	2.901 [0.812, 10.365]	3.110 [0.813, 11.893]	6.5 [-1.8, 14.8]	0.133
			Irbesartan	88	3 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Geographic region	Double-blind period	Sparsentan					Interaction test:	0.524
	North America	Double-blind period	Sparsentan	35	3 (8.6)	3.943 [0.428, 36.305]	4.219 [0.420, 42.421]	6.4 [-6.3, 19.1]	0.311
			Irbesartan	46	1 (2.2)				
	Europe	Double-blind period	Sparsentan	98	12 (12.2)	4.694 [1.364, 16.157]	5.209 [1.425, 19.038]	9.6 [1.6, 17.7]	0.007 *
			Irbesartan	115	3 (2.6)				
	Asia Pacific	Double-blind period	Sparsentan	69	5 (7.2)	1.486 [0.302, 7.311]	1.523 [0.282, 8.235]	2.4 [-8.6, 13.3]	1.000
			Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Baseline BMI	Double-blind period	Sparsentan					Interaction test:	0.613
	< 27 kg/m**2	Double-blind period	Sparsentan	84	10 (11.9)	2.798 [0.911, 8.588]	3.041 [0.916, 10.091]	7.6 [-1.5, 16.8]	0.092
			Irbesartan	94	4 (4.3)				
	>= 27 kg/m**2	Double-blind period	Sparsentan	118	10 (8.5)	4.534 [1.016, 20.228]	4.861 [1.040, 22.715]	6.6 [0.1, 13.1]	0.036 *
			Irbesartan	107	2 (1.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Randomization strata	Double-blind period	Sparsentan					Interaction test:	0.666
	eGFR Low and UP High	Double-blind period	Sparsentan	71	7 (9.9)	2.432 [0.654, 9.038]	2.589 [0.642, 10.435]	5.8 [-3.8, 15.4]	0.203
			Irbesartan	74	3 (4.1)				
	eGFR Low and UP Low	Double-blind period	Sparsentan	55	7 (12.7)	3.500 [0.761, 16.105]	3.865 [0.765, 19.514]	9.1 [-2.8, 21.0]	0.161
			Irbesartan	55	2 (3.6)				
	eGFR High and UP High	Double-blind period	Sparsentan	37	5 (13.5)	10.711 + [0.614, 186.919]	12.354 + [0.657, 232.145]	13.5 [-0.2, 27.3]	0.054
			Irbesartan	36	0 (0.0)				
	eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)	0.949 [0.062, 14.620]	0.947 [0.057, 15.721]	-0.1 [-10.0, 9.7]	1.000
			Irbesartan	37	1 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.802
	< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	15 (11.8)	3.047 [1.141, 8.136]	3.321 [1.169, 9.433]	7.9 [0.6, 15.2]	0.020 *
			Irbesartan	129	5 (3.9)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	3 (6.1)	6.860 + [0.364, 129.358]	7.301 + [0.367, 145.240]	6.1 [-2.7, 14.9]	0.242
			Irbesartan	48	0 (0.0)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)	1.923 [0.186, 19.901]	2.000 [0.170, 23.556]	3.7 [-13.0, 20.4]	1.000
			Irbesartan	25	1 (4.0)				

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N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
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p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Baseline eGFR Group 2	Double-blind period	Sparsentan					Interaction test:	0.396
	< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	6 (7.3)	1.463 [0.429, 4.992]	1.500 [0.407, 5.529]	2.3 [-6.3, 10.9]	0.746
			Irbesartan	80	4 (5.0)				
	45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	9 (20.0)	9.800 [1.292, 74.317]	12.000 [1.454, 99.048]	18.0 [3.5, 32.4]	0.006 *
			Irbesartan	49	1 (2.0)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	3 (6.1)	6.860 + [0.364, 129.358]	7.301 + [0.367, 145.240]	6.1 [-2.7, 14.9]	0.242
			Irbesartan	48	0 (0.0)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)	1.923 [0.186, 19.901]	2.000 [0.170, 23.556]	3.7 [-13.0, 20.4]	1.000
			Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.665
	<= 1.75 g/day	Double-blind period	Sparsentan	98	9 (9.2)	4.316 [0.958, 19.457]	4.652 [0.978, 22.129]	7.1 [-0.4, 14.5]	0.059
			Irbesartan	94	2 (2.1)				
	> 1.75 g/day	Double-blind period	Sparsentan	104	11 (10.6)	2.856 [0.939, 8.685]	3.075 [0.947, 9.989]	6.9 [-1.0, 14.7]	0.063
			Irbesartan	108	4 (3.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
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p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.433
	Yes	Double-blind period	Sparsentan	88	7 (8.0)	2.201 [0.589, 8.229]	2.305 [0.576, 9.228]	4.3 [-3.8, 12.4]	0.331
			Irbesartan	83	3 (3.6)				
	No	Double-blind period	Sparsentan	114	13 (11.4)	4.523 [1.324, 15.457]	4.977 [1.379, 17.961]	8.9 [1.5, 16.2]	0.009 *
			Irbesartan	119	3 (2.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Time since renal biopsy	Double-blind period	Sparsentan					Interaction test:	0.671
	<= 5 years	Double-blind period	Sparsentan	113	10 (8.8)	2.810 [0.906, 8.711]	2.985 [0.909, 9.801]	5.7 [-1.2, 12.6]	0.095
			Irbesartan	127	4 (3.1)				
	> 5 years	Double-blind period	Sparsentan	89	10 (11.2)	4.213 [0.953, 18.635]	4.620 [0.979, 21.794]	8.6 [-0.2, 17.3]	0.040 *
			Irbesartan	75	2 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs										
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
PT: Hypotension	History of hypertension	Double-blind period	Interaction test:							0.119
			Yes	Double-blind period	Sparsentan	153	16 (10.5)	5.473 [1.627, 18.405]	5.995 [1.710, 21.018]	8.5 [2.6, 14.5]
				Irbesartan	157	3 (1.9)				
	No	Double-blind period	Interaction test:							1.000
			Sparsentan	49	4 (8.2)	1.224 [0.290, 5.174]	1.244 [0.263, 5.892]	1.5 [-11.2, 14.2]		
			Irbesartan	45	3 (6.7)					

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AC\_SSSM: Incidence of severe TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

Not done: No significant soc/pts found.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AS\_SSSM: Incidence of serious TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

Not done: No significant soc/pts found.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEFC\_SSIM: Incidence of AESI abnormal liver function during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEFS\_SSIM: Incidence of AESI abnormal liver function during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.326
Male	Double-blind period	Sparsentan	139	4 (2.9)	0.514 [0.158, 1.670]	0.500 [0.147, 1.700]	-2.7 [-8.1, 2.7]	0.378
		Irbesartan	143	8 (5.6)				
Female	Double-blind period	Sparsentan	63	1 (1.6)	2.813 + [0.117, 67.711]	2.856 + [0.114, 71.497]	1.6 [-3.1, 6.3]	1.000
		Irbesartan	59	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.218
<= 45 years	Double-blind period	Sparsentan	96	5 (5.2)	1.031 [0.308, 3.449]	1.033 [0.289, 3.688]	0.2 [-7.1, 7.4]	1.000
		Irbesartan	99	5 (5.1)				
> 45 years	Double-blind period	Sparsentan	106	0 (0.0)	0.139 + [0.007, 2.655]	0.135 + [0.007, 2.643]	-2.9 [-7.1, 1.3]	0.118
		Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: NE
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)				NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	5 (4.9)				NE
		Irbesartan	109	4 (3.7)				
> 40 years	Double-blind period	Sparsentan	91	0 (0.0)				NE
		Irbesartan	88	4 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.902
North America	Double-blind period	Sparsentan	35	1 (2.9)	1.314 [0.085, 20.288]	1.324 [0.080, 21.926]	0.7 [-8.8, 10.1]	1.000
		Irbesartan	46	1 (2.2)				
Europe	Double-blind period	Sparsentan	98	4 (4.1)	0.671 [0.202, 2.223]	0.657 [0.186, 2.313]	-2.0 [-8.8, 4.8]	0.553
		Irbesartan	115	7 (6.1)				
Asia Pacific	Double-blind period	Sparsentan	69	0 (0.0)	0.600 + [0.012, 29.678]	0.597 + [0.012, 30.665]	0.0 [NE, NE]	NE
		Irbesartan	41	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19						RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]		p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test:	NE
< 27 kg/m**2	Double-blind period	Sparsentan	84	1 (1.2)					NE
		Irbesartan	94	3 (3.2)					
>= 27 kg/m**2	Double-blind period	Sparsentan	118	4 (3.4)					NE
		Irbesartan	107	5 (4.7)					

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	2 (2.8)				NE
		Irbesartan	74	2 (2.7)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	1 (1.8)				NE
		Irbesartan	55	0 (0.0)				
eGFR High and UP High	Double-blind period	Sparsentan	37	1 (2.7)				NE
		Irbesartan	36	1 (2.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)				NE
		Irbesartan	37	5 (13.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: NE
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	3 (2.4)				NE
		Irbesartan	129	5 (3.9)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	1 (2.0)				NE
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	2 (2.4)				NE
		Irbesartan	80	1 (1.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	1 (2.2)				NE
		Irbesartan	49	4 (8.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	1 (2.0)				NE
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: NE
<= 1.75 g/day	Double-blind period	Sparsentan	98	3 (3.1)				NE
		Irbesartan	94	5 (5.3)				
> 1.75 g/day	Double-blind period	Sparsentan	104	2 (1.9)				NE
		Irbesartan	108	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: NE
Yes	Double-blind period	Sparsentan	88	1 (1.1) all n<10				NE
		Irbesartan	83	3 (3.6)				
No	Double-blind period	Sparsentan	114	4 (3.5) all n<10				NE
		Irbesartan	119	5 (4.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: NE
<= 5 years	Double-blind period	Sparsentan	113	2 (1.8)				NE
		Irbesartan	127	5 (3.9)				
> 5 years	Double-blind period	Sparsentan	89	3 (3.4)				NE
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.138
Yes	Double-blind period	Sparsentan	153	3 (2.0)	0.385 [0.104, 1.423]	0.373 [0.097, 1.431]	-3.1 [-7.9, 1.6]	0.219
		Irbesartan	157	8 (5.1)				
No	Double-blind period	Sparsentan	49	2 (4.1)	4.600 + [0.227, 93.307]	4.789 + [0.224, 102.502]	4.1 [-3.6, 11.8]	0.496
		Irbesartan	45	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in every subgroup level.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEVC\_SSIM: Incidence of AESI COVID-19 during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
 Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.051
Male	Double-blind period	Sparsentan	139	30 (21.6)	0.964 [0.621, 1.498]	0.955 [0.543, 1.678]	-0.8 [-11.2, 9.6]	0.887
		Irbesartan	143	32 (22.4)				
Female	Double-blind period	Sparsentan	63	21 (33.3)	2.185 [1.090, 4.381]	2.778 [1.150, 6.711]	18.1 [1.6, 34.5]	0.022 *
		Irbesartan	59	9 (15.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.939
<= 45 years	Double-blind period	Sparsentan	96	22 (22.9)	1.260 [0.723, 2.198]	1.338 [0.666, 2.689]	4.7 [-7.6, 17.1]	0.479
		Irbesartan	99	18 (18.2)				
> 45 years	Double-blind period	Sparsentan	106	29 (27.4)	1.225 [0.762, 1.971]	1.310 [0.697, 2.461]	5.0 [-7.6, 17.7]	0.427
		Irbesartan	103	23 (22.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.493
<= 18 years	Double-blind period	Sparsentan	9	4 (44.4)	2.222 [0.333, 14.845]	3.200 [0.248, 41.208]	24.4 [-38.9, 87.8]	0.580
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	19 (18.6)	0.967 [0.553, 1.691]	0.959 [0.482, 1.911]	-0.6 [-12.2, 10.9]	1.000
		Irbesartan	109	21 (19.3)				
> 40 years	Double-blind period	Sparsentan	91	28 (30.8)	1.425 [0.861, 2.358]	1.614 [0.822, 3.171]	9.2 [-4.7, 23.1]	0.178
		Irbesartan	88	19 (21.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
 Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.737
North America	Double-blind period	Sparsentan	35	5 (14.3)	0.939 [0.325, 2.710]	0.929 [0.268, 3.217]	-0.9 [-19.0, 17.1]	1.000
		Irbesartan	46	7 (15.2)				
Europe	Double-blind period	Sparsentan	98	28 (28.6)	1.369 [0.852, 2.199]	1.517 [0.809, 2.842]	7.7 [-4.9, 20.3]	0.204
		Irbesartan	115	24 (20.9)				
Asia Pacific	Double-blind period	Sparsentan	69	18 (26.1)	1.070 [0.548, 2.089]	1.094 [0.448, 2.671]	1.7 [-17.0, 20.4]	1.000
		Irbesartan	41	10 (24.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.359
< 27 kg/m**2	Double-blind period	Sparsentan	84	18 (21.4)	1.007 [0.573, 1.771]	1.009 [0.492, 2.069]	0.2 [-13.0, 13.3]	1.000
		Irbesartan	94	20 (21.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	33 (28.0)	1.425 [0.881, 2.304]	1.590 [0.852, 2.967]	8.3 [-3.6, 20.3]	0.161
		Irbesartan	107	21 (19.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.663
eGFR Low and UP High	Double-blind period	Sparsentan	71	19 (26.8)	1.414 [0.770, 2.600]	1.566 [0.715, 3.429]	7.8 [-7.2, 22.8]	0.323
		Irbesartan	74	14 (18.9)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	16 (29.1)	1.333 [0.697, 2.550]	1.470 [0.619, 3.491]	7.3 [-10.8, 25.3]	0.512
		Irbesartan	55	12 (21.8)				
eGFR High and UP High	Double-blind period	Sparsentan	37	10 (27.0)	1.390 [0.594, 3.252]	1.534 [0.511, 4.605]	7.6 [-14.4, 29.6]	0.581
		Irbesartan	36	7 (19.4)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	6 (15.4)	0.712 [0.273, 1.855]	0.659 [0.205, 2.124]	-6.2 [-26.3, 13.8]	0.561
		Irbesartan	37	8 (21.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.793
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	35 (27.6)	1.270 [0.824, 1.956]	1.372 [0.775, 2.430]	5.9 [-5.5, 17.2]	0.311
		Irbesartan	129	28 (21.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	12 (24.5)	1.069 [0.523, 2.184]	1.091 [0.428, 2.783]	1.6 [-17.4, 20.6]	1.000
		Irbesartan	48	11 (22.9)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	4 (15.4)	1.923 [0.386, 9.584]	2.091 [0.347, 12.589]	7.4 [-14.0, 28.8]	0.668
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.926
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	22 (26.8)	1.263 [0.726, 2.195]	1.359 [0.658, 2.806]	5.6 [-8.8, 19.9]	0.464
		Irbesartan	80	17 (21.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	13 (28.9)	1.287 [0.643, 2.575]	1.403 [0.553, 3.559]	6.4 [-13.4, 26.2]	0.489
		Irbesartan	49	11 (22.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	12 (24.5)	1.069 [0.523, 2.184]	1.091 [0.428, 2.783]	1.6 [-17.4, 20.6]	1.000
		Irbesartan	48	11 (22.9)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	4 (15.4)	1.923 [0.386, 9.584]	2.091 [0.347, 12.589]	7.4 [-14.0, 28.8]	0.668
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.039 #
<= 1.75 g/day	Double-blind period	Sparsentan	98	27 (27.6)	1.992 [1.095, 3.623]	2.369 [1.137, 4.938]	13.7 [1.4, 26.0]	0.021 *
		Irbesartan	94	13 (13.8)				
> 1.75 g/day	Double-blind period	Sparsentan	104	24 (23.1)	0.890 [0.554, 1.430]	0.857 [0.458, 1.605]	-2.8 [-15.4, 9.7]	0.636
		Irbesartan	108	28 (25.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.644
Yes	Double-blind period	Sparsentan	88	24 (27.3)	1.132 [0.678, 1.889]	1.181 [0.594, 2.350]	3.2 [-11.1, 17.4]	0.727
		Irbesartan	83	20 (24.1)				
No	Double-blind period	Sparsentan	114	27 (23.7)	1.342 [0.807, 2.233]	1.448 [0.764, 2.744]	6.0 [-5.2, 17.3]	0.262
		Irbesartan	119	21 (17.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.918
<= 5 years	Double-blind period	Sparsentan	113	28 (24.8)	1.259 [0.782, 2.026]	1.344 [0.729, 2.477]	5.1 [-6.3, 16.5]	0.354
		Irbesartan	127	25 (19.7)				
> 5 years	Double-blind period	Sparsentan	89	23 (25.8)	1.211 [0.692, 2.119]	1.285 [0.620, 2.662]	4.5 [-9.7, 18.7]	0.582
		Irbesartan	75	16 (21.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.324
Yes	Double-blind period	Sparsentan	153	45 (29.4)	1.358 [0.923, 1.997]	1.507 [0.901, 2.523]	7.8 [-2.6, 18.1]	0.120
		Irbesartan	157	34 (21.7)				
No	Double-blind period	Sparsentan	49	6 (12.2)	0.787 [0.286, 2.167]	0.757 [0.234, 2.452]	-3.3 [-19.5, 12.8]	0.768
		Irbesartan	45	7 (15.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.082
Male	Double-blind period	Sparsentan	139	30 (21.6)	0.996 [0.638, 1.553]	0.994 [0.564, 1.753]	-0.1 [-10.4, 10.2]	1.000
		Irbesartan	143	31 (21.7)				
Female	Double-blind period	Sparsentan	63	20 (31.7)	2.081 [1.031, 4.199]	2.584 [1.065, 6.267]	16.5 [0.1, 32.8]	0.036 *
		Irbesartan	59	9 (15.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.750
<= 45 years	Double-blind period	Sparsentan	96	22 (22.9)	1.335 [0.757, 2.353]	1.434 [0.707, 2.907]	5.7 [-6.5, 18.0]	0.372
		Irbesartan	99	17 (17.2)				
> 45 years	Double-blind period	Sparsentan	106	28 (26.4)	1.183 [0.732, 1.912]	1.249 [0.663, 2.353]	4.1 [-8.5, 16.7]	0.523
		Irbesartan	103	23 (22.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.608
<= 18 years	Double-blind period	Sparsentan	9	4 (44.4)	2.222 [0.333, 14.845]	3.200 [0.248, 41.208]	24.4 [-38.9, 87.8]	0.580
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	19 (18.6)	1.015 [0.576, 1.790]	1.019 [0.508, 2.042]	0.3 [-11.2, 11.7]	1.000
		Irbesartan	109	20 (18.3)				
> 40 years	Double-blind period	Sparsentan	91	27 (29.7)	1.374 [0.826, 2.285]	1.532 [0.777, 3.019]	8.1 [-5.8, 21.9]	0.235
		Irbesartan	88	19 (21.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.733
North America	Double-blind period	Sparsentan	35	5 (14.3)	0.939 [0.325, 2.710]	0.929 [0.268, 3.217]	-0.9 [-19.0, 17.1]	1.000
		Irbesartan	46	7 (15.2)				
Europe	Double-blind period	Sparsentan	98	27 (27.6)	1.378 [0.847, 2.241]	1.521 [0.805, 2.875]	7.6 [-4.9, 20.0]	0.200
		Irbesartan	115	23 (20.0)				
Asia Pacific	Double-blind period	Sparsentan	69	18 (26.1)	1.070 [0.548, 2.089]	1.094 [0.448, 2.671]	1.7 [-17.0, 20.4]	1.000
		Irbesartan	41	10 (24.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.240
< 27 kg/m**2	Double-blind period	Sparsentan	84	17 (20.2)	0.951 [0.535, 1.692]	0.939 [0.454, 1.941]	-1.0 [-14.1, 12.0]	1.000
		Irbesartan	94	20 (21.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	33 (28.0)	1.496 [0.917, 2.442]	1.689 [0.899, 3.173]	9.3 [-2.6, 21.1]	0.117
		Irbesartan	107	20 (18.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.629
eGFR Low and UP High	Double-blind period	Sparsentan	71	18 (25.4)	1.340 [0.722, 2.486]	1.456 [0.660, 3.208]	6.4 [-8.4, 21.3]	0.424
		Irbesartan	74	14 (18.9)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	16 (29.1)	1.333 [0.697, 2.550]	1.470 [0.619, 3.491]	7.3 [-10.8, 25.3]	0.512
		Irbesartan	55	12 (21.8)				
eGFR High and UP High	Double-blind period	Sparsentan	37	10 (27.0)	1.622 [0.658, 3.997]	1.852 [0.594, 5.778]	10.4 [-11.2, 31.9]	0.398
		Irbesartan	36	6 (16.7)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	6 (15.4)	0.712 [0.273, 1.855]	0.659 [0.205, 2.124]	-6.2 [-26.3, 13.8]	0.561
		Irbesartan	37	8 (21.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.790
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	34 (26.8)	1.279 [0.822, 1.990]	1.381 [0.775, 2.462]	5.8 [-5.4, 17.0]	0.306
		Irbesartan	129	27 (20.9)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	12 (24.5)	1.069 [0.523, 2.184]	1.091 [0.428, 2.783]	1.6 [-17.4, 20.6]	1.000
		Irbesartan	48	11 (22.9)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	4 (15.4)	1.923 [0.386, 9.584]	2.091 [0.347, 12.589]	7.4 [-14.0, 28.8]	0.668
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.924
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	22 (26.8)	1.263 [0.726, 2.195]	1.359 [0.658, 2.806]	5.6 [-8.8, 19.9]	0.464
		Irbesartan	80	17 (21.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	12 (26.7)	1.307 [0.626, 2.726]	1.418 [0.544, 3.699]	6.3 [-13.0, 25.5]	0.626
		Irbesartan	49	10 (20.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	12 (24.5)	1.069 [0.523, 2.184]	1.091 [0.428, 2.783]	1.6 [-17.4, 20.6]	1.000
		Irbesartan	48	11 (22.9)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	4 (15.4)	1.923 [0.386, 9.584]	2.091 [0.347, 12.589]	7.4 [-14.0, 28.8]	0.668
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.039 #
<= 1.75 g/day	Double-blind period	Sparsentan	98	27 (27.6)	1.992 [1.095, 3.623]	2.369 [1.137, 4.938]	13.7 [1.4, 26.0]	0.021 *
		Irbesartan	94	13 (13.8)				
> 1.75 g/day	Double-blind period	Sparsentan	104	23 (22.1)	0.885 [0.544, 1.439]	0.852 [0.451, 1.609]	-2.9 [-15.2, 9.5]	0.632
		Irbesartan	108	27 (25.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.484
Yes	Double-blind period	Sparsentan	88	23 (26.1)	1.085 [0.646, 1.822]	1.115 [0.558, 2.227]	2.0 [-12.1, 16.2]	0.860
		Irbesartan	83	20 (24.1)				
No	Double-blind period	Sparsentan	114	27 (23.7)	1.409 [0.839, 2.366]	1.536 [0.805, 2.931]	6.9 [-4.3, 18.0]	0.197
		Irbesartan	119	20 (16.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.745
<= 5 years	Double-blind period	Sparsentan	113	28 (24.8)	1.311 [0.809, 2.125]	1.414 [0.763, 2.618]	5.9 [-5.4, 17.2]	0.277
		Irbesartan	127	24 (18.9)				
> 5 years	Double-blind period	Sparsentan	89	22 (24.7)	1.159 [0.658, 2.041]	1.211 [0.582, 2.520]	3.4 [-10.7, 17.5]	0.711
		Irbesartan	75	16 (21.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.318
Yes	Double-blind period	Sparsentan	153	44 (28.8)	1.368 [0.924, 2.026]	1.517 [0.902, 2.550]	7.7 [-2.5, 18.0]	0.148
		Irbesartan	157	33 (21.0)				
No	Double-blind period	Sparsentan	49	6 (12.2)	0.787 [0.286, 2.167]	0.757 [0.234, 2.452]	-3.3 [-19.5, 12.8]	0.768
		Irbesartan	45	7 (15.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AECC\_SSIM: Incidence of AESI cardiovascular system during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AECS\_SSIM: Incidence of AESI cardiovascular system during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
 Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.757
Male	Double-blind period	Sparsentan	139	28 (20.1)	2.216 [1.198, 4.098]	2.523 [1.247, 5.105]	11.1 [2.2, 19.9]	0.011 *
		Irbesartan	143	13 (9.1)				
Female	Double-blind period	Sparsentan	63	22 (34.9)	2.575 [1.245, 5.329]	3.421 [1.380, 8.478]	21.4 [5.1, 37.7]	0.007 *
		Irbesartan	59	8 (13.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.760
<= 45 years	Double-blind period	Sparsentan	96	26 (27.1)	2.234 [1.197, 4.169]	2.693 [1.268, 5.718]	15.0 [3.0, 27.0]	0.011 *
		Irbesartan	99	12 (12.1)				
> 45 years	Double-blind period	Sparsentan	106	24 (22.6)	2.591 [1.266, 5.305]	3.057 [1.345, 6.950]	13.9 [3.3, 24.5]	0.007 *
		Irbesartan	103	9 (8.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
 Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.933
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	3.000 + [0.171, 52.527]	3.667 + [0.145, 92.653]	22.2 [-20.5, 64.9]	0.505
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	29 (28.4)	2.214 [1.242, 3.945]	2.696 [1.329, 5.467]	15.6 [3.9, 27.3]	0.006 *
		Irbesartan	109	14 (12.8)				
> 40 years	Double-blind period	Sparsentan	91	19 (20.9)	2.625 [1.161, 5.933]	3.054 [1.213, 7.685]	12.9 [1.7, 24.1]	0.019 *
		Irbesartan	88	7 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.114
North America	Double-blind period	Sparsentan	35	6 (17.1)	2.629 [0.706, 9.784]	2.966 [0.686, 12.817]	10.6 [-6.3, 27.5]	0.165
		Irbesartan	46	3 (6.5)				
Europe	Double-blind period	Sparsentan	98	21 (21.4)	3.520 [1.563, 7.928]	4.208 [1.704, 10.391]	15.3 [5.2, 25.5]	0.001 *
		Irbesartan	115	7 (6.1)				
Asia Pacific	Double-blind period	Sparsentan	69	23 (33.3)	1.242 [0.678, 2.277]	1.364 [0.581, 3.200]	6.5 [-13.0, 26.0]	0.528
		Irbesartan	41	11 (26.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
 Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.873
< 27 kg/m**2	Double-blind period	Sparsentan	84	25 (29.8)	2.331 [1.251, 4.344]	2.895 [1.347, 6.224]	17.0 [4.0, 30.0]	0.006 *
		Irbesartan	94	12 (12.8)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	25 (21.2)	2.519 [1.231, 5.152]	2.927 [1.298, 6.600]	12.8 [2.8, 22.7]	0.009 *
		Irbesartan	107	9 (8.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
 Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.598
eGFR Low and UP High	Double-blind period	Sparsentan	71	16 (22.5)	1.853 [0.876, 3.918]	2.101 [0.861, 5.127]	10.4 [-3.3, 24.0]	0.125
		Irbesartan	74	9 (12.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	16 (29.1)	2.000 [0.934, 4.285]	2.410 [0.933, 6.226]	14.5 [-2.5, 31.6]	0.105
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	9 (24.3)	4.378 [1.015, 18.888]	5.464 [1.090, 27.385]	18.8 [0.3, 37.2]	0.046 *
		Irbesartan	36	2 (5.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	9 (23.1)	4.269 [0.987, 18.469]	5.250 [1.052, 26.210]	17.7 [-0.1, 35.4]	0.048 *
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.545
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	34 (26.8)	2.031 [1.198, 3.445]	2.409 [1.265, 4.586]	13.6 [3.1, 24.0]	0.008 *
		Irbesartan	129	17 (13.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	11 (22.4)	3.592 [1.068, 12.081]	4.342 [1.128, 16.710]	16.2 [0.6, 31.8]	0.040 *
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	5 (19.2)	4.808 [0.603, 38.316]	5.714 [0.617, 52.902]	15.2 [-5.7, 36.1]	0.191
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.589
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	17 (20.7)	1.659 [0.809, 3.400]	1.831 [0.782, 4.287]	8.2 [-4.4, 20.8]	0.206
		Irbesartan	80	10 (12.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	17 (37.8)	2.644 [1.210, 5.778]	3.643 [1.338, 9.919]	23.5 [4.1, 42.8]	0.017 *
		Irbesartan	49	7 (14.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	11 (22.4)	3.592 [1.068, 12.081]	4.342 [1.128, 16.710]	16.2 [0.6, 31.8]	0.040 *
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	5 (19.2)	4.808 [0.603, 38.316]	5.714 [0.617, 52.902]	15.2 [-5.7, 36.1]	0.191
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
 Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.080
<= 1.75 g/day	Double-blind period	Sparsentan	98	26 (26.5)	4.156 [1.792, 9.641]	5.296 [2.067, 13.569]	20.1 [9.1, 31.2]	<0.001 *
		Irbesartan	94	6 (6.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	24 (23.1)	1.662 [0.925, 2.986]	1.860 [0.914, 3.787]	9.2 [-2.2, 20.5]	0.110
		Irbesartan	108	15 (13.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.252
Yes	Double-blind period	Sparsentan	88	16 (18.2)	1.677 [0.784, 3.584]	1.827 [0.759, 4.400]	7.3 [-4.3, 19.0]	0.199
		Irbesartan	83	9 (10.8)				
No	Double-blind period	Sparsentan	114	34 (29.8)	2.958 [1.614, 5.421]	3.790 [1.846, 7.778]	19.7 [8.9, 30.6]	<0.001 *
		Irbesartan	119	12 (10.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.518
<= 5 years	Double-blind period	Sparsentan	113	26 (23.0)	2.087 [1.148, 3.797]	2.412 [1.189, 4.893]	12.0 [1.7, 22.3]	0.015 *
		Irbesartan	127	14 (11.0)				
> 5 years	Double-blind period	Sparsentan	89	24 (27.0)	2.889 [1.319, 6.327]	3.587 [1.447, 8.893]	17.6 [5.1, 30.2]	0.005 *
		Irbesartan	75	7 (9.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.936
Yes	Double-blind period	Sparsentan	153	35 (22.9)	2.394 [1.364, 4.202]	2.808 [1.463, 5.391]	13.3 [4.6, 22.1]	0.002 *
		Irbesartan	157	15 (9.6)				
No	Double-blind period	Sparsentan	49	15 (30.6)	2.296 [0.976, 5.403]	2.868 [1.001, 8.216]	17.3 [-1.1, 35.7]	0.051
		Irbesartan	45	6 (13.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
 Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.832
Male	Double-blind period	Sparsentan	139	28 (20.1)	2.216 [1.198, 4.098]	2.523 [1.247, 5.105]	11.1 [2.2, 19.9]	0.011 *
		Irbesartan	143	13 (9.1)				
Female	Double-blind period	Sparsentan	63	21 (33.3)	2.458 [1.181, 5.116]	3.188 [1.282, 7.927]	19.8 [3.6, 36.0]	0.011 *
		Irbesartan	59	8 (13.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
 Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.828
<= 45 years	Double-blind period	Sparsentan	96	26 (27.1)	2.234 [1.197, 4.169]	2.693 [1.268, 5.718]	15.0 [3.0, 27.0]	0.011 *
		Irbesartan	99	12 (12.1)				
> 45 years	Double-blind period	Sparsentan	106	23 (21.7)	2.483 [1.207, 5.108]	2.894 [1.268, 6.606]	13.0 [2.4, 23.5]	0.012 *
		Irbesartan	103	9 (8.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.959
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	3.000 + [0.171, 52.527]	3.667 + [0.145, 92.653]	22.2 [-20.5, 64.9]	0.505
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	29 (28.4)	2.214 [1.242, 3.945]	2.696 [1.329, 5.467]	15.6 [3.9, 27.3]	0.006 *
		Irbesartan	109	14 (12.8)				
> 40 years	Double-blind period	Sparsentan	91	18 (19.8)	2.487 [1.093, 5.659]	2.853 [1.127, 7.221]	11.8 [0.8, 22.9]	0.030 *
		Irbesartan	88	7 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
 Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.138
North America	Double-blind period	Sparsentan	35	6 (17.1)	2.629 [0.706, 9.784]	2.966 [0.686, 12.817]	10.6 [-6.3, 27.5]	0.165
		Irbesartan	46	3 (6.5)				
Europe	Double-blind period	Sparsentan	98	20 (20.4)	3.353 [1.480, 7.593]	3.956 [1.595, 9.815]	14.3 [4.3, 24.4]	0.002 *
		Irbesartan	115	7 (6.1)				
Asia Pacific	Double-blind period	Sparsentan	69	23 (33.3)	1.242 [0.678, 2.277]	1.364 [0.581, 3.200]	6.5 [-13.0, 26.0]	0.528
		Irbesartan	41	11 (26.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
 Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.808
< 27 kg/m**2	Double-blind period	Sparsentan	84	24 (28.6)	2.238 [1.195, 4.191]	2.733 [1.267, 5.896]	15.8 [2.9, 28.7]	0.014 *
		Irbesartan	94	12 (12.8)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	25 (21.2)	2.519 [1.231, 5.152]	2.927 [1.298, 6.600]	12.8 [2.8, 22.7]	0.009 *
		Irbesartan	107	9 (8.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.559
eGFR Low and UP High	Double-blind period	Sparsentan	71	15 (21.1)	1.737 [0.813, 3.713]	1.935 [0.786, 4.760]	9.0 [-4.5, 22.4]	0.182
		Irbesartan	74	9 (12.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	16 (29.1)	2.000 [0.934, 4.285]	2.410 [0.933, 6.226]	14.5 [-2.5, 31.6]	0.105
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	9 (24.3)	4.378 [1.015, 18.888]	5.464 [1.090, 27.385]	18.8 [0.3, 37.2]	0.046 *
		Irbesartan	36	2 (5.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	9 (23.1)	4.269 [0.987, 18.469]	5.250 [1.052, 26.210]	17.7 [-0.1, 35.4]	0.048 *
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.516
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	33 (26.0)	1.972 [1.159, 3.354]	2.313 [1.212, 4.414]	12.8 [2.4, 23.2]	0.012 *
		Irbesartan	129	17 (13.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	11 (22.4)	3.592 [1.068, 12.081]	4.342 [1.128, 16.710]	16.2 [0.6, 31.8]	0.040 *
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	5 (19.2)	4.808 [0.603, 38.316]	5.714 [0.617, 52.902]	15.2 [-5.7, 36.1]	0.191
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.607
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	17 (20.7)	1.659 [0.809, 3.400]	1.831 [0.782, 4.287]	8.2 [-4.4, 20.8]	0.206
		Irbesartan	80	10 (12.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	16 (35.6)	2.489 [1.129, 5.488]	3.310 [1.210, 9.056]	21.3 [2.1, 40.5]	0.029 *
		Irbesartan	49	7 (14.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	11 (22.4)	3.592 [1.068, 12.081]	4.342 [1.128, 16.710]	16.2 [0.6, 31.8]	0.040 *
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	5 (19.2)	4.808 [0.603, 38.316]	5.714 [0.617, 52.902]	15.2 [-5.7, 36.1]	0.191
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
 Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.068
<= 1.75 g/day	Double-blind period	Sparsentan	98	26 (26.5)	4.156 [1.792, 9.641]	5.296 [2.067, 13.569]	20.1 [9.1, 31.2]	<0.001 *
		Irbesartan	94	6 (6.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	23 (22.1)	1.592 [0.881, 2.879]	1.760 [0.861, 3.600]	8.2 [-3.0, 19.5]	0.152
		Irbesartan	108	15 (13.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
 Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.206
Yes	Double-blind period	Sparsentan	88	15 (17.0)	1.572 [0.728, 3.395]	1.689 [0.696, 4.103]	6.2 [-5.3, 17.7]	0.277
		Irbesartan	83	9 (10.8)				
No	Double-blind period	Sparsentan	114	34 (29.8)	2.958 [1.614, 5.421]	3.790 [1.846, 7.778]	19.7 [8.9, 30.6]	<0.001 *
		Irbesartan	119	12 (10.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.576
<= 5 years	Double-blind period	Sparsentan	113	26 (23.0)	2.087 [1.148, 3.797]	2.412 [1.189, 4.893]	12.0 [1.7, 22.3]	0.015 *
		Irbesartan	127	14 (11.0)				
> 5 years	Double-blind period	Sparsentan	89	23 (25.8)	2.769 [1.259, 6.091]	3.385 [1.361, 8.421]	16.5 [4.1, 29.0]	0.008 *
		Irbesartan	75	7 (9.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.980
Yes	Double-blind period	Sparsentan	153	34 (22.2)	2.326 [1.322, 4.093]	2.705 [1.406, 5.205]	12.7 [4.0, 21.3]	0.003 *
		Irbesartan	157	15 (9.6)				
No	Double-blind period	Sparsentan	49	15 (30.6)	2.296 [0.976, 5.403]	2.868 [1.001, 8.216]	17.3 [-1.1, 35.7]	0.051
		Irbesartan	45	6 (13.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEHC\_SSIM: Incidence of AESI hypotension during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEHS\_SSIM: Incidence of AESI hypotension during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.207
Male	Double-blind period	Sparsentan	139	12 (8.6)	2.469 [0.893, 6.825]	2.608 [0.894, 7.609]	5.1 [-1.1, 11.4]	0.083
		Irbesartan	143	5 (3.5)				
Female	Double-blind period	Sparsentan	63	1 (1.6)	0.468 [0.044, 5.029]	0.460 [0.041, 5.207]	-1.8 [-9.0, 5.4]	0.610
		Irbesartan	59	2 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
 Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.099
<= 45 years	Double-blind period	Sparsentan	96	4 (4.2)	0.825 [0.228, 2.980]	0.817 [0.213, 3.140]	-0.9 [-7.8, 6.0]	1.000
		Irbesartan	99	5 (5.1)				
> 45 years	Double-blind period	Sparsentan	106	9 (8.5)	4.373 [0.968, 19.754]	4.686 [0.987, 22.238]	6.5 [-0.3, 13.4]	0.059
		Irbesartan	103	2 (1.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.677
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.600 + [0.014, 26.475]	0.579 + [0.010, 33.502]	0.0 [NE, NE]	NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	7 (6.9)	1.496 [0.490, 4.564]	1.533 [0.471, 4.992]	2.3 [-5.0, 9.5]	0.559
		Irbesartan	109	5 (4.6)				
> 40 years	Double-blind period	Sparsentan	91	6 (6.6)	2.901 [0.602, 13.990]	3.035 [0.596, 15.463]	4.3 [-2.8, 11.4]	0.278
		Irbesartan	88	2 (2.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

<u>AESI hepatic disorders</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: NE
North America	Double-blind period	Sparsentan	35	3 (8.6)				NE
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	7 (7.1)				NE
		Irbesartan	115	2 (1.7)				
Asia Pacific	Double-blind period	Sparsentan	69	3 (4.3)				NE
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.976
< 27 kg/m**2	Double-blind period	Sparsentan	84	5 (6.0)	1.865 [0.460, 7.569]	1.920 [0.445, 8.289]	2.8 [-4.5, 10.1]	0.478
		Irbesartan	94	3 (3.2)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	8 (6.8)	1.814 [0.562, 5.851]	1.873 [0.547, 6.407]	3.0 [-3.6, 9.7]	0.382
		Irbesartan	107	4 (3.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	6 (8.5)				NE
		Irbesartan	74	2 (2.7)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)				NE
		Irbesartan	55	2 (3.6)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)				NE
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)				NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.413
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	8 (6.3)	2.031 [0.627, 6.578]	2.101 [0.616, 7.160]	3.2 [-2.8, 9.2]	0.253
		Irbesartan	129	4 (3.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	3.918 [0.454, 33.802]	4.178 [0.450, 38.818]	6.1 [-4.6, 16.8]	0.362
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.481 [0.046, 4.975]	0.460 [0.039, 5.418]	-4.2 [-21.0, 12.7]	0.610
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	6 (7.3)				NE
		Irbesartan	80	3 (3.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	2 (4.4)				NE
		Irbesartan	49	1 (2.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)				NE
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)				NE
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.526
<= 1.75 g/day	Double-blind period	Sparsentan	98	4 (4.1)	1.279 [0.294, 5.562]	1.291 [0.281, 5.928]	0.9 [-5.4, 7.2]	1.000
		Irbesartan	94	3 (3.2)				
> 1.75 g/day	Double-blind period	Sparsentan	104	9 (8.7)	2.337 [0.742, 7.354]	2.463 [0.734, 8.262]	5.0 [-2.5, 12.4]	0.160
		Irbesartan	108	4 (3.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.502
Yes	Double-blind period	Sparsentan	88	6 (6.8)	2.830 [0.588, 13.627]	2.963 [0.581, 15.117]	4.4 [-3.0, 11.8]	0.279
		Irbesartan	83	2 (2.4)				
No	Double-blind period	Sparsentan	114	7 (6.1)	1.461 [0.478, 4.472]	1.492 [0.459, 4.842]	1.9 [-4.6, 8.5]	0.564
		Irbesartan	119	5 (4.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.166
<= 5 years	Double-blind period	Sparsentan	113	6 (5.3)	1.124 [0.373, 3.386]	1.131 [0.354, 3.611]	0.6 [-5.8, 7.0]	1.000
		Irbesartan	127	6 (4.7)				
> 5 years	Double-blind period	Sparsentan	89	7 (7.9)	5.899 [0.742, 46.871]	6.317 [0.759, 52.561]	6.5 [-0.9, 13.9]	0.072
		Irbesartan	75	1 (1.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.149
Yes	Double-blind period	Sparsentan	153	11 (7.2)	2.822 [0.918, 8.670]	2.963 [0.922, 9.518]	4.6 [-0.8, 10.1]	0.067
		Irbesartan	157	4 (2.5)				
No	Double-blind period	Sparsentan	49	2 (4.1)	0.612 [0.107, 3.498]	0.596 [0.095, 3.740]	-2.6 [-13.9, 8.7]	0.668
		Irbesartan	45	3 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
 Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.207
Male	Double-blind period	Sparsentan	139	12 (8.6)	2.469 [0.893, 6.825]	2.608 [0.894, 7.609]	5.1 [-1.1, 11.4]	0.083
		Irbesartan	143	5 (3.5)				
Female	Double-blind period	Sparsentan	63	1 (1.6)	0.468 [0.044, 5.029]	0.460 [0.041, 5.207]	-1.8 [-9.0, 5.4]	0.610
		Irbesartan	59	2 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
 Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.099
<= 45 years	Double-blind period	Sparsentan	96	4 (4.2)	0.825 [0.228, 2.980]	0.817 [0.213, 3.140]	-0.9 [-7.8, 6.0]	1.000
		Irbesartan	99	5 (5.1)				
> 45 years	Double-blind period	Sparsentan	106	9 (8.5)	4.373 [0.968, 19.754]	4.686 [0.987, 22.238]	6.5 [-0.3, 13.4]	0.059
		Irbesartan	103	2 (1.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
 Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.677
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.600 + [0.014, 26.475]	0.579 + [0.010, 33.502]	0.0 [NE, NE]	NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	7 (6.9)	1.496 [0.490, 4.564]	1.533 [0.471, 4.992]	2.3 [-5.0, 9.5]	0.559
		Irbesartan	109	5 (4.6)				
> 40 years	Double-blind period	Sparsentan	91	6 (6.6)	2.901 [0.602, 13.990]	3.035 [0.596, 15.463]	4.3 [-2.8, 11.4]	0.278
		Irbesartan	88	2 (2.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
 Safety Set

<u>AESI hepatic disorders - non-severe</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: NE
North America	Double-blind period	Sparsentan	35	3 (8.6)				NE
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	7 (7.1)				NE
		Irbesartan	115	2 (1.7)				
Asia Pacific	Double-blind period	Sparsentan	69	3 (4.3)				NE
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.976
< 27 kg/m**2	Double-blind period	Sparsentan	84	5 (6.0)	1.865 [0.460, 7.569]	1.920 [0.445, 8.289]	2.8 [-4.5, 10.1]	0.478
		Irbesartan	94	3 (3.2)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	8 (6.8)	1.814 [0.562, 5.851]	1.873 [0.547, 6.407]	3.0 [-3.6, 9.7]	0.382
		Irbesartan	107	4 (3.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	6 (8.5)				NE
		Irbesartan	74	2 (2.7)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)				NE
		Irbesartan	55	2 (3.6)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)				NE
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)				NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
 Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.413
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	8 (6.3)	2.031 [0.627, 6.578]	2.101 [0.616, 7.160]	3.2 [-2.8, 9.2]	0.253
		Irbesartan	129	4 (3.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	3.918 [0.454, 33.802]	4.178 [0.450, 38.818]	6.1 [-4.6, 16.8]	0.362
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.481 [0.046, 4.975]	0.460 [0.039, 5.418]	-4.2 [-21.0, 12.7]	0.610
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	6 (7.3)				NE
		Irbesartan	80	3 (3.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	2 (4.4)				NE
		Irbesartan	49	1 (2.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)				NE
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)				NE
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.526
<= 1.75 g/day	Double-blind period	Sparsentan	98	4 (4.1)	1.279 [0.294, 5.562]	1.291 [0.281, 5.928]	0.9 [-5.4, 7.2]	1.000
		Irbesartan	94	3 (3.2)				
> 1.75 g/day	Double-blind period	Sparsentan	104	9 (8.7)	2.337 [0.742, 7.354]	2.463 [0.734, 8.262]	5.0 [-2.5, 12.4]	0.160
		Irbesartan	108	4 (3.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
 Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.502
Yes	Double-blind period	Sparsentan	88	6 (6.8)	2.830 [0.588, 13.627]	2.963 [0.581, 15.117]	4.4 [-3.0, 11.8]	0.279
		Irbesartan	83	2 (2.4)				
No	Double-blind period	Sparsentan	114	7 (6.1)	1.461 [0.478, 4.472]	1.492 [0.459, 4.842]	1.9 [-4.6, 8.5]	0.564
		Irbesartan	119	5 (4.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.166
<= 5 years	Double-blind period	Sparsentan	113	6 (5.3)	1.124 [0.373, 3.386]	1.131 [0.354, 3.611]	0.6 [-5.8, 7.0]	1.000
		Irbesartan	127	6 (4.7)				
> 5 years	Double-blind period	Sparsentan	89	7 (7.9)	5.899 [0.742, 46.871]	6.317 [0.759, 52.561]	6.5 [-0.9, 13.9]	0.072
		Irbesartan	75	1 (1.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
 Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.149
Yes	Double-blind period	Sparsentan	153	11 (7.2)	2.822 [0.918, 8.670]	2.963 [0.922, 9.518]	4.6 [-0.8, 10.1]	0.067
		Irbesartan	157	4 (2.5)				
No	Double-blind period	Sparsentan	49	2 (4.1)	0.612 [0.107, 3.498]	0.596 [0.095, 3.740]	-2.6 [-13.9, 8.7]	0.668
		Irbesartan	45	3 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AELC\_SSIM: Incidence of AESI hepatic disorders during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AELS\_SSIM: Incidence of AESI hepatic disorders during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.600
Male	Double-blind period	Sparsentan	139	10 (7.2)	1.470 [0.576, 3.752]	1.506 [0.557, 4.075]	2.3 [-4.0, 8.6]	0.462
		Irbesartan	143	7 (4.9)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	2.810 [0.301, 26.263]	2.900 [0.293, 28.688]	3.1 [-4.8, 10.9]	0.620
		Irbesartan	59	1 (1.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
 Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.642
<= 45 years	Double-blind period	Sparsentan	96	6 (6.3)	2.063 [0.531, 8.013]	2.133 [0.518, 8.786]	3.2 [-3.7, 10.1]	0.326
		Irbesartan	99	3 (3.0)				
> 45 years	Double-blind period	Sparsentan	106	7 (6.6)	1.360 [0.446, 4.149]	1.386 [0.425, 4.515]	1.7 [-5.5, 9.0]	0.768
		Irbesartan	103	5 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.682
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	8 (7.8)	2.137 [0.664, 6.883]	2.234 [0.652, 7.659]	4.2 [-3.1, 11.4]	0.240
		Irbesartan	109	4 (3.7)				
> 40 years	Double-blind period	Sparsentan	91	4 (4.4)	0.967 [0.250, 3.747]	0.966 [0.234, 3.986]	-0.1 [-7.3, 7.0]	1.000
		Irbesartan	88	4 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.410
North America	Double-blind period	Sparsentan	35	2 (5.7)	1.314 [0.195, 8.876]	1.333 [0.178, 9.964]	1.4 [-10.8, 13.6]	1.000
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	9 (9.2)	2.640 [0.839, 8.310]	2.806 [0.836, 9.414]	5.7 [-1.9, 13.3]	0.094
		Irbesartan	115	4 (3.5)				
Asia Pacific	Double-blind period	Sparsentan	69	2 (2.9)	0.594 [0.087, 4.059]	0.582 [0.079, 4.298]	-2.0 [-11.6, 7.7]	0.628
		Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.949
< 27 kg/m**2	Double-blind period	Sparsentan	84	6 (7.1)	1.679 [0.490, 5.745]	1.731 [0.471, 6.357]	2.9 [-5.1, 10.9]	0.520
		Irbesartan	94	4 (4.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	7 (5.9)	1.587 [0.478, 5.270]	1.624 [0.462, 5.710]	2.2 [-4.3, 8.7]	0.544
		Irbesartan	107	4 (3.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

<u>AESI acute pancreatitis</u>					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	6 (8.5)				NE
		Irbesartan	74	2 (2.7)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5)				NE
		Irbesartan	55	4 (7.3)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)				NE
		Irbesartan	36	0 (0.0)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)				NE
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.640
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	9 (7.1)	1.306 [0.502, 3.400]	1.329 [0.480, 3.685]	1.7 [-5.1, 8.4]	0.616
		Irbesartan	129	7 (5.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	3.918 [0.454, 33.802]	4.178 [0.450, 38.818]	6.1 [-4.6, 16.8]	0.362
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

<u>AESI acute pancreatitis</u>					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	3 (3.7)				NE
		Irbesartan	80	4 (5.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	6 (13.3)				NE
		Irbesartan	49	3 (6.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)				NE
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)				NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
 Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.241
<= 1.75 g/day	Double-blind period	Sparsentan	98	5 (5.1)	0.959 [0.287, 3.207]	0.957 [0.268, 3.419]	-0.2 [-7.5, 7.1]	1.000
		Irbesartan	94	5 (5.3)				
> 1.75 g/day	Double-blind period	Sparsentan	104	8 (7.7)	2.769 [0.755, 10.154]	2.917 [0.752, 11.312]	4.9 [-2.0, 11.8]	0.129
		Irbesartan	108	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.952
Yes	Double-blind period	Sparsentan	88	7 (8.0)	1.651 [0.501, 5.433]	1.707 [0.481, 6.059]	3.1 [-5.3, 11.6]	0.537
		Irbesartan	83	4 (4.8)				
No	Double-blind period	Sparsentan	114	6 (5.3)	1.566 [0.454, 5.404]	1.597 [0.439, 5.815]	1.9 [-4.2, 8.0]	0.532
		Irbesartan	119	4 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.939
<= 5 years	Double-blind period	Sparsentan	113	7 (6.2)	1.573 [0.514, 4.819]	1.611 [0.497, 5.227]	2.3 [-4.2, 8.7]	0.556
		Irbesartan	127	5 (3.9)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	1.685 [0.436, 6.511]	1.735 [0.419, 7.188]	2.7 [-5.3, 10.8]	0.510
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
 Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.095
Yes	Double-blind period	Sparsentan	153	12 (7.8)	2.463 [0.889, 6.824]	2.587 [0.889, 7.529]	4.7 [-1.1, 10.4]	0.084
		Irbesartan	157	5 (3.2)				
No	Double-blind period	Sparsentan	49	1 (2.0)	0.306 [0.033, 2.838]	0.292 [0.029, 2.911]	-4.6 [-15.1, 5.8]	0.346
		Irbesartan	45	3 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.600
Male	Double-blind period	Sparsentan	139	10 (7.2)	1.470 [0.576, 3.752]	1.506 [0.557, 4.075]	2.3 [-4.0, 8.6]	0.462
		Irbesartan	143	7 (4.9)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	2.810 [0.301, 26.263]	2.900 [0.293, 28.688]	3.1 [-4.8, 10.9]	0.620
		Irbesartan	59	1 (1.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.642
<= 45 years	Double-blind period	Sparsentan	96	6 (6.3)	2.063 [0.531, 8.013]	2.133 [0.518, 8.786]	3.2 [-3.7, 10.1]	0.326
		Irbesartan	99	3 (3.0)				
> 45 years	Double-blind period	Sparsentan	106	7 (6.6)	1.360 [0.446, 4.149]	1.386 [0.425, 4.515]	1.7 [-5.5, 9.0]	0.768
		Irbesartan	103	5 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
 Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.682
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	8 (7.8)	2.137 [0.664, 6.883]	2.234 [0.652, 7.659]	4.2 [-3.1, 11.4]	0.240
		Irbesartan	109	4 (3.7)				
> 40 years	Double-blind period	Sparsentan	91	4 (4.4)	0.967 [0.250, 3.747]	0.966 [0.234, 3.986]	-0.1 [-7.3, 7.0]	1.000
		Irbesartan	88	4 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.410
North America	Double-blind period	Sparsentan	35	2 (5.7)	1.314 [0.195, 8.876]	1.333 [0.178, 9.964]	1.4 [-10.8, 13.6]	1.000
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	9 (9.2)	2.640 [0.839, 8.310]	2.806 [0.836, 9.414]	5.7 [-1.9, 13.3]	0.094
		Irbesartan	115	4 (3.5)				
Asia Pacific	Double-blind period	Sparsentan	69	2 (2.9)	0.594 [0.087, 4.059]	0.582 [0.079, 4.298]	-2.0 [-11.6, 7.7]	0.628
		Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.949
< 27 kg/m**2	Double-blind period	Sparsentan	84	6 (7.1)	1.679 [0.490, 5.745]	1.731 [0.471, 6.357]	2.9 [-5.1, 10.9]	0.520
		Irbesartan	94	4 (4.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	7 (5.9)	1.587 [0.478, 5.270]	1.624 [0.462, 5.710]	2.2 [-4.3, 8.7]	0.544
		Irbesartan	107	4 (3.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	6 (8.5)				NE
		Irbesartan	74	2 (2.7)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5)				NE
		Irbesartan	55	4 (7.3)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)				NE
		Irbesartan	36	0 (0.0)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)				NE
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.640
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	9 (7.1)	1.306 [0.502, 3.400]	1.329 [0.480, 3.685]	1.7 [-5.1, 8.4]	0.616
		Irbesartan	129	7 (5.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	3.918 [0.454, 33.802]	4.178 [0.450, 38.818]	6.1 [-4.6, 16.8]	0.362
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
 Safety Set

<u>AESI acute pancreatitis - non-severe</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	3 (3.7)				NE
		Irbesartan	80	4 (5.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	6 (13.3)				NE
		Irbesartan	49	3 (6.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)				NE
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)				NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
 Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.241
<= 1.75 g/day	Double-blind period	Sparsentan	98	5 (5.1)	0.959 [0.287, 3.207]	0.957 [0.268, 3.419]	-0.2 [-7.5, 7.1]	1.000
		Irbesartan	94	5 (5.3)				
> 1.75 g/day	Double-blind period	Sparsentan	104	8 (7.7)	2.769 [0.755, 10.154]	2.917 [0.752, 11.312]	4.9 [-2.0, 11.8]	0.129
		Irbesartan	108	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.952
Yes	Double-blind period	Sparsentan	88	7 (8.0)	1.651 [0.501, 5.433]	1.707 [0.481, 6.059]	3.1 [-5.3, 11.6]	0.537
		Irbesartan	83	4 (4.8)				
No	Double-blind period	Sparsentan	114	6 (5.3)	1.566 [0.454, 5.404]	1.597 [0.439, 5.815]	1.9 [-4.2, 8.0]	0.532
		Irbesartan	119	4 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.939
<= 5 years	Double-blind period	Sparsentan	113	7 (6.2)	1.573 [0.514, 4.819]	1.611 [0.497, 5.227]	2.3 [-4.2, 8.7]	0.556
		Irbesartan	127	5 (3.9)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	1.685 [0.436, 6.511]	1.735 [0.419, 7.188]	2.7 [-5.3, 10.8]	0.510
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
 Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.095
Yes	Double-blind period	Sparsentan	153	12 (7.8)	2.463 [0.889, 6.824]	2.587 [0.889, 7.529]	4.7 [-1.1, 10.4]	0.084
		Irbesartan	157	5 (3.2)				
No	Double-blind period	Sparsentan	49	1 (2.0)	0.306 [0.033, 2.838]	0.292 [0.029, 2.911]	-4.6 [-15.1, 5.8]	0.346
		Irbesartan	45	3 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEPC\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEPS\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.985
Male	Double-blind period	Sparsentan	139	18 (12.9)	1.543 [0.772, 3.083]	1.624 [0.751, 3.511]	4.6 [-3.3, 12.5]	0.249
		Irbesartan	143	12 (8.4)				
Female	Double-blind period	Sparsentan	63	10 (15.9)	1.561 [0.605, 4.027]	1.667 [0.565, 4.915]	5.7 [-7.8, 19.2]	0.426
		Irbesartan	59	6 (10.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
 Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.322
<= 45 years	Double-blind period	Sparsentan	96	16 (16.7)	2.063 [0.926, 4.594]	2.275 [0.925, 5.597]	8.6 [-1.6, 18.8]	0.082
		Irbesartan	99	8 (8.1)				
> 45 years	Double-blind period	Sparsentan	106	12 (11.3)	1.166 [0.527, 2.580]	1.187 [0.489, 2.882]	1.6 [-7.7, 10.9]	0.823
		Irbesartan	103	10 (9.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.636
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.600 + [0.014, 26.475]	0.579 + [0.010, 33.502]	0.0 [NE, NE]	NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	15 (14.7)	2.004 [0.887, 4.524]	2.177 [0.881, 5.379]	7.4 [-2.0, 16.8]	0.121
		Irbesartan	109	8 (7.3)				
> 40 years	Double-blind period	Sparsentan	91	13 (14.3)	1.257 [0.582, 2.717]	1.300 [0.538, 3.141]	2.9 [-8.0, 13.8]	0.657
		Irbesartan	88	10 (11.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
 Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.951
North America	Double-blind period	Sparsentan	35	1 (2.9)	1.314 [0.085, 20.288]	1.324 [0.080, 21.926]	0.7 [-8.8, 10.1]	1.000
		Irbesartan	46	1 (2.2)				
Europe	Double-blind period	Sparsentan	98	18 (18.4)	1.625 [0.839, 3.145]	1.765 [0.816, 3.817]	7.1 [-3.5, 17.6]	0.174
		Irbesartan	115	13 (11.3)				
Asia Pacific	Double-blind period	Sparsentan	69	9 (13.0)	1.337 [0.439, 4.067]	1.388 [0.399, 4.828]	3.3 [-10.7, 17.3]	0.764
		Irbesartan	41	4 (9.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
 Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.198
< 27 kg/m**2	Double-blind period	Sparsentan	84	9 (10.7)	1.007 [0.430, 2.359]	1.008 [0.389, 2.614]	0.1 [-10.1, 10.3]	1.000
		Irbesartan	94	10 (10.6)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	19 (16.1)	2.154 [0.984, 4.715]	2.375 [0.993, 5.679]	8.6 [-0.6, 17.8]	0.064
		Irbesartan	107	8 (7.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.388
eGFR Low and UP High	Double-blind period	Sparsentan	71	10 (14.1)	1.303 [0.545, 3.112]	1.352 [0.501, 3.650]	3.3 [-8.9, 15.4]	0.620
		Irbesartan	74	8 (10.8)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	9 (16.4)	1.800 [0.644, 5.029]	1.957 [0.611, 6.268]	7.3 [-6.9, 21.5]	0.392
		Irbesartan	55	5 (9.1)				
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	0.778 [0.227, 2.669]	0.752 [0.185, 3.057]	-3.1 [-20.9, 14.8]	0.736
		Irbesartan	36	5 (13.9)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	5 (12.8)	10.450 + [0.598, 182.621]	11.957 + [0.637, 224.309]	12.8 [-0.3, 25.9]	0.055
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.677
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	19 (15.0)	1.608 [0.815, 3.174]	1.715 [0.795, 3.699]	5.7 [-3.1, 14.4]	0.184
		Irbesartan	129	12 (9.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.980 [0.260, 3.695]	0.978 [0.230, 4.155]	-0.2 [-13.2, 12.8]	1.000
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	5 (19.2)	2.404 [0.513, 11.271]	2.738 [0.479, 15.651]	11.2 [-11.2, 33.7]	0.419
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.366
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	8 (9.8)	0.976 [0.385, 2.473]	0.973 [0.347, 2.732]	-0.2 [-10.7, 10.2]	1.000
		Irbesartan	80	8 (10.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	11 (24.4)	2.994 [1.027, 8.734]	3.640 [1.066, 12.427]	16.3 [-0.6, 33.1]	0.047 *
		Irbesartan	49	4 (8.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.980 [0.260, 3.695]	0.978 [0.230, 4.155]	-0.2 [-13.2, 12.8]	1.000
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	5 (19.2)	2.404 [0.513, 11.271]	2.738 [0.479, 15.651]	11.2 [-11.2, 33.7]	0.419
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.302
<= 1.75 g/day	Double-blind period	Sparsentan	98	14 (14.3)	2.238 [0.898, 5.580]	2.444 [0.897, 6.658]	7.9 [-1.6, 17.5]	0.098
		Irbesartan	94	6 (6.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	14 (13.5)	1.212 [0.588, 2.495]	1.244 [0.546, 2.834]	2.4 [-7.4, 12.1]	0.678
		Irbesartan	108	12 (11.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
 Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.115
Yes	Double-blind period	Sparsentan	88	13 (14.8)	1.022 [0.495, 2.110]	1.026 [0.439, 2.397]	0.3 [-11.4, 12.1]	1.000
		Irbesartan	83	12 (14.5)				
No	Double-blind period	Sparsentan	114	15 (13.2)	2.610 [1.049, 6.491]	2.854 [1.066, 7.637]	8.1 [-0.1, 16.3]	0.039 *
		Irbesartan	119	6 (5.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
 Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.033 #
<= 5 years	Double-blind period	Sparsentan	113	19 (16.8)	2.669 [1.216, 5.859]	3.007 [1.261, 7.171]	10.5 [1.6, 19.4]	0.013 *
		Irbesartan	127	8 (6.3)				
> 5 years	Double-blind period	Sparsentan	89	9 (10.1)	0.758 [0.325, 1.768]	0.731 [0.281, 1.906]	-3.2 [-14.4, 7.9]	0.626
		Irbesartan	75	10 (13.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
 Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.713
Yes	Double-blind period	Sparsentan	153	24 (15.7)	1.642 [0.896, 3.008]	1.761 [0.885, 3.503]	6.1 [-1.9, 14.1]	0.124
		Irbesartan	157	15 (9.6)				
No	Double-blind period	Sparsentan	49	4 (8.2)	1.224 [0.290, 5.174]	1.244 [0.263, 5.892]	1.5 [-11.2, 14.2]	1.000
		Irbesartan	45	3 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
 Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.985
Male	Double-blind period	Sparsentan	139	18 (12.9)	1.543 [0.772, 3.083]	1.624 [0.751, 3.511]	4.6 [-3.3, 12.5]	0.249
		Irbesartan	143	12 (8.4)				
Female	Double-blind period	Sparsentan	63	10 (15.9)	1.561 [0.605, 4.027]	1.667 [0.565, 4.915]	5.7 [-7.8, 19.2]	0.426
		Irbesartan	59	6 (10.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
 Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.322
<= 45 years	Double-blind period	Sparsentan	96	16 (16.7)	2.063 [0.926, 4.594]	2.275 [0.925, 5.597]	8.6 [-1.6, 18.8]	0.082
		Irbesartan	99	8 (8.1)				
> 45 years	Double-blind period	Sparsentan	106	12 (11.3)	1.166 [0.527, 2.580]	1.187 [0.489, 2.882]	1.6 [-7.7, 10.9]	0.823
		Irbesartan	103	10 (9.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.636
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.600 + [0.014, 26.475]	0.579 + [0.010, 33.502]	0.0 [NE, NE]	NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	15 (14.7)	2.004 [0.887, 4.524]	2.177 [0.881, 5.379]	7.4 [-2.0, 16.8]	0.121
		Irbesartan	109	8 (7.3)				
> 40 years	Double-blind period	Sparsentan	91	13 (14.3)	1.257 [0.582, 2.717]	1.300 [0.538, 3.141]	2.9 [-8.0, 13.8]	0.657
		Irbesartan	88	10 (11.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.951
North America	Double-blind period	Sparsentan	35	1 (2.9)	1.314 [0.085, 20.288]	1.324 [0.080, 21.926]	0.7 [-8.8, 10.1]	1.000
		Irbesartan	46	1 (2.2)				
Europe	Double-blind period	Sparsentan	98	18 (18.4)	1.625 [0.839, 3.145]	1.765 [0.816, 3.817]	7.1 [-3.5, 17.6]	0.174
		Irbesartan	115	13 (11.3)				
Asia Pacific	Double-blind period	Sparsentan	69	9 (13.0)	1.337 [0.439, 4.067]	1.388 [0.399, 4.828]	3.3 [-10.7, 17.3]	0.764
		Irbesartan	41	4 (9.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.198
< 27 kg/m**2	Double-blind period	Sparsentan	84	9 (10.7)	1.007 [0.430, 2.359]	1.008 [0.389, 2.614]	0.1 [-10.1, 10.3]	1.000
		Irbesartan	94	10 (10.6)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	19 (16.1)	2.154 [0.984, 4.715]	2.375 [0.993, 5.679]	8.6 [-0.6, 17.8]	0.064
		Irbesartan	107	8 (7.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.388
eGFR Low and UP High	Double-blind period	Sparsentan	71	10 (14.1)	1.303 [0.545, 3.112]	1.352 [0.501, 3.650]	3.3 [-8.9, 15.4]	0.620
		Irbesartan	74	8 (10.8)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	9 (16.4)	1.800 [0.644, 5.029]	1.957 [0.611, 6.268]	7.3 [-6.9, 21.5]	0.392
		Irbesartan	55	5 (9.1)				
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	0.778 [0.227, 2.669]	0.752 [0.185, 3.057]	-3.1 [-20.9, 14.8]	0.736
		Irbesartan	36	5 (13.9)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	5 (12.8)	10.450 + [0.598, 182.621]	11.957 + [0.637, 224.309]	12.8 [-0.3, 25.9]	0.055
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.677
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	19 (15.0)	1.608 [0.815, 3.174]	1.715 [0.795, 3.699]	5.7 [-3.1, 14.4]	0.184
		Irbesartan	129	12 (9.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.980 [0.260, 3.695]	0.978 [0.230, 4.155]	-0.2 [-13.2, 12.8]	1.000
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	5 (19.2)	2.404 [0.513, 11.271]	2.738 [0.479, 15.651]	11.2 [-11.2, 33.7]	0.419
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
 Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.366
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	8 (9.8)	0.976 [0.385, 2.473]	0.973 [0.347, 2.732]	-0.2 [-10.7, 10.2]	1.000
		Irbesartan	80	8 (10.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	11 (24.4)	2.994 [1.027, 8.734]	3.640 [1.066, 12.427]	16.3 [-0.6, 33.1]	0.047 *
		Irbesartan	49	4 (8.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.980 [0.260, 3.695]	0.978 [0.230, 4.155]	-0.2 [-13.2, 12.8]	1.000
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	5 (19.2)	2.404 [0.513, 11.271]	2.738 [0.479, 15.651]	11.2 [-11.2, 33.7]	0.419
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
 Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.302
<= 1.75 g/day	Double-blind period	Sparsentan	98	14 (14.3)	2.238 [0.898, 5.580]	2.444 [0.897, 6.658]	7.9 [-1.6, 17.5]	0.098
		Irbesartan	94	6 (6.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	14 (13.5)	1.212 [0.588, 2.495]	1.244 [0.546, 2.834]	2.4 [-7.4, 12.1]	0.678
		Irbesartan	108	12 (11.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
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 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
 Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.115
Yes	Double-blind period	Sparsentan	88	13 (14.8)	1.022 [0.495, 2.110]	1.026 [0.439, 2.397]	0.3 [-11.4, 12.1]	1.000
		Irbesartan	83	12 (14.5)				
No	Double-blind period	Sparsentan	114	15 (13.2)	2.610 [1.049, 6.491]	2.854 [1.066, 7.637]	8.1 [-0.1, 16.3]	0.039 *
		Irbesartan	119	6 (5.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
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 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.033 #
<= 5 years	Double-blind period	Sparsentan	113	19 (16.8)	2.669 [1.216, 5.859]	3.007 [1.261, 7.171]	10.5 [1.6, 19.4]	0.013 *
		Irbesartan	127	8 (6.3)				
> 5 years	Double-blind period	Sparsentan	89	9 (10.1)	0.758 [0.325, 1.768]	0.731 [0.281, 1.906]	-3.2 [-14.4, 7.9]	0.626
		Irbesartan	75	10 (13.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
 Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.713
Yes	Double-blind period	Sparsentan	153	24 (15.7)	1.642 [0.896, 3.008]	1.761 [0.885, 3.503]	6.1 [-1.9, 14.1]	0.124
		Irbesartan	157	15 (9.6)				
No	Double-blind period	Sparsentan	49	4 (8.2)	1.224 [0.290, 5.174]	1.244 [0.263, 5.892]	1.5 [-11.2, 14.2]	1.000
		Irbesartan	45	3 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEOC\_SSIM: Incidence of AESI fluid retention during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEOS\_SSIM: Incidence of AESI fluid retention during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
 Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.692
Male	Double-blind period	Sparsentan	139	10 (7.2)	1.715 [0.640, 4.591]	1.770 [0.625, 5.009]	3.0 [-3.1, 9.1]	0.312
		Irbesartan	143	6 (4.2)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	2.810 [0.301, 26.263]	2.900 [0.293, 28.688]	3.1 [-4.8, 10.9]	0.620
		Irbesartan	59	1 (1.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.454
<= 45 years	Double-blind period	Sparsentan	96	5 (5.2)	1.289 [0.357, 4.657]	1.305 [0.340, 5.013]	1.2 [-5.8, 8.1]	0.745
		Irbesartan	99	4 (4.0)				
> 45 years	Double-blind period	Sparsentan	106	8 (7.5)	2.591 [0.707, 9.497]	2.721 [0.701, 10.558]	4.6 [-2.3, 11.6]	0.215
		Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
 Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.585
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.600 + [0.014, 26.475]	0.579 + [0.010, 33.502]	0.0 [NE, NE]	NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	6 (5.9)	3.206 [0.662, 15.523]	3.344 [0.659, 16.961]	4.0 [-2.1, 10.2]	0.159
		Irbesartan	109	2 (1.8)				
> 40 years	Double-blind period	Sparsentan	91	7 (7.7)	1.354 [0.446, 4.106]	1.383 [0.422, 4.534]	2.0 [-6.4, 10.4]	0.767
		Irbesartan	88	5 (5.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
 Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.335
North America	Double-blind period	Sparsentan	35	0 (0.0)	1.306 + [0.027, 64.228]	1.310 + [0.025, 67.634]	0.0 [NE, NE]	NE
		Irbesartan	46	0 (0.0)				
Europe	Double-blind period	Sparsentan	98	6 (6.1)	3.520 [0.727, 17.048]	3.685 [0.726, 18.690]	4.4 [-1.9, 10.6]	0.147
		Irbesartan	115	2 (1.7)				
Asia Pacific	Double-blind period	Sparsentan	69	7 (10.1)	0.832 [0.282, 2.451]	0.813 [0.240, 2.750]	-2.1 [-16.3, 12.2]	0.759
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.812
< 27 kg/m**2	Double-blind period	Sparsentan	84	8 (9.5)	1.790 [0.609, 5.261]	1.874 [0.588, 5.968]	4.2 [-4.7, 13.1]	0.389
		Irbesartan	94	5 (5.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	5 (4.2)	2.267 [0.449, 11.442]	2.323 [0.441, 12.232]	2.4 [-3.0, 7.7]	0.450
		Irbesartan	107	2 (1.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
 Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.798
eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	2.085 [0.657, 6.618]	2.222 [0.638, 7.737]	5.9 [-4.5, 16.2]	0.238
		Irbesartan	74	4 (5.4)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)	1.000 [0.146, 6.848]	1.000 [0.136, 7.364]	0.0 [-8.8, 8.8]	1.000
		Irbesartan	55	2 (3.6)				
eGFR High and UP High	Double-blind period	Sparsentan	37	1 (2.7)	0.973 [0.063, 14.972]	0.972 [0.058, 16.158]	-0.1 [-10.3, 10.2]	1.000
		Irbesartan	36	1 (2.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)	4.750 + [0.236, 95.763]	5.000 + [0.232, 107.700]	5.1 [-4.4, 14.7]	0.494
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.379
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	10 (7.9)	1.693 [0.634, 4.520]	1.752 [0.617, 4.974]	3.2 [-3.5, 9.9]	0.314
		Irbesartan	129	6 (4.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	3 (6.1)	6.860 + [0.364, 129.358]	7.301 + [0.367, 145.240]	6.1 [-2.7, 14.9]	0.242
		Irbesartan	48	0 (0.0)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.321 + [0.014, 7.528]	0.308 + [0.012, 7.927]	-4.0 [-15.6, 7.6]	0.490
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
 Safety Set

AESI anemia					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	6 (7.3)				NE
		Irbesartan	80	3 (3.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)				NE
		Irbesartan	49	3 (6.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	3 (6.1)				NE
		Irbesartan	48	0 (0.0)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.712
<= 1.75 g/day	Double-blind period	Sparsentan	98	5 (5.1)	2.398	2.473	3.0	0.445
		Irbesartan	94	2 (2.1)	[0.477, 12.059]	[0.468, 13.072]	[-3.3, 9.3]	
> 1.75 g/day	Double-blind period	Sparsentan	104	8 (7.7)	1.662	1.717	3.1	0.402
		Irbesartan	108	5 (4.6)	[0.562, 4.914]	[0.543, 5.429]	[-4.4, 10.5]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
 Safety Set

AESI anemia					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.207
Yes	Double-blind period	Sparsentan	88	4 (4.5)	0.943 [0.244, 3.649]	0.940 [0.227, 3.889]	-0.3 [-7.8, 7.2]	1.000
		Irbesartan	83	4 (4.8)				
No	Double-blind period	Sparsentan	114	9 (7.9)	3.132 [0.870, 11.276]	3.314 [0.874, 12.570]	5.4 [-1.2, 11.9]	0.079
		Irbesartan	119	3 (2.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.260
<= 5 years	Double-blind period	Sparsentan	113	7 (6.2)	1.311 [0.454, 3.787]	1.332 [0.434, 4.086]	1.5 [-5.1, 8.1]	0.777
		Irbesartan	127	6 (4.7)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	5.056 [0.623, 41.067]	5.349 [0.629, 45.470]	5.4 [-1.6, 12.5]	0.127
		Irbesartan	75	1 (1.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
 Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.701
Yes	Double-blind period	Sparsentan	153	10 (6.5)	1.710 [0.637, 4.590]	1.760 [0.624, 4.967]	2.7 [-2.9, 8.3]	0.314
		Irbesartan	157	6 (3.8)				
No	Double-blind period	Sparsentan	49	3 (6.1)	2.755 [0.297, 25.538]	2.870 [0.288, 28.639]	3.9 [-6.2, 14.0]	0.618
		Irbesartan	45	1 (2.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.692
Male	Double-blind period	Sparsentan	139	10 (7.2)	1.715 [0.640, 4.591]	1.770 [0.625, 5.009]	3.0 [-3.1, 9.1]	0.312
		Irbesartan	143	6 (4.2)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	2.810 [0.301, 26.263]	2.900 [0.293, 28.688]	3.1 [-4.8, 10.9]	0.620
		Irbesartan	59	1 (1.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.454
<= 45 years	Double-blind period	Sparsentan	96	5 (5.2)	1.289 [0.357, 4.657]	1.305 [0.340, 5.013]	1.2 [-5.8, 8.1]	0.745
		Irbesartan	99	4 (4.0)				
> 45 years	Double-blind period	Sparsentan	106	8 (7.5)	2.591 [0.707, 9.497]	2.721 [0.701, 10.558]	4.6 [-2.3, 11.6]	0.215
		Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.585
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.600 + [0.014, 26.475]	0.579 + [0.010, 33.502]	0.0 [NE, NE]	NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	6 (5.9)	3.206 [0.662, 15.523]	3.344 [0.659, 16.961]	4.0 [-2.1, 10.2]	0.159
		Irbesartan	109	2 (1.8)				
> 40 years	Double-blind period	Sparsentan	91	7 (7.7)	1.354 [0.446, 4.106]	1.383 [0.422, 4.534]	2.0 [-6.4, 10.4]	0.767
		Irbesartan	88	5 (5.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.335
North America	Double-blind period	Sparsentan	35	0 (0.0)	1.306 + [0.027, 64.228]	1.310 + [0.025, 67.634]	0.0 [NE, NE]	NE
		Irbesartan	46	0 (0.0)				
Europe	Double-blind period	Sparsentan	98	6 (6.1)	3.520 [0.727, 17.048]	3.685 [0.726, 18.690]	4.4 [-1.9, 10.6]	0.147
		Irbesartan	115	2 (1.7)				
Asia Pacific	Double-blind period	Sparsentan	69	7 (10.1)	0.832 [0.282, 2.451]	0.813 [0.240, 2.750]	-2.1 [-16.3, 12.2]	0.759
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.812
< 27 kg/m**2	Double-blind period	Sparsentan	84	8 (9.5)	1.790 [0.609, 5.261]	1.874 [0.588, 5.968]	4.2 [-4.7, 13.1]	0.389
		Irbesartan	94	5 (5.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	5 (4.2)	2.267 [0.449, 11.442]	2.323 [0.441, 12.232]	2.4 [-3.0, 7.7]	0.450
		Irbesartan	107	2 (1.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.798
eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	2.085 [0.657, 6.618]	2.222 [0.638, 7.737]	5.9 [-4.5, 16.2]	0.238
		Irbesartan	74	4 (5.4)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)	1.000 [0.146, 6.848]	1.000 [0.136, 7.364]	0.0 [-8.8, 8.8]	1.000
		Irbesartan	55	2 (3.6)				
eGFR High and UP High	Double-blind period	Sparsentan	37	1 (2.7)	0.973 [0.063, 14.972]	0.972 [0.058, 16.158]	-0.1 [-10.3, 10.2]	1.000
		Irbesartan	36	1 (2.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)	4.750 + [0.236, 95.763]	5.000 + [0.232, 107.700]	5.1 [-4.4, 14.7]	0.494
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.379
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	10 (7.9)	1.693 [0.634, 4.520]	1.752 [0.617, 4.974]	3.2 [-3.5, 9.9]	0.314
		Irbesartan	129	6 (4.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	3 (6.1)	6.860 + [0.364, 129.358]	7.301 + [0.367, 145.240]	6.1 [-2.7, 14.9]	0.242
		Irbesartan	48	0 (0.0)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.321 + [0.014, 7.528]	0.308 + [0.012, 7.927]	-4.0 [-15.6, 7.6]	0.490
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	6 (7.3)				NE
		Irbesartan	80	3 (3.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)				NE
		Irbesartan	49	3 (6.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	3 (6.1)				NE
		Irbesartan	48	0 (0.0)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.712
<= 1.75 g/day	Double-blind period	Sparsentan	98	5 (5.1)	2.398 [0.477, 12.059]	2.473 [0.468, 13.072]	3.0 [-3.3, 9.3]	0.445
		Irbesartan	94	2 (2.1)				
> 1.75 g/day	Double-blind period	Sparsentan	104	8 (7.7)	1.662 [0.562, 4.914]	1.717 [0.543, 5.429]	3.1 [-4.4, 10.5]	0.402
		Irbesartan	108	5 (4.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.207
Yes	Double-blind period	Sparsentan	88	4 (4.5)	0.943 [0.244, 3.649]	0.940 [0.227, 3.889]	-0.3 [-7.8, 7.2]	1.000
		Irbesartan	83	4 (4.8)				
No	Double-blind period	Sparsentan	114	9 (7.9)	3.132 [0.870, 11.276]	3.314 [0.874, 12.570]	5.4 [-1.2, 11.9]	0.079
		Irbesartan	119	3 (2.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.260
<= 5 years	Double-blind period	Sparsentan	113	7 (6.2)	1.311 [0.454, 3.787]	1.332 [0.434, 4.086]	1.5 [-5.1, 8.1]	0.777
		Irbesartan	127	6 (4.7)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	5.056 [0.623, 41.067]	5.349 [0.629, 45.470]	5.4 [-1.6, 12.5]	0.127
		Irbesartan	75	1 (1.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.701
Yes	Double-blind period	Sparsentan	153	10 (6.5)	1.710 [0.637, 4.590]	1.760 [0.624, 4.967]	2.7 [-2.9, 8.3]	0.314
		Irbesartan	157	6 (3.8)				
No	Double-blind period	Sparsentan	49	3 (6.1)	2.755 [0.297, 25.538]	2.870 [0.288, 28.639]	3.9 [-6.2, 14.0]	0.618
		Irbesartan	45	1 (2.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEAC\_SSIM: Incidence of AESI anemia during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEAS\_SSIM: Incidence of AESI anemia during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
 Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.205
Male	Double-blind period	Sparsentan	139	18 (12.9)	1.543 [0.772, 3.083]	1.624 [0.751, 3.511]	4.6 [-3.3, 12.5]	0.249
		Irbesartan	143	12 (8.4)				
Female	Double-blind period	Sparsentan	63	5 (7.9)	0.669 [0.225, 1.992]	0.640 [0.192, 2.141]	-3.9 [-16.2, 8.3]	0.551
		Irbesartan	59	7 (11.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.771
<= 45 years	Double-blind period	Sparsentan	96	12 (12.5)	1.125 [0.522, 2.426]	1.143 [0.478, 2.731]	1.4 [-8.7, 11.5]	0.826
		Irbesartan	99	11 (11.1)				
> 45 years	Double-blind period	Sparsentan	106	11 (10.4)	1.336 [0.560, 3.187]	1.375 [0.530, 3.570]	2.6 [-6.1, 11.3]	0.632
		Irbesartan	103	8 (7.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.339
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.600 + [0.014, 26.475]	0.579 + [0.010, 33.502]	0.0 [NE, NE]	NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	13 (12.7)	0.926 [0.464, 1.850]	0.915 [0.412, 2.032]	-1.0 [-11.1, 9.1]	0.842
		Irbesartan	109	15 (13.8)				
> 40 years	Double-blind period	Sparsentan	91	10 (11.0)	2.418 [0.787, 7.424]	2.593 [0.782, 8.599]	6.4 [-2.4, 15.3]	0.163
		Irbesartan	88	4 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.371
North America	Double-blind period	Sparsentan	35	5 (14.3)	3.286 [0.677, 15.949]	3.667 [0.667, 20.156]	9.9 [-5.6, 25.5]	0.230
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	10 (10.2)	0.978 [0.442, 2.165]	0.975 [0.402, 2.366]	-0.2 [-9.4, 8.9]	1.000
		Irbesartan	115	12 (10.4)				
Asia Pacific	Double-blind period	Sparsentan	69	8 (11.6)	0.951 [0.333, 2.712]	0.944 [0.287, 3.107]	-0.6 [-15.1, 13.9]	1.000
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.976
< 27 kg/m**2	Double-blind period	Sparsentan	84	11 (13.1)	1.231 [0.551, 2.751]	1.266 [0.508, 3.151]	2.5 [-8.2, 13.1]	0.648
		Irbesartan	94	10 (10.6)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	12 (10.2)	1.209 [0.531, 2.755]	1.233 [0.498, 3.053]	1.8 [-6.7, 10.2]	0.819
		Irbesartan	107	9 (8.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.350
eGFR Low and UP High	Double-blind period	Sparsentan	71	14 (19.7)	1.621 [0.750, 3.507]	1.774 [0.714, 4.406]	7.6 [-5.7, 20.8]	0.259
		Irbesartan	74	9 (12.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	7 (12.7)	0.875 [0.341, 2.247]	0.857 [0.288, 2.551]	-1.8 [-16.5, 12.8]	1.000
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4)	4.868 + [0.242, 98.023]	5.141 + [0.238, 110.889]	5.4 [-4.6, 15.4]	0.493
		Irbesartan	36	0 (0.0)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0)	0.190 + [0.009, 3.831]	0.180 + [0.008, 3.872]	-5.4 [-15.3, 4.5]	0.234
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.745
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	21 (16.5)	1.333 [0.730, 2.435]	1.399 [0.693, 2.824]	4.1 [-5.3, 13.5]	0.378
		Irbesartan	129	16 (12.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.653 [0.114, 3.737]	0.638 [0.102, 4.000]	-2.2 [-13.0, 8.7]	0.678
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.805
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	17 (20.7)	1.185 [0.627, 2.239]	1.233 [0.562, 2.706]	3.2 [-10.1, 16.6]	0.691
		Irbesartan	80	14 (17.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	2.178 [0.419, 11.322]	2.293 [0.399, 13.171]	4.8 [-7.3, 16.9]	0.421
		Irbesartan	49	2 (4.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.653 [0.114, 3.737]	0.638 [0.102, 4.000]	-2.2 [-13.0, 8.7]	0.678
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
 Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.192
<= 1.75 g/day	Double-blind period	Sparsentan	98	7 (7.1)	0.746 [0.290, 1.922]	0.726 [0.259, 2.037]	-2.4 [-11.3, 6.4]	0.608
		Irbesartan	94	9 (9.6)				
> 1.75 g/day	Double-blind period	Sparsentan	104	16 (15.4)	1.662 [0.791, 3.492]	1.782 [0.769, 4.131]	6.1 [-3.6, 15.9]	0.211
		Irbesartan	108	10 (9.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.043 #
Yes	Double-blind period	Sparsentan	88	16 (18.2)	2.156 [0.934, 4.974]	2.413 [0.938, 6.207]	9.7 [-1.5, 21.0]	0.074
		Irbesartan	83	7 (8.4)				
No	Double-blind period	Sparsentan	114	7 (6.1)	0.609 [0.249, 1.492]	0.583 [0.221, 1.539]	-3.9 [-11.8, 3.9]	0.341
		Irbesartan	119	12 (10.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.089
<= 5 years	Double-blind period	Sparsentan	113	15 (13.3)	1.873 [0.853, 4.113]	2.007 [0.842, 4.784]	6.2 [-2.3, 14.7]	0.133
		Irbesartan	127	9 (7.1)				
> 5 years	Double-blind period	Sparsentan	89	8 (9.0)	0.674 [0.280, 1.621]	0.642 [0.240, 1.720]	-4.3 [-15.3, 6.6]	0.455
		Irbesartan	75	10 (13.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
 Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.049 #
Yes	Double-blind period	Sparsentan	153	21 (13.7)	1.658 [0.861, 3.191]	1.762 [0.848, 3.660]	5.4 [-2.2, 13.0]	0.147
		Irbesartan	157	13 (8.3)				
No	Double-blind period	Sparsentan	49	2 (4.1)	0.306 [0.065, 1.440]	0.277 [0.053, 1.448]	-9.3 [-22.8, 4.3]	0.147
		Irbesartan	45	6 (13.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.205
Male	Double-blind period	Sparsentan	139	18 (12.9)	1.543 [0.772, 3.083]	1.624 [0.751, 3.511]	4.6 [-3.3, 12.5]	0.249
		Irbesartan	143	12 (8.4)				
Female	Double-blind period	Sparsentan	63	5 (7.9)	0.669 [0.225, 1.992]	0.640 [0.192, 2.141]	-3.9 [-16.2, 8.3]	0.551
		Irbesartan	59	7 (11.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.771
<= 45 years	Double-blind period	Sparsentan	96	12 (12.5)	1.125 [0.522, 2.426]	1.143 [0.478, 2.731]	1.4 [-8.7, 11.5]	0.826
		Irbesartan	99	11 (11.1)				
> 45 years	Double-blind period	Sparsentan	106	11 (10.4)	1.336 [0.560, 3.187]	1.375 [0.530, 3.570]	2.6 [-6.1, 11.3]	0.632
		Irbesartan	103	8 (7.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.339
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.600 + [0.014, 26.475]	0.579 + [0.010, 33.502]	0.0 [NE, NE]	NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	13 (12.7)	0.926 [0.464, 1.850]	0.915 [0.412, 2.032]	-1.0 [-11.1, 9.1]	0.842
		Irbesartan	109	15 (13.8)				
> 40 years	Double-blind period	Sparsentan	91	10 (11.0)	2.418 [0.787, 7.424]	2.593 [0.782, 8.599]	6.4 [-2.4, 15.3]	0.163
		Irbesartan	88	4 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.371
North America	Double-blind period	Sparsentan	35	5 (14.3)	3.286 [0.677, 15.949]	3.667 [0.667, 20.156]	9.9 [-5.6, 25.5]	0.230
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	10 (10.2)	0.978 [0.442, 2.165]	0.975 [0.402, 2.366]	-0.2 [-9.4, 8.9]	1.000
		Irbesartan	115	12 (10.4)				
Asia Pacific	Double-blind period	Sparsentan	69	8 (11.6)	0.951 [0.333, 2.712]	0.944 [0.287, 3.107]	-0.6 [-15.1, 13.9]	1.000
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.976
< 27 kg/m**2	Double-blind period	Sparsentan	84	11 (13.1)	1.231 [0.551, 2.751]	1.266 [0.508, 3.151]	2.5 [-8.2, 13.1]	0.648
		Irbesartan	94	10 (10.6)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	12 (10.2)	1.209 [0.531, 2.755]	1.233 [0.498, 3.053]	1.8 [-6.7, 10.2]	0.819
		Irbesartan	107	9 (8.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.350
eGFR Low and UP High	Double-blind period	Sparsentan	71	14 (19.7)	1.621 [0.750, 3.507]	1.774 [0.714, 4.406]	7.6 [-5.7, 20.8]	0.259
		Irbesartan	74	9 (12.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	7 (12.7)	0.875 [0.341, 2.247]	0.857 [0.288, 2.551]	-1.8 [-16.5, 12.8]	1.000
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4)	4.868 + [0.242, 98.023]	5.141 + [0.238, 110.889]	5.4 [-4.6, 15.4]	0.493
		Irbesartan	36	0 (0.0)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0)	0.190 + [0.009, 3.831]	0.180 + [0.008, 3.872]	-5.4 [-15.3, 4.5]	0.234
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.745
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	21 (16.5)	1.333 [0.730, 2.435]	1.399 [0.693, 2.824]	4.1 [-5.3, 13.5]	0.378
		Irbesartan	129	16 (12.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.653 [0.114, 3.737]	0.638 [0.102, 4.000]	-2.2 [-13.0, 8.7]	0.678
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.805
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	17 (20.7)	1.185 [0.627, 2.239]	1.233 [0.562, 2.706]	3.2 [-10.1, 16.6]	0.691
		Irbesartan	80	14 (17.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	2.178 [0.419, 11.322]	2.293 [0.399, 13.171]	4.8 [-7.3, 16.9]	0.421
		Irbesartan	49	2 (4.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.653 [0.114, 3.737]	0.638 [0.102, 4.000]	-2.2 [-13.0, 8.7]	0.678
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.192
<= 1.75 g/day	Double-blind period	Sparsentan	98	7 (7.1)	0.746 [0.290, 1.922]	0.726 [0.259, 2.037]	-2.4 [-11.3, 6.4]	0.608
		Irbesartan	94	9 (9.6)				
> 1.75 g/day	Double-blind period	Sparsentan	104	16 (15.4)	1.662 [0.791, 3.492]	1.782 [0.769, 4.131]	6.1 [-3.6, 15.9]	0.211
		Irbesartan	108	10 (9.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.043 #
Yes	Double-blind period	Sparsentan	88	16 (18.2)	2.156 [0.934, 4.974]	2.413 [0.938, 6.207]	9.7 [-1.5, 21.0]	0.074
		Irbesartan	83	7 (8.4)				
No	Double-blind period	Sparsentan	114	7 (6.1)	0.609 [0.249, 1.492]	0.583 [0.221, 1.539]	-3.9 [-11.8, 3.9]	0.341
		Irbesartan	119	12 (10.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.089
<= 5 years	Double-blind period	Sparsentan	113	15 (13.3)	1.873 [0.853, 4.113]	2.007 [0.842, 4.784]	6.2 [-2.3, 14.7]	0.133
		Irbesartan	127	9 (7.1)				
> 5 years	Double-blind period	Sparsentan	89	8 (9.0)	0.674 [0.280, 1.621]	0.642 [0.240, 1.720]	-4.3 [-15.3, 6.6]	0.455
		Irbesartan	75	10 (13.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.049 #
Yes	Double-blind period	Sparsentan	153	21 (13.7)	1.658 [0.861, 3.191]	1.762 [0.848, 3.660]	5.4 [-2.2, 13.0]	0.147
		Irbesartan	157	13 (8.3)				
No	Double-blind period	Sparsentan	49	2 (4.1)	0.306 [0.065, 1.440]	0.277 [0.053, 1.448]	-9.3 [-22.8, 4.3]	0.147
		Irbesartan	45	6 (13.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEUC\_SSIM: Incidence of AESI hyperkalemia during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEUS\_SSIM: Incidence of AESI hyperkalemia during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
 Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.238
Male	Double-blind period	Sparsentan	139	5 (3.6)	0.429 [0.155, 1.185]	0.407 [0.140, 1.188]	-4.8 [-11.0, 1.4]	0.132
		Irbesartan	143	12 (8.4)				
Female	Double-blind period	Sparsentan	63	4 (6.3)	1.249 [0.292, 5.346]	1.266 [0.271, 5.909]	1.3 [-8.6, 11.1]	1.000
		Irbesartan	59	3 (5.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
 Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.493
<= 45 years	Double-blind period	Sparsentan	96	4 (4.2)	0.458 [0.146, 1.439]	0.435 [0.129, 1.463]	-4.9 [-12.9, 3.0]	0.251
		Irbesartan	99	9 (9.1)				
> 45 years	Double-blind period	Sparsentan	106	5 (4.7)	0.810 [0.255, 2.571]	0.800 [0.236, 2.708]	-1.1 [-8.1, 5.9]	0.766
		Irbesartan	103	6 (5.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.661
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	0.556 [0.044, 7.095]	0.500 [0.024, 10.251]	-8.9 [-65.1, 47.3]	1.000
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	4 (3.9)	0.427 [0.138, 1.320]	0.404 [0.123, 1.332]	-5.3 [-12.8, 2.3]	0.168
		Irbesartan	109	10 (9.2)				
> 40 years	Double-blind period	Sparsentan	91	4 (4.4)	0.967 [0.250, 3.747]	0.966 [0.234, 3.986]	-0.1 [-7.3, 7.0]	1.000
		Irbesartan	88	4 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
 Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.229
North America	Double-blind period	Sparsentan	35	2 (5.7)	1.314 [0.195, 8.876]	1.333 [0.178, 9.964]	1.4 [-10.8, 13.6]	1.000
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	5 (5.1)	0.838 [0.275, 2.558]	0.829 [0.255, 2.701]	-1.0 [-8.1, 6.1]	1.000
		Irbesartan	115	7 (6.1)				
Asia Pacific	Double-blind period	Sparsentan	69	2 (2.9)	0.198 [0.042, 0.936]	0.174 [0.033, 0.908]	-11.7 [-25.2, 1.7]	0.050
		Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.470
< 27 kg/m**2	Double-blind period	Sparsentan	84	3 (3.6)	0.420 [0.115, 1.530]	0.398 [0.102, 1.553]	-4.9 [-13.0, 3.1]	0.220
		Irbesartan	94	8 (8.5)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	6 (5.1)	0.777 [0.270, 2.240]	0.765 [0.249, 2.353]	-1.5 [-8.5, 5.6]	0.777
		Irbesartan	107	7 (6.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
 Safety Set

<u>AESI cardiac arrhythmias</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	5 (7.0) all n<10				NE
		Irbesartan	74	4 (5.4)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5) all n<10				NE
		Irbesartan	55	6 (10.9)				
eGFR High and UP High	Double-blind period	Sparsentan	37	1 (2.7) all n<10				NE
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0) all n<10				NE
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.934
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	7 (5.5)	0.646 [0.259, 1.615]	0.626 [0.235, 1.669]	-3.0 [-10.0, 4.0]	0.465
		Irbesartan	129	11 (8.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.490 [0.094, 2.550]	0.468 [0.082, 2.684]	-4.3 [-15.9, 7.4]	0.436
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
 Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.762
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	3 (3.7)	0.418 [0.112, 1.560]	0.396 [0.099, 1.589]	-5.1 [-13.7, 3.5]	0.208
		Irbesartan	80	7 (8.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	1.089 [0.289, 4.099]	1.098 [0.258, 4.675]	0.7 [-12.7, 14.2]	1.000
		Irbesartan	49	4 (8.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.490 [0.094, 2.550]	0.468 [0.082, 2.684]	-4.3 [-15.9, 7.4]	0.436
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
 Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.235
<= 1.75 g/day	Double-blind period	Sparsentan	98	6 (6.1)	0.959 [0.321, 2.869]	0.957 [0.297, 3.078]	-0.3 [-8.2, 7.6]	1.000
		Irbesartan	94	6 (6.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	3 (2.9)	0.346 [0.096, 1.243]	0.327 [0.086, 1.242]	-5.4 [-12.5, 1.6]	0.135
		Irbesartan	108	9 (8.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.304
Yes	Double-blind period	Sparsentan	88	5 (5.7)	0.429 [0.156, 1.181]	0.394 [0.131, 1.188]	-7.6 [-17.5, 2.4]	0.116
		Irbesartan	83	11 (13.3)				
No	Double-blind period	Sparsentan	114	4 (3.5)	1.044 [0.267, 4.075]	1.045 [0.255, 4.284]	0.1 [-5.4, 5.7]	1.000
		Irbesartan	119	4 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.113
<= 5 years	Double-blind period	Sparsentan	113	3 (2.7)	0.307 [0.088, 1.071]	0.288 [0.078, 1.058]	-6.0 [-12.6, 0.5]	0.056
		Irbesartan	127	11 (8.7)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	1.264 [0.370, 4.313]	1.283 [0.348, 4.728]	1.4 [-7.1, 9.9]	0.756
		Irbesartan	75	4 (5.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.647
Yes	Double-blind period	Sparsentan	153	9 (5.9)	0.660 [0.294, 1.479]	0.638 [0.268, 1.522]	-3.0 [-9.5, 3.4]	0.387
		Irbesartan	157	14 (8.9)				
No	Double-blind period	Sparsentan	49	0 (0.0)	0.307 + [0.013, 7.341]	0.300 + [0.012, 7.546]	-2.2 [-8.7, 4.2]	0.479
		Irbesartan	45	1 (2.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.238
Male	Double-blind period	Sparsentan	139	5 (3.6)	0.429 [0.155, 1.185]	0.407 [0.140, 1.188]	-4.8 [-11.0, 1.4]	0.132
		Irbesartan	143	12 (8.4)				
Female	Double-blind period	Sparsentan	63	4 (6.3)	1.249 [0.292, 5.346]	1.266 [0.271, 5.909]	1.3 [-8.6, 11.1]	1.000
		Irbesartan	59	3 (5.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.493
<= 45 years	Double-blind period	Sparsentan	96	4 (4.2)	0.458 [0.146, 1.439]	0.435 [0.129, 1.463]	-4.9 [-12.9, 3.0]	0.251
		Irbesartan	99	9 (9.1)				
> 45 years	Double-blind period	Sparsentan	106	5 (4.7)	0.810 [0.255, 2.571]	0.800 [0.236, 2.708]	-1.1 [-8.1, 5.9]	0.766
		Irbesartan	103	6 (5.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.661
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	0.556 [0.044, 7.095]	0.500 [0.024, 10.251]	-8.9 [-65.1, 47.3]	1.000
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	4 (3.9)	0.427 [0.138, 1.320]	0.404 [0.123, 1.332]	-5.3 [-12.8, 2.3]	0.168
		Irbesartan	109	10 (9.2)				
> 40 years	Double-blind period	Sparsentan	91	4 (4.4)	0.967 [0.250, 3.747]	0.966 [0.234, 3.986]	-0.1 [-7.3, 7.0]	1.000
		Irbesartan	88	4 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.229
North America	Double-blind period	Sparsentan	35	2 (5.7)	1.314 [0.195, 8.876]	1.333 [0.178, 9.964]	1.4 [-10.8, 13.6]	1.000
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	5 (5.1)	0.838 [0.275, 2.558]	0.829 [0.255, 2.701]	-1.0 [-8.1, 6.1]	1.000
		Irbesartan	115	7 (6.1)				
Asia Pacific	Double-blind period	Sparsentan	69	2 (2.9)	0.198 [0.042, 0.936]	0.174 [0.033, 0.908]	-11.7 [-25.2, 1.7]	0.050
		Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.470
< 27 kg/m**2	Double-blind period	Sparsentan	84	3 (3.6)	0.420 [0.115, 1.530]	0.398 [0.102, 1.553]	-4.9 [-13.0, 3.1]	0.220
		Irbesartan	94	8 (8.5)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	118	6 (5.1)	0.777 [0.270, 2.240]	0.765 [0.249, 2.353]	-1.5 [-8.5, 5.6]	0.777
		Irbesartan	107	7 (6.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
 Safety Set

<u>AESI cardiac arrhythmias - non-severe</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	5 (7.0) all n<10				NE
		Irbesartan	74	4 (5.4)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5) all n<10				NE
		Irbesartan	55	6 (10.9)				
eGFR High and UP High	Double-blind period	Sparsentan	37	1 (2.7) all n<10				NE
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0) all n<10				NE
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.934
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	7 (5.5)	0.646 [0.259, 1.615]	0.626 [0.235, 1.669]	-3.0 [-10.0, 4.0]	0.465
		Irbesartan	129	11 (8.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.490 [0.094, 2.550]	0.468 [0.082, 2.684]	-4.3 [-15.9, 7.4]	0.436
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.762
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	3 (3.7)	0.418 [0.112, 1.560]	0.396 [0.099, 1.589]	-5.1 [-13.7, 3.5]	0.208
		Irbesartan	80	7 (8.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	1.089 [0.289, 4.099]	1.098 [0.258, 4.675]	0.7 [-12.7, 14.2]	1.000
		Irbesartan	49	4 (8.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.490 [0.094, 2.550]	0.468 [0.082, 2.684]	-4.3 [-15.9, 7.4]	0.436
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.235
<= 1.75 g/day	Double-blind period	Sparsentan	98	6 (6.1)	0.959 [0.321, 2.869]	0.957 [0.297, 3.078]	-0.3 [-8.2, 7.6]	1.000
		Irbesartan	94	6 (6.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	3 (2.9)	0.346 [0.096, 1.243]	0.327 [0.086, 1.242]	-5.4 [-12.5, 1.6]	0.135
		Irbesartan	108	9 (8.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.304
Yes	Double-blind period	Sparsentan	88	5 (5.7)	0.429 [0.156, 1.181]	0.394 [0.131, 1.188]	-7.6 [-17.5, 2.4]	0.116
		Irbesartan	83	11 (13.3)				
No	Double-blind period	Sparsentan	114	4 (3.5)	1.044 [0.267, 4.075]	1.045 [0.255, 4.284]	0.1 [-5.4, 5.7]	1.000
		Irbesartan	119	4 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.113
<= 5 years	Double-blind period	Sparsentan	113	3 (2.7)	0.307 [0.088, 1.071]	0.288 [0.078, 1.058]	-6.0 [-12.6, 0.5]	0.056
		Irbesartan	127	11 (8.7)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	1.264 [0.370, 4.313]	1.283 [0.348, 4.728]	1.4 [-7.1, 9.9]	0.756
		Irbesartan	75	4 (5.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT1AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.647
Yes	Double-blind period	Sparsentan	153	9 (5.9)	0.660 [0.294, 1.479]	0.638 [0.268, 1.522]	-3.0 [-9.5, 3.4]	0.387
		Irbesartan	157	14 (8.9)				
No	Double-blind period	Sparsentan	49	0 (0.0)	0.307 + [0.013, 7.341]	0.300 + [0.012, 7.546]	-2.2 [-8.7, 4.2]	0.479
		Irbesartan	45	1 (2.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEYC\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT1AEYS\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
Safety Set

Not done. Less than 10 patients with events in every subgroup level.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEIC\_SSIM: Incidence of AESI acute kidney injury during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

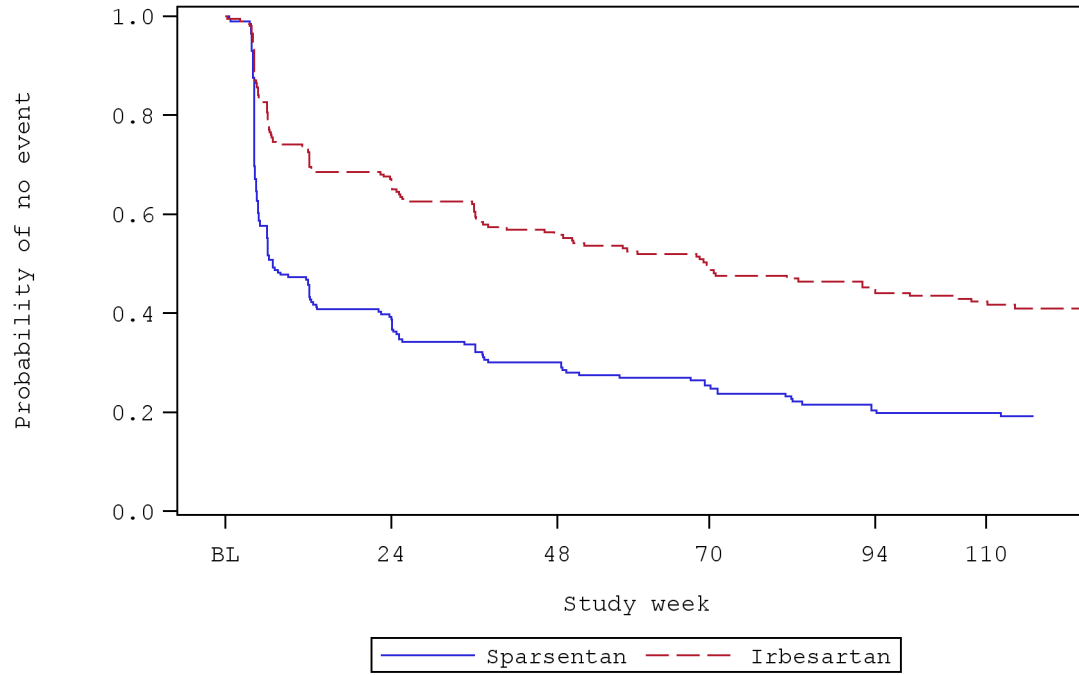
Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT1AEIS\_SSIM: Incidence of AESI acute kidney injury during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

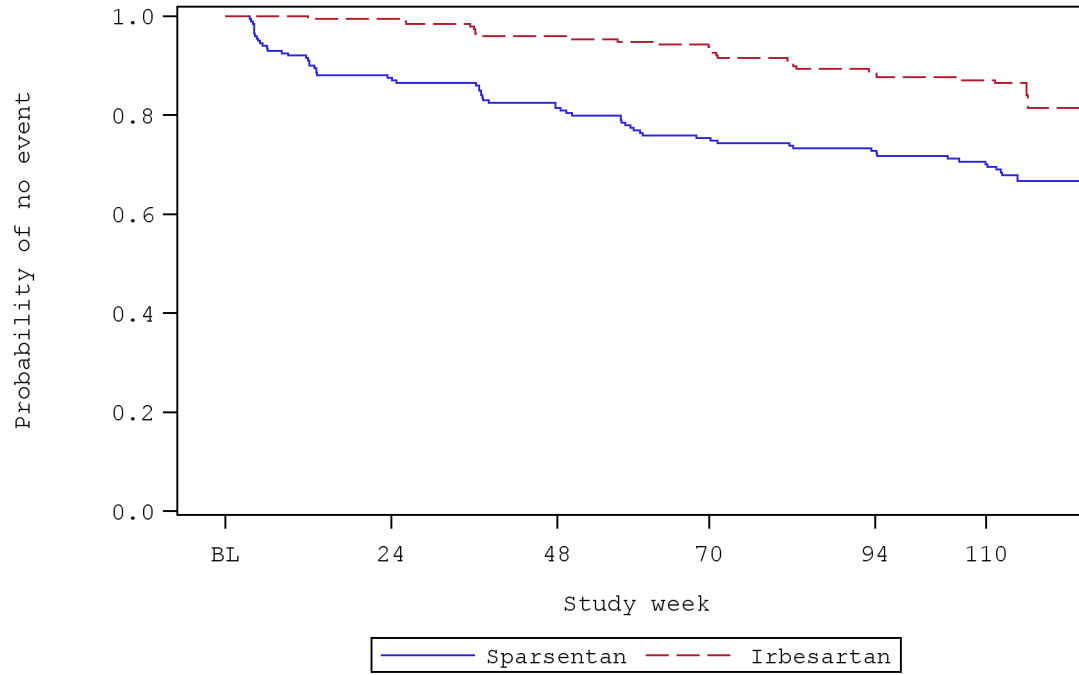
Figure PF2URPT\_FMK0: Time to partial remission - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	78	58	47	35	33
Irbesartan	202	134	104	90	77	71

Partial remission means an urinary protein excretion of <1.0 g/day. No event means no partial remission.  
 Reference table: PT2URPT\_FMT0

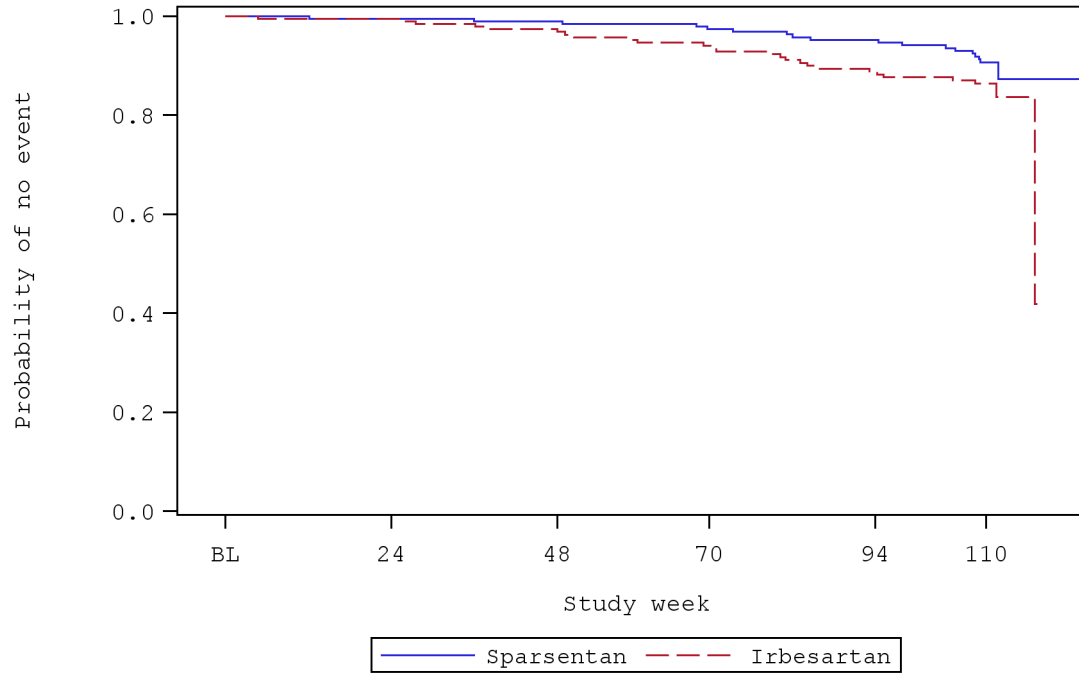
Figure PF2URFT\_FMK0: Time to complete remission - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	175	161	147	137	127
Irbesartan	202	198	181	172	156	148

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no Complete remission.  
 Reference table: PT2URFT\_FMT0

Figure PF2GEDT\_FMK0: Time to confirmed 40% reduction in eGFR, ESRD or death - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT

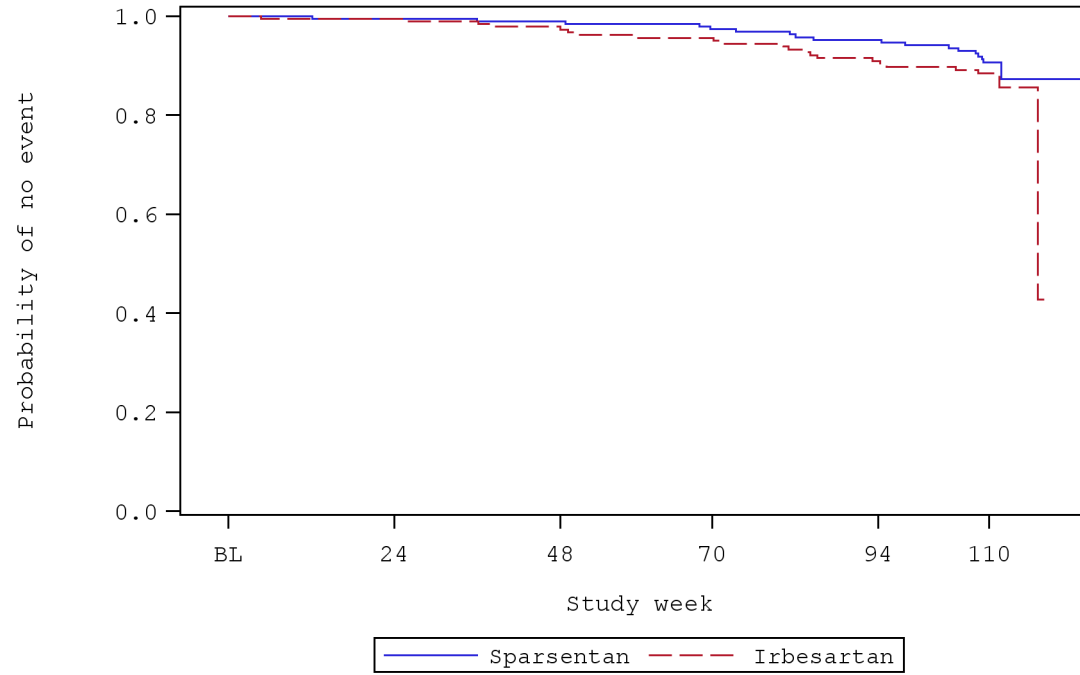


Sparsentan	202	196	190	182	174	123
Irbesartan	202	194	177	165	151	101

ESRD = End-stage renal disease. eGFR = Estimated glomerular filtration rate.  
 Reference table: PT2GEDT\_FMT0



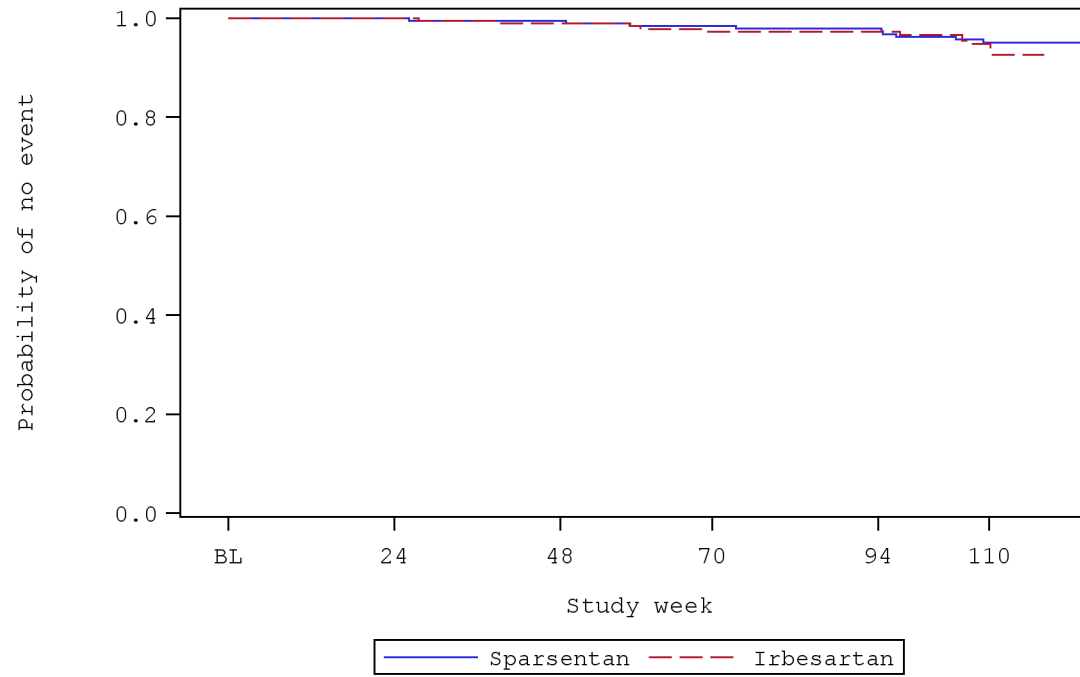
Figure PF2GT\_FMK0: Time to confirmed 40% reduction in eGFR - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	196	190	182	174	123
Irbesartan	202	192	175	165	151	101

eGFR = Estimated glomerular filtration rate.  
 Reference table: PT2GT\_FMT0

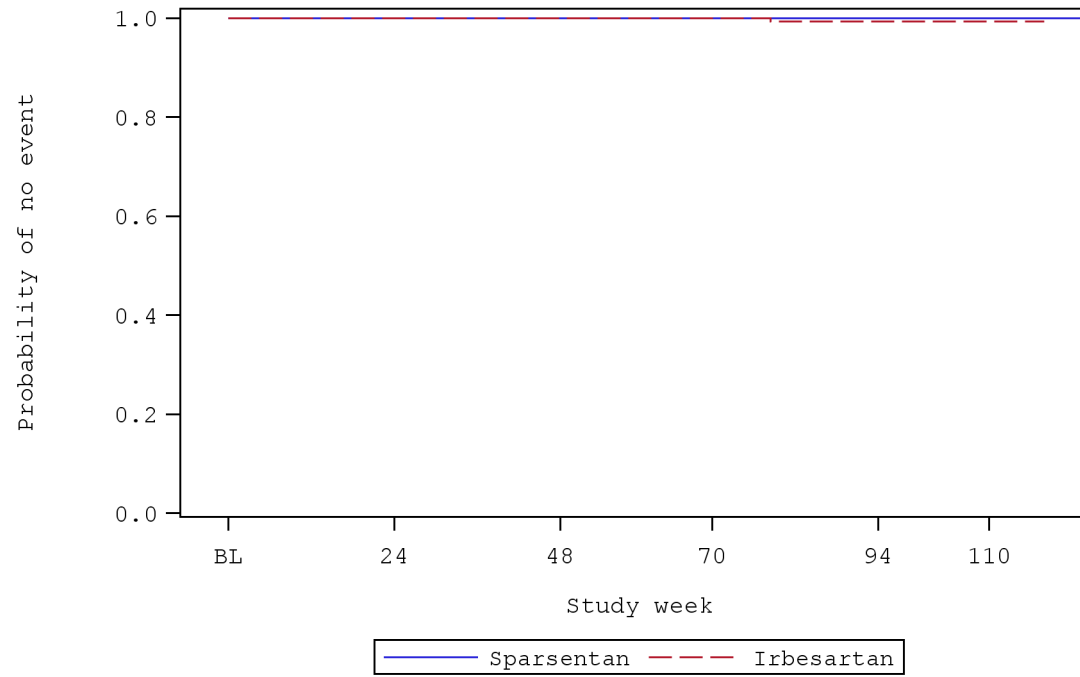
Figure PF2ET\_FMK0: Time to ESRD - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	197	191	184	178	124
Irbesartan	202	195	178	169	163	109

ESRD = End-stage renal disease.  
 Reference table: PT2ET\_FMT0

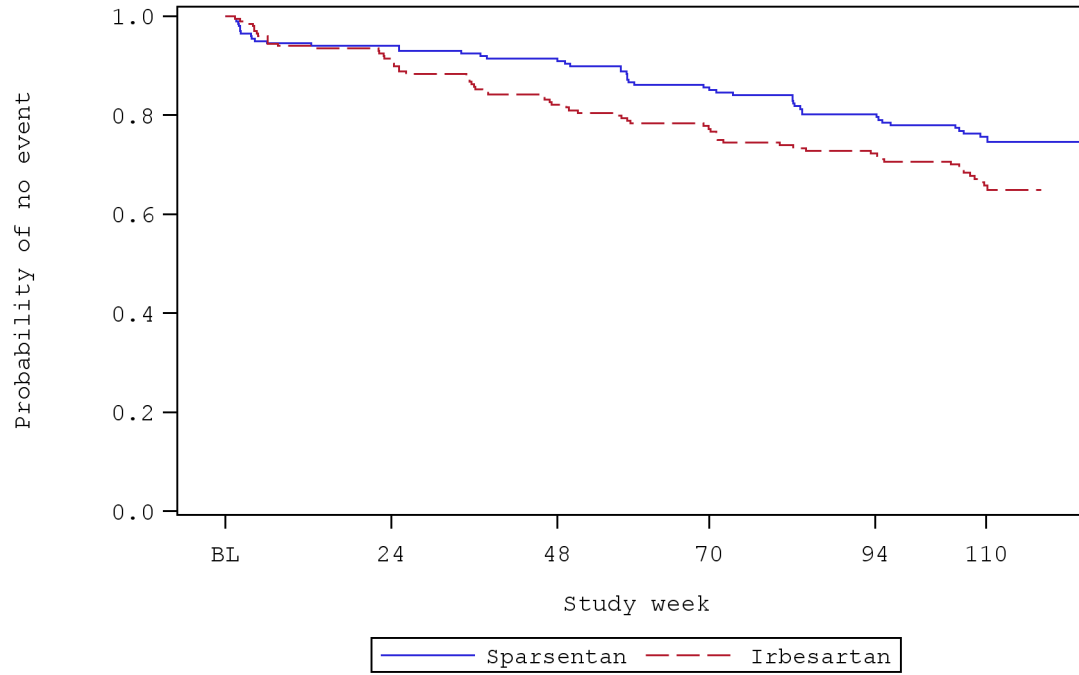
Figure PF2DT\_FMK0: Time to death - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	197	191	186	178	125
Irbesartan	202	193	176	169	163	112

Reference table: PT2DT\_FMT0

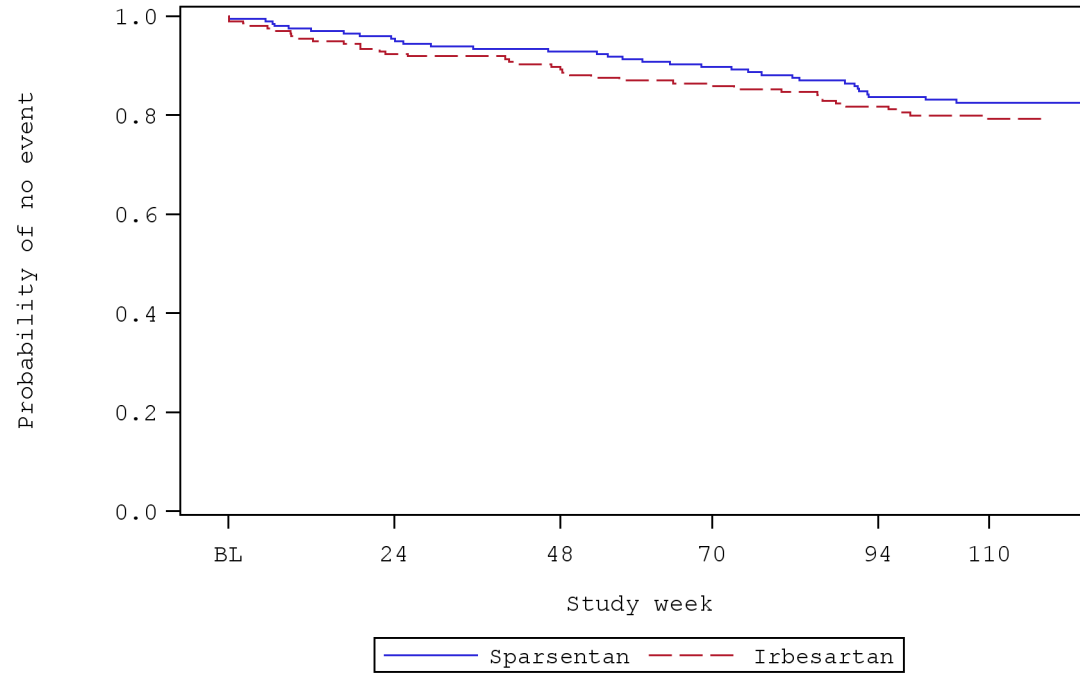
Figure PF2CKDT\_FMK0: Time to CKD stage 4 or 5 - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	185	176	160	146	97
Irbesartan	202	179	155	140	130	84

Reference table: PT2CKDT\_FMT0

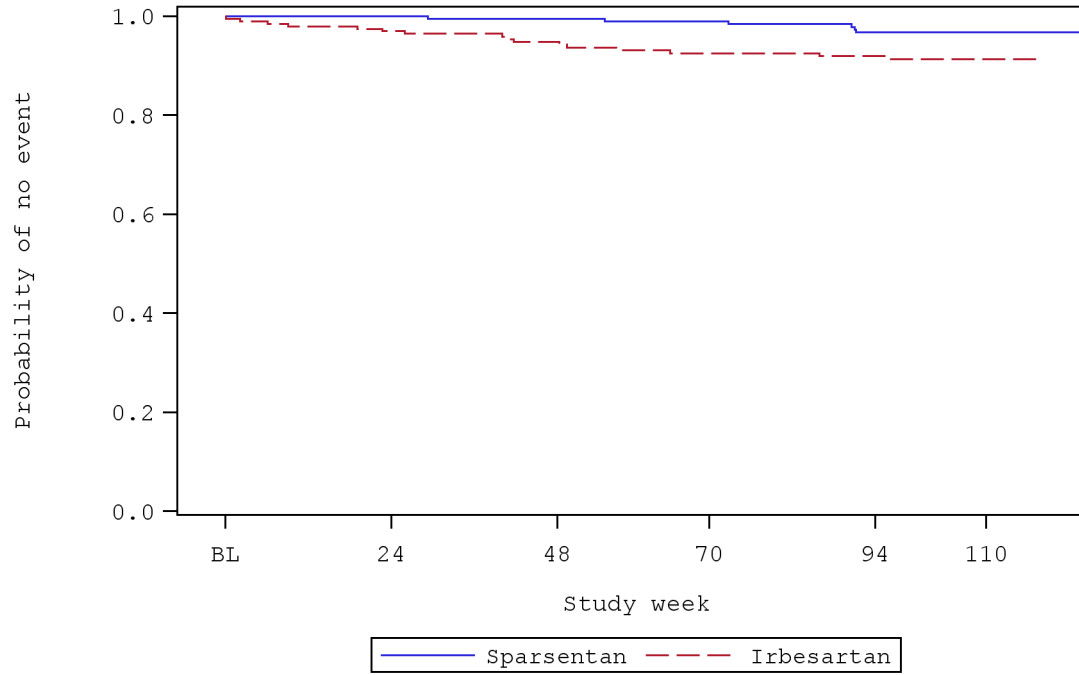
Figure PF2MIST\_FMK0: Time to systemic immunosuppressive medication - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	189	178	168	150	104
Irbesartan	202	181	163	152	138	98

Reference table: PT2MIST\_FMT0

Figure PF2MIKT\_FMK0: Time to systemic immunosuppressive renal medication - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	197	190	185	173	123
Irbesartan	202	188	169	159	152	108

Reference table: PT2MIKT\_FMT0

Table PT2VSC\_FMC0: Change from baseline in EQ-5D VAS  
 Full Analysis Set

Change from baseline in EQ-5D VAS				Repeated measures analysis					
Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Week 24	Sparsentan	202	164 (81.2)	3.45 (0.90)	(1.69, 5.22)	2.84 (1.32)	(0.26, 5.42)	0.031 *	
	Irbesartan	202	143 (70.8)	0.61 (0.96)	(-1.27, 2.50)				
Week 48	Sparsentan	202	168 (83.2)	2.51 (0.89)	(0.77, 4.25)	0.43 (1.32)	(-2.16, 3.03)	0.743	
	Irbesartan	202	132 (65.3)	2.07 (0.98)	(0.14, 4.00)				
Week 70	Sparsentan	202	166 (82.2)	0.55 (0.89)	(-1.20, 2.31)	-0.66 (1.33)	(-3.26, 1.95)	0.621	
	Irbesartan	202	136 (67.3)	1.21 (0.98)	(-0.71, 3.13)				
Week 94	Sparsentan	202	156 (77.2)	1.47 (0.92)	(-0.33, 3.27)	-0.63 (1.35)	(-3.27, 2.01)	0.641	
	Irbesartan	202	136 (67.3)	2.10 (0.99)	(0.17, 4.03)				
Week 110	Sparsentan	202	152 (75.2)	1.74 (0.93)	(-0.10, 3.57)	0.12 (1.40)	(-2.62, 2.86)	0.929	
	Irbesartan	202	121 (59.9)	1.61 (1.04)	(-0.43, 3.65)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

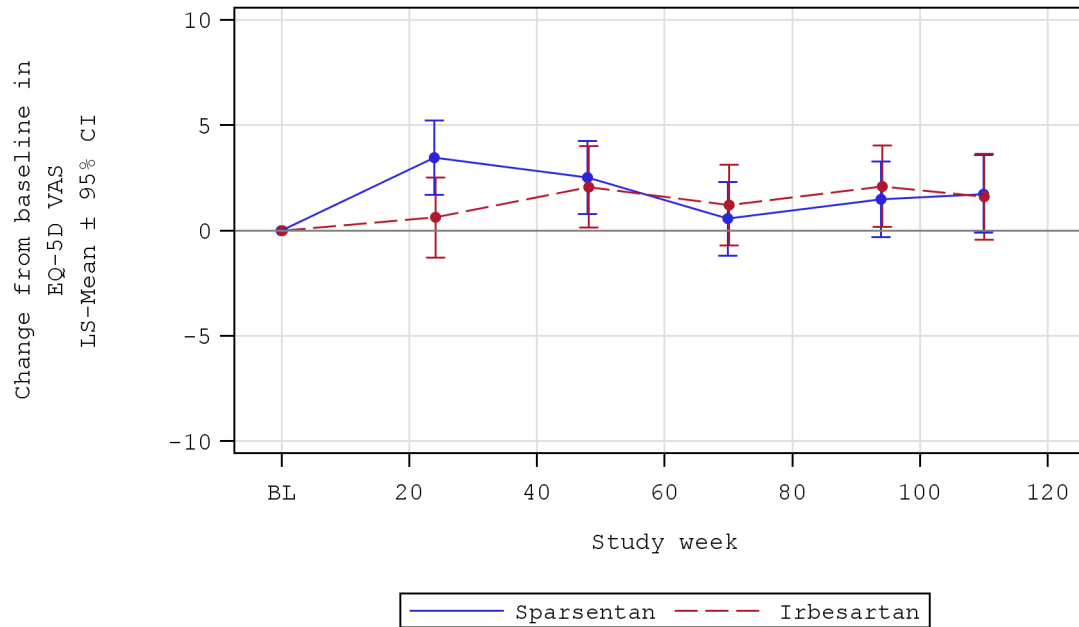
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 08FEB2024

Figure PF2VSC\_FMG0: Change from baseline in EQ-5D VAS  
 Full Analysis Set  
 Study: PROTECT

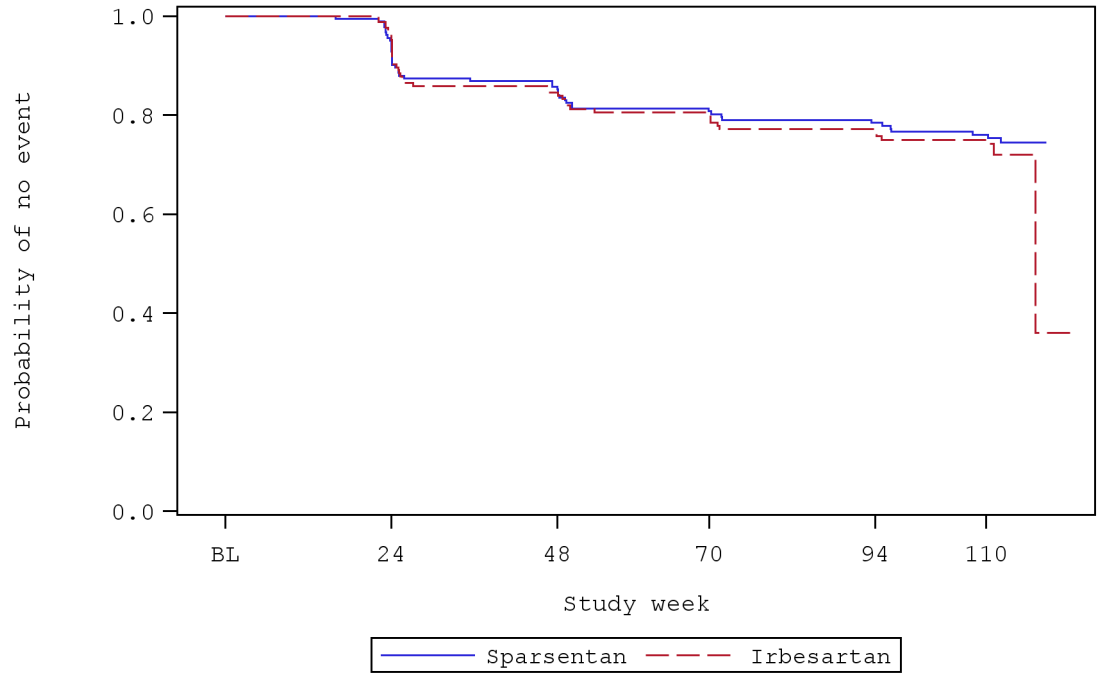


Sparsentan	164	168	166	156	152
Irbesartan	143	132	136	136	121

MMRM = Mixed model Repeated Measures. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. A decrease reflects a worsening of the status of the patient. Reference table: PT2VSC\_FMG0



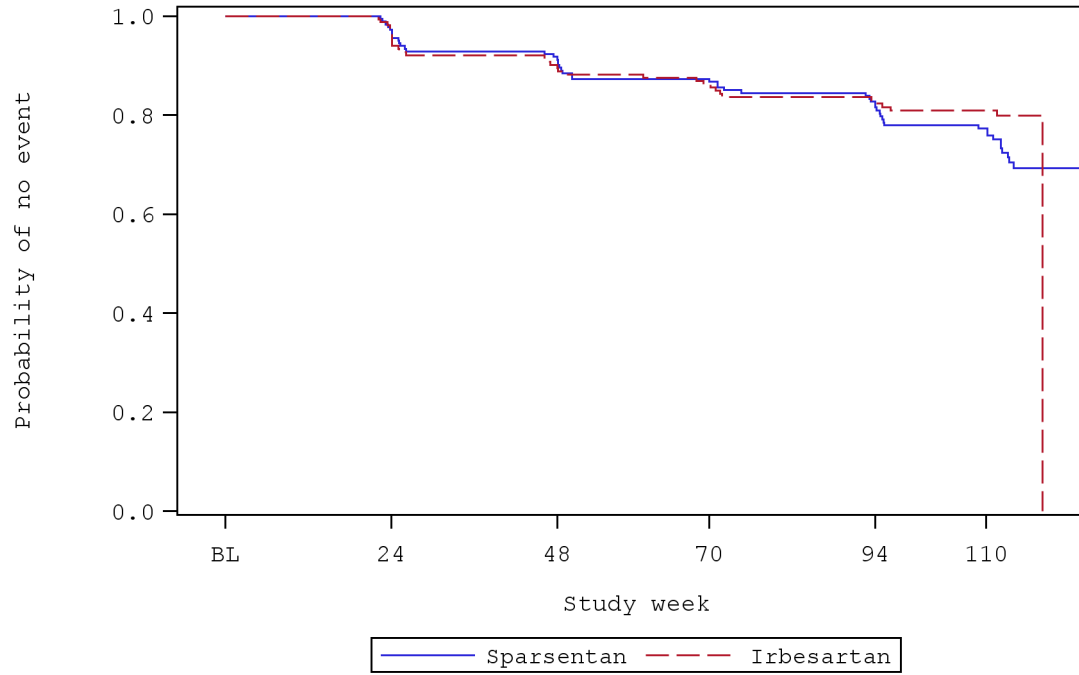
Figure PF2VSIT\_FMK0: Time to increase in EQ-5D VAS by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	174	155	144	131	113
Irbesartan	202	159	131	116	109	90

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2VSIT\_FMT0

Figure PF2VSDT\_FMK0: Time to decrease in EQ-5D VAS by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	179	166	156	138	114
Irbesartan	202	162	141	132	124	104

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2VSDT\_FMT0

Table PT2KBUC\_FMC0: KDQOL: Change from baseline in burden of kidney disease  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease				Repeated measures analysis					
				Change from Baseline		Treatment Difference			
Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Week 24	Sparsentan	202	157 (77.7)	6.02 (1.43)	(3.22, 8.83)	3.16 (2.12)	(-1.00, 7.33)	0.137	
	Irbesartan	202	131 (64.9)	2.86 (1.56)	(-0.21, 5.93)				
Week 48	Sparsentan	202	162 (80.2)	5.90 (1.40)	(3.15, 8.65)	1.60 (2.12)	(-2.55, 5.75)	0.449	
	Irbesartan	202	124 (61.4)	4.30 (1.58)	(1.20, 7.39)				
Week 70	Sparsentan	202	160 (79.2)	6.90 (1.42)	(4.12, 9.67)	3.34 (2.11)	(-0.81, 7.48)	0.114	
	Irbesartan	202	130 (64.4)	3.56 (1.56)	(0.50, 6.62)				
Week 94	Sparsentan	202	151 (74.8)	4.77 (1.45)	(1.93, 7.61)	3.90 (2.14)	(-0.30, 8.09)	0.069	
	Irbesartan	202	130 (64.4)	0.87 (1.57)	(-2.20, 3.95)				
Week 110	Sparsentan	202	149 (73.8)	5.47 (1.47)	(2.59, 8.36)	4.97 (2.20)	(0.66, 9.28)	0.024 *	
	Irbesartan	202	121 (59.9)	0.50 (1.62)	(-2.69, 3.69)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

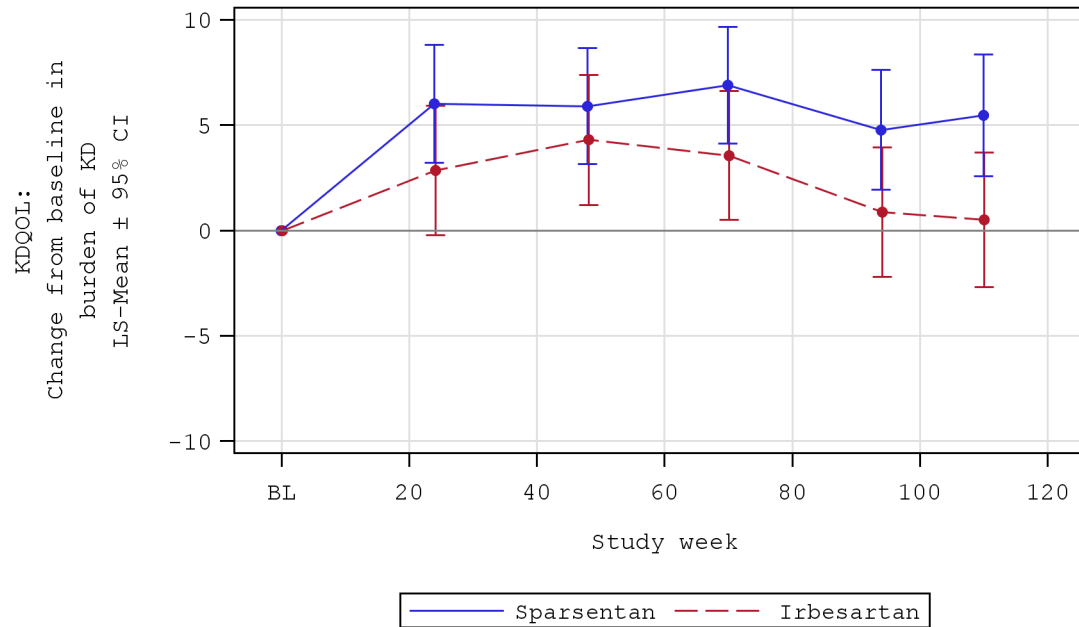
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 08FEB2024

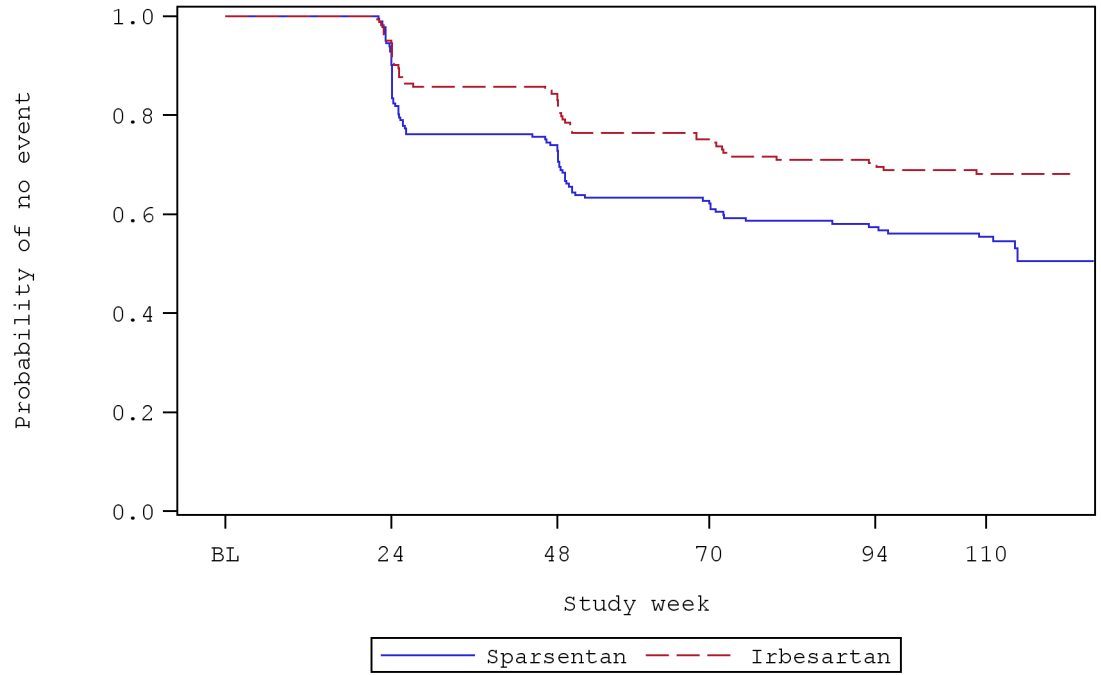
Figure PF2KBUC\_FMG0: KDQOL: Change from baseline in burden of kidney disease  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	157	162	160	151	149
Irbesartan	131	124	130	130	121

MMRM = Mixed model Repeated Measures. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.  
 KD = kidney disease. A decrease reflects a worsening of the status of the patient. Reference table: PT2KBUC\_FMG0

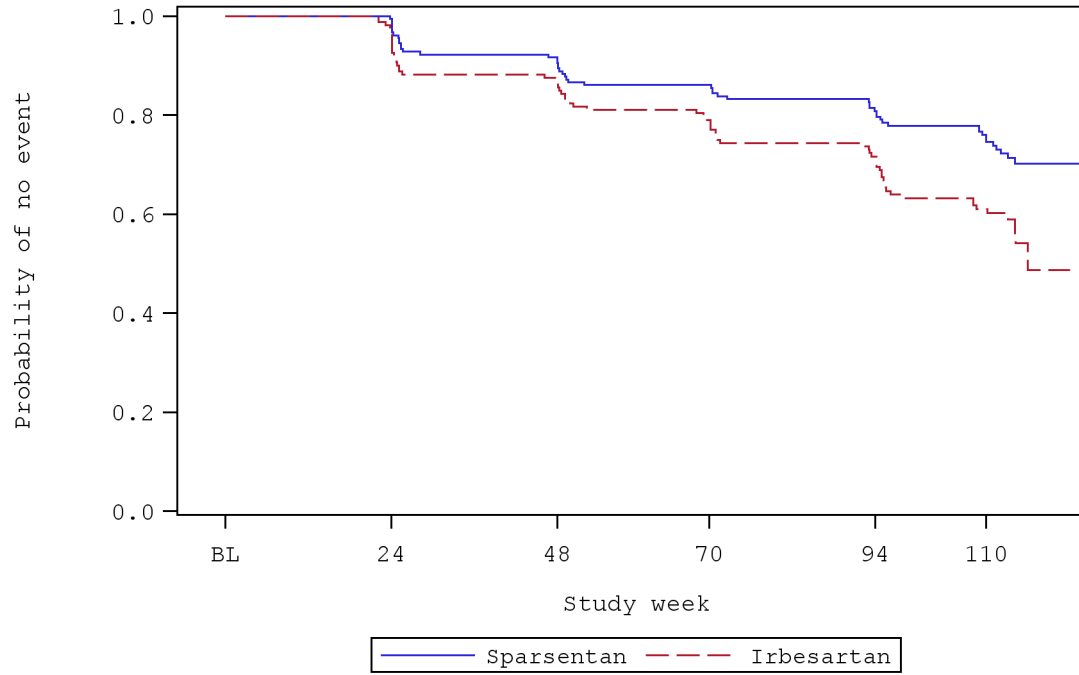
Figure PF2KBUIT\_FMK0: KDQOL: Time to increase in burden of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	168	132	111	92	74
Irbesartan	202	155	129	110	100	79

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KBUIT\_FMT0

Figure PF2KBUDT\_FMK0: KDQOL: Time to decrease in burden of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	181	164	153	136	113
Irbesartan	202	158	136	118	103	78

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KBUDT\_FMT0

Table PT2KEFC\_FMC0: KDQOL: Change from baseline in effect of kidney disease  
 Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease				Repeated measures analysis				
				Change from Baseline		Treatment Difference		
Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Week 24	Sparsentan	202	157 (77.7)	1.55 (0.85)	(-0.11, 3.22)	2.11 (1.26)	(-0.36, 4.57)	0.094
	Irbesartan	202	131 (64.9)	-0.55 (0.93)	(-2.37, 1.27)			
Week 48	Sparsentan	202	162 (80.2)	1.67 (0.83)	(0.03, 3.30)	0.77 (1.25)	(-1.69, 3.23)	0.539
	Irbesartan	202	124 (61.4)	0.90 (0.94)	(-0.94, 2.73)			
Week 70	Sparsentan	202	160 (79.2)	0.86 (0.84)	(-0.78, 2.51)	0.11 (1.25)	(-2.34, 2.56)	0.929
	Irbesartan	202	130 (64.4)	0.75 (0.92)	(-1.06, 2.57)			
Week 94	Sparsentan	202	151 (74.8)	1.58 (0.86)	(-0.11, 3.26)	0.16 (1.26)	(-2.32, 2.64)	0.899
	Irbesartan	202	130 (64.4)	1.42 (0.93)	(-0.40, 3.24)			
Week 110	Sparsentan	202	149 (73.8)	1.50 (0.87)	(-0.21, 3.21)	0.56 (1.30)	(-1.98, 3.11)	0.664
	Irbesartan	202	121 (59.9)	0.93 (0.96)	(-0.95, 2.82)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

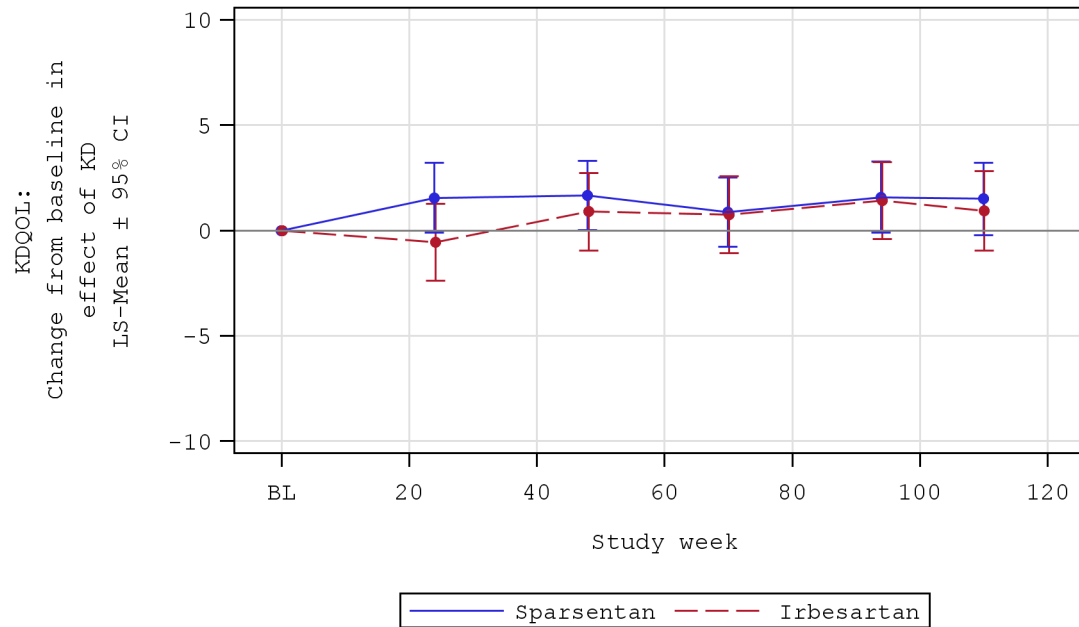
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 08FEB2024

Figure PF2KEFC\_FMG0: KDQOL: Change from baseline in effect of kidney disease  
 Full Analysis Set  
 Study: PROTECT

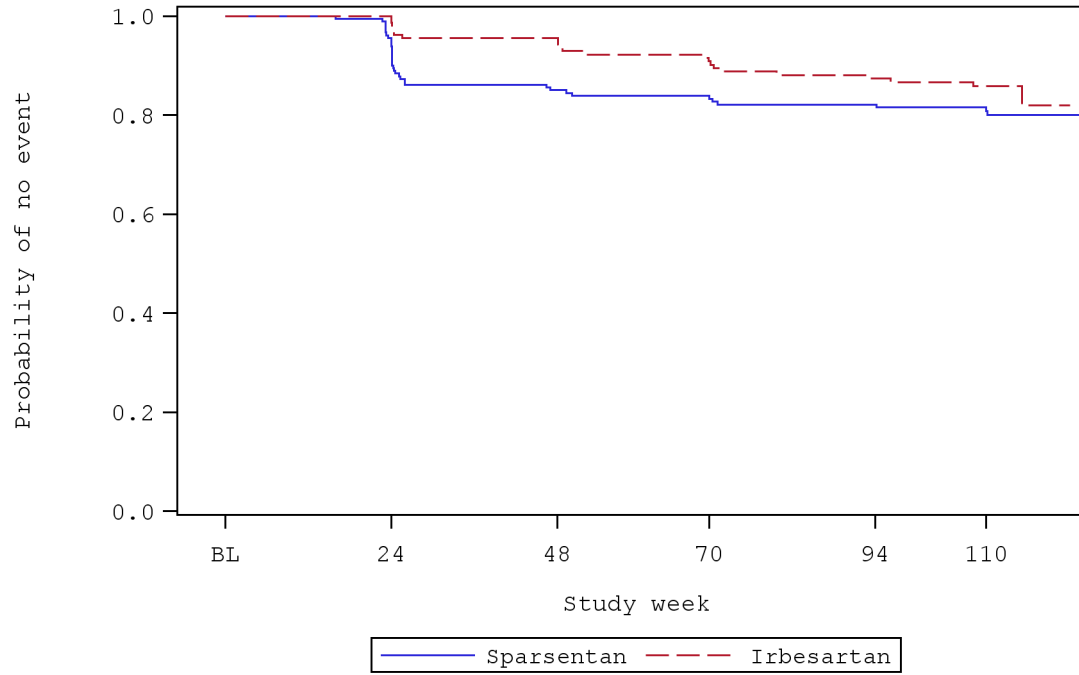


Sparsentan	157	162	160	151	149
Irbesartan	131	124	130	130	121

MMRM = Mixed model Repeated Measures. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.  
 KD = kidney disease. A decrease reflects a worsening of the status of the patient. Reference table: PT2KEFC\_FMG0



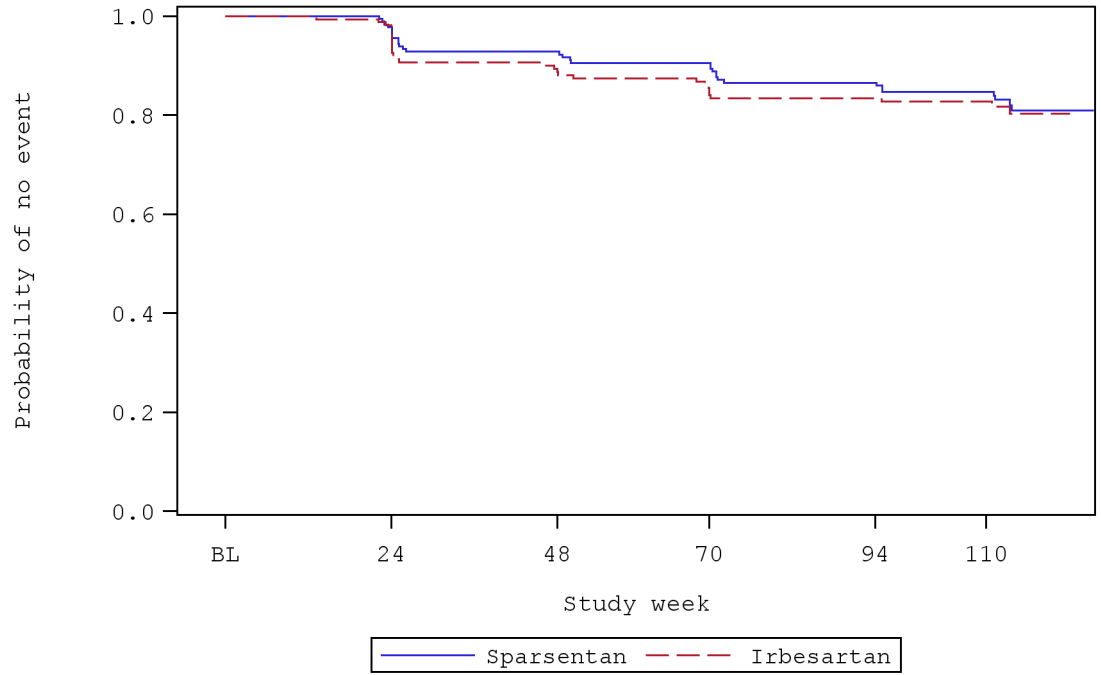
Figure PF2KEFIT\_FMK0: KDQOL: Time to increase in effect of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	173	151	147	133	112
Irbesartan	202	161	145	131	122	103

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KEFIT\_FMT0

Figure PF2KEFDT\_FMK0: KDQOL: Time to decrease in effect of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	177	166	160	144	123
Irbesartan	202	159	138	127	119	102

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KEFDT\_FMT0

Table PT2KSYC\_FMC0: KDQOL: Change from baseline in symptom/problems of kidney disease  
 Full Analysis Set

				Repeated measures analysis					
KDQOL: Change from baseline in symptom/problems of kidney disease				Change from Baseline		Treatment Difference			
Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Week 24	Sparsentan	202	157 (77.7)	-0.13 (0.95)	(-2.00, 1.74)	1.49 (1.41)	(-1.28, 4.27)	0.291	
	Irbesartan	202	131 (64.9)	-1.62 (1.04)	(-3.67, 0.43)				
Week 48	Sparsentan	202	162 (80.2)	-0.70 (0.94)	(-2.54, 1.14)	1.10 (1.42)	(-1.68, 3.87)	0.438	
	Irbesartan	202	124 (61.4)	-1.80 (1.06)	(-3.88, 0.28)				
Week 70	Sparsentan	202	160 (79.2)	-1.37 (0.95)	(-3.22, 0.49)	-0.04 (1.41)	(-2.80, 2.73)	0.979	
	Irbesartan	202	130 (64.4)	-1.33 (1.04)	(-3.38, 0.72)				
Week 94	Sparsentan	202	151 (74.8)	-1.28 (0.97)	(-3.19, 0.62)	0.98 (1.43)	(-1.82, 3.78)	0.494	
	Irbesartan	202	130 (64.4)	-2.26 (1.05)	(-4.32, -0.21)				
Week 110	Sparsentan	202	149 (73.8)	-1.14 (0.98)	(-3.07, 0.79)	0.12 (1.46)	(-2.75, 3.00)	0.932	
	Irbesartan	202	121 (59.9)	-1.27 (1.09)	(-3.40, 0.86)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

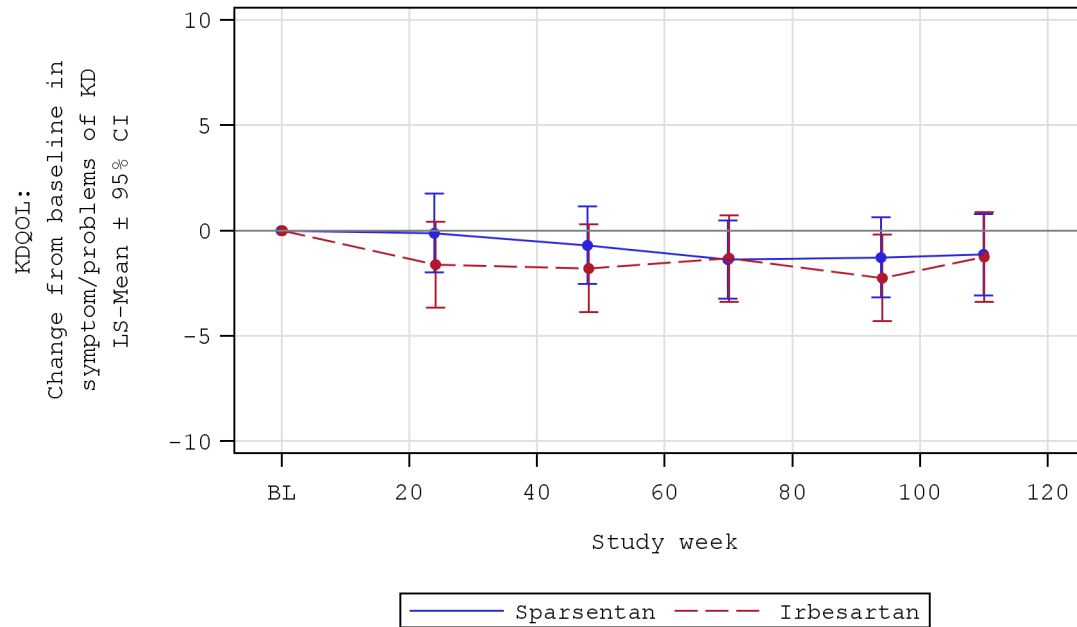
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 08FEB2024

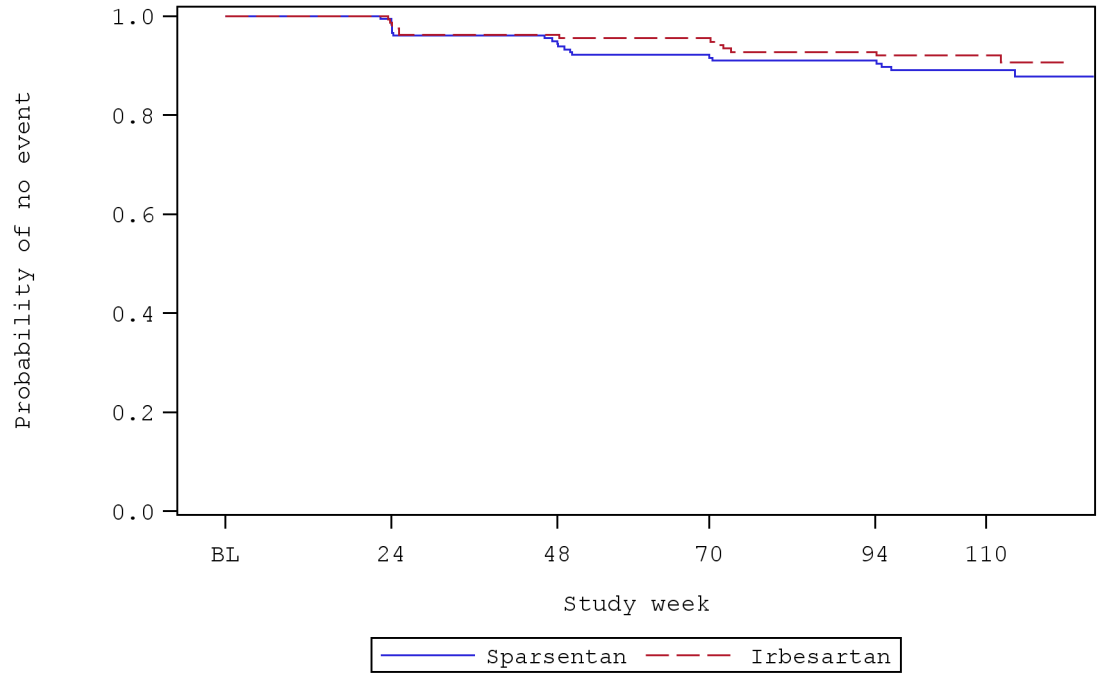
Figure PF2KSYC\_FMG0: KDQOL: Change from baseline in symptom/problems of kidney disease  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	157	162	160	151	149
Irbesartan	131	124	130	130	121

MMRM = Mixed model Repeated Measures. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.  
 KD = kidney disease. A decrease reflects a worsening of the status of the patient. Reference table: PT2KSYC\_FMC0

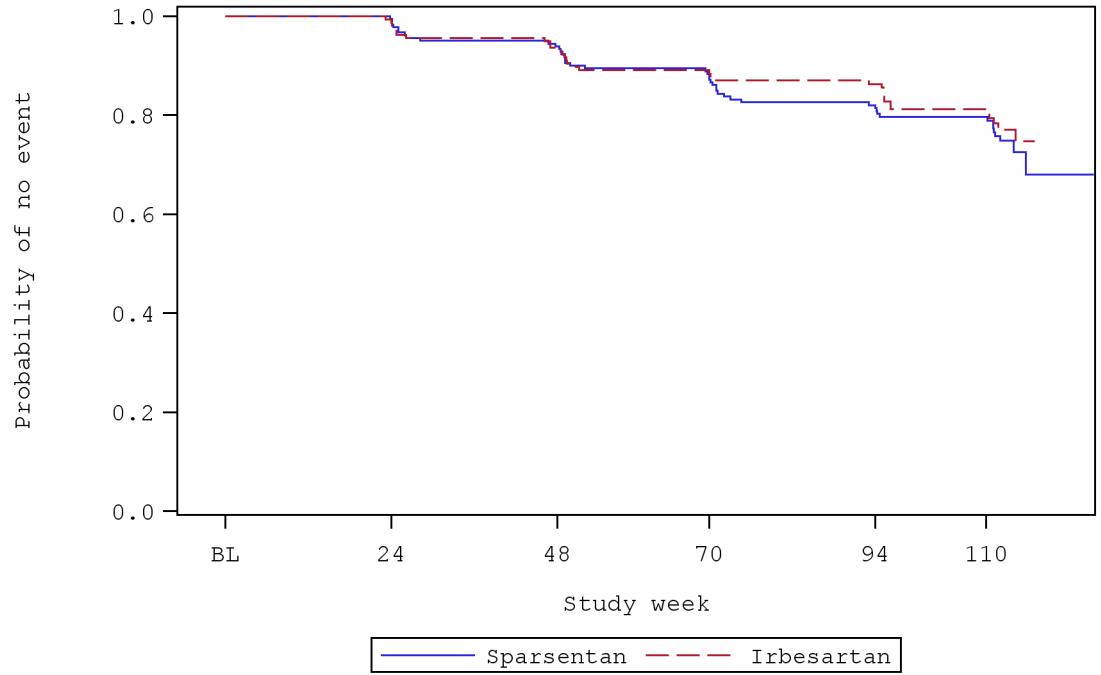
Figure PF2KSYIT\_FMK0: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	180	169	162	148	124
Irbesartan	202	159	146	138	129	110

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KSYIT\_FMT0

Figure PF2KSYDT\_FMK0: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	181	169	157	137	114
Irbesartan	202	161	144	131	122	98

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KSYDT\_FMT0

Table PT2KPSC\_FMC0: KDQOL-SF12: Change from baseline in PCS  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS				Repeated measures analysis				
Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Week 24	Sparsentan	202	159 (78.7)	0.39 (0.49)	(-0.58, 1.36)	0.43 (0.72)	(-0.99, 1.85)	0.555
	Irbesartan	202	138 (68.3)	-0.04 (0.53)	(-1.08, 1.00)			
Week 48	Sparsentan	202	165 (81.7)	0.06 (0.48)	(-0.89, 1.01)	0.12 (0.73)	(-1.31, 1.54)	0.873
	Irbesartan	202	127 (62.9)	-0.06 (0.54)	(-1.12, 1.01)			
Week 70	Sparsentan	202	163 (80.7)	-0.58 (0.49)	(-1.53, 0.38)	0.35 (0.72)	(-1.07, 1.77)	0.627
	Irbesartan	202	135 (66.8)	-0.93 (0.53)	(-1.97, 0.12)			
Week 94	Sparsentan	202	155 (76.7)	-0.45 (0.50)	(-1.42, 0.53)	-0.11 (0.73)	(-1.55, 1.33)	0.885
	Irbesartan	202	133 (65.8)	-0.34 (0.54)	(-1.40, 0.72)			
Week 110	Sparsentan	202	150 (74.3)	-0.60 (0.51)	(-1.60, 0.40)	-0.19 (0.76)	(-1.68, 1.30)	0.804
	Irbesartan	202	121 (59.9)	-0.41 (0.56)	(-1.52, 0.69)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

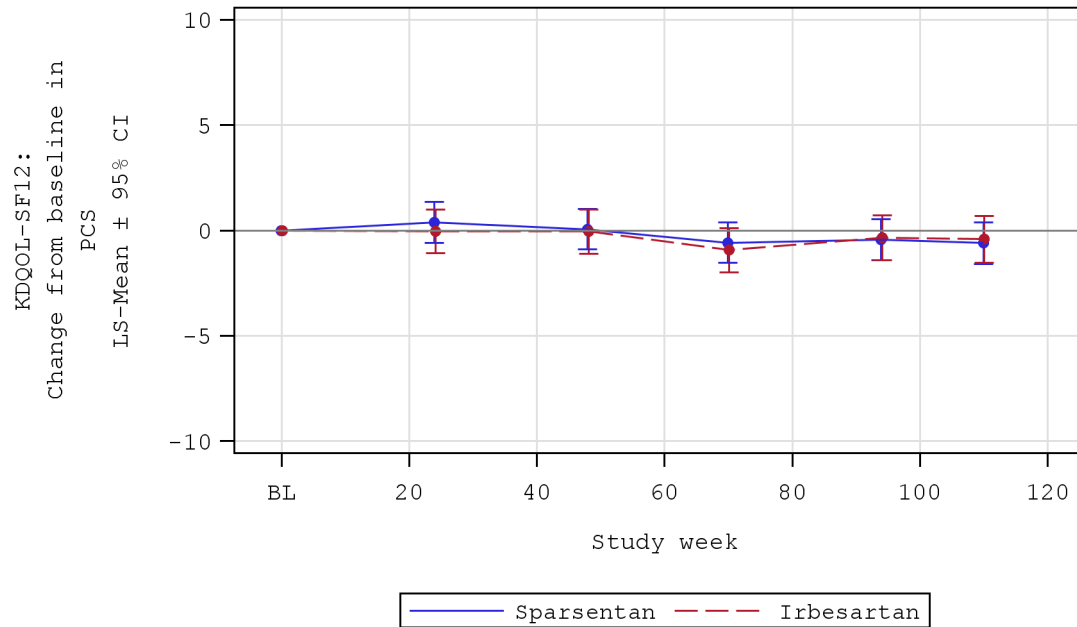
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 08FEB2024

Figure PF2KPSC\_FMG0: KDQOL-SF12: Change from baseline in PCS  
 Full Analysis Set  
 Study: PROTECT

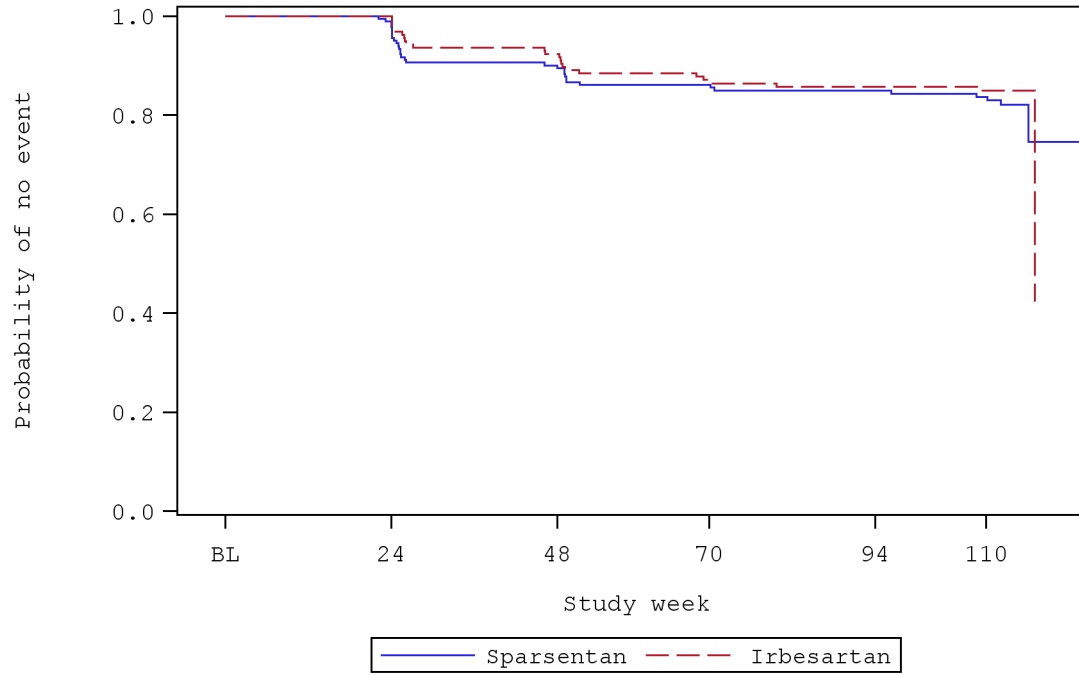


Sparsentan	159	165	163	155	150
Irbesartan	138	127	135	133	121

MMRM = Mixed model Repeated Measures. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. A decrease reflects a worsening of the status of the patient. Reference table: PT2KPSC\_FMG0



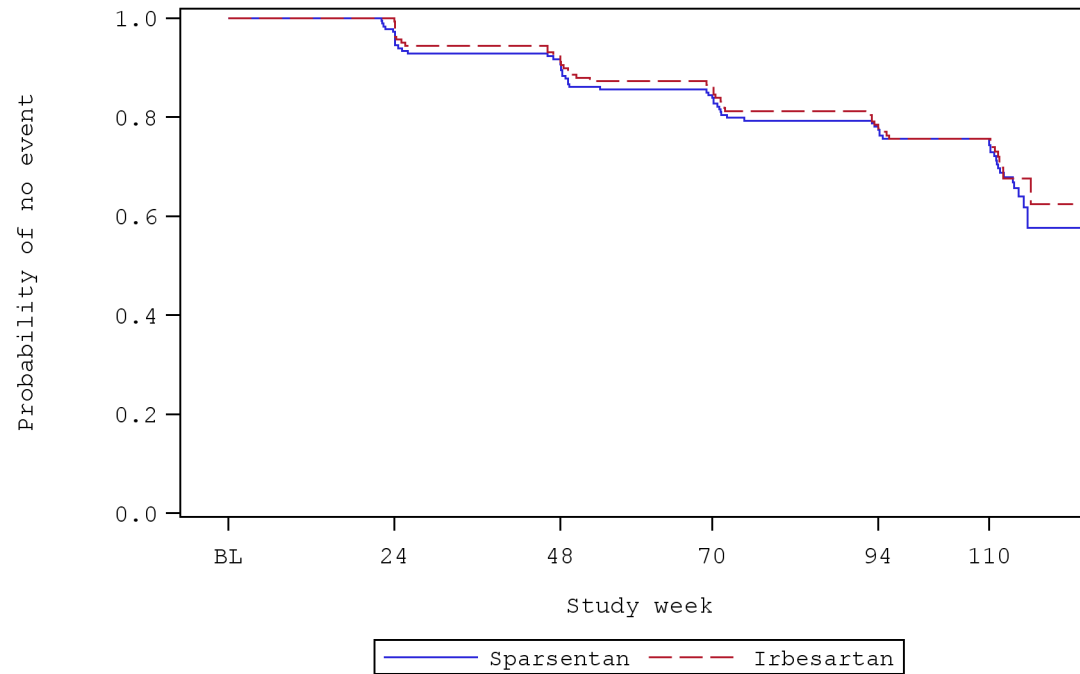
Figure PF2KPSIT\_FMK0: KDQOL-SF12: Time to increase in PCS by at least 8.4 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	180	161	152	140	119
Irbesartan	202	163	143	129	123	105

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FMT0

Figure PF2KPSDT\_FMK0: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	177	164	149	130	109
Irbesartan	202	163	143	128	113	94

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KPSDT\_FMT0

Table PT2KMSC\_FMC0: KDQOL-SF12: Change from baseline in MCS  
 Full Analysis Set

KDQOL-SF12: change from baseline in MCS				Repeated measures analysis					
Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Week 24	Sparsentan	202	159 (78.7)	0.85 (0.57)	(-0.26, 1.97)	-0.12 (0.83)	(-1.75, 1.52)	0.889	
	Irbesartan	202	138 (68.3)	0.97 (0.61)	(-0.23, 2.17)				
Week 48	Sparsentan	202	165 (81.7)	0.11 (0.56)	(-0.98, 1.20)	1.39 (0.84)	(-0.25, 3.04)	0.097	
	Irbesartan	202	127 (62.9)	-1.28 (0.63)	(-2.51, -0.06)				
Week 70	Sparsentan	202	163 (80.7)	-0.05 (0.56)	(-1.15, 1.06)	0.39 (0.83)	(-1.24, 2.03)	0.636	
	Irbesartan	202	135 (66.8)	-0.44 (0.62)	(-1.65, 0.77)				
Week 94	Sparsentan	202	155 (76.7)	-0.33 (0.57)	(-1.46, 0.80)	1.07 (0.85)	(-0.59, 2.73)	0.205	
	Irbesartan	202	133 (65.8)	-1.40 (0.62)	(-2.62, -0.19)				
Week 110	Sparsentan	202	150 (74.3)	-0.85 (0.59)	(-2.00, 0.30)	1.00 (0.87)	(-0.72, 2.72)	0.253	
	Irbesartan	202	121 (59.9)	-1.85 (0.65)	(-3.12, -0.57)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect.

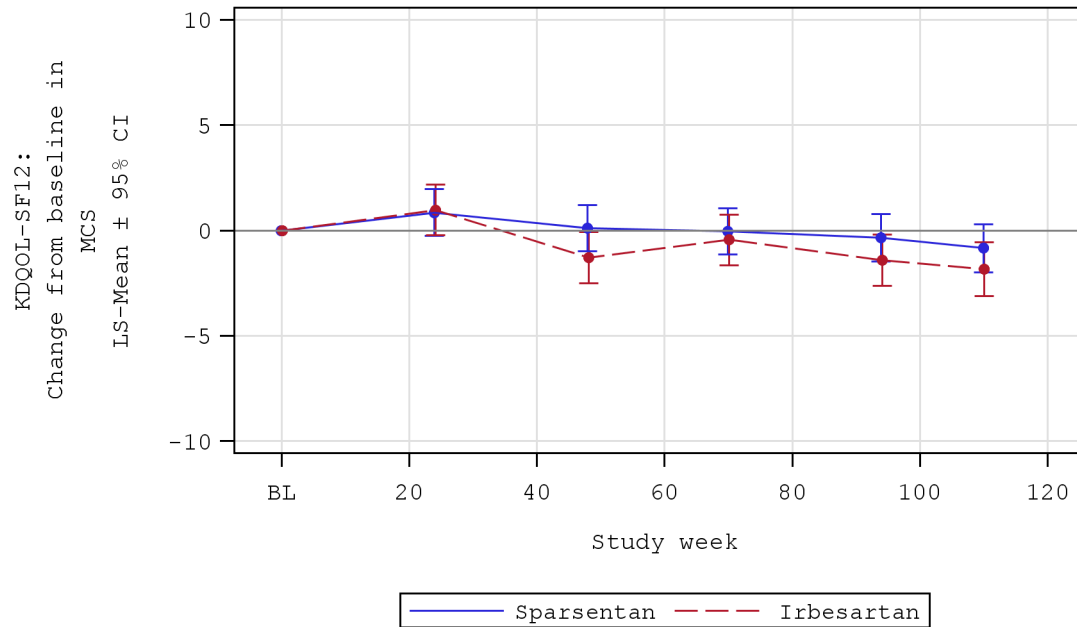
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate.

A decrease reflects a worsening of the status of the patient.

A first order regressive covariance structure was used.

Source Data: aqs, created on: 08FEB2024

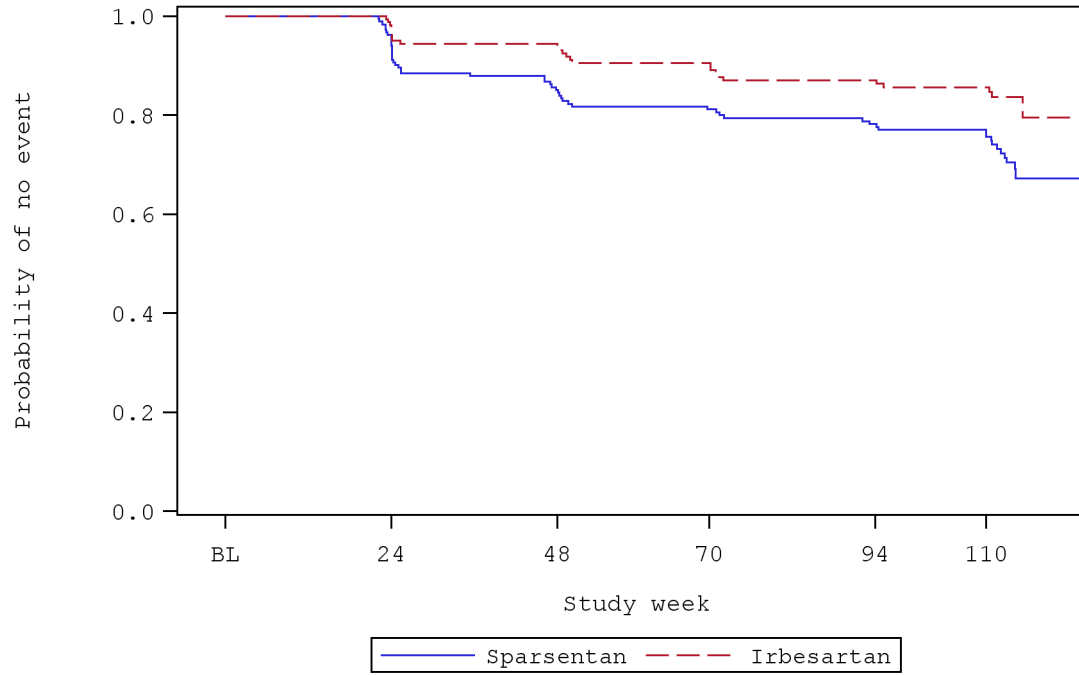
Figure PF2KMSC\_FMG0: KDQOL-SF12: Change from baseline in MCS  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	159	165	163	155	150
Irbesartan	138	127	135	133	121

MMRM = Mixed model Repeated Measures. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. A decrease reflects a worsening of the status of the patient. Reference table: PT2KMSC\_FMG0

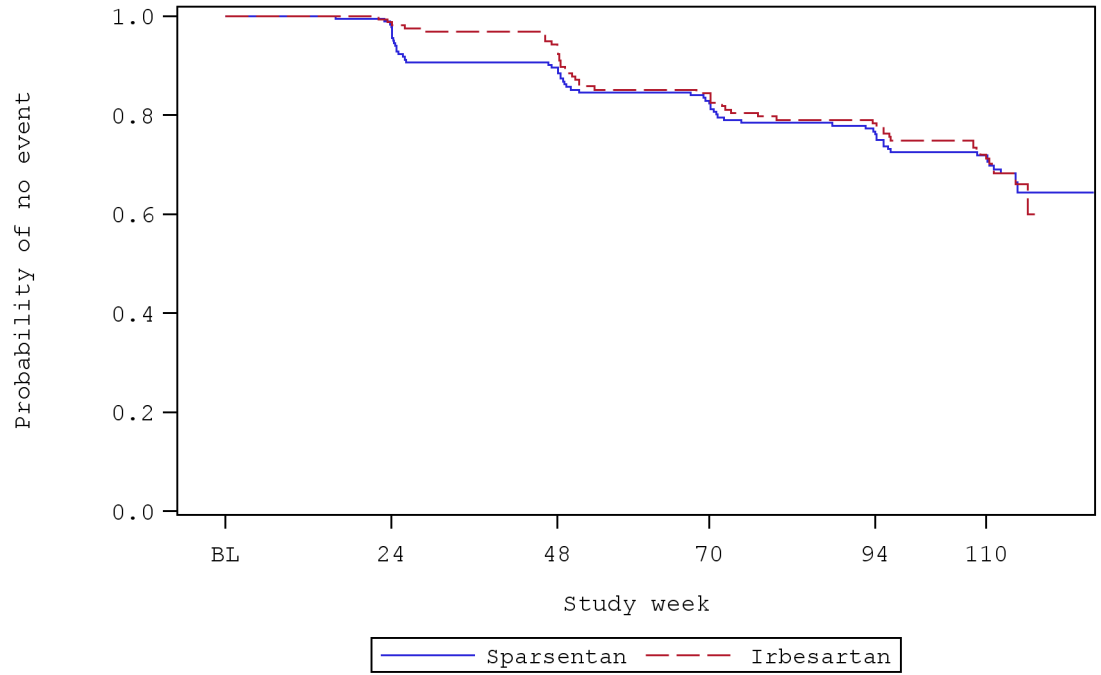
Figure PF2KMSIT\_FMK0: KDQOL-SF12: Time to increase in MCS by at least 9.0 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	175	152	143	129	110
Irbesartan	202	160	146	133	123	103

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KMSIT\_FMT0

Figure PF2KMSDT\_FMK0: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points - Kaplan-Meier plot  
 Full Analysis Set  
 Study: PROTECT



Sparsentan	202	180	162	149	131	109
Irbesartan	202	162	146	126	114	91

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KMSDT\_FMT0

Table PT2AC\_SMS0: Incidence of severe TEAEs during double-blind period by SOC/PT  
Safety Set

Not done. No valid soc/pt combination with sufficient high incidence found.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher was used. \* = significant treatment effect.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Only events are reported with at least 5% of affected patients in at least 1 treatment arm.  
Source Data: aae, created on: 30JAN2024

Table PT2AD\_SMSD: Incidence of fatal TEAEs during double-blind period by SOC/PT Safety Set

Fatal TEAEs				
SOC/PT	Time	Treatment	N	n (%)
SOC: Cardiac disorders	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)
PT: Cardio-respiratory arrest	Double-blind period	Sparsentan	202	0 (0.0)
		Irbesartan	202	1 (0.5)

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Source Data: aae, created on: 30JAN2024



Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Sex	Overall	Sparsentan						Interaction:	0.015	#
Male	Week 6	Sparsentan	139	134 (96.4)	-40.75	(-47.48, -33.15)	0.597	(0.504, 0.707)	<0.001	*
		Irbesartan	143	138 (96.5)	-0.76	(-11.89, 11.77)				
	Week 36	Sparsentan	139	135 (97.1)	-47.93	(-53.84, -41.25)	0.560	(0.472, 0.664)	<0.001	*
		Irbesartan	143	132 (92.3)	-6.98	(-17.51, 4.90)				
Week 58	Sparsentan	139	129 (92.8)	-43.32	(-49.82, -35.96)	0.597	(0.502, 0.709)	<0.001	*	
	Irbesartan	143	127 (88.8)	-5.01	(-15.93, 7.33)					
Week 110	Sparsentan	139	116 (83.5)	-41.90	(-48.82, -34.06)	0.608	(0.507, 0.730)	<0.001	*	
	Irbesartan	143	107 (74.8)	-4.48	(-16.15, 8.81)					
Female	Week 6	Sparsentan	63	59 (93.7)	-34.29	(-44.52, -22.17)	0.808	(0.633, 1.031)	0.087	
		Irbesartan	59	56 (94.9)	-18.64	(-31.73, -3.04)				
	Week 36	Sparsentan	63	59 (93.7)	-47.56	(-55.75, -37.85)	0.710	(0.556, 0.907)	0.006	*
		Irbesartan	59	55 (93.2)	-26.13	(-38.00, -11.99)				
	Week 58	Sparsentan	63	59 (93.7)	-45.59	(-54.10, -35.50)	0.733	(0.573, 0.938)	0.014	*
		Irbesartan	59	52 (88.1)	-25.80	(-37.89, -11.35)				
	Week 110	Sparsentan	63	52 (82.5)	-42.59	(-51.88, -31.52)	0.670	(0.515, 0.872)	0.003	*
		Irbesartan	59	42 (71.2)	-14.28	(-29.46, 4.18)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Age	Overall	Sparsentan						Interaction:	0.876	
<= 45 years	Week 6	Sparsentan	96	92 (95.8)	-43.69	(-51.43, -34.71)	0.616	(0.500, 0.759)	<0.001	*
		Irbesartan	99	96 (97.0)	-8.61	(-21.03, 5.77)				
	Week 36	Sparsentan	96	91 (94.8)	-50.29	(-57.16, -42.33)	0.628	(0.510, 0.774)	<0.001	*
		Irbesartan	99	91 (91.9)	-20.86	(-31.68, -8.33)				
Week 58	Sparsentan	96	89 (92.7)	-42.51	(-50.52, -33.21)	0.693	(0.561, 0.857)	<0.001	*	
	Irbesartan	99	85 (85.9)	-17.07	(-28.63, -3.64)					
Week 110	Sparsentan	96	76 (79.2)	-41.78	(-50.28, -31.82)	0.660	(0.525, 0.830)	<0.001	*	
	Irbesartan	99	69 (69.7)	-11.78	(-25.23, 4.09)					
> 45 years	Week 6	Sparsentan	106	101 (95.3)	-33.86	(-41.98, -24.60)	0.692	(0.575, 0.835)	<0.001	*
		Irbesartan	103	98 (95.1)	-4.48	(-16.38, 9.11)				
	Week 36	Sparsentan	106	103 (97.2)	-45.30	(-52.00, -37.66)	0.577	(0.478, 0.696)	<0.001	*
		Irbesartan	103	96 (93.2)	-5.17	(-17.08, 8.45)				
Week 58	Sparsentan	106	99 (93.4)	-45.21	(-51.97, -37.49)	0.584	(0.484, 0.706)	<0.001	*	
	Irbesartan	103	94 (91.3)	-6.26	(-18.15, 7.36)					
Week 110	Sparsentan	106	92 (86.8)	-42.16	(-49.48, -33.79)	0.599	(0.492, 0.730)	<0.001	*	
	Irbesartan	103	80 (77.7)	-3.47	(-16.32, 11.35)					

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.698	
<= 18 years	Week 6	Sparsentan	9	8 (88.9)	-50.70	(-73.06, -9.78)	0.533	(0.190, 1.499)	0.220	
		Irbesartan	5	5 (100.0)	-7.56	(-58.32, 105.01)				
	Week 36	Sparsentan	9	8 (88.9)	-37.36	(-65.94, 15.21)	0.741	(0.263, 2.086)	0.554	
		Irbesartan	5	5 (100.0)	-15.49	(-61.90, 87.43)				
	Week 58	Sparsentan	9	7 (77.8)	-42.84	(-69.46, 6.99)	0.773	(0.269, 2.219)	0.618	
		Irbesartan	5	4 (80.0)	-26.05	(-67.11, 66.29)				
	Week 110	Sparsentan	9	5 (55.6)	10.68	(-44.69, 121.46)	1.617	(0.526, 4.976)	0.387	
		Irbesartan	5	4 (80.0)	-31.57	(-70.60, 59.25)				
> 18 to 40 years	Week 6	Sparsentan	102	99 (97.1)	-42.81	(-50.17, -34.35)	0.638	(0.526, 0.773)	<0.001	*
		Irbesartan	109	105 (96.3)	-10.33	(-21.59, 2.54)				
	Week 36	Sparsentan	102	97 (95.1)	-50.04	(-56.50, -42.62)	0.612	(0.504, 0.743)	<0.001	*
		Irbesartan	109	100 (91.7)	-18.37	(-28.69, -6.55)				
	Week 58	Sparsentan	102	95 (93.1)	-45.49	(-52.60, -37.31)	0.628	(0.516, 0.765)	<0.001	*
		Irbesartan	109	93 (85.3)	-13.19	(-24.40, -0.32)				
	Week 110	Sparsentan	102	82 (80.4)	-43.36	(-51.11, -34.37)	0.637	(0.516, 0.786)	<0.001	*
		Irbesartan	109	78 (71.6)	-11.05	(-23.44, 3.35)				

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
> 40 years	Week 6	Sparsentan	91	86 (94.5)	-31.60	(-40.72, -21.06)	0.709	(0.578, 0.869)	<0.001	*
		Irbesartan	88	84 (95.5)	-3.45	(-16.52, 11.66)				
	Week 36	Sparsentan	91	89 (97.8)	-45.02	(-52.33, -36.59)	0.597	(0.487, 0.732)	<0.001	*
		Irbesartan	88	82 (93.2)	-7.90	(-20.45, 6.62)				
	Week 58	Sparsentan	91	86 (94.5)	-41.25	(-49.09, -32.21)	0.656	(0.534, 0.805)	<0.001	*
		Irbesartan	88	82 (93.2)	-10.38	(-22.68, 3.87)				
	Week 110	Sparsentan	91	81 (89.0)	-42.07	(-49.93, -32.99)	0.598	(0.483, 0.741)	<0.001	*
		Irbesartan	88	67 (76.1)	-3.15	(-17.24, 13.34)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Geographic region	Overall	Sparsentan						Interaction:	0.901	
North America	Week 6	Sparsentan	35	33 (94.3)	-28.23	(-42.14, -10.98)	0.684	(0.514, 0.910)	0.009	*
		Irbesartan	46	45 (97.8)	4.94	(-12.97, 26.53)				
	Week 36	Sparsentan	35	33 (94.3)	-48.43	(-58.45, -36.00)	0.477	(0.358, 0.637)	<0.001	*
		Irbesartan	46	43 (93.5)	7.99	(-10.65, 30.53)				
Week 58	Sparsentan	35	28 (80.0)	-38.72	(-51.02, -23.33)	0.607	(0.451, 0.817)	0.001	*	
	Irbesartan	46	40 (87.0)	0.98	(-16.84, 22.63)					
Week 110	Sparsentan	35	27 (77.1)	-42.20	(-54.25, -26.98)	0.571	(0.414, 0.786)	<0.001	*	
	Irbesartan	46	31 (67.4)	1.29	(-18.52, 25.92)					
Europe	Week 6	Sparsentan	98	92 (93.9)	-42.24	(-49.83, -33.50)	0.632	(0.521, 0.766)	<0.001	*
		Irbesartan	115	109 (94.8)	-8.56	(-19.78, 4.22)				
	Week 36	Sparsentan	98	94 (95.9)	-47.63	(-54.51, -39.71)	0.621	(0.512, 0.752)	<0.001	*
		Irbesartan	115	108 (93.9)	-15.61	(-25.97, -3.81)				
Week 58	Sparsentan	98	93 (94.9)	-41.54	(-49.26, -32.65)	0.668	(0.550, 0.811)	<0.001	*	
	Irbesartan	115	101 (87.8)	-12.47	(-23.38, -0.00)					
Week 110	Sparsentan	98	79 (80.6)	-42.38	(-50.32, -33.17)	0.637	(0.519, 0.782)	<0.001	*	
	Irbesartan	115	87 (75.7)	-9.51	(-21.44, 4.24)					

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Asia Pacific	Week 6	Sparsentan	69	68 (98.6)	-40.09	(-49.70, -28.64)	0.653	(0.490, 0.869)	0.004	*
		Irbesartan	41	40 (97.6)	-8.19	(-26.77, 15.11)				
	Week 36	Sparsentan	69	67 (97.1)	-48.88	(-57.10, -39.09)	0.664	(0.497, 0.888)	0.006	*
		Irbesartan	41	36 (87.8)	-23.07	(-38.91, -3.11)				
	Week 58	Sparsentan	69	67 (97.1)	-51.02	(-58.91, -41.60)	0.603	(0.450, 0.807)	<0.001	*
		Irbesartan	41	38 (92.7)	-18.73	(-35.54, 2.48)				
	Week 110	Sparsentan	69	62 (89.9)	-43.42	(-52.74, -32.28)	0.603	(0.443, 0.820)	0.001	*
		Irbesartan	41	31 (75.6)	-6.18	(-26.85, 20.33)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Baseline BMI	Overall	Sparsentan						Interaction:	0.968	
< 27 kg/m**2	Week 6	Sparsentan	83	77 (92.8)	-42.10	(-50.22, -32.66)	0.658	(0.535, 0.810)	<0.001	*
		Irbesartan	94	90 (95.7)	-12.06	(-23.70, 1.36)				
	Week 36	Sparsentan	83	80 (96.4)	-51.66	(-58.41, -43.81)	0.629	(0.511, 0.774)	<0.001	*
		Irbesartan	94	85 (90.4)	-23.12	(-33.37, -11.29)				
	Week 58	Sparsentan	83	76 (91.6)	-50.83	(-57.76, -42.77)	0.622	(0.503, 0.768)	<0.001	*
		Irbesartan	94	80 (85.1)	-20.93	(-31.69, -8.47)				
	Week 110	Sparsentan	83	70 (84.3)	-45.20	(-53.18, -35.86)	0.670	(0.535, 0.837)	<0.001	*
		Irbesartan	94	66 (70.2)	-18.16	(-30.17, -4.07)				
≥ 27 kg/m**2	Week 6	Sparsentan	119	116 (97.5)	-35.81	(-43.49, -27.08)	0.655	(0.544, 0.788)	<0.001	*
		Irbesartan	107	103 (96.3)	-1.97	(-14.33, 12.17)				
	Week 36	Sparsentan	119	114 (95.8)	-44.44	(-51.13, -36.84)	0.580	(0.481, 0.699)	<0.001	*
		Irbesartan	107	101 (94.4)	-4.15	(-16.31, 9.78)				
	Week 58	Sparsentan	119	112 (94.1)	-38.27	(-45.75, -29.76)	0.641	(0.531, 0.773)	<0.001	*
		Irbesartan	107	99 (92.5)	-3.64	(-15.97, 10.51)				
	Week 110	Sparsentan	119	98 (82.4)	-39.52	(-47.12, -30.82)	0.593	(0.486, 0.723)	<0.001	*
		Irbesartan	107	82 (76.6)	2.02	(-11.88, 18.12)				

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

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Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Randomization strata	Overall	Sparsentan						Interaction:	0.126	
eGFR Low and UP High	Week 6	Sparsentan	71	68 (95.8)	-31.96	(-42.23, -19.87)	0.670	(0.533, 0.842)	<0.001	*
		Irbesartan	74	73 (98.6)	1.58	(-13.43, 19.20)				
	Week 36	Sparsentan	71	68 (95.8)	-43.21	(-51.80, -33.08)	0.608	(0.483, 0.766)	<0.001	*
		Irbesartan	74	68 (91.9)	-6.62	(-20.55, 9.74)				
	Week 58	Sparsentan	71	67 (94.4)	-42.12	(-50.93, -31.73)	0.640	(0.507, 0.807)	<0.001	*
		Irbesartan	74	65 (87.8)	-9.50	(-23.18, 6.63)				
	Week 110	Sparsentan	71	59 (83.1)	-34.89	(-45.20, -22.66)	0.761	(0.593, 0.976)	0.032	*
		Irbesartan	74	53 (71.6)	-14.44	(-28.51, 2.39)				
eGFR Low and UP Low	Week 6	Sparsentan	55	51 (92.7)	-37.13	(-47.91, -24.11)	0.724	(0.555, 0.944)	0.017	*
		Irbesartan	55	53 (96.4)	-13.15	(-27.98, 4.74)				
	Week 36	Sparsentan	55	51 (92.7)	-42.54	(-52.42, -30.62)	0.624	(0.478, 0.814)	<0.001	*
		Irbesartan	55	51 (92.7)	-7.89	(-23.68, 11.17)				
	Week 58	Sparsentan	55	52 (94.5)	-37.61	(-48.34, -24.65)	0.720	(0.551, 0.940)	0.016	*
		Irbesartan	55	51 (92.7)	-13.37	(-28.27, 4.63)				
	Week 110	Sparsentan	55	46 (83.6)	-31.16	(-43.35, -16.35)	0.751	(0.568, 0.993)	0.045	*
		Irbesartan	55	42 (76.4)	-8.33	(-25.00, 12.04)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024



Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
eGFR High and UP High	Week 6	Sparsentan	37	35 (94.6)	-44.91	(-57.14, -29.18)	0.587	(0.408, 0.842)	0.004	*
		Irbesartan	36	32 (88.9)	-6.08	(-27.57, 21.80)				
	Week 36	Sparsentan	37	37 (100.0)	-53.20	(-63.53, -39.95)	0.580	(0.404, 0.833)	0.003	*
		Irbesartan	36	32 (88.9)	-19.35	(-37.92, 4.77)				
	Week 58	Sparsentan	37	32 (86.5)	-51.25	(-62.24, -37.07)	0.546	(0.377, 0.790)	0.001	*
		Irbesartan	36	31 (86.1)	-10.64	(-31.59, 16.71)				
	Week 110	Sparsentan	37	30 (81.1)	-59.64	(-69.07, -47.34)	0.402	(0.273, 0.594)	<0.001	*
		Irbesartan	36	27 (75.0)	0.28	(-24.43, 33.08)				
eGFR High and UP Low	Week 6	Sparsentan	39	39 (100.0)	-46.47	(-56.60, -33.97)	0.606	(0.448, 0.819)	0.001	*
		Irbesartan	37	36 (97.3)	-11.63	(-28.82, 9.72)				
	Week 36	Sparsentan	39	38 (97.4)	-56.62	(-64.88, -46.43)	0.584	(0.431, 0.790)	<0.001	*
		Irbesartan	37	36 (97.3)	-25.70	(-40.19, -7.71)				
	Week 58	Sparsentan	39	37 (94.9)	-49.18	(-58.93, -37.12)	0.591	(0.434, 0.806)	<0.001	*
		Irbesartan	37	32 (86.5)	-14.03	(-31.30, 7.58)				
	Week 110	Sparsentan	39	33 (84.6)	-49.45	(-59.44, -36.99)	0.512	(0.369, 0.708)	<0.001	*
		Irbesartan	37	27 (73.0)	-1.17	(-22.23, 25.60)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	<0.001	#
< 60 mL/min/1.73 m**2	Week 6	Sparsentan	127	120 (94.5)	-34.74	(-42.25, -26.25)	0.683	(0.575, 0.811)	<0.001	*
		Irbesartan	129	126 (97.7)	-4.38	(-15.28, 7.91)				
	Week 36	Sparsentan	127	120 (94.5)	-42.19	(-48.86, -34.65)	0.637	(0.535, 0.757)	<0.001	*
		Irbesartan	129	119 (92.2)	-9.20	(-19.61, 2.57)				
	Week 58	Sparsentan	127	120 (94.5)	-39.68	(-46.66, -31.78)	0.678	(0.570, 0.808)	<0.001	*
		Irbesartan	129	116 (89.9)	-11.09	(-21.40, 0.56)				
	Week 110	Sparsentan	127	105 (82.7)	-31.37	(-39.62, -22.00)	0.756	(0.629, 0.910)	0.003	*
		Irbesartan	129	96 (74.4)	-9.27	(-20.53, 3.59)				
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	48 (98.0)	-46.38	(-56.23, -34.32)	0.525	(0.393, 0.702)	<0.001	*
		Irbesartan	48	45 (93.8)	2.08	(-16.99, 25.54)				
	Week 36	Sparsentan	49	48 (98.0)	-54.63	(-62.98, -44.41)	0.445	(0.332, 0.596)	<0.001	*
		Irbesartan	48	45 (93.8)	2.05	(-17.21, 25.79)				
	Week 58	Sparsentan	49	43 (87.8)	-50.86	(-60.10, -39.47)	0.480	(0.356, 0.647)	<0.001	*
		Irbesartan	48	42 (87.5)	2.45	(-17.23, 26.81)				
	Week 110	Sparsentan	49	42 (85.7)	-58.08	(-66.13, -48.12)	0.381	(0.279, 0.521)	<0.001	*
		Irbesartan	48	36 (75.0)	9.98	(-12.31, 37.94)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	25 (96.2)	-42.16	(-55.91, -24.12)	0.806	(0.544, 1.193)	0.279	
		Irbesartan	25	23 (92.0)	-28.21	(-45.84, -4.86)				
	Week 36	Sparsentan	26	26 (100.0)	-58.02	(-67.96, -45.01)	0.809	(0.548, 1.195)	0.285	
		Irbesartan	25	23 (92.0)	-48.11	(-60.77, -31.37)				
	Week 58	Sparsentan	26	25 (96.2)	-50.80	(-62.49, -35.45)	0.748	(0.502, 1.113)	0.151	
		Irbesartan	25	21 (84.0)	-34.19	(-50.71, -12.13)				
	Week 110	Sparsentan	26	21 (80.8)	-53.94	(-65.37, -38.75)	0.635	(0.415, 0.974)	0.038	*
		Irbesartan	25	17 (68.0)	-27.53	(-47.20, -0.53)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	<0.001	#
< 45 mL/min/1.73 m**2	Week 6	Sparsentan	82	78 (95.1)	-29.26	(-39.49, -17.31)	0.756	(0.606, 0.944)	0.014	*
		Irbesartan	80	78 (97.5)	-6.49	(-20.14, 9.51)				
	Week 36	Sparsentan	82	78 (95.1)	-37.10	(-46.21, -26.44)	0.700	(0.560, 0.876)	0.002	*
		Irbesartan	80	72 (90.0)	-10.19	(-23.43, 5.34)				
	Week 58	Sparsentan	82	77 (93.9)	-36.45	(-45.70, -25.62)	0.743	(0.593, 0.930)	0.010	*
		Irbesartan	80	70 (87.5)	-14.42	(-27.15, 0.54)				
	Week 110	Sparsentan	82	67 (81.7)	-27.68	(-38.62, -14.80)	0.803	(0.632, 1.019)	0.071	
		Irbesartan	80	59 (73.8)	-9.90	(-24.25, 7.17)				
45 to < 60 mL/min/1.73 m**2	Week 6	Sparsentan	45	42 (93.3)	-43.33	(-53.36, -31.15)	0.572	(0.437, 0.749)	<0.001	*
		Irbesartan	49	48 (98.0)	-0.92	(-17.68, 19.24)				
	Week 36	Sparsentan	45	42 (93.3)	-50.30	(-59.12, -39.59)	0.537	(0.410, 0.704)	<0.001	*
		Irbesartan	49	47 (95.9)	-7.48	(-23.16, 11.41)				
	Week 58	Sparsentan	45	43 (95.6)	-44.99	(-54.75, -33.13)	0.583	(0.444, 0.766)	<0.001	*
		Irbesartan	49	46 (93.9)	-5.67	(-21.85, 13.85)				
	Week 110	Sparsentan	45	38 (84.4)	-37.49	(-48.96, -23.43)	0.683	(0.513, 0.911)	0.010	*
		Irbesartan	49	37 (75.5)	-8.53	(-25.30, 11.99)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	48 (98.0)	-46.38	(-56.23, -34.32)	0.525	(0.393, 0.702)	<0.001	*
		Irbesartan	48	45 (93.8)	2.08	(-16.99, 25.54)				
	Week 36	Sparsentan	49	48 (98.0)	-54.63	(-62.98, -44.41)	0.445	(0.332, 0.596)	<0.001	*
		Irbesartan	48	45 (93.8)	2.05	(-17.21, 25.79)				
	Week 58	Sparsentan	49	43 (87.8)	-50.86	(-60.10, -39.47)	0.480	(0.356, 0.647)	<0.001	*
		Irbesartan	48	42 (87.5)	2.45	(-17.23, 26.81)				
	Week 110	Sparsentan	49	42 (85.7)	-58.08	(-66.13, -48.12)	0.381	(0.279, 0.521)	<0.001	*
		Irbesartan	48	36 (75.0)	9.98	(-12.31, 37.94)				
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	25 (96.2)	-42.16	(-55.91, -24.12)	0.806	(0.544, 1.193)	0.279	
		Irbesartan	25	23 (92.0)	-28.21	(-45.84, -4.86)				
	Week 36	Sparsentan	26	26 (100.0)	-58.02	(-67.96, -45.01)	0.809	(0.548, 1.195)	0.285	
		Irbesartan	25	23 (92.0)	-48.11	(-60.77, -31.37)				
	Week 58	Sparsentan	26	25 (96.2)	-50.80	(-62.49, -35.45)	0.748	(0.502, 1.113)	0.151	
		Irbesartan	25	21 (84.0)	-34.19	(-50.71, -12.13)				
	Week 110	Sparsentan	26	21 (80.8)	-53.94	(-65.37, -38.75)	0.635	(0.415, 0.974)	0.038	*
		Irbesartan	25	17 (68.0)	-27.53	(-47.20, -0.53)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

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Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.372	
<= 1.75 g/day	Week 6	Sparsentan	98	94 (95.9)	-33.40	(-42.39, -23.02)	0.699	(0.567, 0.860)	<0.001	*
		Irbesartan	93	89 (95.7)	-4.68	(-17.86, 10.62)				
	Week 36	Sparsentan	98	94 (95.9)	-44.02	(-51.58, -35.28)	0.600	(0.487, 0.740)	<0.001	*
		Irbesartan	93	87 (93.5)	-6.77	(-19.74, 8.29)				
	Week 58	Sparsentan	98	93 (94.9)	-36.30	(-44.93, -26.32)	0.669	(0.542, 0.826)	<0.001	*
		Irbesartan	93	82 (88.2)	-4.82	(-18.26, 10.85)				
	Week 110	Sparsentan	98	84 (85.7)	-33.39	(-42.65, -22.64)	0.657	(0.526, 0.819)	<0.001	*
		Irbesartan	93	70 (75.3)	1.41	(-13.81, 19.33)				
> 1.75 g/day	Week 6	Sparsentan	104	99 (95.2)	-43.48	(-50.56, -35.39)	0.619	(0.513, 0.746)	<0.001	*
		Irbesartan	109	105 (96.3)	-8.66	(-19.86, 4.11)				
	Week 36	Sparsentan	104	100 (96.2)	-51.09	(-57.22, -44.08)	0.603	(0.500, 0.728)	<0.001	*
		Irbesartan	109	100 (91.7)	-18.91	(-28.93, -7.48)				
	Week 58	Sparsentan	104	95 (91.3)	-50.50	(-56.78, -43.31)	0.603	(0.498, 0.730)	<0.001	*
		Irbesartan	109	97 (89.0)	-17.93	(-28.21, -6.18)				
	Week 110	Sparsentan	104	84 (80.8)	-49.39	(-56.09, -41.66)	0.599	(0.489, 0.734)	<0.001	*
		Irbesartan	109	79 (72.5)	-15.51	(-26.91, -2.34)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

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Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.939	
Yes	Week 6	Sparsentan	90	84 (93.3)	-39.34	(-47.70, -29.64)	0.651	(0.527, 0.805)	<0.001	*
		Irbesartan	88	82 (93.2)	-6.84	(-19.91, 8.35)				
	Week 36	Sparsentan	90	87 (96.7)	-47.26	(-54.51, -38.85)	0.583	(0.472, 0.720)	<0.001	*
		Irbesartan	88	82 (93.2)	-9.51	(-22.21, 5.27)				
	Week 58	Sparsentan	90	83 (92.2)	-42.31	(-50.32, -33.02)	0.669	(0.540, 0.829)	<0.001	*
		Irbesartan	88	80 (90.9)	-13.82	(-26.06, 0.45)				
	Week 110	Sparsentan	90	74 (82.2)	-45.14	(-53.04, -35.93)	0.601	(0.480, 0.752)	<0.001	*
		Irbesartan	88	68 (77.3)	-8.70	(-22.39, 7.41)				
No	Week 6	Sparsentan	112	109 (97.3)	-38.12	(-45.76, -29.41)	0.661	(0.549, 0.795)	<0.001	*
		Irbesartan	114	112 (98.2)	-6.33	(-17.76, 6.68)				
	Week 36	Sparsentan	112	107 (95.5)	-47.99	(-54.43, -40.64)	0.620	(0.514, 0.747)	<0.001	*
		Irbesartan	114	105 (92.1)	-16.10	(-26.45, -4.30)				
	Week 58	Sparsentan	112	105 (93.8)	-45.20	(-52.03, -37.40)	0.611	(0.505, 0.738)	<0.001	*
		Irbesartan	114	99 (86.8)	-10.26	(-21.51, 2.60)				
	Week 110	Sparsentan	112	94 (83.9)	-39.08	(-46.94, -30.06)	0.655	(0.536, 0.800)	<0.001	*
		Irbesartan	114	81 (71.1)	-6.98	(-19.58, 7.59)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
Time since renal biopsy Overall		Sparsentan						Interaction:	0.555	
<= 5 years	Week 6	Sparsentan	113	106 (93.8)	-40.00	(-47.67, -31.21)	0.663	(0.549, 0.800)	<0.001	*
		Irbesartan	127	124 (97.6)	-9.48	(-20.38, 2.91)				
	Week 36	Sparsentan	113	107 (94.7)	-51.62	(-57.80, -44.53)	0.595	(0.493, 0.718)	<0.001	*
		Irbesartan	127	117 (92.1)	-18.68	(-28.57, -7.42)				
Week 58	Sparsentan	113	104 (92.0)	-41.87	(-49.35, -33.30)	0.741	(0.613, 0.896)	0.002	*	
	Irbesartan	127	117 (92.1)	-21.57	(-31.17, -10.62)					
Week 110	Sparsentan	113	96 (85.0)	-48.23	(-55.09, -40.33)	0.605	(0.495, 0.740)	<0.001	*	
	Irbesartan	127	93 (73.2)	-14.50	(-25.81, -1.47)					
> 5 years	Week 6	Sparsentan	89	87 (97.8)	-36.98	(-45.15, -27.59)	0.639	(0.519, 0.786)	<0.001	*
		Irbesartan	75	70 (93.3)	-1.35	(-15.35, 14.96)				
	Week 36	Sparsentan	89	87 (97.8)	-42.29	(-49.79, -33.66)	0.595	(0.484, 0.732)	<0.001	*
		Irbesartan	75	70 (93.3)	-3.00	(-16.78, 13.06)				
	Week 58	Sparsentan	89	84 (94.4)	-46.36	(-53.40, -38.26)	0.494	(0.399, 0.610)	<0.001	*
		Irbesartan	75	62 (82.7)	8.64	(-7.24, 27.24)				
	Week 110	Sparsentan	89	72 (80.9)	-32.70	(-41.91, -22.03)	0.642	(0.514, 0.803)	<0.001	*
		Irbesartan	75	56 (74.7)	4.76	(-11.35, 23.78)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024



Table PT2RPR\_FSLM: Percent change from baseline in UP/C by subgroup  
Full Analysis Set

Percent change from baseline in UP/C					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Percentage change from Baseline		Ratio (Sparsentan/Irbesartan)			
					Geometric LS-Mean	95% CI	LS-Mean Ratio	95% CI	p-value	
History of hypertension Overall		Sparsentan						Interaction:	0.539	
Yes	Week 6	Sparsentan	155	147 (94.8)	-37.69	(-44.25, -30.37)	0.673	(0.576, 0.786)	<0.001	*
		Irbesartan	161	153 (95.0)	-7.41	(-17.00, 3.28)				
	Week 36	Sparsentan	155	149 (96.1)	-44.44	(-50.28, -37.92)	0.622	(0.532, 0.727)	<0.001	*
		Irbesartan	161	150 (93.2)	-10.71	(-19.99, -0.34)				
Week 58	Sparsentan	155	145 (93.5)	-39.97	(-46.32, -32.88)	0.680	(0.580, 0.796)	<0.001	*	
	Irbesartan	161	145 (90.1)	-11.68	(-20.99, -1.27)					
Week 110	Sparsentan	155	130 (83.9)	-40.05	(-46.59, -32.71)	0.642	(0.544, 0.758)	<0.001	*	
	Irbesartan	161	121 (75.2)	-6.65	(-17.13, 5.17)					
No	Week 6	Sparsentan	47	46 (97.9)	-41.41	(-52.54, -27.67)	0.618	(0.454, 0.843)	0.002	*
		Irbesartan	41	41 (100.0)	-5.25	(-24.38, 18.72)				
	Week 36	Sparsentan	47	45 (95.7)	-56.72	(-65.01, -46.46)	0.569	(0.416, 0.779)	<0.001	*
		Irbesartan	41	37 (90.2)	-23.93	(-39.52, -4.32)				
	Week 58	Sparsentan	47	43 (91.5)	-54.92	(-63.68, -44.04)	0.521	(0.378, 0.717)	<0.001	*
		Irbesartan	41	34 (82.9)	-13.49	(-31.55, 9.35)				
	Week 110	Sparsentan	47	38 (80.9)	-47.17	(-57.91, -33.69)	0.610	(0.432, 0.860)	0.005	*
		Irbesartan	41	28 (68.3)	-13.35	(-33.00, 12.06)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. Analysis was performed using the natural logarithm of the dependent variable. Reported estimates are back-transformed to the ratio scale. LS-Mean = Least square mean. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. Only key visits up to Week 110 are displayed.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and log(baseline) as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed values up to Week 110 are included in the model. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Estimated LS Mean and 95% CIs were converted to percentage change as follows:  $[\exp(\text{least squares mean change from baseline in natural log(UP/C)}) - 1] \times 100$ . The Ratio (Sparsentan/Irbesartan) can be interpreted in an analogous way as other common summarizing ratios (e.g. RR).

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio. A first order autoregressive covariance structure was used.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
Subgroup: Sex													
Male	UP/C	Baseline	Sparsentan	139	139 (100.0)	1.30 (0.79)	0.1	0.72	1.09	1.77	4.2		
			Irbesartan	143	143 (100.0)	1.33 (0.85)	0.2	0.85	1.13	1.57	6.9		
		Week 6	Sparsentan	139	134 (96.4)	0.91 (0.82)	0.1	0.38	0.68	1.11	5.6		
			Irbesartan	143	138 (96.5)	1.37 (0.87)	0.1	0.76	1.15	1.70	5.4		
		Week 36	Sparsentan	139	135 (97.1)	0.92 (0.94)	0.1	0.29	0.56	1.32	5.2		
			Irbesartan	143	132 (92.3)	1.36 (0.98)	0.1	0.65	1.17	1.80	6.2		
		Week 58	Sparsentan	139	129 (92.8)	0.92 (0.91)	0.1	0.28	0.70	1.22	5.9		
			Irbesartan	143	127 (88.8)	1.40 (1.03)	0.1	0.72	1.09	1.87	5.6		
		Week 110	Sparsentan	139	116 (83.5)	0.93 (0.87)	0.0	0.31	0.69	1.30	5.3		
			Irbesartan	143	107 (74.8)	1.50 (1.32)	0.1	0.66	1.22	1.91	7.8		
		Percent change from baseline in UP/C	Week 6	Sparsentan	Sparsentan	139	134 (96.4)	-0.40 (0.70)	-2.3	-0.76	-0.38	-0.05	2.3
					Irbesartan	143	138 (96.5)	0.03 (0.72)	-4.3	-0.29	-0.03	0.33	3.7
				Week 36	Sparsentan	139	135 (97.1)	-0.39 (0.82)	-3.1	-0.74	-0.42	-0.07	3.6
					Irbesartan	143	132 (92.3)	0.02 (0.86)	-4.3	-0.44	-0.01	0.39	2.6
				Week 58	Sparsentan	139	129 (92.8)	-0.38 (0.90)	-3.0	-0.82	-0.36	0.02	5.7
					Irbesartan	143	127 (88.8)	0.06 (0.85)	-1.8	-0.37	-0.06	0.42	3.5
				Week 110	Sparsentan	139	116 (83.5)	-0.31 (0.82)	-3.3	-0.73	-0.36	0.04	2.5
					Irbesartan	143	107 (74.8)	0.16 (1.19)	-5.5	-0.35	0.02	0.61	5.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
 Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	
Female	UP/C	Baseline	Sparsentan	63	63 (100.0)	1.71 (1.06)	0.6	1.03	1.40	2.13	7.0	
			Irbesartan	59	59 (100.0)	1.72 (0.94)	0.5	1.14	1.50	2.04	4.5	
		Week 6	Sparsentan	63	59 (93.7)	1.10 (0.73)	0.1	0.57	0.92	1.39	3.3	
			Irbesartan	59	56 (94.9)	1.54 (1.08)	0.4	0.75	1.15	2.06	6.2	
		Week 36	Sparsentan	63	59 (93.7)	0.95 (0.69)	0.1	0.50	0.80	1.26	3.7	
			Irbesartan	59	55 (93.2)	1.44 (1.18)	0.2	0.62	1.02	2.06	6.6	
		Week 58	Sparsentan	63	59 (93.7)	0.99 (0.68)	0.1	0.49	0.77	1.55	3.0	
			Irbesartan	59	52 (88.1)	1.39 (1.06)	0.2	0.60	1.07	1.77	5.5	
		Week 110	Sparsentan	63	52 (82.5)	1.22 (1.23)	0.1	0.41	0.96	1.57	6.7	
			Irbesartan	59	42 (71.2)	1.60 (1.05)	0.2	0.81	1.36	2.13	4.2	
		Percent change from baseline in UP/C	Week 6	Sparsentan	63	59 (93.7)	-0.48 (0.82)	-2.3	-0.83	-0.48	-0.14	1.9
				Irbesartan	59	56 (94.9)	-0.19 (0.69)	-1.5	-0.57	-0.22	0.13	1.9
	Week 36		Sparsentan	63	59 (93.7)	-0.68 (0.85)	-3.2	-1.09	-0.60	-0.16	1.1	
			Irbesartan	59	55 (93.2)	-0.22 (0.79)	-1.9	-0.62	-0.21	0.08	2.4	
	Week 58		Sparsentan	63	59 (93.7)	-0.61 (0.93)	-2.8	-1.00	-0.54	-0.14	1.7	
			Irbesartan	59	52 (88.1)	-0.29 (0.77)	-3.2	-0.72	-0.25	0.12	1.3	
	Week 110	Sparsentan	63	52 (82.5)	-0.38 (1.17)	-2.9	-1.04	-0.38	0.17	3.5		
		Irbesartan	59	42 (71.2)	0.00 (0.79)	-1.3	-0.50	-0.06	0.43	2.7		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
Subgroup: Age													
<= 45 years	UP/C	Baseline	Sparsentan	96	96 (100.0)	1.41 (0.96)	0.2	0.79	1.15	1.76	7.0		
			Irbesartan	99	99 (100.0)	1.44 (0.76)	0.2	0.91	1.30	1.77	4.2		
		Week 6	Sparsentan	96	92 (95.8)	0.91 (0.83)	0.1	0.37	0.76	1.11	5.6		
			Irbesartan	99	96 (97.0)	1.43 (1.01)	0.1	0.70	1.15	1.89	6.2		
		Week 36	Sparsentan	96	91 (94.8)	0.94 (0.93)	0.1	0.28	0.58	1.32	5.2		
			Irbesartan	99	91 (91.9)	1.33 (1.11)	0.1	0.58	0.93	2.02	6.6		
		Week 58	Sparsentan	96	89 (92.7)	0.93 (0.78)	0.1	0.29	0.71	1.22	3.5		
			Irbesartan	99	85 (85.9)	1.34 (1.04)	0.1	0.57	1.05	1.86	5.6		
		Week 110	Sparsentan	96	76 (79.2)	0.95 (0.88)	0.0	0.35	0.78	1.26	5.3		
			Irbesartan	99	69 (69.7)	1.43 (1.20)	0.1	0.65	1.11	1.80	6.4		
		Percent change from baseline in UP/C	Week 6	Sparsentan	96	92 (95.8)	-0.47 (0.76)	-2.3	-0.86	-0.46	-0.14	2.3	
					Irbesartan	99	96 (97.0)	-0.01 (0.71)	-1.4	-0.45	-0.18	0.33	3.7
				Week 36	Sparsentan	96	91 (94.8)	-0.43 (0.89)	-2.6	-0.73	-0.46	-0.07	3.6
					Irbesartan	99	91 (91.9)	-0.07 (0.81)	-1.9	-0.55	-0.16	0.27	2.4
Week 58	Sparsentan			96	89 (92.7)	-0.43 (0.79)	-2.8	-0.83	-0.36	0.04	1.7		
	Irbesartan			99	85 (85.9)	-0.07 (0.99)	-3.2	-0.48	-0.24	0.25	3.5		
Week 110	Sparsentan			96	76 (79.2)	-0.33 (0.91)	-3.3	-0.79	-0.31	0.08	2.5		
	Irbesartan			99	69 (69.7)	0.10 (1.07)	-1.8	-0.50	-0.06	0.46	4.3		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
 Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
> 45 years	UP/C	Baseline	Sparsentan	106	106 (100.0)	1.44 (0.84)	0.1	0.74	1.31	1.88	4.2		
			Irbesartan	103	103 (100.0)	1.44 (1.01)	0.4	0.85	1.17	1.72	6.9		
		Week 6	Sparsentan	106	101 (95.3)	1.03 (0.77)	0.1	0.47	0.82	1.33	3.5		
			Irbesartan	103	98 (95.1)	1.40 (0.87)	0.4	0.80	1.15	1.70	5.4		
		Week 36	Sparsentan	106	103 (97.2)	0.93 (0.81)	0.1	0.40	0.69	1.28	4.4		
			Irbesartan	103	96 (93.2)	1.44 (0.97)	0.2	0.78	1.29	1.79	6.2		
		Week 58	Sparsentan	106	99 (93.4)	0.95 (0.90)	0.1	0.32	0.75	1.29	5.9		
			Irbesartan	103	94 (91.3)	1.45 (1.03)	0.1	0.82	1.18	1.84	5.5		
		Week 110	Sparsentan	106	92 (86.8)	1.08 (1.09)	0.1	0.35	0.75	1.48	6.7		
			Irbesartan	103	80 (77.7)	1.61 (1.29)	0.1	0.81	1.36	2.06	7.8		
		Percent change from baseline in UP/C	Week 6	Sparsentan	106	101 (95.3)	-0.38 (0.71)	-2.3	-0.71	-0.36	-0.01	1.7	
					Irbesartan	103	98 (95.1)	-0.06 (0.73)	-4.3	-0.31	-0.08	0.29	1.9
				Week 36	Sparsentan	106	103 (97.2)	-0.52 (0.79)	-3.2	-1.01	-0.52	-0.15	1.7
					Irbesartan	103	96 (93.2)	-0.04 (0.88)	-4.3	-0.48	-0.05	0.33	2.6
				Week 58	Sparsentan	106	99 (93.4)	-0.47 (1.01)	-3.0	-1.01	-0.45	-0.03	5.7
					Irbesartan	103	94 (91.3)	-0.01 (0.67)	-1.8	-0.31	-0.03	0.35	1.7
				Week 110	Sparsentan	106	92 (86.8)	-0.33 (0.96)	-2.9	-0.75	-0.40	0.04	3.5
					Irbesartan	103	80 (77.7)	0.13 (1.12)	-5.5	-0.36	0.09	0.56	5.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
Subgroup: Age at IgAN diagnosis													
<= 18 years	UP/C	Baseline	Sparsentan	9	9 (100.0)	2.47 (1.88)	0.7	1.33	1.86	2.43	7.0		
			Irbesartan	5	5 (100.0)	1.35 (0.44)	0.7	1.30	1.41	1.49	1.9		
		Week 6	Sparsentan	9	8 (88.9)	1.09 (0.79)	0.2	0.46	0.96	1.54	2.6		
			Irbesartan	5	5 (100.0)	1.40 (0.78)	0.8	0.79	0.95	2.05	2.4		
		Week 36	Sparsentan	9	8 (88.9)	1.29 (0.85)	0.4	0.58	1.19	1.73	2.9		
			Irbesartan	5	5 (100.0)	1.23 (0.56)	0.7	0.84	0.92	1.84	1.8		
		Week 58	Sparsentan	9	7 (77.8)	1.37 (1.06)	0.1	0.71	1.10	2.73	2.9		
			Irbesartan	5	4 (80.0)	1.07 (0.69)	0.4	0.50	1.04	1.65	1.9		
		Week 110	Sparsentan	9	5 (55.6)	2.40 (1.98)	0.1	1.31	2.09	3.20	5.3		
			Irbesartan	5	4 (80.0)	1.07 (0.69)	0.3	0.54	1.08	1.61	1.9		
		Percent change from baseline in UP/C		Week 6	Sparsentan	9	8 (88.9)	-0.82 (0.88)	-2.2	-1.36	-0.80	-0.19	0.4
					Irbesartan	5	5 (100.0)	0.06 (0.77)	-0.9	-0.51	0.13	0.64	0.9
				Week 36	Sparsentan	9	8 (88.9)	-0.61 (1.06)	-2.5	-1.22	-0.37	0.13	0.5
					Irbesartan	5	5 (100.0)	-0.11 (0.67)	-1.1	-0.46	0.25	0.35	0.4
				Week 58	Sparsentan	9	7 (77.8)	-0.62 (1.23)	-2.8	-1.15	-0.65	0.49	0.9
					Irbesartan	5	4 (80.0)	-0.24 (0.87)	-1.5	-0.77	0.05	0.29	0.5
				Week 110	Sparsentan	9	5 (55.6)	0.25 (1.07)	-0.9	-0.64	0.38	0.77	1.7
					Irbesartan	5	4 (80.0)	-0.29 (0.90)	-1.6	-0.83	0.05	0.26	0.4

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
 Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
> 18 to 40 years	UP/C	Baseline	Sparsentan	102	102 (100.0)	1.34 (0.75)	0.2	0.79	1.17	1.76	3.6		
			Irbesartan	109	109 (100.0)	1.46 (0.77)	0.2	0.93	1.33	1.75	4.5		
		Week 6	Sparsentan	102	99 (97.1)	0.92 (0.82)	0.1	0.37	0.75	1.12	5.6		
			Irbesartan	109	105 (96.3)	1.43 (0.97)	0.1	0.72	1.14	1.87	6.2		
		Week 36	Sparsentan	102	97 (95.1)	0.91 (0.92)	0.1	0.28	0.58	1.24	5.2		
			Irbesartan	109	100 (91.7)	1.37 (1.06)	0.1	0.63	1.04	1.79	6.6		
		Week 58	Sparsentan	102	95 (93.1)	0.85 (0.71)	0.1	0.29	0.67	1.11	3.5		
			Irbesartan	109	93 (85.3)	1.41 (1.02)	0.1	0.70	1.12	1.75	5.6		
		Week 110	Sparsentan	102	82 (80.4)	0.94 (0.95)	0.0	0.37	0.73	1.24	6.7		
			Irbesartan	109	78 (71.6)	1.57 (1.38)	0.1	0.70	1.18	1.93	7.8		
		Percent change from baseline in UP/C	Week 6	Sparsentan	102	99 (97.1)	-0.45 (0.72)	-2.3	-0.79	-0.45	-0.14	2.3	
					Irbesartan	109	105 (96.3)	-0.04 (0.62)	-1.5	-0.47	-0.18	0.34	1.9
				Week 36	Sparsentan	102	97 (95.1)	-0.44 (0.86)	-2.6	-0.81	-0.51	-0.07	3.6
					Irbesartan	109	100 (91.7)	-0.08 (0.83)	-1.9	-0.60	-0.13	0.27	2.6
				Week 58	Sparsentan	102	95 (93.1)	-0.49 (0.73)	-2.8	-0.93	-0.38	-0.02	1.2
					Irbesartan	109	93 (85.3)	-0.06 (0.95)	-3.2	-0.47	-0.24	0.35	3.5
				Week 110	Sparsentan	102	82 (80.4)	-0.35 (0.98)	-3.3	-0.83	-0.38	0.02	3.5
					Irbesartan	109	78 (71.6)	0.18 (1.16)	-1.6	-0.50	-0.05	0.46	5.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
> 40 years	UP/C	Baseline	Sparsentan	91	91 (100.0)	1.42 (0.86)	0.1	0.73	1.27	1.79	4.2		
			Irbesartan	88	88 (100.0)	1.42 (1.05)	0.4	0.82	1.11	1.72	6.9		
		Week 6	Sparsentan	91	86 (94.5)	1.03 (0.78)	0.1	0.48	0.82	1.36	3.5		
			Irbesartan	88	84 (95.5)	1.41 (0.91)	0.4	0.79	1.17	1.76	5.4		
		Week 36	Sparsentan	91	89 (97.8)	0.92 (0.81)	0.1	0.42	0.69	1.28	4.4		
			Irbesartan	88	82 (93.2)	1.42 (1.04)	0.2	0.62	1.16	2.14	6.2		
		Week 58	Sparsentan	91	86 (94.5)	1.00 (0.94)	0.1	0.35	0.76	1.40	5.9		
			Irbesartan	88	82 (93.2)	1.40 (1.07)	0.1	0.73	1.05	1.94	5.5		
		Week 110	Sparsentan	91	81 (89.0)	1.02 (0.92)	0.1	0.34	0.76	1.47	5.3		
			Irbesartan	88	67 (76.1)	1.51 (1.12)	0.1	0.68	1.35	2.06	7.3		
		Percent change from baseline in UP/C	Week 6	Sparsentan	91	86 (94.5)	-0.35 (0.72)	-2.3	-0.71	-0.32	-0.00	1.7	
					Irbesartan	88	84 (95.5)	-0.03 (0.82)	-4.3	-0.28	-0.07	0.27	3.7
				Week 36	Sparsentan	91	89 (97.8)	-0.50 (0.80)	-3.2	-0.85	-0.51	-0.15	1.7
					Irbesartan	88	82 (93.2)	-0.01 (0.88)	-4.3	-0.38	-0.05	0.35	2.3
				Week 58	Sparsentan	91	86 (94.5)	-0.39 (1.07)	-3.0	-0.94	-0.44	0.05	5.7
					Irbesartan	88	82 (93.2)	-0.00 (0.70)	-1.8	-0.31	-0.04	0.28	2.4
				Week 110	Sparsentan	91	81 (89.0)	-0.35 (0.89)	-2.9	-0.72	-0.38	0.05	2.0
					Irbesartan	88	67 (76.1)	0.07 (1.03)	-5.5	-0.36	0.03	0.81	1.8

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
 Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024



Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max
Subgroup: Geographic region											
North America	UP/C	Baseline	Sparsentan	35	35 (100.0)	1.56 (0.97)	0.1	0.90	1.36	2.20	4.0
			Irbesartan	46	46 (100.0)	1.65 (1.05)	0.4	1.14	1.43	1.77	6.9
	Week 6	Sparsentan	35	33 (94.3)	1.12 (0.73)	0.2	0.52	1.00	1.40	3.3	
		Irbesartan	46	45 (97.8)	1.75 (1.05)	0.4	1.04	1.47	2.22	6.2	
	Week 36	Sparsentan	35	33 (94.3)	0.90 (0.68)	0.2	0.39	0.73	1.12	2.6	
		Irbesartan	46	43 (93.5)	1.74 (1.07)	0.3	1.03	1.47	2.46	6.6	
	Week 58	Sparsentan	35	28 (80.0)	1.08 (1.17)	0.1	0.41	0.67	1.37	5.9	
		Irbesartan	46	40 (87.0)	1.77 (1.23)	0.3	0.82	1.46	2.53	5.6	
	Week 110	Sparsentan	35	27 (77.1)	1.03 (1.00)	0.1	0.39	0.76	1.31	5.3	
		Irbesartan	46	31 (67.4)	1.65 (0.94)	0.2	0.89	1.38	2.40	3.7	
	Percent change from baseline in UP/C	Week 6	Sparsentan	35	33 (94.3)	-0.42 (1.04)	-2.3	-0.70	-0.18	0.16	1.9
			Irbesartan	46	45 (97.8)	0.09 (0.98)	-4.3	-0.27	0.06	0.51	1.9
		Week 36	Sparsentan	35	33 (94.3)	-0.68 (1.09)	-3.2	-1.18	-0.46	0.02	1.2
			Irbesartan	46	43 (93.5)	0.09 (1.03)	-4.3	-0.46	0.06	0.69	2.4
		Week 58	Sparsentan	35	28 (80.0)	-0.44 (1.58)	-2.8	-1.03	-0.48	0.13	5.7
			Irbesartan	46	40 (87.0)	0.07 (1.06)	-2.1	-0.55	-0.08	0.63	3.5
		Week 110	Sparsentan	35	27 (77.1)	-0.40 (1.00)	-2.8	-0.94	-0.20	0.08	1.7
			Irbesartan	46	31 (67.4)	-0.07 (1.27)	-5.5	-0.56	-0.10	0.79	2.0

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	
Europe	UP/C	Baseline	Sparsentan	98	98 (100.0)	1.39 (0.94)	0.3	0.75	1.20	1.82	7.0	
			Irbesartan	115	115 (100.0)	1.40 (0.80)	0.4	0.85	1.17	1.77	4.5	
		Week 6	Sparsentan	98	92 (93.9)	0.88 (0.68)	0.1	0.39	0.75	1.08	3.5	
			Irbesartan	115	109 (94.8)	1.33 (0.85)	0.3	0.71	1.04	1.66	4.2	
		Week 36	Sparsentan	98	94 (95.9)	0.95 (0.96)	0.1	0.31	0.65	1.35	5.2	
			Irbesartan	115	108 (93.9)	1.29 (0.95)	0.2	0.57	1.01	1.73	4.6	
		Week 58	Sparsentan	98	93 (94.9)	0.91 (0.74)	0.1	0.32	0.75	1.24	4.1	
			Irbesartan	115	101 (87.8)	1.31 (0.94)	0.1	0.65	0.99	1.75	5.5	
		Week 110	Sparsentan	98	79 (80.6)	0.96 (0.93)	0.1	0.31	0.69	1.31	5.3	
			Irbesartan	115	87 (75.7)	1.50 (1.31)	0.1	0.62	1.19	1.96	7.8	
		Percent change from baseline in UP/C	Week 6	Sparsentan	98	92 (93.9)	-0.45 (0.66)	-2.3	-0.79	-0.43	-0.11	1.6
				Irbesartan	115	109 (94.8)	-0.08 (0.65)	-1.5	-0.48	-0.12	0.25	3.7
	Week 36		Sparsentan	98	94 (95.9)	-0.38 (0.82)	-3.1	-0.75	-0.47	-0.10	3.6	
			Irbesartan	115	108 (93.9)	-0.08 (0.81)	-1.7	-0.53	-0.15	0.27	2.6	
	Week 58		Sparsentan	98	93 (94.9)	-0.41 (0.68)	-2.2	-0.88	-0.36	-0.02	1.2	
			Irbesartan	115	101 (87.8)	-0.08 (0.83)	-3.2	-0.41	-0.10	0.23	3.0	
	Week 110	Sparsentan	98	79 (80.6)	-0.33 (0.87)	-2.9	-0.76	-0.44	0.05	2.5		
		Irbesartan	115	87 (75.7)	0.16 (1.11)	-1.6	-0.44	-0.01	0.61	5.3		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
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Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	
Asia Pacific	UP/C	Baseline	Sparsentan	69	69 (100.0)	1.41 (0.81)	0.2	0.79	1.22	1.76	4.2	
			Irbesartan	41	41 (100.0)	1.32 (0.94)	0.2	0.80	1.02	1.54	5.5	
		Week 6	Sparsentan	69	68 (98.6)	1.03 (0.95)	0.1	0.39	0.77	1.29	5.6	
			Irbesartan	41	40 (97.6)	1.30 (0.98)	0.1	0.66	1.04	1.68	5.4	
		Week 36	Sparsentan	69	67 (97.1)	0.92 (0.83)	0.1	0.30	0.69	1.24	4.0	
			Irbesartan	41	36 (87.8)	1.26 (1.19)	0.1	0.54	0.90	1.58	6.2	
		Week 58	Sparsentan	69	67 (97.1)	0.91 (0.82)	0.1	0.26	0.71	1.29	3.5	
			Irbesartan	41	38 (92.7)	1.26 (0.97)	0.1	0.67	1.11	1.60	5.2	
		Week 110	Sparsentan	69	62 (89.9)	1.10 (1.08)	0.0	0.37	0.83	1.54	6.7	
			Irbesartan	41	31 (75.6)	1.48 (1.37)	0.1	0.68	1.22	1.80	7.3	
		Percent change from baseline in UP/C	Week 6	Sparsentan	69	68 (98.6)	-0.38 (0.66)	-2.1	-0.71	-0.41	-0.10	2.3
				Irbesartan	41	40 (97.6)	-0.04 (0.52)	-1.4	-0.32	-0.11	0.32	1.1
	Week 36		Sparsentan	69	67 (97.1)	-0.50 (0.69)	-2.6	-0.84	-0.51	-0.16	1.1	
			Irbesartan	41	36 (87.8)	-0.13 (0.70)	-1.9	-0.55	-0.15	0.15	2.2	
	Week 58		Sparsentan	69	67 (97.1)	-0.51 (0.84)	-3.0	-0.97	-0.48	-0.07	1.7	
			Irbesartan	41	38 (92.7)	-0.05 (0.54)	-1.3	-0.36	-0.07	0.28	1.3	
	Week 110	Sparsentan	69	62 (89.9)	-0.31 (1.01)	-3.3	-0.68	-0.31	0.06	3.5		
		Irbesartan	41	31 (75.6)	0.17 (0.85)	-1.8	-0.21	0.15	0.35	2.7		

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
Subgroup: Baseline BMI													
< 27 kg/m**2	UP/C	Baseline	Sparsentan	83	83 (100.0)	1.40 (0.74)	0.3	0.79	1.27	1.79	3.6		
			Irbesartan	94	94 (100.0)	1.51 (0.87)	0.4	0.92	1.31	1.80	5.5		
		Week 6	Sparsentan	83	77 (92.8)	0.95 (0.74)	0.1	0.38	0.76	1.27	3.3		
			Irbesartan	94	90 (95.7)	1.44 (0.96)	0.3	0.75	1.14	1.90	5.4		
		Week 36	Sparsentan	83	80 (96.4)	0.93 (0.91)	0.1	0.34	0.63	1.27	5.2		
			Irbesartan	94	85 (90.4)	1.29 (0.97)	0.2	0.62	1.03	1.61	6.2		
		Week 58	Sparsentan	83	76 (91.6)	0.80 (0.60)	0.1	0.29	0.73	1.11	2.9		
			Irbesartan	94	80 (85.1)	1.33 (1.05)	0.1	0.61	0.97	1.78	5.6		
		Week 110	Sparsentan	83	70 (84.3)	1.09 (1.19)	0.1	0.30	0.79	1.31	6.7		
			Irbesartan	94	66 (70.2)	1.50 (1.29)	0.1	0.68	1.24	1.84	7.3		
		Percent change from baseline in UP/C	Week 6	Sparsentan	83	77 (92.8)	-0.45 (0.72)	-2.2	-0.88	-0.48	-0.12	1.9	
					Irbesartan	94	90 (95.7)	-0.08 (0.57)	-1.5	-0.42	-0.19	0.25	1.5
				Week 36	Sparsentan	83	80 (96.4)	-0.49 (0.87)	-2.6	-0.85	-0.56	-0.17	3.6
					Irbesartan	94	85 (90.4)	-0.19 (0.64)	-1.9	-0.53	-0.18	0.22	1.5
				Week 58	Sparsentan	83	76 (91.6)	-0.61 (0.80)	-2.8	-1.02	-0.52	-0.07	1.1
					Irbesartan	94	80 (85.1)	-0.16 (0.83)	-3.2	-0.49	-0.24	0.16	3.5
				Week 110	Sparsentan	83	70 (84.3)	-0.29 (1.02)	-2.9	-0.74	-0.38	0.06	3.5
					Irbesartan	94	66 (70.2)	0.01 (0.85)	-1.8	-0.44	-0.09	0.43	3.6

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

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Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
>= 27 kg/m**2	UP/C	Baseline	Sparsentan	119	119 (100.0)	1.44 (1.00)	0.1	0.74	1.22	1.86	7.0		
			Irbesartan	107	107 (100.0)	1.39 (0.92)	0.2	0.84	1.16	1.66	6.9		
		Week 6	Sparsentan	119	116 (97.5)	0.99 (0.83)	0.1	0.46	0.80	1.17	5.6		
			Irbesartan	107	103 (96.3)	1.41 (0.92)	0.1	0.79	1.16	1.75	6.2		
		Week 36	Sparsentan	119	114 (95.8)	0.93 (0.84)	0.1	0.33	0.70	1.32	4.4		
			Irbesartan	107	101 (94.4)	1.48 (1.09)	0.1	0.68	1.24	2.06	6.6		
		Week 58	Sparsentan	119	112 (94.1)	1.04 (0.96)	0.1	0.36	0.73	1.34	5.9		
			Irbesartan	107	99 (92.5)	1.46 (1.02)	0.1	0.73	1.18	1.94	5.5		
		Week 110	Sparsentan	119	98 (82.4)	0.97 (0.83)	0.0	0.39	0.75	1.39	5.3		
			Irbesartan	107	82 (76.6)	1.56 (1.22)	0.1	0.79	1.25	2.10	7.8		
		Percent change from baseline in UP/C	Week 6	Sparsentan	119	116 (97.5)	-0.40 (0.75)	-2.3	-0.72	-0.33	0.02	2.3	
					Irbesartan	107	103 (96.3)	0.02 (0.82)	-4.3	-0.29	0.05	0.34	3.7
				Week 36	Sparsentan	119	114 (95.8)	-0.46 (0.81)	-3.2	-0.85	-0.42	-0.07	1.7
					Irbesartan	107	101 (94.4)	0.07 (0.98)	-4.3	-0.46	-0.00	0.49	2.6
				Week 58	Sparsentan	119	112 (94.1)	-0.34 (0.97)	-3.0	-0.79	-0.33	0.06	5.7
					Irbesartan	107	99 (92.5)	0.06 (0.83)	-2.1	-0.36	-0.02	0.46	3.0
				Week 110	Sparsentan	119	98 (82.4)	-0.37 (0.88)	-3.3	-0.76	-0.37	0.04	2.0
					Irbesartan	107	82 (76.6)	0.21 (1.26)	-5.5	-0.35	0.05	0.84	5.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
 Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	
Subgroup: Randomization strata												
eGFR Low and UP High	UP/C	Baseline	Sparsentan	71	71 (100.0)	1.91 (1.07)	0.1	1.26	1.74	2.39	7.0	
			Irbesartan	74	74 (100.0)	1.97 (1.06)	0.5	1.25	1.69	2.51	6.9	
		Week 6	Sparsentan	71	68 (95.8)	1.31 (0.80)	0.3	0.75	1.06	2.06	3.5	
			Irbesartan	74	73 (98.6)	1.96 (1.00)	0.6	1.19	1.90	2.60	6.2	
		Week 36	Sparsentan	71	68 (95.8)	1.30 (1.01)	0.1	0.55	0.97	1.79	4.4	
			Irbesartan	74	68 (91.9)	1.92 (1.13)	0.2	1.19	1.64	2.60	6.6	
		Week 58	Sparsentan	71	67 (94.4)	1.23 (0.99)	0.1	0.54	0.93	1.68	5.9	
			Irbesartan	74	65 (87.8)	1.86 (1.19)	0.1	0.94	1.63	2.48	5.6	
		Week 110	Sparsentan	71	59 (83.1)	1.45 (1.24)	0.1	0.53	1.21	2.00	6.7	
			Irbesartan	74	53 (71.6)	1.96 (1.46)	0.1	1.04	1.49	2.48	7.8	
		Percent change from baseline in UP/C	Week 6	Sparsentan	71	68 (95.8)	-0.51 (0.82)	-2.3	-1.00	-0.51	0.16	1.5
				Irbesartan	74	73 (98.6)	-0.00 (0.92)	-4.3	-0.50	0.07	0.47	1.9
			Week 36	Sparsentan	71	68 (95.8)	-0.54 (0.98)	-3.2	-1.11	-0.58	0.22	1.7
				Irbesartan	74	68 (91.9)	-0.04 (1.12)	-4.3	-0.76	-0.00	0.64	2.4
			Week 58	Sparsentan	71	67 (94.4)	-0.57 (1.17)	-3.0	-1.25	-0.59	-0.07	5.7
				Irbesartan	74	65 (87.8)	-0.09 (1.06)	-3.2	-0.85	0.03	0.51	3.5
			Week 110	Sparsentan	71	59 (83.1)	-0.27 (1.16)	-2.9	-0.94	-0.44	0.38	3.5
				Irbesartan	74	53 (71.6)	0.06 (1.50)	-5.5	-0.74	0.02	0.71	5.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
eGFR Low and UP Low	UP/C	Baseline	Sparsentan	55	55 (100.0)	1.02 (0.51)	0.3	0.66	0.93	1.31	2.8		
			Irbesartan	55	55 (100.0)	1.01 (0.38)	0.4	0.71	0.92	1.32	1.9		
		Week 6	Sparsentan	55	51 (92.7)	0.73 (0.53)	0.1	0.41	0.60	0.91	2.8		
			Irbesartan	55	53 (96.4)	0.91 (0.40)	0.4	0.63	0.80	1.06	2.3		
		Week 36	Sparsentan	55	51 (92.7)	0.72 (0.54)	0.1	0.32	0.58	1.00	2.2		
			Irbesartan	55	51 (92.7)	1.02 (0.66)	0.2	0.49	0.88	1.52	3.1		
		Week 58	Sparsentan	55	52 (94.5)	0.76 (0.57)	0.1	0.26	0.67	1.06	2.4		
			Irbesartan	55	51 (92.7)	0.98 (0.61)	0.1	0.55	0.85	1.28	2.7		
		Week 110	Sparsentan	55	46 (83.6)	0.87 (0.67)	0.1	0.35	0.75	1.31	3.4		
			Irbesartan	55	42 (76.4)	1.10 (0.72)	0.1	0.49	0.88	1.80	2.6		
		Percent change from baseline in UP/C	Week 6	Sparsentan	55	51 (92.7)	-0.31 (0.66)	-2.3	-0.58	-0.26	0.02	1.7	
					Irbesartan	55	53 (96.4)	-0.11 (0.42)	-1.1	-0.30	-0.12	0.09	1.2
				Week 36	Sparsentan	55	51 (92.7)	-0.30 (0.66)	-1.9	-0.62	-0.30	0.02	1.2
					Irbesartan	55	51 (92.7)	-0.01 (0.66)	-1.2	-0.36	-0.14	0.26	2.6
				Week 58	Sparsentan	55	52 (94.5)	-0.25 (0.60)	-1.6	-0.54	-0.28	0.08	1.1
					Irbesartan	55	51 (92.7)	-0.03 (0.59)	-1.0	-0.37	-0.10	0.28	1.7
				Week 110	Sparsentan	55	46 (83.6)	-0.14 (0.73)	-1.7	-0.59	-0.23	0.20	2.0
					Irbesartan	55	42 (76.4)	0.09 (0.62)	-1.3	-0.35	0.02	0.51	1.2

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
eGFR High and UP High	UP/C	Baseline	Sparsentan	37	37 (100.0)	1.71 (0.76)	0.6	1.22	1.45	2.13	3.6		
			Irbesartan	36	36 (100.0)	1.57 (0.83)	0.5	1.14	1.47	1.83	5.5		
		Week 6	Sparsentan	37	35 (94.6)	1.19 (1.08)	0.1	0.48	1.05	1.47	5.6		
			Irbesartan	36	32 (88.9)	1.66 (1.05)	0.6	1.02	1.39	1.88	5.4		
		Week 36	Sparsentan	37	37 (100.0)	1.04 (1.02)	0.1	0.49	0.80	1.31	5.2		
			Irbesartan	36	32 (88.9)	1.53 (1.21)	0.2	0.77	1.05	2.15	6.2		
		Week 58	Sparsentan	37	32 (86.5)	1.04 (0.89)	0.1	0.38	0.91	1.18	3.5		
			Irbesartan	36	31 (86.1)	1.67 (1.14)	0.3	0.75	1.35	2.39	5.2		
		Week 110	Sparsentan	37	30 (81.1)	0.87 (1.00)	0.0	0.34	0.62	1.14	5.3		
			Irbesartan	36	27 (75.0)	1.92 (1.54)	0.3	0.95	1.42	2.33	7.3		
		Percent change from baseline in UP/C	Percent change from baseline in UP/C	Week 6	Sparsentan	37	35 (94.6)	-0.50 (0.97)	-2.2	-0.94	-0.62	-0.18	2.3
					Irbesartan	36	32 (88.9)	0.04 (0.85)	-0.9	-0.49	-0.25	0.28	3.7
				Week 36	Sparsentan	37	37 (100.0)	-0.68 (1.07)	-2.6	-1.16	-0.65	-0.26	3.6
					Irbesartan	36	32 (88.9)	-0.06 (0.78)	-1.3	-0.60	-0.03	0.25	2.2
				Week 58	Sparsentan	37	32 (86.5)	-0.73 (1.01)	-2.8	-1.17	-0.63	0.00	1.2
					Irbesartan	36	31 (86.1)	0.08 (0.94)	-1.2	-0.51	-0.23	0.25	3.0
				Week 110	Sparsentan	37	30 (81.1)	-0.83 (1.02)	-3.3	-1.41	-0.61	-0.22	1.7
					Irbesartan	36	27 (75.0)	0.31 (1.16)	-1.1	-0.50	-0.01	0.91	4.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024



Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
eGFR High and UP Low	UP/C	Baseline	Sparsentan	39	39 (100.0)	0.84 (0.38)	0.2	0.48	0.80	1.08	1.8		
			Irbesartan	37	37 (100.0)	0.89 (0.31)	0.2	0.69	0.90	1.10	1.6		
		Week 6	Sparsentan	39	39 (100.0)	0.49 (0.31)	0.1	0.29	0.38	0.60	1.4		
			Irbesartan	37	36 (97.3)	0.85 (0.43)	0.1	0.58	0.81	1.14	1.9		
		Week 36	Sparsentan	39	38 (97.4)	0.46 (0.36)	0.1	0.23	0.33	0.65	1.3		
			Irbesartan	37	36 (97.3)	0.78 (0.48)	0.1	0.39	0.70	1.09	2.3		
		Week 58	Sparsentan	39	37 (94.9)	0.59 (0.60)	0.1	0.20	0.32	0.75	3.0		
			Irbesartan	37	32 (86.5)	0.86 (0.51)	0.1	0.41	0.80	1.07	1.9		
		Week 110	Sparsentan	39	33 (84.6)	0.60 (0.54)	0.1	0.19	0.43	0.86	2.5		
			Irbesartan	37	27 (73.0)	0.96 (0.52)	0.1	0.52	0.84	1.35	2.1		
		Percent change from baseline in UP/C	Week 6	Sparsentan	39	39 (100.0)	-0.35 (0.26)	-1.0	-0.58	-0.31	-0.15	0.2	
					Irbesartan	37	36 (97.3)	-0.04 (0.42)	-1.2	-0.29	-0.08	0.25	0.7
				Week 36	Sparsentan	39	38 (97.4)	-0.39 (0.32)	-1.0	-0.66	-0.40	-0.19	0.4
					Irbesartan	37	36 (97.3)	-0.13 (0.49)	-1.3	-0.40	-0.13	0.14	1.3
				Week 58	Sparsentan	39	37 (94.9)	-0.27 (0.53)	-1.1	-0.64	-0.34	-0.03	1.7
					Irbesartan	37	32 (86.5)	-0.05 (0.50)	-1.3	-0.28	-0.10	0.10	1.3
				Week 110	Sparsentan	39	33 (84.6)	-0.26 (0.43)	-0.9	-0.52	-0.30	-0.11	1.0
					Irbesartan	37	27 (73.0)	0.07 (0.53)	-1.1	-0.24	-0.04	0.44	1.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
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Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
Subgroup: Baseline eGFR Group 1													
< 60 mL/min/1.73 m**2	UP/C	Baseline	Sparsentan	127	127 (100.0)	1.48 (0.98)	0.1	0.77	1.28	1.88	7.0		
			Irbesartan	129	129 (100.0)	1.52 (0.95)	0.4	0.88	1.32	1.77	6.9		
		Week 6	Sparsentan	127	120 (94.5)	1.02 (0.74)	0.1	0.48	0.84	1.31	3.5		
			Irbesartan	129	126 (97.7)	1.48 (0.94)	0.4	0.80	1.16	1.99	6.2		
		Week 36	Sparsentan	127	120 (94.5)	1.04 (0.89)	0.1	0.39	0.73	1.51	4.4		
			Irbesartan	129	119 (92.2)	1.46 (1.04)	0.2	0.65	1.28	1.82	6.6		
		Week 58	Sparsentan	127	120 (94.5)	0.99 (0.86)	0.1	0.40	0.74	1.41	5.9		
			Irbesartan	129	116 (89.9)	1.45 (1.07)	0.1	0.72	1.15	1.94	5.6		
		Week 110	Sparsentan	127	105 (82.7)	1.18 (1.05)	0.1	0.46	0.87	1.62	6.7		
			Irbesartan	129	96 (74.4)	1.56 (1.25)	0.1	0.80	1.28	2.06	7.8		
		Percent change from baseline in UP/C	Week 6	Sparsentan	127	120 (94.5)	-0.42 (0.73)	-2.3	-0.86	-0.38	0.03	1.7	
					Irbesartan	129	126 (97.7)	-0.05 (0.74)	-4.3	-0.37	-0.11	0.34	1.9
				Week 36	Sparsentan	127	120 (94.5)	-0.41 (0.85)	-3.2	-0.85	-0.43	0.10	1.7
					Irbesartan	129	119 (92.2)	-0.05 (0.93)	-4.3	-0.54	-0.10	0.35	2.6
				Week 58	Sparsentan	127	120 (94.5)	-0.41 (0.97)	-3.0	-0.97	-0.39	0.03	5.7
					Irbesartan	129	116 (89.9)	-0.05 (0.84)	-3.2	-0.47	-0.10	0.35	3.5
				Week 110	Sparsentan	127	105 (82.7)	-0.19 (0.97)	-2.9	-0.72	-0.28	0.26	3.5
					Irbesartan	129	96 (74.4)	0.11 (1.17)	-5.5	-0.43	0.03	0.66	5.3

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max
60 to < 90 mL/min/1.73 m**2	UP/C	Baseline	Sparsentan	49	49 (100.0)	1.25 (0.59)	0.3	0.79	1.22	1.66	2.8
		Week 6	Irbesartan	48	48 (100.0)	1.34 (0.63)	0.5	0.87	1.20	1.70	3.4
			Sparsentan	49	48 (98.0)	0.88 (0.78)	0.1	0.32	0.58	1.21	3.5
		Week 36	Irbesartan	48	45 (93.8)	1.49 (0.84)	0.4	0.84	1.35	1.78	4.2
			Sparsentan	49	48 (98.0)	0.77 (0.81)	0.1	0.27	0.59	1.04	5.2
		Week 58	Irbesartan	48	45 (93.8)	1.47 (0.85)	0.3	0.84	1.32	2.07	4.1
			Sparsentan	49	43 (87.8)	0.83 (0.79)	0.1	0.18	0.77	1.16	3.1
		Week 110	Irbesartan	48	42 (87.5)	1.50 (0.86)	0.3	0.85	1.39	1.86	3.8
			Sparsentan	49	42 (85.7)	0.71 (0.64)	0.1	0.21	0.44	1.13	2.9
		Percent change from baseline in UP/C	Week 6	Sparsentan	49	48 (98.0)	-0.38 (0.69)	-2.3	-0.70	-0.43	-0.15
	Irbesartan			48	45 (93.8)	0.12 (0.77)	-1.4	-0.34	0.07	0.35	3.7
	Week 36		Sparsentan	49	48 (98.0)	-0.50 (0.77)	-1.8	-0.76	-0.54	-0.26	3.6
			Irbesartan	48	45 (93.8)	0.12 (0.72)	-1.3	-0.27	0.04	0.40	2.2
	Week 58		Sparsentan	49	43 (87.8)	-0.44 (0.67)	-1.8	-0.94	-0.52	-0.17	1.7
			Irbesartan	48	42 (87.5)	0.14 (0.94)	-2.1	-0.26	-0.02	0.47	3.0
	Week 110		Sparsentan	49	42 (85.7)	-0.57 (0.66)	-1.7	-1.01	-0.52	-0.19	1.0
			Irbesartan	48	36 (75.0)	0.20 (1.07)	-1.6	-0.31	0.03	0.62	4.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
>= 90 mL/min/1.73 m**2	UP/C	Baseline	Sparsentan	26	26 (100.0)	1.49 (0.98)	0.4	0.79	1.10	2.14	3.6		
			Irbesartan	25	25 (100.0)	1.24 (0.99)	0.2	0.73	1.10	1.46	5.5		
		Week 6	Sparsentan	26	25 (96.2)	0.91 (1.06)	0.2	0.37	0.63	0.93	5.6		
			Irbesartan	25	23 (92.0)	0.96 (1.03)	0.1	0.59	0.79	1.00	5.4		
		Week 36	Sparsentan	26	26 (100.0)	0.76 (0.83)	0.1	0.26	0.45	1.07	4.0		
			Irbesartan	25	23 (92.0)	0.85 (1.22)	0.1	0.24	0.58	0.92	6.2		
		Week 58	Sparsentan	26	25 (96.2)	0.86 (0.85)	0.2	0.29	0.55	0.85	3.5		
			Irbesartan	25	21 (84.0)	0.92 (1.06)	0.1	0.41	0.64	0.99	5.2		
		Week 110	Sparsentan	26	21 (80.8)	0.87 (1.17)	0.0	0.32	0.47	0.87	5.3		
			Irbesartan	25	17 (68.0)	1.22 (1.65)	0.1	0.45	0.87	1.24	7.3		
			Percent change from baseline in UP/C	Week 6	Sparsentan	26	25 (96.2)	-0.52 (0.85)	-2.2	-0.67	-0.50	-0.15	2.3
					Irbesartan	25	23 (92.0)	-0.26 (0.31)	-0.9	-0.45	-0.21	-0.09	0.4
				Week 36	Sparsentan	26	26 (100.0)	-0.73 (0.88)	-2.6	-0.95	-0.57	-0.19	0.6
					Irbesartan	25	23 (92.0)	-0.39 (0.47)	-1.3	-0.62	-0.44	-0.08	0.7
				Week 58	Sparsentan	26	25 (96.2)	-0.64 (1.01)	-2.8	-0.93	-0.36	0.04	0.7
					Irbesartan	25	21 (84.0)	-0.31 (0.44)	-1.2	-0.51	-0.31	-0.03	0.7
				Week 110	Sparsentan	26	21 (80.8)	-0.57 (1.15)	-3.3	-0.71	-0.37	-0.11	1.7
					Irbesartan	25	17 (68.0)	-0.05 (0.66)	-1.1	-0.33	-0.13	0.14	1.8

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
Subgroup: Baseline eGFR Group 2													
< 45 mL/min/1.73 m**2	UP/C	Baseline	Sparsentan	82	82 (100.0)	1.53 (1.05)	0.1	0.78	1.30	1.91	7.0		
			Irbesartan	80	80 (100.0)	1.52 (0.95)	0.5	0.97	1.34	1.77	6.9		
		Week 6	Sparsentan	82	78 (95.1)	1.08 (0.76)	0.1	0.51	0.85	1.33	3.5		
			Irbesartan	80	78 (97.5)	1.42 (0.83)	0.4	0.72	1.16	2.06	4.1		
		Week 36	Sparsentan	82	78 (95.1)	1.08 (0.86)	0.1	0.39	0.80	1.58	4.4		
			Irbesartan	80	72 (90.0)	1.44 (0.92)	0.2	0.58	1.37	1.84	4.0		
		Week 58	Sparsentan	82	77 (93.9)	1.09 (0.98)	0.1	0.47	0.76	1.52	5.9		
			Irbesartan	80	70 (87.5)	1.38 (0.94)	0.1	0.71	1.27	1.88	5.1		
		Week 110	Sparsentan	82	67 (81.7)	1.18 (0.94)	0.1	0.51	0.88	1.67	5.3		
			Irbesartan	80	59 (73.8)	1.52 (1.22)	0.1	0.81	1.25	2.06	7.8		
		Percent change from baseline in UP/C	Percent change from baseline in UP/C	Week 6	Sparsentan	82	78 (95.1)	-0.38 (0.79)	-2.3	-0.80	-0.31	0.05	1.7
					Irbesartan	80	78 (97.5)	-0.09 (0.79)	-4.3	-0.45	-0.12	0.33	1.8
				Week 36	Sparsentan	82	78 (95.1)	-0.37 (0.90)	-3.2	-0.93	-0.36	0.25	1.2
					Irbesartan	80	72 (90.0)	-0.07 (1.03)	-4.3	-0.65	-0.12	0.40	2.6
				Week 58	Sparsentan	82	77 (93.9)	-0.34 (1.07)	-3.0	-0.97	-0.36	0.09	5.7
					Irbesartan	80	70 (87.5)	-0.12 (0.86)	-3.2	-0.61	-0.11	0.36	2.8
				Week 110	Sparsentan	82	67 (81.7)	-0.21 (0.89)	-2.6	-0.72	-0.28	0.38	2.0
					Irbesartan	80	59 (73.8)	0.10 (1.28)	-5.5	-0.49	0.07	0.61	5.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max
45 to < 60 mL/min/1.73 m**2	UP/C	Baseline	Sparsentan	45	45 (100.0)	1.40 (0.83)	0.2	0.77	1.22	1.75	3.6
		Week 6	Irbesartan	49	49 (100.0)	1.51 (0.97)	0.4	0.85	1.25	1.82	4.5
			Sparsentan	45	42 (93.3)	0.92 (0.71)	0.1	0.38	0.75	1.20	2.6
		Week 36	Irbesartan	49	48 (98.0)	1.56 (1.09)	0.4	0.82	1.17	1.95	6.2
			Sparsentan	45	42 (93.3)	0.95 (0.93)	0.1	0.33	0.62	1.08	3.7
		Week 58	Irbesartan	49	47 (95.9)	1.48 (1.21)	0.2	0.73	1.12	1.62	6.6
			Sparsentan	45	43 (95.6)	0.81 (0.54)	0.1	0.39	0.72	1.11	2.4
		Week 110	Irbesartan	49	46 (93.9)	1.56 (1.25)	0.2	0.75	1.08	1.97	5.6
			Sparsentan	45	38 (84.4)	1.17 (1.24)	0.1	0.40	0.86	1.54	6.7
		Percent change from baseline in UP/C	Week 6	Sparsentan	45	42 (93.3)	-0.48 (0.61)	-1.6	-0.94	-0.51	-0.07
	Irbesartan			49	48 (98.0)	0.03 (0.66)	-1.5	-0.28	0.02	0.37	1.9
	Week 36		Sparsentan	45	42 (93.3)	-0.48 (0.74)	-2.2	-0.85	-0.57	-0.15	1.7
			Irbesartan	49	47 (95.9)	-0.01 (0.74)	-1.7	-0.36	-0.05	0.32	2.4
	Week 58		Sparsentan	45	43 (95.6)	-0.56 (0.74)	-2.1	-1.00	-0.44	-0.04	1.1
			Irbesartan	49	46 (93.9)	0.04 (0.81)	-1.1	-0.45	-0.09	0.31	3.5
	Week 110		Sparsentan	45	38 (84.4)	-0.16 (1.11)	-2.9	-0.74	-0.26	0.24	3.5
			Irbesartan	49	37 (75.5)	0.13 (0.98)	-1.8	-0.36	-0.02	0.81	3.6

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	
60 to < 90 mL/min/1.73 m**2	UP/C	Baseline	Sparsentan	49	49 (100.0)	1.25 (0.59)	0.3	0.79	1.22	1.66	2.8	
		Week 6	Irbesartan	48	48 (100.0)	1.34 (0.63)	0.5	0.87	1.20	1.70	3.4	
			Sparsentan	49	48 (98.0)	0.88 (0.78)	0.1	0.32	0.58	1.21	3.5	
		Week 36	Irbesartan	48	45 (93.8)	1.49 (0.84)	0.4	0.84	1.35	1.78	4.2	
			Sparsentan	49	48 (98.0)	0.77 (0.81)	0.1	0.27	0.59	1.04	5.2	
		Week 58	Irbesartan	48	45 (93.8)	1.47 (0.85)	0.3	0.84	1.32	2.07	4.1	
			Sparsentan	49	43 (87.8)	0.83 (0.79)	0.1	0.18	0.77	1.16	3.1	
		Week 110	Irbesartan	48	42 (87.5)	1.50 (0.86)	0.3	0.85	1.39	1.86	3.8	
			Sparsentan	49	42 (85.7)	0.71 (0.64)	0.1	0.21	0.44	1.13	2.9	
		Percent change from baseline in UP/C	Week 6	Sparsentan	49	48 (98.0)	-0.38 (0.69)	-2.3	-0.70	-0.43	-0.15	1.9
				Irbesartan	48	45 (93.8)	0.12 (0.77)	-1.4	-0.34	0.07	0.35	3.7
			Week 36	Sparsentan	49	48 (98.0)	-0.50 (0.77)	-1.8	-0.76	-0.54	-0.26	3.6
				Irbesartan	48	45 (93.8)	0.12 (0.72)	-1.3	-0.27	0.04	0.40	2.2
	Week 58		Sparsentan	49	43 (87.8)	-0.44 (0.67)	-1.8	-0.94	-0.52	-0.17	1.7	
			Irbesartan	48	42 (87.5)	0.14 (0.94)	-2.1	-0.26	-0.02	0.47	3.0	
	Week 110		Sparsentan	49	42 (85.7)	-0.57 (0.66)	-1.7	-1.01	-0.52	-0.19	1.0	
		Irbesartan	48	36 (75.0)	0.20 (1.07)	-1.6	-0.31	0.03	0.62	4.3		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
>= 90 mL/min/1.73 m**2	UP/C	Baseline	Sparsentan	26	26 (100.0)	1.49 (0.98)	0.4	0.79	1.10	2.14	3.6		
			Irbesartan	25	25 (100.0)	1.24 (0.99)	0.2	0.73	1.10	1.46	5.5		
		Week 6	Sparsentan	26	25 (96.2)	0.91 (1.06)	0.2	0.37	0.63	0.93	5.6		
			Irbesartan	25	23 (92.0)	0.96 (1.03)	0.1	0.59	0.79	1.00	5.4		
		Week 36	Sparsentan	26	26 (100.0)	0.76 (0.83)	0.1	0.26	0.45	1.07	4.0		
			Irbesartan	25	23 (92.0)	0.85 (1.22)	0.1	0.24	0.58	0.92	6.2		
		Week 58	Sparsentan	26	25 (96.2)	0.86 (0.85)	0.2	0.29	0.55	0.85	3.5		
			Irbesartan	25	21 (84.0)	0.92 (1.06)	0.1	0.41	0.64	0.99	5.2		
		Week 110	Sparsentan	26	21 (80.8)	0.87 (1.17)	0.0	0.32	0.47	0.87	5.3		
			Irbesartan	25	17 (68.0)	1.22 (1.65)	0.1	0.45	0.87	1.24	7.3		
			Percent change from baseline in UP/C	Week 6	Sparsentan	26	25 (96.2)	-0.52 (0.85)	-2.2	-0.67	-0.50	-0.15	2.3
					Irbesartan	25	23 (92.0)	-0.26 (0.31)	-0.9	-0.45	-0.21	-0.09	0.4
				Week 36	Sparsentan	26	26 (100.0)	-0.73 (0.88)	-2.6	-0.95	-0.57	-0.19	0.6
					Irbesartan	25	23 (92.0)	-0.39 (0.47)	-1.3	-0.62	-0.44	-0.08	0.7
				Week 58	Sparsentan	26	25 (96.2)	-0.64 (1.01)	-2.8	-0.93	-0.36	0.04	0.7
					Irbesartan	25	21 (84.0)	-0.31 (0.44)	-1.2	-0.51	-0.31	-0.03	0.7
				Week 110	Sparsentan	26	21 (80.8)	-0.57 (1.15)	-3.3	-0.71	-0.37	-0.11	1.7
					Irbesartan	25	17 (68.0)	-0.05 (0.66)	-1.1	-0.33	-0.13	0.14	1.8

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024



Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
Subgroup: Baseline urine protein excretion													
<= 1.75 g/day	UP/C	Baseline	Sparsentan	98	98 (100.0)	0.87 (0.41)	0.1	0.60	0.78	1.06	2.8		
			Irbesartan	93	93 (100.0)	0.93 (0.42)	0.2	0.67	0.88	1.09	3.4		
		Week 6	Sparsentan	98	94 (95.9)	0.69 (0.56)	0.1	0.32	0.52	0.86	3.3		
			Irbesartan	93	89 (95.7)	0.95 (0.50)	0.1	0.62	0.81	1.18	2.8		
		Week 36	Sparsentan	98	94 (95.9)	0.65 (0.53)	0.1	0.26	0.44	0.94	2.2		
			Irbesartan	93	87 (93.5)	1.03 (0.77)	0.1	0.46	0.82	1.33	3.3		
		Week 58	Sparsentan	98	93 (94.9)	0.75 (0.78)	0.1	0.22	0.58	0.93	5.9		
			Irbesartan	93	82 (88.2)	1.04 (0.72)	0.1	0.48	0.85	1.43	3.8		
		Week 110	Sparsentan	98	84 (85.7)	0.82 (0.71)	0.1	0.30	0.58	1.25	3.4		
			Irbesartan	93	70 (75.3)	1.12 (0.83)	0.1	0.52	0.86	1.54	5.1		
			Percent change from baseline in UP/C	Week 6	Sparsentan	98	94 (95.9)	-0.18 (0.51)	-1.7	-0.47	-0.24	-0.01	1.9
					Irbesartan	93	89 (95.7)	0.01 (0.51)	-1.4	-0.27	-0.10	0.32	1.8
				Week 36	Sparsentan	98	94 (95.9)	-0.23 (0.56)	-2.3	-0.59	-0.28	-0.05	1.2
					Irbesartan	93	87 (93.5)	0.08 (0.71)	-1.3	-0.35	-0.05	0.35	2.6
Week 58	Sparsentan			98	93 (94.9)	-0.12 (0.83)	-2.2	-0.51	-0.23	0.09	5.7		
	Irbesartan			93	82 (88.2)	0.09 (0.70)	-2.1	-0.29	-0.04	0.46	3.0		
Week 110	Sparsentan			98	84 (85.7)	-0.03 (0.64)	-1.1	-0.44	-0.19	0.15	2.5		
	Irbesartan			93	70 (75.3)	0.21 (0.79)	-1.3	-0.24	0.07	0.57	4.3		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
> 1.75 g/day	UP/C	Baseline	Sparsentan	104	104 (100.0)	1.95 (0.92)	0.7	1.33	1.76	2.28	7.0		
			Irbesartan	109	109 (100.0)	1.88 (0.96)	0.5	1.35	1.59	2.04	6.9		
		Week 6	Sparsentan	104	99 (95.2)	1.24 (0.89)	0.1	0.59	1.03	1.62	5.6		
			Irbesartan	109	105 (96.3)	1.82 (1.03)	0.5	1.05	1.61	2.38	6.2		
		Week 36	Sparsentan	104	100 (96.2)	1.20 (1.03)	0.1	0.50	0.83	1.53	5.2		
			Irbesartan	109	100 (91.7)	1.70 (1.14)	0.2	0.91	1.44	2.33	6.6		
		Week 58	Sparsentan	104	95 (91.3)	1.13 (0.85)	0.1	0.48	0.89	1.61	4.1		
			Irbesartan	109	97 (89.0)	1.71 (1.16)	0.1	0.92	1.48	2.21	5.6		
		Week 110	Sparsentan	104	84 (80.8)	1.23 (1.19)	0.0	0.44	0.88	1.60	6.7		
			Irbesartan	109	79 (72.5)	1.88 (1.44)	0.1	0.95	1.42	2.48	7.8		
		Percent change from baseline in UP/C	Week 6	Sparsentan	104	99 (95.2)	-0.65 (0.84)	-2.3	-1.07	-0.71	-0.10	2.3	
					Irbesartan	109	105 (96.3)	-0.07 (0.86)	-4.3	-0.53	-0.12	0.33	3.7
				Week 36	Sparsentan	104	100 (96.2)	-0.71 (0.98)	-3.2	-1.17	-0.71	-0.24	3.6
					Irbesartan	109	100 (91.7)	-0.16 (0.94)	-4.3	-0.71	-0.16	0.27	2.4
				Week 58	Sparsentan	104	95 (91.3)	-0.77 (0.88)	-3.0	-1.25	-0.82	-0.21	1.4
					Irbesartan	109	97 (89.0)	-0.15 (0.93)	-3.2	-0.74	-0.24	0.23	3.5
				Week 110	Sparsentan	104	84 (80.8)	-0.63 (1.08)	-3.3	-1.27	-0.68	-0.11	3.5
					Irbesartan	109	79 (72.5)	0.03 (1.30)	-5.5	-0.65	-0.06	0.54	5.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
Subgroup: Baseline use of antihypertensives													
Yes	UP/C	Baseline	Sparsentan	90	90 (100.0)	1.60 (1.04)	0.3	0.80	1.39	1.93	7.0		
			Irbesartan	88	88 (100.0)	1.52 (1.08)	0.4	0.86	1.22	1.81	6.9		
		Week 6	Sparsentan	90	84 (93.3)	1.08 (0.96)	0.1	0.47	0.81	1.32	5.6		
			Irbesartan	88	82 (93.2)	1.45 (0.96)	0.4	0.79	1.13	1.75	5.4		
		Week 36	Sparsentan	90	87 (96.7)	0.99 (0.95)	0.1	0.39	0.71	1.32	5.2		
			Irbesartan	88	82 (93.2)	1.49 (1.12)	0.2	0.68	1.17	1.82	6.2		
		Week 58	Sparsentan	90	83 (92.2)	1.01 (0.82)	0.1	0.47	0.71	1.52	4.1		
			Irbesartan	88	80 (90.9)	1.49 (1.15)	0.1	0.76	1.18	1.85	5.5		
		Week 110	Sparsentan	90	74 (82.2)	1.02 (0.89)	0.0	0.33	0.80	1.39	5.3		
			Irbesartan	88	68 (77.3)	1.64 (1.55)	0.1	0.69	1.25	2.01	7.8		
		Percent change from baseline in UP/C		Week 6	Sparsentan	90	84 (93.3)	-0.46 (0.86)	-2.3	-0.92	-0.48	-0.12	2.3
					Irbesartan	88	82 (93.2)	-0.09 (0.84)	-4.3	-0.39	-0.11	0.27	3.7
				Week 36	Sparsentan	90	87 (96.7)	-0.54 (0.92)	-3.2	-1.03	-0.53	-0.20	3.6
					Irbesartan	88	82 (93.2)	-0.04 (0.99)	-4.3	-0.52	-0.06	0.41	2.6
		Week 58	Sparsentan	90	83 (92.2)	-0.50 (0.78)	-3.0	-0.93	-0.44	-0.03	1.4		
			Irbesartan	88	80 (90.9)	-0.03 (0.86)	-2.1	-0.43	-0.10	0.38	3.0		
		Week 110	Sparsentan	90	74 (82.2)	-0.46 (0.89)	-3.3	-0.77	-0.44	0.05	1.2		
			Irbesartan	88	68 (77.3)	0.13 (1.39)	-5.5	-0.56	-0.08	0.68	5.3		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
 Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
No	UP/C	Baseline	Sparsentan	112	112 (100.0)	1.29 (0.74)	0.1	0.74	1.14	1.75	3.6		
			Irbesartan	114	114 (100.0)	1.38 (0.72)	0.2	0.88	1.24	1.66	4.2		
		Week 6	Sparsentan	112	109 (97.3)	0.89 (0.64)	0.1	0.38	0.79	1.15	3.3		
			Irbesartan	114	112 (98.2)	1.40 (0.93)	0.1	0.72	1.15	1.89	6.2		
		Week 36	Sparsentan	112	107 (95.5)	0.88 (0.80)	0.1	0.29	0.58	1.30	3.7		
			Irbesartan	114	105 (92.1)	1.31 (0.97)	0.1	0.59	1.04	1.80	6.6		
		Week 58	Sparsentan	112	105 (93.8)	0.89 (0.85)	0.1	0.27	0.75	1.21	5.9		
			Irbesartan	114	99 (86.8)	1.33 (0.93)	0.1	0.64	1.03	1.86	5.6		
		Week 110	Sparsentan	112	94 (83.9)	1.03 (1.08)	0.1	0.35	0.74	1.31	6.7		
			Irbesartan	114	81 (71.1)	1.43 (0.93)	0.1	0.78	1.22	1.93	4.2		
		Percent change from baseline in UP/C	Percent change from baseline in UP/C	Week 6	Sparsentan	112	109 (97.3)	-0.39 (0.62)	-2.2	-0.69	-0.38	-0.01	1.9
					Irbesartan	114	112 (98.2)	0.01 (0.61)	-1.4	-0.35	-0.11	0.34	1.9
				Week 36	Sparsentan	112	107 (95.5)	-0.42 (0.76)	-2.6	-0.78	-0.39	-0.07	1.7
					Irbesartan	114	105 (92.1)	-0.06 (0.72)	-1.9	-0.54	-0.10	0.27	2.4
				Week 58	Sparsentan	112	105 (93.8)	-0.41 (1.01)	-2.8	-0.95	-0.39	0.02	5.7
					Irbesartan	114	99 (86.8)	-0.05 (0.82)	-3.2	-0.43	-0.07	0.28	3.5
				Week 110	Sparsentan	112	94 (83.9)	-0.23 (0.97)	-2.9	-0.72	-0.29	0.06	3.5
					Irbesartan	114	81 (71.1)	0.10 (0.77)	-1.8	-0.33	0.03	0.48	2.7

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
Subgroup: Time since renal biopsy													
<= 5 years	UP/C	Baseline	Sparsentan	113	113 (100.0)	1.41 (0.87)	0.1	0.78	1.14	1.82	4.2		
			Irbesartan	127	127 (100.0)	1.45 (0.97)	0.2	0.88	1.17	1.72	6.9		
		Week 6	Sparsentan	113	106 (93.8)	0.99 (0.89)	0.1	0.38	0.78	1.30	5.6		
			Irbesartan	127	124 (97.6)	1.36 (0.96)	0.1	0.72	1.06	1.67	6.2		
		Week 36	Sparsentan	113	107 (94.7)	0.92 (0.92)	0.1	0.28	0.61	1.30	5.2		
			Irbesartan	127	117 (92.1)	1.36 (1.15)	0.1	0.57	1.02	1.80	6.6		
		Week 58	Sparsentan	113	104 (92.0)	1.00 (0.92)	0.1	0.33	0.74	1.52	5.9		
			Irbesartan	127	117 (92.1)	1.33 (1.14)	0.1	0.64	0.93	1.67	5.6		
		Week 110	Sparsentan	113	96 (85.0)	0.86 (0.72)	0.0	0.34	0.67	1.25	3.4		
			Irbesartan	127	93 (73.2)	1.42 (1.23)	0.1	0.65	1.16	1.80	7.3		
		Percent change from baseline in UP/C	Week 6	Sparsentan	113	106 (93.8)	-0.42 (0.79)	-2.3	-0.81	-0.44	-0.07	2.3	
					Irbesartan	127	124 (97.6)	-0.09 (0.69)	-4.3	-0.37	-0.11	0.24	1.9
				Week 36	Sparsentan	113	107 (94.7)	-0.51 (0.88)	-3.2	-0.84	-0.52	-0.14	3.6
					Irbesartan	127	117 (92.1)	-0.09 (0.85)	-4.3	-0.50	-0.12	0.27	2.4
				Week 58	Sparsentan	113	104 (92.0)	-0.39 (1.02)	-3.0	-0.80	-0.49	0.05	5.7
					Irbesartan	127	117 (92.1)	-0.11 (0.90)	-3.2	-0.48	-0.19	0.20	3.5
				Week 110	Sparsentan	113	96 (85.0)	-0.46 (0.91)	-3.3	-0.76	-0.45	-0.04	2.0
					Irbesartan	127	93 (73.2)	-0.00 (1.11)	-5.5	-0.56	-0.07	0.46	4.3

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	
> 5 years	UP/C	Baseline	Sparsentan	89	89 (100.0)	1.45 (0.94)	0.3	0.80	1.32	1.81	7.0	
			Irbesartan	75	75 (100.0)	1.42 (0.76)	0.4	0.88	1.30	1.77	4.5	
		Week 6	Sparsentan	89	87 (97.8)	0.96 (0.67)	0.1	0.49	0.81	1.20	3.3	
			Irbesartan	75	70 (93.3)	1.52 (0.89)	0.3	0.92	1.30	2.05	4.2	
		Week 36	Sparsentan	89	87 (97.8)	0.95 (0.80)	0.1	0.38	0.70	1.32	4.4	
			Irbesartan	75	70 (93.3)	1.43 (0.83)	0.2	0.79	1.34	1.84	3.5	
		Week 58	Sparsentan	89	84 (94.4)	0.87 (0.73)	0.1	0.29	0.72	1.18	4.1	
			Irbesartan	75	62 (82.7)	1.53 (0.79)	0.2	0.94	1.53	1.94	4.0	
		Week 110	Sparsentan	89	72 (80.9)	1.23 (1.25)	0.1	0.41	0.87	1.61	6.7	
			Irbesartan	75	56 (74.7)	1.70 (1.28)	0.2	0.87	1.36	2.18	7.8	
		Percent change from baseline in UP/C	Week 6	Sparsentan	89	87 (97.8)	-0.43 (0.66)	-2.2	-0.73	-0.36	-0.03	1.9
				Irbesartan	75	70 (93.3)	0.08 (0.76)	-1.5	-0.39	-0.08	0.45	3.7
	Week 36		Sparsentan	89	87 (97.8)	-0.44 (0.78)	-2.6	-0.93	-0.42	0.03	1.7	
			Irbesartan	75	70 (93.3)	0.01 (0.85)	-1.6	-0.54	-0.02	0.35	2.6	
	Week 58		Sparsentan	89	84 (94.4)	-0.52 (0.76)	-2.8	-1.00	-0.37	-0.02	1.2	
			Irbesartan	75	62 (82.7)	0.10 (0.69)	-1.5	-0.29	0.11	0.47	2.4	
	Week 110	Sparsentan	89	72 (80.9)	-0.16 (0.96)	-2.4	-0.68	-0.21	0.23	3.5		
		Irbesartan	75	56 (74.7)	0.31 (1.05)	-1.6	-0.23	0.18	0.84	5.3		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
 Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
Subgroup: History of hypertension													
Yes	UP/C	Baseline	Sparsentan	155	155 (100.0)	1.42 (0.82)	0.1	0.78	1.25	1.86	4.2		
			Irbesartan	161	161 (100.0)	1.47 (0.93)	0.2	0.89	1.25	1.77	6.9		
		Week 6	Sparsentan	155	147 (94.8)	1.01 (0.85)	0.1	0.44	0.82	1.27	5.6		
			Irbesartan	161	153 (95.0)	1.43 (0.95)	0.1	0.76	1.16	1.80	6.2		
		Week 36	Sparsentan	155	149 (96.1)	0.99 (0.89)	0.1	0.39	0.71	1.35	5.2		
			Irbesartan	161	150 (93.2)	1.45 (1.09)	0.1	0.65	1.22	2.02	6.6		
		Week 58	Sparsentan	155	145 (93.5)	1.01 (0.88)	0.1	0.39	0.76	1.48	5.9		
			Irbesartan	161	145 (90.1)	1.47 (1.09)	0.1	0.72	1.12	1.94	5.6		
		Week 110	Sparsentan	155	130 (83.9)	1.05 (0.95)	0.0	0.39	0.78	1.36	5.3		
			Irbesartan	161	121 (75.2)	1.57 (1.31)	0.1	0.79	1.28	2.05	7.8		
		Percent change from baseline in UP/C		Week 6	Sparsentan	155	147 (94.8)	-0.40 (0.79)	-2.3	-0.77	-0.38	-0.00	2.3
					Irbesartan	161	153 (95.0)	-0.05 (0.76)	-4.3	-0.39	-0.11	0.27	3.7
				Week 36	Sparsentan	155	149 (96.1)	-0.44 (0.86)	-3.2	-0.84	-0.47	-0.10	3.6
					Irbesartan	161	150 (93.2)	-0.03 (0.88)	-4.3	-0.53	-0.11	0.39	2.6
		Week 58	Sparsentan	155	145 (93.5)	-0.39 (0.95)	-3.0	-0.88	-0.36	-0.01	5.7		
			Irbesartan	161	145 (90.1)	-0.00 (0.85)	-2.1	-0.45	-0.09	0.31	3.5		
		Week 110	Sparsentan	155	130 (83.9)	-0.34 (0.93)	-3.3	-0.76	-0.38	0.05	2.5		
			Irbesartan	161	121 (75.2)	0.11 (1.16)	-5.5	-0.50	-0.01	0.57	5.3		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
 Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024

Table PT2RPR\_FSHM: Course of UP/C (percentage) by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max		
No	UP/C	Baseline	Sparsentan	47	47 (100.0)	1.46 (1.13)	0.2	0.74	1.26	1.82	7.0		
			Irbesartan	41	41 (100.0)	1.34 (0.73)	0.4	0.85	1.15	1.56	3.8		
		Week 6	Sparsentan	47	46 (97.9)	0.84 (0.59)	0.1	0.37	0.71	1.15	2.6		
			Irbesartan	41	41 (100.0)	1.39 (0.91)	0.3	0.79	1.09	1.68	3.6		
		Week 36	Sparsentan	47	45 (95.7)	0.75 (0.76)	0.1	0.25	0.49	1.00	3.4		
			Irbesartan	41	37 (90.2)	1.13 (0.77)	0.2	0.56	0.99	1.33	3.5		
		Week 58	Sparsentan	47	43 (91.5)	0.71 (0.65)	0.1	0.18	0.58	0.95	2.9		
			Irbesartan	41	34 (82.9)	1.12 (0.67)	0.3	0.58	0.95	1.60	3.0		
		Week 110	Sparsentan	47	38 (80.9)	0.93 (1.16)	0.1	0.25	0.64	1.22	6.7		
			Irbesartan	41	28 (68.3)	1.34 (0.95)	0.3	0.62	1.12	1.82	4.2		
		Percent change from baseline in UP/C	Percent change from baseline in UP/C	Week 6	Sparsentan	47	46 (97.9)	-0.50 (0.52)	-2.1	-0.73	-0.44	-0.18	0.4
					Irbesartan	41	41 (100.0)	0.05 (0.55)	-0.9	-0.30	-0.07	0.39	1.2
				Week 36	Sparsentan	47	45 (95.7)	-0.59 (0.76)	-2.6	-0.93	-0.54	-0.17	1.0
					Irbesartan	41	37 (90.2)	-0.14 (0.67)	-1.9	-0.44	-0.08	0.15	1.8
				Week 58	Sparsentan	47	43 (91.5)	-0.64 (0.78)	-2.8	-1.08	-0.54	-0.17	0.7
					Irbesartan	41	34 (82.9)	-0.19 (0.78)	-3.2	-0.37	-0.08	0.25	1.1
				Week 110	Sparsentan	47	38 (80.9)	-0.31 (0.98)	-2.4	-0.72	-0.36	0.06	3.5
					Irbesartan	41	28 (68.3)	0.12 (0.76)	-1.6	-0.33	0.03	0.41	2.7

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
Only key visits up to Week 110 are displayed. An increase reflects a worsening of the status of the patient.  
Results are presented in g/g.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

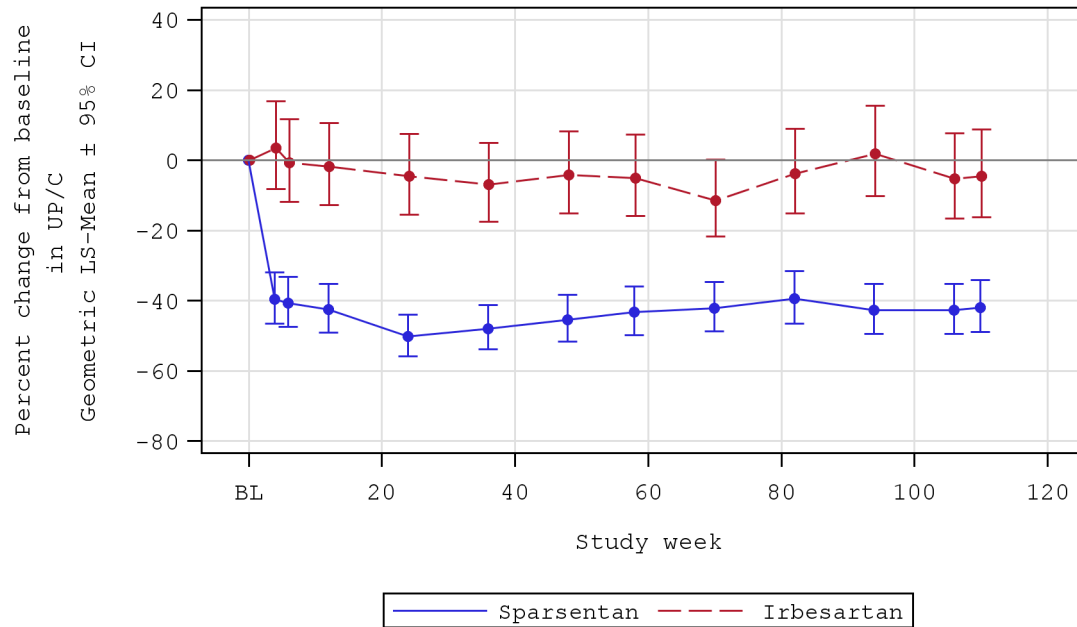
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Source Data: alb, created on: 19FEB2024



Figure PF2RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Sex: Male



Sparsentan	134	135	129	116
Irbesartan	138	132	127	107

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data.

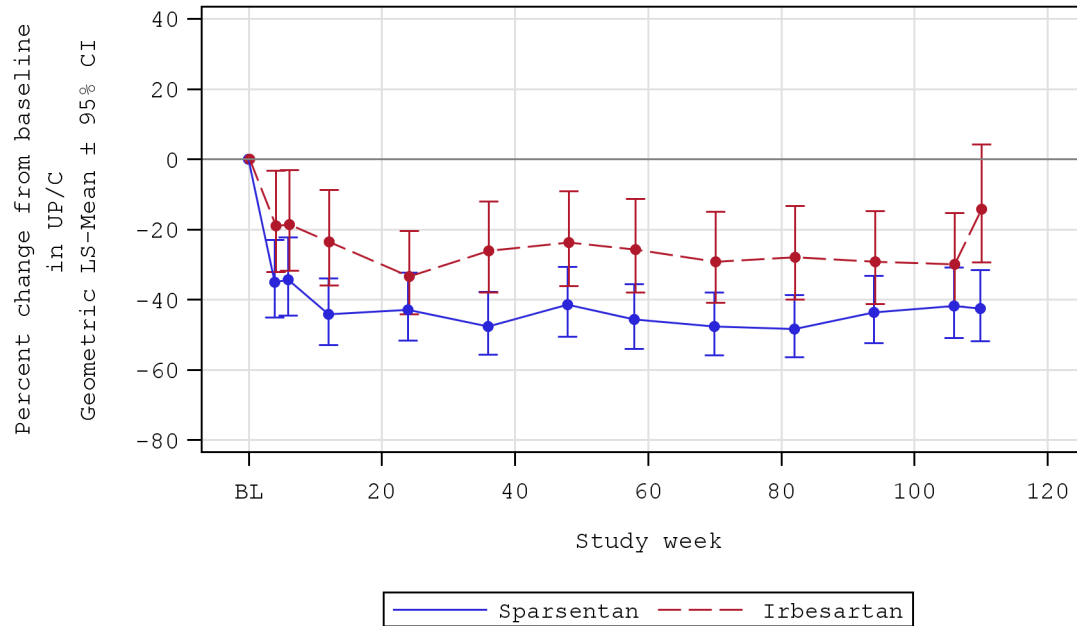
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Reference table: PT2RPR\_FSLM

Figure PF2RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Sex: Female



Sparsentan	59	59	59	52
Irbesartan	56	55	52	42

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data.

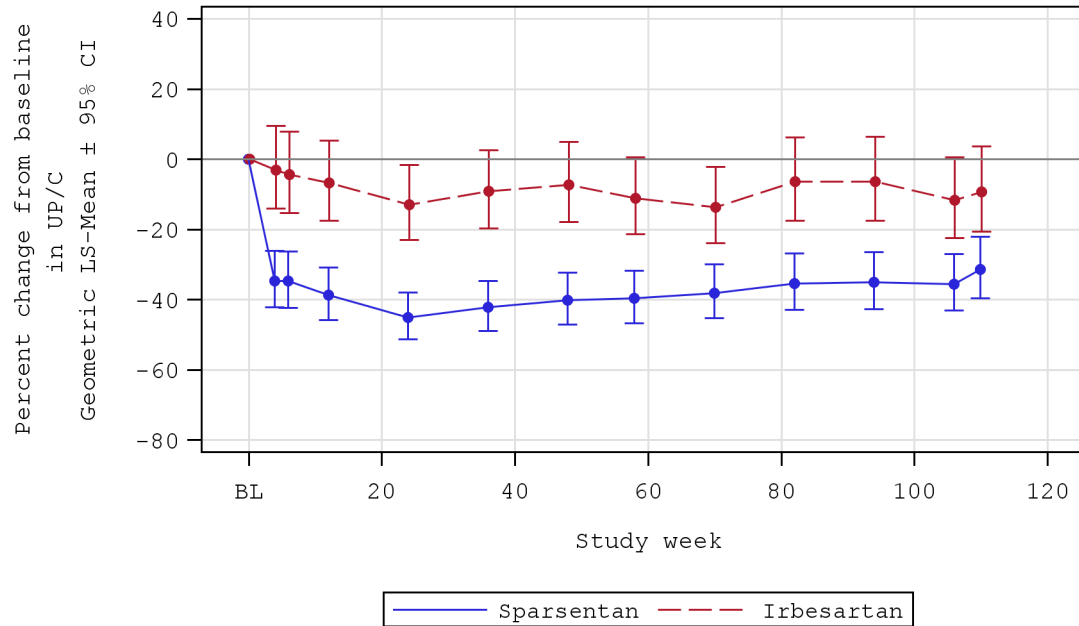
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Reference table: PT2RPR\_FSLM

Figure PF2RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1: < 60 mL/min/1.73 m\*\*2



Sparsentan	120	120	120	105
Irbesartan	126	119	116	96

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data.

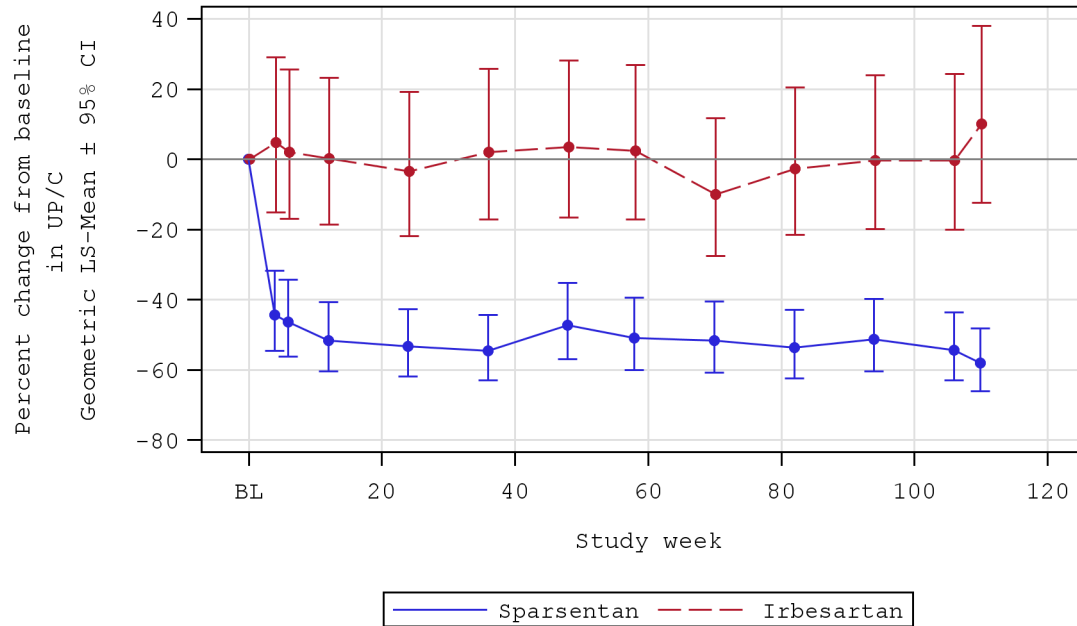
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Reference table: PT2RPR\_FSLM

Figure PF2RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1: 60 to < 90 mL/min/1.73 m\*\*2



Sparsentan	48	48	43	42
Irbesartan	45	45	42	36

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data.

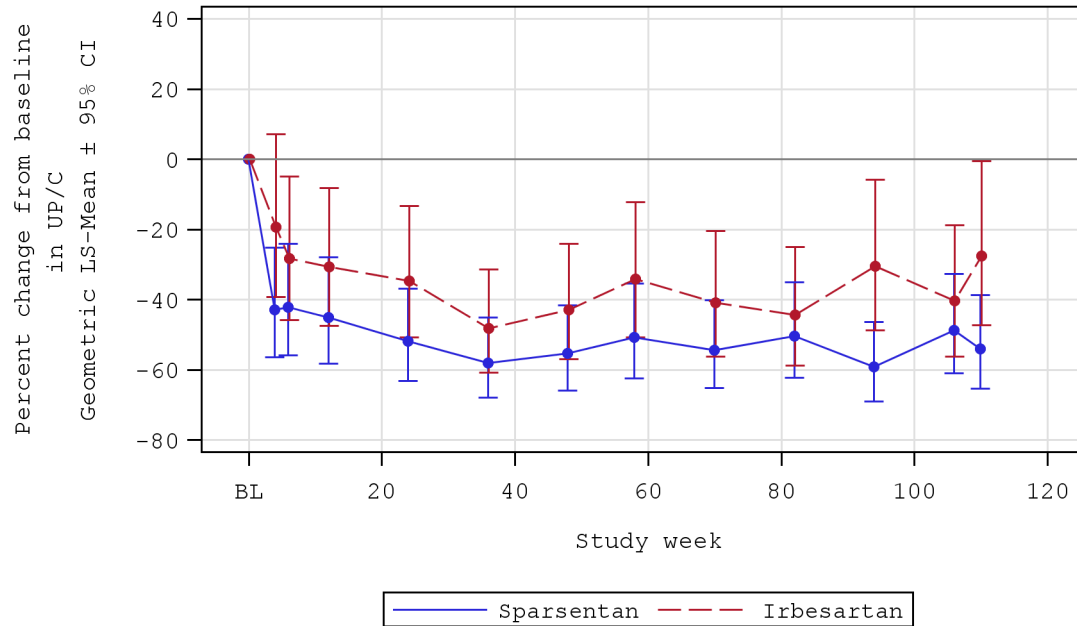
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Reference table: PT2RPR\_FSLM

Figure PF2RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1:  $\geq 90$  mL/min/1.73 m<sup>2</sup>



Sparsentan	25	26	25	21
Irbesartan	23	23	21	17

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data.

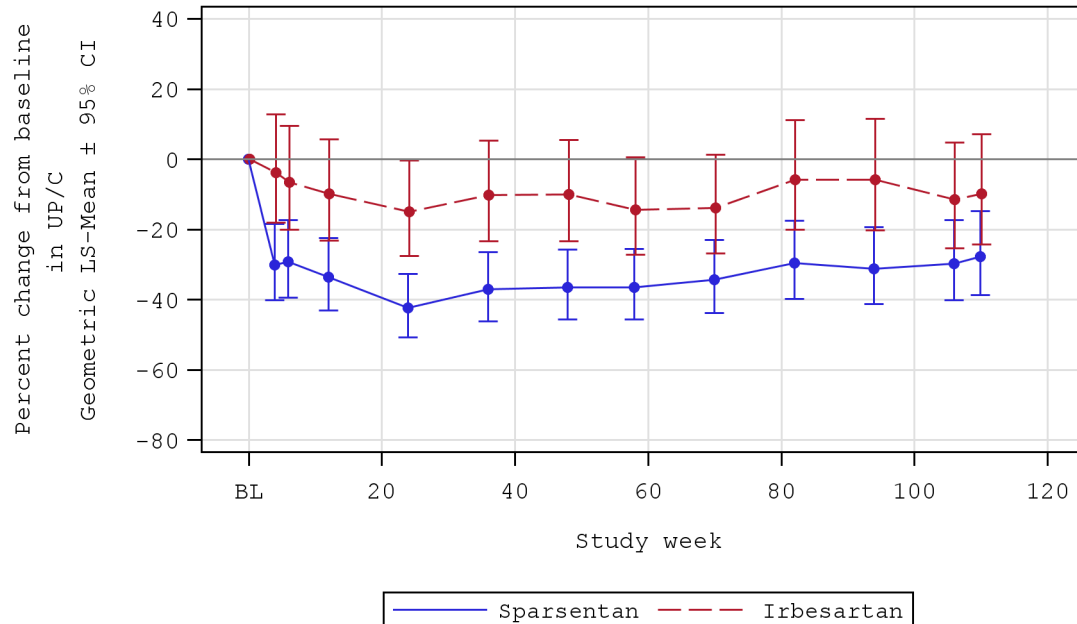
eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High  $\geq 60$  ml/min/1.73m<sup>2</sup>, UP Low =  $\leq 1.75$  g/day, UP High =  $> 1.75$  g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Reference table: PT2RPR\_FSLM

Figure PF2RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: < 45 mL/min/1.73 m\*\*2



Sparsentan	78	78	77	67
Irbesartan	78	72	70	59

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data.

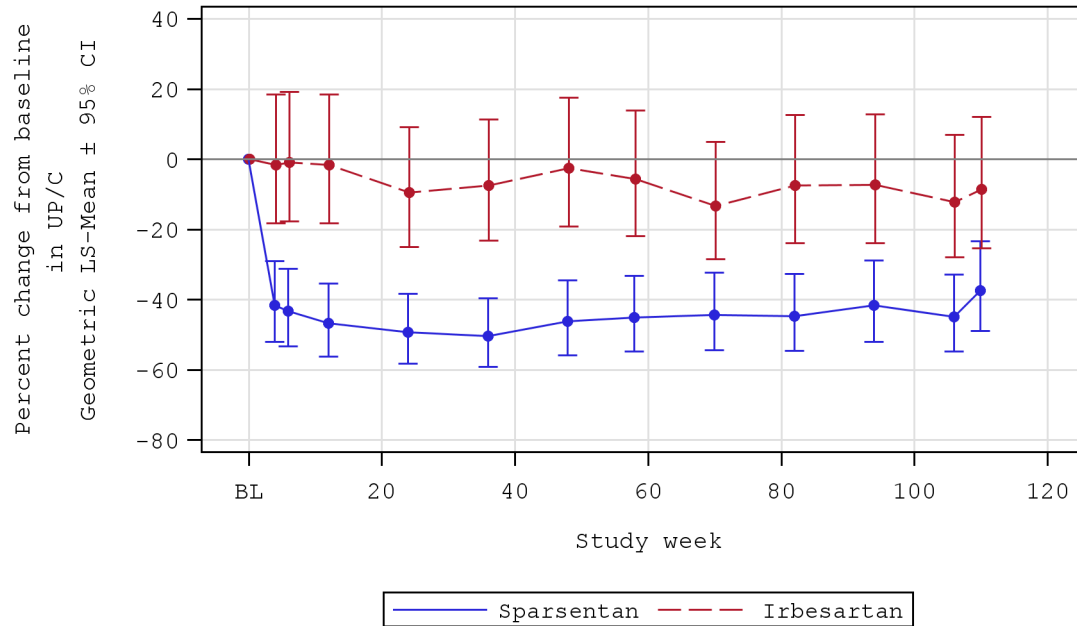
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Reference table: PT2RPR\_FSLM

Figure PF2RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: 45 to < 60 mL/min/1.73 m\*\*2



Sparsentan	42	42	43	38
Irbesartan	48	47	46	37

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data.

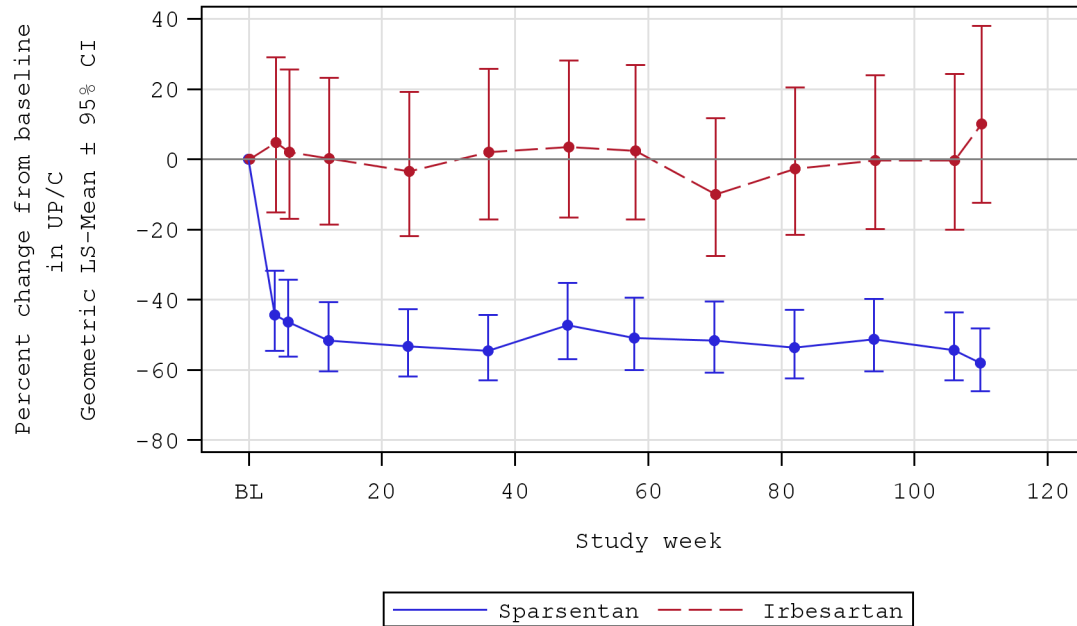
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Reference table: PT2RPR\_FSLM

Figure PF2RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: 60 to < 90 mL/min/1.73 m\*\*2



Sparsentan	48	48	43	42
Irbesartan	45	45	42	36

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

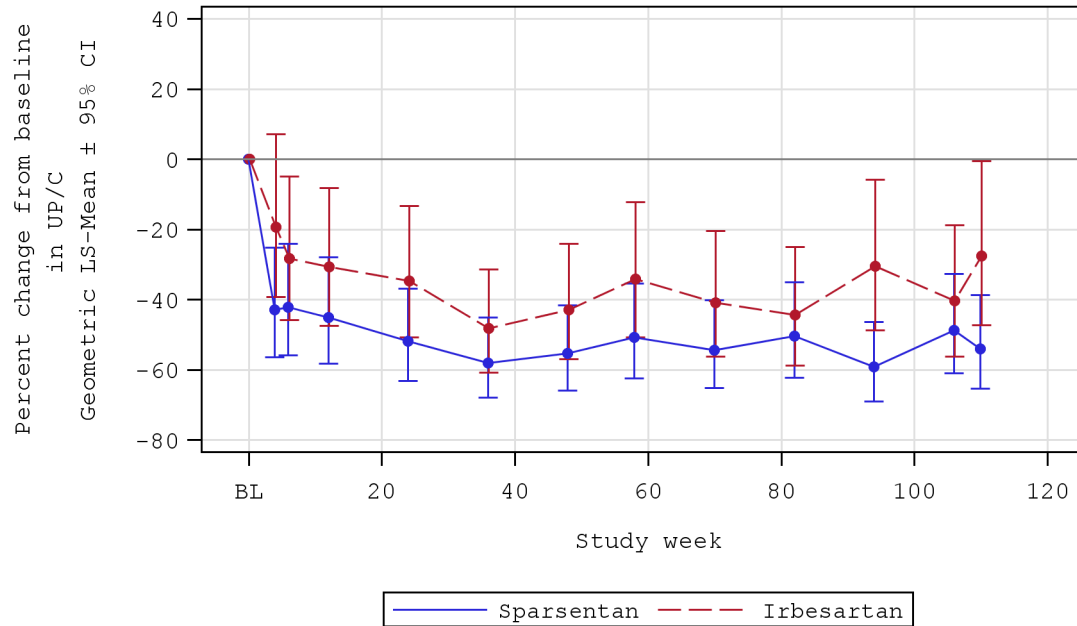
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Reference table: PT2RPR\_FSLM



Figure PF2RPR\_FSGM: Percent change from baseline in UP/C by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2:  $\geq 90$  mL/min/1.73 m<sup>2</sup>



Sparsentan	25	26	25	21
Irbesartan	23	23	21	17

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/g. Analysis was performed on log-transformed data.

eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High  $\geq 60$  ml/min/1.73m<sup>2</sup>, UP Low =  $\leq 1.75$  g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

UP/C = urine protein/creatinine ratio.

Reference table: PT2RPR\_FSLM

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Sex													
Male	Urine protein excretion	Baseline	Sparsentan	139	139 (100.0)	2.29 (1.38)	0.1	1.21	2.07	3.14	7.2		
			Irbesartan	143	143 (100.0)	2.25 (1.33)	0.5	1.34	1.85	2.85	7.5		
		Week 6	Sparsentan	139	135 (97.1)	1.59 (1.33)	0.2	0.64	1.17	2.03	6.5		
			Irbesartan	143	137 (95.8)	2.37 (1.70)	0.4	1.25	1.91	3.07	10.3		
		Week 36	Sparsentan	139	135 (97.1)	1.54 (1.60)	0.1	0.55	0.96	2.03	9.4		
			Irbesartan	143	132 (92.3)	2.33 (1.81)	0.2	1.01	1.89	3.09	9.2		
		Week 58	Sparsentan	139	129 (92.8)	1.60 (1.56)	0.1	0.57	1.15	1.99	7.5		
			Irbesartan	143	127 (88.8)	2.34 (1.85)	0.1	1.08	1.80	3.06	12.8		
		Week 110	Sparsentan	139	116 (83.5)	1.52 (1.36)	0.1	0.51	1.13	2.14	6.0		
			Irbesartan	143	108 (75.5)	2.49 (2.00)	0.1	1.11	1.94	3.29	9.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	139	135 (97.1)	-0.72 (1.20)	-4.2	-1.41	-0.71	-0.09	5.2	-0.64 [-0.88, -0.39]
				Irbesartan	143	137 (95.8)	0.11 (1.39)	-3.7	-0.57	-0.10	0.72	8.4	
	Week 36		Sparsentan	139	135 (97.1)	-0.76 (1.34)	-4.7	-1.54	-0.77	-0.11	4.6	-0.56 [-0.81, -0.32]	
			Irbesartan	143	132 (92.3)	0.06 (1.57)	-3.3	-0.78	-0.15	0.76	6.5		
	Week 58		Sparsentan	139	129 (92.8)	-0.67 (1.49)	-4.4	-1.45	-0.72	0.04	6.4	-0.49 [-0.74, -0.24]	
			Irbesartan	143	127 (88.8)	0.08 (1.57)	-3.3	-0.74	-0.17	0.58	6.7		
Week 110	Sparsentan	139	116 (83.5)	-0.69 (1.46)	-7.1	-1.46	-0.71	-0.00	3.5	-0.54 [-0.80, -0.27]			
	Irbesartan	143	108 (75.5)	0.18 (1.79)	-6.0	-0.67	0.02	1.13	5.9				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Female	Urine protein excretion	Baseline	Sparsentan	63	63 (100.0)	2.09 (2.14)	0.5	1.15	1.45	2.32	14.7		
			Irbesartan	59	59 (100.0)	1.95 (0.97)	0.6	1.33	1.77	2.33	5.7		
		Week 6	Sparsentan	63	60 (95.2)	1.28 (0.83)	0.2	0.69	1.03	1.72	3.9		
			Irbesartan	59	56 (94.9)	1.72 (1.20)	0.4	0.93	1.37	2.26	7.6		
		Week 36	Sparsentan	63	59 (93.7)	1.06 (0.79)	0.1	0.53	0.81	1.35	3.8		
			Irbesartan	59	55 (93.2)	1.75 (1.43)	0.2	0.92	1.22	2.44	7.1		
		Week 58	Sparsentan	63	59 (93.7)	1.05 (0.71)	0.1	0.57	0.91	1.40	3.4		
			Irbesartan	59	51 (86.4)	1.62 (1.56)	0.2	0.69	1.22	1.97	9.4		
		Week 110	Sparsentan	63	53 (84.1)	1.26 (1.21)	0.1	0.42	0.96	1.56	6.0		
			Irbesartan	59	42 (71.2)	1.75 (1.11)	0.3	0.94	1.60	2.38	4.8		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	63	60 (95.2)	-0.67 (1.84)	-13.0	-0.89	-0.45	-0.01	1.8	-0.28 [-0.64, 0.09]
				Irbesartan	59	56 (94.9)	-0.26 (0.84)	-1.8	-0.82	-0.20	0.06	3.0	
			Week 36	Sparsentan	63	59 (93.7)	-0.91 (1.95)	-13.7	-1.07	-0.66	-0.20	1.7	-0.46 [-0.83, -0.09]
				Irbesartan	59	55 (93.2)	-0.19 (1.02)	-2.0	-0.91	-0.28	0.39	2.6	
			Week 58	Sparsentan	63	59 (93.7)	-0.88 (1.95)	-13.3	-1.29	-0.71	-0.03	1.3	-0.38 [-0.76, -0.00]
				Irbesartan	59	51 (86.4)	-0.28 (0.97)	-2.1	-0.90	-0.34	0.16	3.6	
			Week 110	Sparsentan	63	53 (84.1)	-0.68 (2.21)	-13.5	-1.29	-0.34	0.32	3.2	-0.34 [-0.75, 0.07]
				Irbesartan	59	42 (71.2)	-0.08 (1.02)	-2.4	-0.69	-0.05	0.36	2.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Age													
<= 45 years	Urine protein excretion	Baseline	Sparsentan	96	96 (100.0)	2.39 (2.00)	0.4	1.23	1.81	2.91	14.7		
			Irbesartan	99	99 (100.0)	2.18 (1.06)	0.5	1.41	1.92	2.70	5.9		
		Week 6	Sparsentan	96	93 (96.9)	1.45 (1.17)	0.2	0.60	1.08	1.95	6.2		
			Irbesartan	99	95 (96.0)	2.18 (1.63)	0.4	1.08	1.84	2.67	10.3		
		Week 36	Sparsentan	96	91 (94.8)	1.43 (1.40)	0.1	0.38	0.91	2.14	8.3		
			Irbesartan	99	91 (91.9)	2.03 (1.68)	0.2	0.81	1.44	2.92	9.2		
		Week 58	Sparsentan	96	89 (92.7)	1.54 (1.44)	0.1	0.59	1.09	2.03	7.5		
			Irbesartan	99	84 (84.8)	2.01 (1.68)	0.1	0.90	1.50	2.45	8.6		
		Week 110	Sparsentan	96	76 (79.2)	1.47 (1.30)	0.1	0.56	1.11	2.08	6.0		
			Irbesartan	99	69 (69.7)	2.23 (1.89)	0.1	0.94	1.72	2.75	8.8		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	96	93 (96.9)	-0.88 (1.68)	-13.0	-1.22	-0.71	-0.03	2.1	-0.56 [-0.85, -0.27]
				Irbesartan	99	95 (96.0)	-0.02 (1.39)	-3.2	-0.70	-0.20	0.44	8.4	
			Week 36	Sparsentan	96	91 (94.8)	-0.89 (1.86)	-13.7	-1.45	-0.66	0.01	2.6	-0.45 [-0.75, -0.16]
				Irbesartan	99	91 (91.9)	-0.13 (1.47)	-3.3	-1.10	-0.29	0.62	6.5	
	Week 58		Sparsentan	96	89 (92.7)	-0.79 (1.93)	-13.3	-1.36	-0.59	0.21	3.3	-0.37 [-0.67, -0.07]	
			Irbesartan	99	84 (84.8)	-0.14 (1.58)	-3.3	-0.92	-0.43	0.38	6.7		
	Week 110	Sparsentan	96	76 (79.2)	-0.75 (2.13)	-13.5	-1.40	-0.55	0.25	3.5	-0.43 [-0.76, -0.10]		
		Irbesartan	99	69 (69.7)	0.09 (1.76)	-3.6	-0.74	-0.08	0.93	5.9			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 45 years	Urine protein excretion	Baseline	Sparsentan	106	106 (100.0)	2.09 (1.24)	0.1	1.12	1.76	2.83	6.3	
			Irbesartan	103	103 (100.0)	2.16 (1.40)	0.6	1.25	1.77	2.43	7.5	
		Week 6	Sparsentan	106	102 (96.2)	1.53 (1.24)	0.2	0.73	1.09	2.01	6.5	
			Irbesartan	103	98 (95.1)	2.19 (1.57)	0.4	1.13	1.70	2.74	9.5	
		Week 36	Sparsentan	106	103 (97.2)	1.37 (1.44)	0.1	0.59	0.91	1.54	9.4	
			Irbesartan	103	96 (93.2)	2.28 (1.77)	0.2	1.07	1.71	2.68	9.2	
		Week 58	Sparsentan	106	99 (93.4)	1.33 (1.32)	0.1	0.49	0.94	1.58	6.8	
			Irbesartan	103	94 (91.3)	2.24 (1.89)	0.1	1.10	1.78	2.91	12.8	
		Week 110	Sparsentan	106	93 (87.7)	1.41 (1.34)	0.1	0.43	0.97	1.78	6.0	
	Irbesartan		103	81 (78.6)	2.33 (1.77)	0.1	1.16	1.78	2.87	9.2		
	Change from baseline in urine protein excretion	Week 6	Sparsentan	106	102 (96.2)	-0.54 (1.13)	-3.8	-1.05	-0.56	-0.10	5.2	-0.50 [-0.78, -0.21]
			Irbesartan	103	98 (95.1)	0.02 (1.13)	-3.7	-0.56	-0.08	0.61	4.2	
		Week 36	Sparsentan	106	103 (97.2)	-0.73 (1.23)	-4.2	-1.36	-0.76	-0.27	4.6	-0.64 [-0.92, -0.35]
			Irbesartan	103	96 (93.2)	0.10 (1.39)	-2.5	-0.63	-0.13	0.62	5.0	
		Week 58	Sparsentan	106	99 (93.4)	-0.69 (1.35)	-4.4	-1.44	-0.74	-0.08	6.4	-0.58 [-0.87, -0.29]
			Irbesartan	103	94 (91.3)	0.08 (1.29)	-2.1	-0.63	-0.10	0.48	5.8	
		Week 110	Sparsentan	106	93 (87.7)	-0.64 (1.30)	-4.6	-1.34	-0.65	-0.06	3.4	-0.55 [-0.85, -0.24]
			Irbesartan	103	81 (78.6)	0.13 (1.49)	-6.0	-0.59	0.00	0.95	3.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age at IgAN diagnosis														
<= 18 years	Urine protein excretion	Baseline	Sparsentan	9	9 (100.0)	3.85 (2.41)	1.2	2.78	3.07	4.00	9.7			
			Irbesartan	5	5 (100.0)	1.91 (1.00)	0.6	1.22	2.14	2.56	3.1			
		Week 6	Sparsentan	9	8 (88.9)	2.24 (1.77)	0.4	0.81	1.66	3.84	4.9			
			Irbesartan	5	5 (100.0)	1.91 (1.46)	0.6	1.18	1.25	2.27	4.3			
		Week 36	Sparsentan	9	8 (88.9)	2.28 (1.55)	0.5	1.07	1.76	3.74	4.6			
			Irbesartan	5	5 (100.0)	1.99 (1.26)	0.6	1.11	1.65	2.94	3.6			
		Week 58	Sparsentan	9	7 (77.8)	2.42 (2.13)	0.1	0.87	1.55	4.99	5.6			
			Irbesartan	5	4 (80.0)	1.20 (0.76)	0.6	0.73	0.98	1.67	2.3			
		Week 110	Sparsentan	9	5 (55.6)	3.20 (2.45)	0.1	1.58	3.08	5.26	6.0			
			Irbesartan	5	4 (80.0)	1.48 (0.94)	0.4	0.83	1.44	2.14	2.7			
			Change from baseline in urine protein excretion	Week 6	Sparsentan	9	8 (88.9)	-0.87 (1.57)	-2.5	-2.18	-1.03	-0.11	2.1	-0.59 [-1.73, 0.55]
					Irbesartan	5	5 (100.0)	-0.00 (1.28)	-1.9	-0.02	0.03	0.13	1.7	
			Week 36	Sparsentan	9	8 (88.9)	-0.84 (1.43)	-3.1	-2.00	-0.50	0.26	0.9	-0.68 [-1.83, 0.47]	
				Irbesartan	5	5 (100.0)	0.08 (1.20)	-1.9	0.03	0.44	0.80	1.1		
		Week 58	Sparsentan	9	7 (77.8)	-0.75 (1.80)	-3.7	-1.52	-1.10	0.99	1.7	-0.12 [-1.35, 1.11]		
			Irbesartan	5	4 (80.0)	-0.55 (1.31)	-2.5	-1.33	-0.01	0.23	0.3			
		Week 110	Sparsentan	9	5 (55.6)	0.14 (1.34)	-1.5	-1.09	0.68	1.26	1.3	0.68 [-0.67, 2.03]		
			Irbesartan	5	4 (80.0)	-0.76 (1.30)	-2.7	-1.59	-0.24	0.07	0.1			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 18 to 40 years	Urine protein excretion	Baseline	Sparsentan	102	102 (100.0)	2.26 (1.79)	0.4	1.21	1.81	2.81	14.7	
			Irbesartan	109	109 (100.0)	2.21 (1.08)	0.5	1.42	1.96	2.70	5.9	
		Week 6	Sparsentan	102	100 (98.0)	1.40 (1.12)	0.2	0.60	1.07	1.80	6.2	
			Irbesartan	109	105 (96.3)	2.15 (1.48)	0.4	1.17	1.82	2.61	7.7	
		Week 36	Sparsentan	102	97 (95.1)	1.41 (1.53)	0.1	0.51	0.90	1.98	9.4	
			Irbesartan	109	100 (91.7)	2.03 (1.58)	0.2	1.00	1.59	2.57	9.2	
		Week 58	Sparsentan	102	95 (93.1)	1.42 (1.41)	0.1	0.59	0.94	1.67	7.5	
			Irbesartan	109	92 (84.4)	2.00 (1.44)	0.1	0.95	1.69	2.49	6.8	
		Week 110	Sparsentan	102	82 (80.4)	1.44 (1.29)	0.1	0.59	0.99	1.99	5.7	
			Irbesartan	109	79 (72.5)	2.26 (1.81)	0.1	1.00	1.79	2.69	8.8	
	Change from baseline in urine protein excretion	Week 6	Sparsentan	102	100 (98.0)	-0.88 (1.59)	-13.0	-1.21	-0.69	-0.14	1.8	-0.58 [-0.86, -0.30]
			Irbesartan	109	105 (96.3)	-0.08 (1.17)	-3.2	-0.68	-0.20	0.44	4.2	
		Week 36	Sparsentan	102	97 (95.1)	-0.86 (1.85)	-13.7	-1.49	-0.73	-0.09	4.6	-0.40 [-0.68, -0.12]
			Irbesartan	109	100 (91.7)	-0.20 (1.44)	-3.3	-1.15	-0.30	0.41	6.5	
		Week 58	Sparsentan	102	95 (93.1)	-0.85 (1.84)	-13.3	-1.45	-0.71	-0.03	3.3	-0.39 [-0.68, -0.10]
			Irbesartan	109	92 (84.4)	-0.22 (1.38)	-3.3	-0.96	-0.41	0.39	5.6	
		Week 110	Sparsentan	102	82 (80.4)	-0.78 (2.10)	-13.5	-1.57	-0.61	0.18	3.5	-0.45 [-0.76, -0.14]
			Irbesartan	109	79 (72.5)	0.07 (1.65)	-3.6	-0.80	-0.08	0.97	5.9	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	Urine protein excretion	Baseline	Sparsentan	91	91 (100.0)	2.03 (1.29)	0.1	1.06	1.54	2.70	6.3	
			Irbesartan	88	88 (100.0)	2.12 (1.43)	0.6	1.27	1.68	2.37	7.5	
		Week 6	Sparsentan	91	87 (95.6)	1.53 (1.22)	0.2	0.72	1.17	2.08	6.5	
			Irbesartan	88	83 (94.3)	2.24 (1.75)	0.4	1.03	1.79	2.82	10.3	
		Week 36	Sparsentan	91	89 (97.8)	1.30 (1.26)	0.1	0.56	0.91	1.47	6.4	
			Irbesartan	88	82 (93.2)	2.33 (1.91)	0.2	0.96	1.60	3.38	9.2	
		Week 58	Sparsentan	91	86 (94.5)	1.36 (1.25)	0.1	0.49	1.09	1.62	6.6	
			Irbesartan	88	82 (93.2)	2.33 (2.14)	0.1	1.01	1.70	2.95	12.8	
		Week 110	Sparsentan	91	82 (90.1)	1.33 (1.20)	0.1	0.43	0.96	1.78	6.0	
	Irbesartan		88	67 (76.1)	2.36 (1.88)	0.1	1.10	1.77	3.28	9.2		
	Change from baseline in urine protein excretion	Week 6	Sparsentan	91	87 (95.6)	-0.48 (1.17)	-3.8	-1.03	-0.51	0.01	5.2	-0.45 [-0.76, -0.15]
			Irbesartan	88	83 (94.3)	0.10 (1.39)	-3.7	-0.56	-0.06	0.62	8.4	
		Week 36	Sparsentan	91	89 (97.8)	-0.75 (1.18)	-4.2	-1.19	-0.73	-0.27	2.6	-0.74 [-1.05, -0.43]
			Irbesartan	88	82 (93.2)	0.20 (1.42)	-2.5	-0.61	-0.12	0.93	5.1	
		Week 58	Sparsentan	91	86 (94.5)	-0.61 (1.40)	-4.4	-1.29	-0.67	0.04	6.4	-0.58 [-0.89, -0.27]
			Irbesartan	88	82 (93.2)	0.22 (1.47)	-2.1	-0.59	-0.09	0.54	6.7	
		Week 110	Sparsentan	91	82 (90.1)	-0.65 (1.26)	-4.6	-1.29	-0.63	-0.06	3.4	-0.60 [-0.93, -0.27]
			Irbesartan	88	67 (76.1)	0.21 (1.59)	-6.0	-0.59	-0.00	1.05	4.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Geographic region													
North America	Urine protein excretion	Baseline	Sparsentan	35	35 (100.0)	2.73 (2.53)	0.1	1.18	2.19	3.14	14.7		
			Irbesartan	46	46 (100.0)	2.55 (1.46)	0.6	1.68	2.23	3.00	7.5		
		Week 6	Sparsentan	35	35 (100.0)	1.96 (1.33)	0.3	1.08	1.71	2.64	6.5		
			Irbesartan	46	45 (97.8)	2.65 (1.70)	0.5	1.41	2.43	3.66	7.7		
		Week 36	Sparsentan	35	33 (94.3)	1.43 (1.14)	0.2	0.63	1.21	1.61	4.3		
			Irbesartan	46	43 (93.5)	2.85 (1.97)	0.5	1.30	2.52	3.63	9.2		
		Week 58	Sparsentan	35	28 (80.0)	1.67 (1.63)	0.3	0.64	1.14	1.90	6.6		
			Irbesartan	46	40 (87.0)	2.74 (2.32)	0.5	1.19	2.22	3.13	12.8		
		Week 110	Sparsentan	35	27 (77.1)	1.60 (1.40)	0.2	0.70	1.24	2.16	6.0		
			Irbesartan	46	31 (67.4)	2.67 (2.09)	0.4	1.21	2.45	2.94	9.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	35	35 (100.0)	-0.77 (2.71)	-13.0	-1.65	-0.31	0.43	5.2	-0.41 [-0.86, 0.03]
				Irbesartan	46	45 (97.8)	0.11 (1.54)	-3.7	-0.54	-0.13	0.86	4.2	
			Week 36	Sparsentan	35	33 (94.3)	-1.29 (2.72)	-13.7	-1.71	-0.58	-0.11	2.6	-0.76 [-1.23, -0.29]
				Irbesartan	46	43 (93.5)	0.34 (1.56)	-2.5	-0.47	-0.06	1.07	4.3	
	Week 58		Sparsentan	35	28 (80.0)	-1.00 (3.25)	-13.3	-2.37	-0.68	0.39	6.4	-0.48 [-0.97, 0.01]	
			Irbesartan	46	40 (87.0)	0.18 (1.69)	-2.7	-0.87	-0.19	0.95	5.8		
	Week 110	Sparsentan	35	27 (77.1)	-1.09 (2.95)	-13.5	-1.91	-0.64	0.18	3.4	-0.43 [-0.95, 0.09]		
		Irbesartan	46	31 (67.4)	-0.03 (1.98)	-6.0	-0.91	-0.14	1.09	4.2			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Europe	Urine protein excretion	Baseline	Sparsentan	98	98 (100.0)	2.18 (1.41)	0.4	1.21	1.90	2.95	9.7		
			Irbesartan	115	115 (100.0)	2.12 (1.17)	0.5	1.32	1.82	2.60	5.9		
		Week 6	Sparsentan	98	92 (93.9)	1.36 (1.02)	0.2	0.67	1.08	1.91	5.5		
			Irbesartan	115	109 (94.8)	2.08 (1.54)	0.5	1.09	1.64	2.61	10.3		
		Week 36	Sparsentan	98	94 (95.9)	1.42 (1.35)	0.1	0.56	0.96	1.90	6.4		
			Irbesartan	115	108 (93.9)	2.00 (1.48)	0.2	0.90	1.60	2.60	7.0		
		Week 58	Sparsentan	98	93 (94.9)	1.46 (1.29)	0.1	0.64	1.13	1.79	7.5		
			Irbesartan	115	100 (87.0)	2.06 (1.66)	0.1	0.93	1.57	2.75	9.4		
		Week 110	Sparsentan	98	80 (81.6)	1.38 (1.25)	0.1	0.44	0.96	1.93	6.0		
			Irbesartan	115	88 (76.5)	2.24 (1.73)	0.1	1.00	1.74	2.95	8.8		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	98	92 (93.9)	-0.74 (1.06)	-4.2	-1.20	-0.71	-0.12	2.7	-0.59 [-0.87, -0.30]
				Irbesartan	115	109 (94.8)	-0.05 (1.26)	-2.4	-0.78	-0.12	0.44	8.4	
			Week 36	Sparsentan	98	94 (95.9)	-0.71 (1.17)	-4.1	-1.28	-0.77	-0.07	2.6	-0.45 [-0.73, -0.17]
				Irbesartan	115	108 (93.9)	-0.13 (1.38)	-3.3	-1.08	-0.23	0.61	5.1	
			Week 58	Sparsentan	98	93 (94.9)	-0.63 (1.21)	-4.2	-1.27	-0.74	-0.02	2.5	-0.43 [-0.71, -0.14]
				Irbesartan	115	100 (87.0)	-0.05 (1.49)	-3.3	-0.76	-0.25	0.43	6.7	
			Week 110	Sparsentan	98	80 (81.6)	-0.64 (1.29)	-3.6	-1.29	-0.73	-0.06	3.5	-0.53 [-0.83, -0.22]
				Irbesartan	115	88 (76.5)	0.13 (1.60)	-3.4	-0.73	-0.03	0.96	5.9	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Asia Pacific	Urine protein excretion	Baseline	Sparsentan	69	69 (100.0)	2.04 (1.36)	0.4	1.15	1.49	2.51	7.2			
			Irbesartan	41	41 (100.0)	1.87 (1.08)	0.6	1.25	1.59	2.29	6.8			
		Week 6	Sparsentan	69	68 (98.6)	1.43 (1.32)	0.2	0.61	0.95	1.79	6.2			
			Irbesartan	41	39 (95.1)	1.93 (1.56)	0.4	0.93	1.55	2.64	9.5			
		Week 36	Sparsentan	69	67 (97.1)	1.35 (1.63)	0.1	0.50	0.83	1.74	9.4			
			Irbesartan	41	36 (87.8)	1.82 (1.93)	0.2	0.72	1.22	1.80	9.2			
		Week 58	Sparsentan	69	67 (97.1)	1.29 (1.38)	0.1	0.43	0.81	1.60	6.8			
			Irbesartan	41	38 (92.7)	1.69 (1.32)	0.1	0.90	1.52	2.20	6.6			
		Week 110	Sparsentan	69	62 (89.9)	1.44 (1.38)	0.1	0.44	1.07	1.71	5.7			
			Irbesartan	41	31 (75.6)	2.03 (1.78)	0.3	1.04	1.65	2.38	9.2			
		Change from baseline in urine protein excretion		Week 6	Sparsentan	69	68 (98.6)	-0.62 (0.78)	-2.4	-1.01	-0.61	-0.10	0.9	-0.78 [-1.18, -0.37]
					Irbesartan	41	39 (95.1)	0.02 (0.88)	-1.7	-0.50	-0.20	0.42	2.8	
				Week 36	Sparsentan	69	67 (97.1)	-0.71 (1.18)	-3.1	-1.32	-0.73	-0.27	4.6	-0.50 [-0.91, -0.09]
					Irbesartan	41	36 (87.8)	-0.08 (1.41)	-2.0	-0.76	-0.34	-0.03	6.5	
				Week 58	Sparsentan	69	67 (97.1)	-0.77 (1.12)	-4.4	-1.44	-0.63	-0.04	2.0	-0.58 [-0.98, -0.17]
					Irbesartan	41	38 (92.7)	-0.17 (0.89)	-1.7	-0.84	-0.28	0.25	2.2	
				Week 110	Sparsentan	69	62 (89.9)	-0.58 (1.48)	-7.1	-1.29	-0.42	0.16	3.2	-0.54 [-0.98, -0.10]
					Irbesartan	41	31 (75.6)	0.18 (1.23)	-2.2	-0.42	0.00	0.43	3.2	

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An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline BMI													
< 27 kg/m**2	Urine protein excretion	Baseline	Sparsentan	83	83 (100.0)	1.88 (1.03)	0.4	1.15	1.58	2.48	5.0		
			Irbesartan	94	94 (100.0)	2.14 (1.13)	0.6	1.38	1.82	2.56	6.8		
		Week 6	Sparsentan	83	78 (94.0)	1.33 (1.18)	0.2	0.64	0.98	1.72	6.5		
			Irbesartan	94	89 (94.7)	2.06 (1.58)	0.5	1.06	1.55	2.43	9.5		
		Week 36	Sparsentan	83	80 (96.4)	1.17 (1.15)	0.1	0.46	0.80	1.42	6.3		
			Irbesartan	94	85 (90.4)	1.81 (1.32)	0.2	0.96	1.44	2.54	7.7		
		Week 58	Sparsentan	83	76 (91.6)	1.06 (0.89)	0.1	0.38	0.85	1.30	5.0		
			Irbesartan	94	79 (84.0)	1.70 (1.23)	0.1	0.90	1.35	2.20	6.6		
		Week 110	Sparsentan	83	70 (84.3)	1.32 (1.38)	0.1	0.40	0.96	1.67	6.0		
			Irbesartan	94	66 (70.2)	1.96 (1.75)	0.1	0.94	1.46	2.39	9.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	83	78 (94.0)	-0.54 (1.12)	-2.4	-1.18	-0.65	0.01	5.2	-0.39 [-0.70, -0.09]
				Irbesartan	94	89 (94.7)	-0.11 (1.04)	-2.4	-0.73	-0.30	0.29	2.9	
			Week 36	Sparsentan	83	80 (96.4)	-0.75 (1.01)	-3.1	-1.39	-0.75	-0.27	2.2	-0.42 [-0.73, -0.12]
				Irbesartan	94	85 (90.4)	-0.33 (0.97)	-3.3	-0.91	-0.32	0.43	2.0	
	Week 58		Sparsentan	83	76 (91.6)	-0.81 (1.01)	-3.7	-1.44	-0.77	-0.17	1.3	-0.41 [-0.73, -0.09]	
			Irbesartan	94	79 (84.0)	-0.39 (1.02)	-3.3	-0.97	-0.35	0.13	2.8		
	Week 110	Sparsentan	83	70 (84.3)	-0.51 (1.27)	-3.5	-1.32	-0.58	0.18	3.5	-0.30 [-0.64, 0.04]		
		Irbesartan	94	66 (70.2)	-0.12 (1.35)	-3.4	-0.69	-0.12	0.34	3.8			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 27 kg/m**2	Urine protein excretion	Baseline	Sparsentan	119	119 (100.0)	2.47 (1.94)	0.1	1.18	1.92	3.16	14.7		
			Irbesartan	107	107 (100.0)	2.20 (1.33)	0.6	1.33	1.82	2.78	7.5		
		Week 6	Sparsentan	119	117 (98.3)	1.60 (1.21)	0.2	0.70	1.24	2.20	6.2		
			Irbesartan	107	103 (96.3)	2.30 (1.61)	0.4	1.19	1.93	2.82	10.3		
		Week 36	Sparsentan	119	114 (95.8)	1.55 (1.57)	0.1	0.56	1.01	2.00	9.4		
			Irbesartan	107	101 (94.4)	2.47 (1.97)	0.2	1.01	1.96	3.38	9.2		
		Week 58	Sparsentan	119	112 (94.1)	1.68 (1.58)	0.1	0.61	1.14	2.06	7.5		
			Irbesartan	107	99 (92.5)	2.48 (2.08)	0.1	1.05	1.89	3.14	12.8		
		Week 110	Sparsentan	119	99 (83.2)	1.52 (1.27)	0.1	0.53	1.15	2.30	5.5		
			Irbesartan	107	83 (77.6)	2.56 (1.84)	0.1	1.25	2.20	3.36	9.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	119	117 (98.3)	-0.81 (1.59)	-13.0	-1.28	-0.67	-0.09	2.7	-0.60 [-0.87, -0.33]
				Irbesartan	107	103 (96.3)	0.10 (1.43)	-3.7	-0.54	-0.04	0.62	8.4	
			Week 36	Sparsentan	119	114 (95.8)	-0.85 (1.84)	-13.7	-1.37	-0.70	-0.11	4.6	-0.62 [-0.90, -0.35]
				Irbesartan	107	101 (94.4)	0.25 (1.69)	-2.5	-0.78	-0.14	1.00	6.5	
			Week 58	Sparsentan	119	112 (94.1)	-0.68 (1.96)	-13.3	-1.29	-0.63	0.12	6.4	-0.52 [-0.80, -0.25]
				Irbesartan	107	99 (92.5)	0.27 (1.64)	-2.7	-0.73	0.02	0.89	6.7	
			Week 110	Sparsentan	119	99 (83.2)	-0.82 (1.98)	-13.5	-1.48	-0.65	-0.04	3.4	-0.58 [-0.88, -0.29]
				Irbesartan	107	83 (77.6)	0.29 (1.79)	-6.0	-0.65	0.18	1.45	5.9	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Randomization strata													
eGFR Low and UP High	Urine protein excretion	Baseline	Sparsentan	71	71 (100.0)	3.08 (1.57)	0.1	2.07	2.95	3.74	9.7		
		Week 6	Irbesartan	74	74 (100.0)	2.96 (1.43)	0.5	1.86	2.59	3.77	7.5		
			Sparsentan	71	68 (95.8)	2.10 (1.20)	0.5	1.24	1.86	2.49	5.7		
		Week 36	Irbesartan	74	73 (98.6)	2.96 (1.63)	0.8	1.89	2.65	3.75	7.7		
			Sparsentan	71	68 (95.8)	2.05 (1.74)	0.1	0.83	1.48	2.61	9.4		
		Week 58	Irbesartan	74	68 (91.9)	3.04 (1.88)	0.2	1.73	2.68	3.71	9.2		
			Sparsentan	71	67 (94.4)	2.03 (1.65)	0.2	0.82	1.34	2.70	7.5		
		Week 110	Irbesartan	74	64 (86.5)	2.92 (2.13)	0.1	1.63	2.42	3.71	12.8		
			Sparsentan	71	59 (83.1)	2.18 (1.63)	0.2	0.86	1.75	3.08	6.0		
		Week 110	Irbesartan	74	54 (73.0)	2.92 (2.04)	0.1	1.48	2.48	3.72	9.2		
			Change from baseline in urine protein excretion	Week 6	Sparsentan	71	68 (95.8)	-0.88 (1.31)	-4.2	-1.63	-0.84	0.10	1.9
		Week 36	Irbesartan	74	73 (98.6)	0.01 (1.45)	-3.7	-0.84	-0.04	0.72	4.2		
			Sparsentan	71	68 (95.8)	-0.94 (1.66)	-4.7	-1.92	-1.00	-0.02	4.6	-0.59 [-0.94, -0.25]	
		Week 58	Irbesartan	74	68 (91.9)	0.06 (1.72)	-3.3	-1.22	-0.05	1.08	5.0		
Sparsentan	71		67 (94.4)	-0.89 (1.85)	-4.4	-2.13	-1.00	0.04	6.4	-0.50 [-0.84, -0.15]			
Week 110	Irbesartan	74	64 (86.5)	-0.03 (1.63)	-3.3	-1.23	-0.01	0.73	5.8				
	Sparsentan	71	59 (83.1)	-0.64 (1.77)	-4.6	-1.73	-0.66	0.57	3.5	-0.32 [-0.69, 0.05]			
Week 110	Irbesartan	74	54 (73.0)	-0.05 (1.96)	-6.0	-1.23	-0.01	1.60	3.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.  
An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR Low and UP Low	Urine protein excretion	Baseline	Sparsentan	55	55 (100.0)	1.56 (1.91)	0.4	0.97	1.19	1.51	14.7			
			Irbesartan	55	55 (100.0)	1.43 (0.46)	0.6	1.10	1.33	1.76	3.1			
		Week 6	Sparsentan	55	52 (94.5)	1.06 (0.96)	0.2	0.61	0.91	1.28	6.5			
			Irbesartan	55	53 (96.4)	1.32 (0.61)	0.6	0.89	1.29	1.64	4.0			
		Week 36	Sparsentan	55	51 (92.7)	0.99 (0.82)	0.1	0.52	0.65	1.29	4.0			
			Irbesartan	55	51 (92.7)	1.40 (0.99)	0.3	0.69	1.04	1.70	4.4			
		Week 58	Sparsentan	55	52 (94.5)	0.96 (0.75)	0.1	0.47	0.87	1.24	4.7			
			Irbesartan	55	51 (92.7)	1.37 (1.05)	0.1	0.67	1.16	1.72	5.0			
		Week 110	Sparsentan	55	46 (83.6)	1.12 (0.83)	0.1	0.39	0.96	1.58	3.0			
			Irbesartan	55	42 (76.4)	1.48 (0.96)	0.1	0.83	1.22	2.25	4.0			
		Change from baseline in urine protein excretion		Week 6	Sparsentan	55	52 (94.5)	-0.53 (2.06)	-13.0	-0.80	-0.40	0.08	5.2	-0.27 [-0.65, 0.12]
					Irbesartan	55	53 (96.4)	-0.12 (0.70)	-1.5	-0.48	-0.19	0.30	2.9	
				Week 36	Sparsentan	55	51 (92.7)	-0.59 (2.12)	-13.7	-0.88	-0.29	0.11	2.6	-0.32 [-0.71, 0.07]
					Irbesartan	55	51 (92.7)	-0.05 (1.02)	-1.3	-0.75	-0.29	0.26	3.6	
				Week 58	Sparsentan	55	52 (94.5)	-0.58 (2.00)	-13.3	-0.93	-0.39	0.11	3.3	-0.31 [-0.70, 0.07]
					Irbesartan	55	51 (92.7)	-0.08 (1.05)	-1.5	-0.65	-0.34	0.13	3.7	
				Week 110	Sparsentan	55	46 (83.6)	-0.48 (2.18)	-13.5	-0.86	-0.35	0.37	1.9	-0.31 [-0.73, 0.11]
					Irbesartan	55	42 (76.4)	0.05 (0.87)	-1.8	-0.57	-0.08	0.93	1.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR High and UP High	Urine protein excretion	Baseline	Sparsentan	37	37 (100.0)	2.60 (1.20)	1.0	1.96	2.42	2.99	7.2		
			Irbesartan	36	36 (100.0)	2.47 (1.05)	0.6	1.95	2.32	2.84	6.8		
		Week 6	Sparsentan	37	36 (97.3)	1.82 (1.43)	0.2	0.77	1.35	2.43	6.2		
			Irbesartan	36	32 (88.9)	2.72 (2.13)	0.6	1.44	2.29	3.12	10.3		
		Week 36	Sparsentan	37	37 (100.0)	1.53 (1.53)	0.1	0.63	1.05	1.74	8.3		
			Irbesartan	36	32 (88.9)	2.57 (2.05)	0.3	1.36	1.95	2.88	9.2		
		Week 58	Sparsentan	37	32 (86.5)	1.63 (1.58)	0.1	0.54	1.21	1.86	5.6		
			Irbesartan	36	31 (86.1)	2.54 (1.95)	0.7	1.12	1.98	2.98	8.6		
		Week 110	Sparsentan	37	30 (81.1)	1.16 (1.20)	0.1	0.45	0.83	1.42	6.0		
			Irbesartan	36	27 (75.0)	2.92 (2.26)	0.4	1.58	2.22	3.21	9.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	37	36 (97.3)	-0.79 (1.14)	-2.4	-1.44	-0.92	-0.40	2.7	-0.63 [-1.12, -0.14]
				Irbesartan	36	32 (88.9)	0.19 (1.91)	-3.2	-0.80	-0.03	0.83	8.4	
			Week 36	Sparsentan	37	37 (100.0)	-1.07 (1.00)	-3.1	-1.65	-1.09	-0.50	1.0	-0.76 [-1.25, -0.27]
				Irbesartan	36	32 (88.9)	0.07 (1.91)	-2.5	-1.26	-0.19	0.73	6.5	
			Week 58	Sparsentan	37	32 (86.5)	-1.00 (1.20)	-3.7	-1.79	-1.02	-0.28	1.7	-0.65 [-1.16, -0.15]
				Irbesartan	36	31 (86.1)	0.07 (2.00)	-2.7	-1.05	-0.30	0.76	6.7	
			Week 110	Sparsentan	37	30 (81.1)	-1.42 (1.47)	-7.1	-2.06	-1.20	-0.76	1.3	-0.96 [-1.50, -0.41]
				Irbesartan	36	27 (75.0)	0.34 (2.18)	-3.6	-0.90	-0.00	1.53	5.9	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP Low	Urine protein excretion	Baseline	Sparsentan	39	39 (100.0)	1.26 (0.48)	0.4	0.96	1.18	1.51	3.3			
			Irbesartan	37	37 (100.0)	1.37 (0.44)	0.6	1.11	1.41	1.60	2.9			
		Week 6	Sparsentan	39	39 (100.0)	0.72 (0.38)	0.2	0.46	0.64	0.90	1.8			
			Irbesartan	37	35 (94.6)	1.38 (0.79)	0.4	0.87	1.19	1.91	3.6			
		Week 36	Sparsentan	39	38 (97.4)	0.65 (0.49)	0.1	0.30	0.56	0.88	2.3			
			Irbesartan	37	36 (97.3)	1.21 (0.75)	0.2	0.75	1.09	1.57	3.6			
		Week 58	Sparsentan	39	37 (94.9)	0.83 (0.73)	0.1	0.30	0.57	1.05	3.1			
			Irbesartan	37	32 (86.5)	1.38 (0.84)	0.2	0.92	1.12	1.59	3.6			
		Week 110	Sparsentan	39	34 (87.2)	0.83 (0.63)	0.1	0.30	0.70	1.35	2.3			
			Irbesartan	37	27 (73.0)	1.61 (1.10)	0.3	0.90	1.28	2.18	4.6			
		Change from baseline in urine protein excretion		Week 6	Sparsentan	39	39 (100.0)	-0.54 (0.53)	-2.6	-0.79	-0.51	-0.19	0.6	-0.90 [-1.37, -0.42]
					Irbesartan	37	35 (94.6)	-0.01 (0.66)	-1.1	-0.52	-0.25	0.72	1.4	
				Week 36	Sparsentan	39	38 (97.4)	-0.62 (0.58)	-2.4	-0.89	-0.66	-0.31	0.7	-0.72 [-1.19, -0.25]
					Irbesartan	37	36 (97.3)	-0.17 (0.68)	-1.5	-0.64	-0.24	0.39	1.0	
				Week 58	Sparsentan	39	37 (94.9)	-0.43 (0.80)	-2.7	-0.90	-0.45	-0.02	1.4	-0.50 [-0.98, -0.02]
					Irbesartan	37	32 (86.5)	-0.02 (0.83)	-1.5	-0.50	-0.24	0.25	2.6	
				Week 110	Sparsentan	39	34 (87.2)	-0.42 (0.75)	-2.3	-0.96	-0.38	-0.04	1.0	-0.80 [-1.33, -0.28]
					Irbesartan	37	27 (73.0)	0.28 (1.01)	-0.9	-0.56	0.04	0.62	3.6	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
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An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 1														
< 60 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	127	127 (100.0)	2.28 (1.52)	0.1	1.19	1.85	3.12	9.7			
		Week 6	Irbesartan	129	129 (100.0)	2.28 (1.34)	0.5	1.33	1.82	2.80	7.5			
			Sparsentan	127	121 (95.3)	1.59 (1.21)	0.2	0.73	1.25	2.08	6.5			
		Week 36	Irbesartan	129	126 (97.7)	2.22 (1.47)	0.5	1.17	1.86	2.73	7.7			
			Sparsentan	127	120 (94.5)	1.57 (1.51)	0.1	0.60	1.04	2.08	9.4			
		Week 58	Irbesartan	129	119 (92.2)	2.23 (1.74)	0.2	0.96	1.70	3.02	9.2			
			Sparsentan	127	120 (94.5)	1.54 (1.43)	0.1	0.65	1.15	1.93	7.5			
		Week 110	Irbesartan	129	115 (89.1)	2.19 (1.89)	0.1	0.95	1.72	2.80	12.8			
			Sparsentan	127	105 (82.7)	1.69 (1.41)	0.1	0.66	1.40	2.42	6.0			
		Week 110	Irbesartan	129	97 (75.2)	2.28 (1.79)	0.1	1.10	1.77	2.87	9.2			
			Sparsentan	127	105 (82.7)	1.69 (1.41)	0.1	0.66	1.40	2.42	6.0			
			Change from baseline in urine protein excretion	Week 6	Sparsentan	127	121 (95.3)	-0.64 (1.24)	-4.2	-1.29	-0.67	0.08	5.2	-0.47 [-0.72, -0.22]
				Week 36	Irbesartan	129	126 (97.7)	-0.07 (1.17)	-3.7	-0.68	-0.19	0.42	4.2	
					Sparsentan	127	120 (94.5)	-0.66 (1.43)	-4.7	-1.35	-0.68	0.09	4.6	-0.42 [-0.68, -0.16]
				Week 58	Irbesartan	129	119 (92.2)	-0.06 (1.42)	-3.3	-0.92	-0.19	0.53	5.0	
					Sparsentan	127	120 (94.5)	-0.63 (1.52)	-4.4	-1.45	-0.59	0.11	6.4	-0.39 [-0.64, -0.13]
				Week 110	Irbesartan	129	115 (89.1)	-0.08 (1.34)	-2.7	-0.84	-0.30	0.44	5.8	
					Sparsentan	127	105 (82.7)	-0.43 (1.48)	-4.6	-1.29	-0.51	0.41	3.5	-0.29 [-0.57, -0.01]
		Week 110	Irbesartan	129	97 (75.2)	0.00 (1.52)	-6.0	-0.82	-0.08	0.76	3.8			
			Sparsentan	127	105 (82.7)	1.69 (1.41)	0.1	0.66	1.40	2.42	6.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	49	49 (100.0)	2.14 (2.03)	0.6	1.12	1.62	2.62	14.7		
		Week 6	Irbesartan	48	48 (100.0)	1.97 (0.91)	0.6	1.36	1.84	2.45	4.6		
			Sparsentan	49	48 (98.0)	1.33 (1.16)	0.2	0.56	0.86	1.84	5.5		
		Week 36	Irbesartan	48	44 (91.7)	2.30 (1.75)	0.4	1.28	1.88	3.00	10.3		
			Sparsentan	49	48 (98.0)	1.06 (0.92)	0.1	0.46	0.83	1.30	3.9		
		Week 58	Irbesartan	48	45 (93.8)	2.40 (1.69)	0.2	1.30	1.94	2.94	9.2		
			Sparsentan	49	43 (87.8)	1.18 (1.17)	0.1	0.29	0.88	1.50	5.6		
		Week 110	Irbesartan	48	42 (87.5)	2.37 (1.65)	0.2	1.29	2.06	2.98	8.6		
			Sparsentan	49	43 (87.8)	0.95 (0.88)	0.1	0.30	0.74	1.28	4.3		
		Week 110	Irbesartan	48	36 (75.0)	2.53 (1.79)	0.4	1.26	2.12	3.07	7.6		
			Sparsentan	49	48 (98.0)	-0.80 (2.04)	-13.0	-1.09	-0.67	-0.15	2.7	-0.58 [-1.00, -0.17]	
		Change from baseline in urine protein excretion	Week 6	Irbesartan	48	44 (91.7)	0.28 (1.60)	-3.2	-0.52	0.04	0.80	8.4	
				Sparsentan	49	48 (98.0)	-1.10 (2.00)	-13.7	-1.36	-0.71	-0.49	0.9	-0.85 [-1.27, -0.42]
			Week 36	Irbesartan	48	45 (93.8)	0.44 (1.58)	-2.2	-0.42	0.26	0.93	6.5	
				Sparsentan	49	43 (87.8)	-0.97 (2.15)	-13.3	-1.34	-0.76	-0.10	1.7	-0.69 [-1.12, -0.25]
			Week 58	Irbesartan	48	42 (87.5)	0.39 (1.80)	-3.3	-0.42	0.03	1.01	6.7	
				Sparsentan	49	43 (87.8)	-1.17 (2.12)	-13.5	-1.57	-0.82	-0.26	0.9	-0.82 [-1.28, -0.36]
			Week 110	Irbesartan	48	36 (75.0)	0.52 (1.97)	-3.6	-0.58	0.31	1.56	5.9	
	Sparsentan			49	48 (98.0)	-0.80 (2.04)	-13.0	-1.09	-0.67	-0.15	2.7	-0.58 [-1.00, -0.17]	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	26	26 (100.0)	2.15 (1.51)	0.7	1.18	1.64	2.48	7.2	
		Week 6	Irbesartan	25	25 (100.0)	1.94 (1.21)	0.6	1.30	1.63	2.10	6.8	
			Sparsentan	26	26 (100.0)	1.32 (1.24)	0.4	0.57	0.92	1.60	6.2	
		Week 36	Irbesartan	25	23 (92.0)	1.78 (1.96)	0.4	0.71	1.18	1.99	9.5	
			Sparsentan	26	26 (100.0)	1.19 (1.64)	0.1	0.34	0.63	1.50	8.3	
		Week 58	Irbesartan	25	23 (92.0)	1.33 (1.54)	0.2	0.47	0.95	1.56	7.7	
			Sparsentan	26	25 (96.2)	1.32 (1.44)	0.2	0.41	0.83	1.67	5.4	
		Week 110	Irbesartan	25	21 (84.0)	1.36 (1.32)	0.3	0.80	1.05	1.33	6.6	
			Sparsentan	26	21 (80.8)	1.16 (1.32)	0.1	0.42	0.73	1.36	6.0	
		Week 110	Irbesartan	25	17 (68.0)	1.80 (2.08)	0.3	0.81	1.26	1.99	9.2	
			Sparsentan	26	26 (100.0)	-0.82 (0.72)	-2.4	-1.20	-0.68	-0.44	0.4	-0.82 [-1.40, -0.23]
		Week 36	Irbesartan	25	23 (92.0)	-0.14 (0.96)	-1.9	-0.66	-0.32	0.18	2.8	
			Sparsentan	26	26 (100.0)	-0.96 (1.03)	-3.1	-1.57	-0.89	-0.39	1.0	-0.35 [-0.92, 0.22]
		Week 58	Irbesartan	25	23 (92.0)	-0.63 (0.86)	-2.5	-1.27	-0.55	-0.14	1.0	
			Sparsentan	26	25 (96.2)	-0.82 (1.16)	-3.7	-1.29	-0.71	-0.09	1.4	-0.27 [-0.85, 0.31]
		Week 110	Irbesartan	25	21 (84.0)	-0.54 (0.82)	-2.3	-0.92	-0.52	-0.12	1.4	
			Sparsentan	26	21 (80.8)	-1.00 (1.77)	-7.1	-1.57	-0.49	-0.06	1.3	-0.55 [-1.20, 0.10]
		Week 110	Irbesartan	25	17 (68.0)	-0.16 (1.15)	-2.3	-0.67	-0.30	0.05	2.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 2													
< 45 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	82	82 (100.0)	2.39 (1.59)	0.1	1.28	1.99	3.15	9.7		
		Week 6	Irbesartan	80	80 (100.0)	2.32 (1.40)	0.5	1.33	1.95	2.94	7.5		
			Sparsentan	82	78 (95.1)	1.75 (1.34)	0.2	0.92	1.28	2.15	6.5		
		Week 36	Irbesartan	80	78 (97.5)	2.18 (1.37)	0.6	1.04	1.84	2.70	7.0		
			Sparsentan	82	78 (95.1)	1.73 (1.62)	0.1	0.60	1.26	2.38	9.4		
		Week 58	Irbesartan	80	72 (90.0)	2.30 (1.72)	0.2	1.00	2.09	3.14	9.2		
			Sparsentan	82	77 (93.9)	1.75 (1.64)	0.1	0.65	1.19	2.03	7.5		
		Week 110	Irbesartan	80	69 (86.3)	2.15 (1.93)	0.1	0.87	1.79	2.78	12.8		
			Sparsentan	82	67 (81.7)	1.86 (1.49)	0.1	0.69	1.53	2.63	6.0		
		Week 110	Irbesartan	80	59 (73.8)	2.27 (1.70)	0.1	1.10	1.72	3.03	9.2		
			Sparsentan	82	78 (95.1)	-0.57 (1.34)	-4.2	-1.36	-0.65	0.11	5.2	-0.33 [-0.65, -0.02]	
		Change from baseline in urine protein excretion	Week 6	Irbesartan	80	78 (97.5)	-0.14 (1.22)	-3.7	-0.78	-0.21	0.38	3.2	
				Sparsentan	82	78 (95.1)	-0.59 (1.53)	-4.7	-1.45	-0.62	0.29	4.6	-0.35 [-0.67, -0.03]
				Irbesartan	80	72 (90.0)	-0.06 (1.47)	-2.5	-0.95	-0.18	0.62	5.0	
Sparsentan	82			77 (93.9)	-0.53 (1.66)	-4.4	-1.51	-0.49	0.27	6.4	-0.25 [-0.57, 0.08]		
Irbesartan	80			69 (86.3)	-0.15 (1.39)	-2.7	-0.97	-0.26	0.39	5.8			
Sparsentan	82			67 (81.7)	-0.38 (1.45)	-3.6	-1.48	-0.49	0.60	3.4	-0.26 [-0.61, 0.09]		
Week 110	Irbesartan	80	59 (73.8)	0.01 (1.53)	-6.0	-0.82	0.00	0.95	3.2				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
45 to < 60 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	45	45 (100.0)	2.08 (1.39)	0.4	1.18	1.76	2.75	6.4			
			Irbesartan	49	49 (100.0)	2.22 (1.25)	0.6	1.39	1.77	2.64	5.7			
		Week 6	Sparsentan	45	43 (95.6)	1.31 (0.89)	0.2	0.60	1.09	2.01	3.5			
			Irbesartan	49	48 (98.0)	2.28 (1.62)	0.5	1.20	1.86	2.76	7.7			
		Week 36	Sparsentan	45	42 (93.3)	1.28 (1.25)	0.1	0.59	0.83	1.47	6.4			
			Irbesartan	49	47 (95.9)	2.12 (1.79)	0.3	0.93	1.30	2.63	7.8			
		Week 58	Sparsentan	45	43 (95.6)	1.17 (0.84)	0.1	0.59	1.03	1.53	4.5			
			Irbesartan	49	46 (93.9)	2.24 (1.85)	0.3	0.98	1.68	2.91	9.4			
		Week 110	Sparsentan	45	38 (84.4)	1.40 (1.21)	0.1	0.61	1.11	1.75	5.7			
			Irbesartan	49	38 (77.6)	2.29 (1.94)	0.1	1.08	1.78	2.86	8.8			
		Change from baseline in urine protein excretion		Week 6	Sparsentan	45	43 (95.6)	-0.77 (1.04)	-4.0	-1.28	-0.69	-0.09	0.9	-0.76 [-1.19, -0.34]
					Irbesartan	49	48 (98.0)	0.04 (1.09)	-1.9	-0.61	-0.08	0.43	4.2	
				Week 36	Sparsentan	45	42 (93.3)	-0.80 (1.22)	-4.1	-1.26	-0.82	-0.07	2.6	-0.57 [-0.99, -0.14]
					Irbesartan	49	47 (95.9)	-0.07 (1.34)	-3.3	-0.78	-0.21	0.43	4.3	
				Week 58	Sparsentan	45	43 (95.6)	-0.81 (1.23)	-4.2	-1.44	-0.63	-0.03	2.4	-0.67 [-1.10, -0.24]
					Irbesartan	49	46 (93.9)	0.03 (1.26)	-1.9	-0.74	-0.39	0.48	3.7	
				Week 110	Sparsentan	45	38 (84.4)	-0.52 (1.54)	-4.6	-1.20	-0.52	0.04	3.5	-0.34 [-0.79, 0.11]
					Irbesartan	49	38 (77.6)	-0.00 (1.53)	-2.4	-0.90	-0.33	0.70	3.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	49	49 (100.0)	2.14 (2.03)	0.6	1.12	1.62	2.62	14.7		
		Week 6	Irbesartan	48	48 (100.0)	1.97 (0.91)	0.6	1.36	1.84	2.45	4.6		
			Sparsentan	49	48 (98.0)	1.33 (1.16)	0.2	0.56	0.86	1.84	5.5		
		Week 36	Irbesartan	48	44 (91.7)	2.30 (1.75)	0.4	1.28	1.88	3.00	10.3		
			Sparsentan	49	48 (98.0)	1.06 (0.92)	0.1	0.46	0.83	1.30	3.9		
		Week 58	Irbesartan	48	45 (93.8)	2.40 (1.69)	0.2	1.30	1.94	2.94	9.2		
			Sparsentan	49	43 (87.8)	1.18 (1.17)	0.1	0.29	0.88	1.50	5.6		
		Week 110	Irbesartan	48	42 (87.5)	2.37 (1.65)	0.2	1.29	2.06	2.98	8.6		
			Sparsentan	49	43 (87.8)	0.95 (0.88)	0.1	0.30	0.74	1.28	4.3		
		Week 110	Irbesartan	48	36 (75.0)	2.53 (1.79)	0.4	1.26	2.12	3.07	7.6		
			Change from baseline in urine protein excretion	Week 6	Sparsentan	49	48 (98.0)	-0.80 (2.04)	-13.0	-1.09	-0.67	-0.15	2.7
		Week 36	Irbesartan	48	44 (91.7)	0.28 (1.60)	-3.2	-0.52	0.04	0.80	8.4		
			Sparsentan	49	48 (98.0)	-1.10 (2.00)	-13.7	-1.36	-0.71	-0.49	0.9	-0.85 [-1.27, -0.42]	
		Week 58	Irbesartan	48	45 (93.8)	0.44 (1.58)	-2.2	-0.42	0.26	0.93	6.5		
			Sparsentan	49	43 (87.8)	-0.97 (2.15)	-13.3	-1.34	-0.76	-0.10	1.7	-0.69 [-1.12, -0.25]	
		Week 110	Irbesartan	48	42 (87.5)	0.39 (1.80)	-3.3	-0.42	0.03	1.01	6.7		
			Sparsentan	49	43 (87.8)	-1.17 (2.12)	-13.5	-1.57	-0.82	-0.26	0.9	-0.82 [-1.28, -0.36]	
		Week 110	Irbesartan	48	36 (75.0)	0.52 (1.97)	-3.6	-0.58	0.31	1.56	5.9		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 90 mL/min/1.73 m**2	Urine protein excretion	Baseline	Sparsentan	26	26 (100.0)	2.15 (1.51)	0.7	1.18	1.64	2.48	7.2	
		Week 6	Irbesartan	25	25 (100.0)	1.94 (1.21)	0.6	1.30	1.63	2.10	6.8	
			Sparsentan	26	26 (100.0)	1.32 (1.24)	0.4	0.57	0.92	1.60	6.2	
		Week 36	Irbesartan	25	23 (92.0)	1.78 (1.96)	0.4	0.71	1.18	1.99	9.5	
			Sparsentan	26	26 (100.0)	1.19 (1.64)	0.1	0.34	0.63	1.50	8.3	
		Week 58	Irbesartan	25	23 (92.0)	1.33 (1.54)	0.2	0.47	0.95	1.56	7.7	
			Sparsentan	26	25 (96.2)	1.32 (1.44)	0.2	0.41	0.83	1.67	5.4	
		Week 110	Irbesartan	25	21 (84.0)	1.36 (1.32)	0.3	0.80	1.05	1.33	6.6	
			Sparsentan	26	21 (80.8)	1.16 (1.32)	0.1	0.42	0.73	1.36	6.0	
		Week 110	Irbesartan	25	17 (68.0)	1.80 (2.08)	0.3	0.81	1.26	1.99	9.2	
			Sparsentan	26	26 (100.0)	-0.82 (0.72)	-2.4	-1.20	-0.68	-0.44	0.4	-0.82 [-1.40, -0.23]
		Week 36	Irbesartan	25	23 (92.0)	-0.14 (0.96)	-1.9	-0.66	-0.32	0.18	2.8	
			Sparsentan	26	26 (100.0)	-0.96 (1.03)	-3.1	-1.57	-0.89	-0.39	1.0	-0.35 [-0.92, 0.22]
		Week 58	Irbesartan	25	23 (92.0)	-0.63 (0.86)	-2.5	-1.27	-0.55	-0.14	1.0	
			Sparsentan	26	25 (96.2)	-0.82 (1.16)	-3.7	-1.29	-0.71	-0.09	1.4	-0.27 [-0.85, 0.31]
		Week 110	Irbesartan	25	21 (84.0)	-0.54 (0.82)	-2.3	-0.92	-0.52	-0.12	1.4	
			Sparsentan	26	21 (80.8)	-1.00 (1.77)	-7.1	-1.57	-0.49	-0.06	1.3	-0.55 [-1.20, 0.10]
		Week 110	Irbesartan	25	17 (68.0)	-0.16 (1.15)	-2.3	-0.67	-0.30	0.05	2.5	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline urine protein excretion													
<= 1.75 g/day	Urine protein excretion	Baseline	Sparsentan	98	98 (100.0)	1.14 (0.33)	0.1	0.97	1.16	1.40	1.7		
			Irbesartan	93	93 (100.0)	1.26 (0.32)	0.5	1.07	1.33	1.50	1.7		
		Week 6	Sparsentan	98	95 (96.9)	0.97 (0.80)	0.2	0.48	0.77	1.18	6.5		
			Irbesartan	93	88 (94.6)	1.45 (0.89)	0.4	0.89	1.26	1.80	4.8		
		Week 36	Sparsentan	98	94 (95.9)	0.91 (0.80)	0.1	0.34	0.64	1.21	4.0		
			Irbesartan	93	87 (93.5)	1.48 (1.16)	0.2	0.73	1.11	1.77	6.1		
		Week 58	Sparsentan	98	93 (94.9)	1.01 (0.98)	0.1	0.32	0.79	1.22	6.6		
			Irbesartan	93	82 (88.2)	1.50 (1.17)	0.1	0.81	1.17	1.74	6.7		
		Week 110	Sparsentan	98	85 (86.7)	1.09 (0.94)	0.1	0.40	0.88	1.53	4.8		
			Irbesartan	93	70 (75.3)	1.66 (1.22)	0.1	0.87	1.31	2.27	7.0		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	98	95 (96.9)	-0.18 (0.81)	-1.2	-0.67	-0.31	0.08	5.2	-0.44 [-0.74, -0.15]
				Irbesartan	93	88 (94.6)	0.18 (0.84)	-1.1	-0.35	0.01	0.48	3.2	
			Week 36	Sparsentan	98	94 (95.9)	-0.24 (0.79)	-1.4	-0.81	-0.42	0.13	2.6	-0.47 [-0.76, -0.17]
				Irbesartan	93	87 (93.5)	0.21 (1.13)	-1.4	-0.44	-0.13	0.53	5.0	
	Week 58		Sparsentan	98	93 (94.9)	-0.13 (1.01)	-1.4	-0.76	-0.30	0.23	6.4	-0.32 [-0.62, -0.02]	
			Irbesartan	93	82 (88.2)	0.22 (1.17)	-1.5	-0.49	-0.10	0.55	5.6		
	Week 110	Sparsentan	98	85 (86.7)	-0.04 (0.95)	-1.3	-0.71	-0.26	0.36	3.5	-0.42 [-0.74, -0.10]		
		Irbesartan	93	70 (75.3)	0.41 (1.18)	-1.3	-0.30	0.09	0.95	5.9			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 1.75 g/day	Urine protein excretion	Baseline	Sparsentan	104	104 (100.0)	3.25 (1.74)	1.8	2.23	2.82	3.50	14.7	
			Irbesartan	109	109 (100.0)	2.94 (1.21)	1.8	2.06	2.54	3.33	7.5	
		Week 6	Sparsentan	104	100 (96.2)	1.99 (1.31)	0.3	1.04	1.67	2.49	6.2	
			Irbesartan	109	105 (96.3)	2.80 (1.79)	0.7	1.59	2.38	3.31	10.3	
		Week 36	Sparsentan	104	100 (96.2)	1.85 (1.70)	0.1	0.71	1.28	2.52	9.4	
			Irbesartan	109	100 (91.7)	2.75 (1.92)	0.2	1.30	2.45	3.46	9.2	
		Week 58	Sparsentan	104	95 (91.3)	1.84 (1.58)	0.2	0.75	1.29	2.47	7.5	
			Irbesartan	109	96 (88.1)	2.67 (2.05)	0.1	1.40	2.18	3.13	12.8	
		Week 110	Sparsentan	104	84 (80.8)	1.79 (1.54)	0.1	0.67	1.40	2.58	6.0	
			Irbesartan	109	80 (73.4)	2.83 (2.08)	0.1	1.48	2.39	3.57	9.2	
	Change from baseline in urine protein excretion	Week 6	Sparsentan	104	100 (96.2)	-1.20 (1.69)	-13.0	-1.82	-1.19	-0.55	2.7	-0.65 [-0.93, -0.37]
			Irbesartan	109	105 (96.3)	-0.15 (1.52)	-3.7	-1.00	-0.32	0.42	8.4	
		Week 36	Sparsentan	104	100 (96.2)	-1.34 (1.87)	-13.7	-2.14	-1.29	-0.54	4.6	-0.65 [-0.93, -0.36]
			Irbesartan	109	100 (91.7)	-0.21 (1.63)	-3.3	-1.28	-0.44	0.64	6.5	
		Week 58	Sparsentan	104	95 (91.3)	-1.33 (1.91)	-13.3	-2.13	-1.32	-0.52	2.5	-0.62 [-0.91, -0.33]
			Irbesartan	109	96 (88.1)	-0.23 (1.60)	-3.3	-1.24	-0.42	0.35	6.7	
		Week 110	Sparsentan	104	84 (80.8)	-1.35 (2.05)	-13.5	-2.07	-1.36	-0.43	3.2	-0.61 [-0.92, -0.29]
			Irbesartan	109	80 (73.4)	-0.16 (1.88)	-6.0	-1.25	-0.36	0.88	4.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline use of antihypertensives													
Yes	Urine protein excretion	Baseline	Sparsentan	90	90 (100.0)	2.59 (2.05)	0.4	1.31	2.18	3.16	14.7		
			Irbesartan	88	88 (100.0)	2.20 (1.42)	0.5	1.30	1.74	2.57	7.5		
		Week 6	Sparsentan	90	86 (95.6)	1.64 (1.32)	0.2	0.74	1.16	2.16	6.5		
			Irbesartan	88	82 (93.2)	2.29 (1.84)	0.4	1.13	1.64	2.87	10.3		
		Week 36	Sparsentan	90	87 (96.7)	1.49 (1.41)	0.1	0.62	0.96	1.92	8.3		
			Irbesartan	88	82 (93.2)	2.30 (1.85)	0.2	1.01	1.73	2.83	9.2		
		Week 58	Sparsentan	90	83 (92.2)	1.60 (1.43)	0.1	0.69	1.20	1.81	7.5		
			Irbesartan	88	80 (90.9)	2.23 (1.91)	0.1	0.99	1.71	2.76	9.4		
		Week 110	Sparsentan	90	75 (83.3)	1.49 (1.19)	0.1	0.53	1.24	2.26	5.3		
			Irbesartan	88	69 (78.4)	2.32 (2.01)	0.1	0.94	1.71	2.74	9.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	90	86 (95.6)	-0.88 (1.86)	-13.0	-1.45	-0.76	-0.12	5.2	-0.57 [-0.88, -0.26]
				Irbesartan	88	82 (93.2)	0.07 (1.41)	-3.7	-0.57	-0.05	0.46	8.4	
			Week 36	Sparsentan	90	87 (96.7)	-1.01 (1.88)	-13.7	-1.47	-0.76	-0.29	2.6	-0.63 [-0.94, -0.32]
				Irbesartan	88	82 (93.2)	0.09 (1.60)	-3.3	-0.78	-0.14	0.75	6.5	
	Week 58		Sparsentan	90	83 (92.2)	-0.87 (1.96)	-13.3	-1.60	-0.72	-0.02	3.3	-0.52 [-0.83, -0.20]	
			Irbesartan	88	80 (90.9)	0.05 (1.60)	-3.3	-0.74	-0.21	0.53	6.7		
	Week 110	Sparsentan	90	75 (83.3)	-0.99 (2.07)	-13.5	-1.66	-0.71	0.04	1.9	-0.56 [-0.89, -0.22]		
		Irbesartan	88	69 (78.4)	0.12 (1.90)	-6.0	-0.91	-0.11	0.95	5.9			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
No	Urine protein excretion	Baseline	Sparsentan	112	112 (100.0)	1.94 (1.17)	0.1	1.10	1.53	2.54	6.4		
			Irbesartan	114	114 (100.0)	2.14 (1.08)	0.6	1.41	1.93	2.63	6.9		
		Week 6	Sparsentan	112	109 (97.3)	1.37 (1.09)	0.2	0.60	1.06	1.77	5.7		
			Irbesartan	114	111 (97.4)	2.10 (1.39)	0.4	1.08	1.84	2.67	7.7		
		Week 36	Sparsentan	112	107 (95.5)	1.32 (1.43)	0.1	0.50	0.85	1.70	9.4		
			Irbesartan	114	105 (92.1)	2.04 (1.62)	0.2	0.95	1.52	2.69	9.2		
		Week 58	Sparsentan	112	105 (93.8)	1.30 (1.32)	0.1	0.44	0.88	1.58	6.8		
			Irbesartan	114	98 (86.0)	2.05 (1.70)	0.2	0.98	1.56	2.75	12.8		
		Week 110	Sparsentan	112	94 (83.9)	1.40 (1.41)	0.1	0.43	0.96	1.65	6.0		
			Irbesartan	114	81 (71.1)	2.25 (1.66)	0.2	1.08	1.81	2.86	9.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	112	109 (97.3)	-0.56 (0.94)	-4.0	-1.00	-0.58	0.02	2.1	-0.49 [-0.76, -0.22]
				Irbesartan	114	111 (97.4)	-0.05 (1.15)	-3.2	-0.68	-0.20	0.44	4.2	
			Week 36	Sparsentan	112	107 (95.5)	-0.64 (1.21)	-4.1	-1.28	-0.67	-0.04	4.6	-0.44 [-0.71, -0.17]
				Irbesartan	114	105 (92.1)	-0.09 (1.29)	-2.5	-0.78	-0.26	0.58	4.3	
			Week 58	Sparsentan	112	105 (93.8)	-0.63 (1.35)	-4.2	-1.29	-0.71	0.02	6.4	-0.41 [-0.69, -0.13]
				Irbesartan	114	98 (86.0)	-0.09 (1.29)	-2.7	-0.84	-0.28	0.46	5.8	
			Week 110	Sparsentan	112	94 (83.9)	-0.45 (1.34)	-3.5	-1.26	-0.56	0.16	3.5	-0.41 [-0.71, -0.11]
				Irbesartan	114	81 (71.1)	0.10 (1.34)	-2.7	-0.65	0.05	0.93	4.2	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Time since renal biopsy													
<= 5 years	Urine protein excretion	Baseline	Sparsentan	113	113 (100.0)	2.28 (1.84)	0.1	1.17	1.67	2.95	14.7		
			Irbesartan	127	127 (100.0)	2.19 (1.32)	0.5	1.37	1.78	2.63	7.5		
		Week 6	Sparsentan	113	108 (95.6)	1.54 (1.30)	0.2	0.64	1.08	1.98	6.5		
			Irbesartan	127	123 (96.9)	2.11 (1.56)	0.4	1.05	1.55	2.69	9.5		
		Week 36	Sparsentan	113	107 (94.7)	1.35 (1.37)	0.1	0.50	0.88	1.72	8.3		
			Irbesartan	127	117 (92.1)	2.11 (1.84)	0.2	0.81	1.44	2.69	9.2		
		Week 58	Sparsentan	113	104 (92.0)	1.52 (1.46)	0.1	0.58	1.15	1.81	7.5		
			Irbesartan	127	116 (91.3)	2.10 (1.98)	0.1	0.91	1.47	2.65	12.8		
		Week 110	Sparsentan	113	96 (85.0)	1.26 (1.08)	0.1	0.43	0.96	1.88	6.0		
			Irbesartan	127	93 (73.2)	2.19 (1.96)	0.1	0.92	1.58	2.64	9.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	113	108 (95.6)	-0.76 (1.70)	-13.0	-1.21	-0.71	-0.00	5.2	-0.48 [-0.74, -0.22]
				Irbesartan	127	123 (96.9)	-0.09 (1.08)	-3.7	-0.66	-0.18	0.43	3.2	
			Week 36	Sparsentan	113	107 (94.7)	-0.97 (1.74)	-13.7	-1.46	-0.83	-0.20	2.6	-0.57 [-0.84, -0.31]
				Irbesartan	127	117 (92.1)	-0.07 (1.37)	-3.3	-0.78	-0.24	0.62	6.5	
	Week 58		Sparsentan	113	104 (92.0)	-0.74 (1.96)	-13.3	-1.44	-0.69	0.12	6.4	-0.40 [-0.66, -0.13]	
			Irbesartan	127	116 (91.3)	-0.05 (1.49)	-3.3	-0.87	-0.28	0.42	5.8		
	Week 110	Sparsentan	113	96 (85.0)	-0.99 (1.96)	-13.5	-1.65	-0.72	-0.15	3.4	-0.54 [-0.83, -0.25]		
		Irbesartan	127	93 (73.2)	0.01 (1.68)	-6.0	-0.80	-0.13	0.76	5.9			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 5 years	Urine protein excretion	Baseline	Sparsentan	89	89 (100.0)	2.16 (1.39)	0.4	1.18	1.77	2.81	9.7		
			Irbesartan	75	75 (100.0)	2.13 (1.09)	0.6	1.20	1.96	2.60	5.9		
		Week 6	Sparsentan	89	87 (97.8)	1.43 (1.07)	0.2	0.72	1.09	1.95	5.7		
			Irbesartan	75	70 (93.3)	2.30 (1.67)	0.5	1.18	1.91	2.74	10.3		
		Week 36	Sparsentan	89	87 (97.8)	1.46 (1.48)	0.1	0.60	1.00	1.92	9.4		
			Irbesartan	75	70 (93.3)	2.24 (1.54)	0.2	1.12	1.86	2.84	7.8		
		Week 58	Sparsentan	89	84 (94.4)	1.32 (1.27)	0.1	0.52	0.90	1.57	6.8		
			Irbesartan	75	62 (82.7)	2.19 (1.39)	0.3	1.22	1.93	2.91	8.6		
		Week 110	Sparsentan	89	73 (82.0)	1.68 (1.55)	0.1	0.53	1.15	2.12	6.0		
			Irbesartan	75	57 (76.0)	2.43 (1.56)	0.3	1.29	2.24	3.21	6.6		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	89	87 (97.8)	-0.63 (0.99)	-4.0	-1.00	-0.59	-0.12	2.1	-0.62 [-0.95, -0.30]
				Irbesartan	75	70 (93.3)	0.15 (1.53)	-2.4	-0.63	-0.09	0.61	8.4	
			Week 36	Sparsentan	89	87 (97.8)	-0.61 (1.26)	-4.1	-1.30	-0.64	-0.07	4.6	-0.50 [-0.82, -0.18]
				Irbesartan	75	70 (93.3)	0.09 (1.54)	-2.5	-0.78	-0.13	0.60	5.1	
			Week 58	Sparsentan	89	84 (94.4)	-0.73 (1.16)	-4.2	-1.33	-0.74	-0.09	2.0	-0.62 [-0.95, -0.28]
				Irbesartan	75	62 (82.7)	0.03 (1.33)	-2.5	-0.73	-0.14	0.52	6.7	
			Week 110	Sparsentan	89	73 (82.0)	-0.30 (1.24)	-2.8	-1.08	-0.55	0.37	3.5	-0.42 [-0.77, -0.07]
				Irbesartan	75	57 (76.0)	0.27 (1.50)	-2.7	-0.39	0.10	1.05	4.7	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: History of hypertension													
Yes	Urine protein excretion	Baseline	Sparsentan	155	155 (100.0)	2.26 (1.66)	0.1	1.18	1.85	3.07	14.7		
			Irbesartan	161	161 (100.0)	2.23 (1.31)	0.5	1.34	1.82	2.78	7.5		
		Week 6	Sparsentan	155	149 (96.1)	1.53 (1.21)	0.2	0.70	1.09	2.02	6.5		
			Irbesartan	161	153 (95.0)	2.22 (1.66)	0.4	1.17	1.81	2.74	10.3		
		Week 36	Sparsentan	155	149 (96.1)	1.46 (1.37)	0.1	0.60	0.97	1.72	8.3		
			Irbesartan	161	150 (93.2)	2.27 (1.84)	0.2	1.01	1.70	2.85	9.2		
		Week 58	Sparsentan	155	145 (93.5)	1.54 (1.41)	0.1	0.64	1.13	1.81	7.5		
			Irbesartan	161	145 (90.1)	2.27 (1.92)	0.1	0.98	1.74	2.95	12.8		
		Week 110	Sparsentan	155	131 (84.5)	1.50 (1.31)	0.1	0.53	1.12	2.12	6.0		
			Irbesartan	161	122 (75.8)	2.38 (1.92)	0.1	1.11	1.84	2.91	9.2		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	155	149 (96.1)	-0.73 (1.54)	-13.0	-1.29	-0.69	-0.03	5.2	-0.49 [-0.72, -0.27]
				Irbesartan	161	153 (95.0)	-0.02 (1.31)	-3.7	-0.66	-0.18	0.41	8.4	
			Week 36	Sparsentan	155	149 (96.1)	-0.82 (1.62)	-13.7	-1.42	-0.67	-0.07	2.6	-0.55 [-0.78, -0.32]
				Irbesartan	161	150 (93.2)	0.04 (1.51)	-3.3	-0.78	-0.20	0.60	6.5	
	Week 58		Sparsentan	155	145 (93.5)	-0.71 (1.77)	-13.3	-1.32	-0.59	-0.01	6.4	-0.46 [-0.69, -0.23]	
			Irbesartan	161	145 (90.1)	0.05 (1.51)	-2.7	-0.79	-0.23	0.54	6.7		
	Week 110	Sparsentan	155	131 (84.5)	-0.74 (1.84)	-13.5	-1.48	-0.59	0.13	3.5	-0.49 [-0.74, -0.24]		
		Irbesartan	161	122 (75.8)	0.13 (1.70)	-6.0	-0.77	-0.05	1.00	5.9			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSHM: Course of urine protein excretion by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
No	Urine protein excretion	Baseline	Sparsentan	47	47 (100.0)	2.13 (1.62)	0.4	1.15	1.58	2.63	9.7		
			Irbesartan	41	41 (100.0)	1.93 (0.90)	0.6	1.25	1.82	2.43	4.6		
		Week 6	Sparsentan	47	46 (97.9)	1.36 (1.18)	0.2	0.58	1.01	1.60	5.7		
			Irbesartan	41	40 (97.6)	2.03 (1.34)	0.5	0.99	1.76	2.57	7.2		
		Week 36	Sparsentan	47	45 (95.7)	1.17 (1.55)	0.1	0.31	0.69	1.61	9.4		
			Irbesartan	41	37 (90.2)	1.70 (1.10)	0.2	0.92	1.48	2.48	4.4		
		Week 58	Sparsentan	47	43 (91.5)	1.07 (1.19)	0.1	0.23	0.82	1.40	6.8		
			Irbesartan	41	33 (80.5)	1.53 (0.86)	0.3	0.98	1.33	1.98	4.6		
		Week 110	Sparsentan	47	38 (80.9)	1.24 (1.35)	0.1	0.30	0.76	1.67	5.7		
			Irbesartan	41	28 (68.3)	1.86 (1.26)	0.3	0.99	1.39	2.56	4.8		
		Change from baseline in urine protein excretion	Week 6	Sparsentan	47	46 (97.9)	-0.61 (0.95)	-4.0	-1.00	-0.54	-0.22	2.1	-0.69 [-1.13, -0.26]
				Irbesartan	41	40 (97.6)	0.09 (1.07)	-2.4	-0.57	0.04	0.73	2.9	
			Week 36	Sparsentan	47	45 (95.7)	-0.78 (1.32)	-4.1	-1.32	-0.82	-0.39	4.6	-0.46 [-0.90, -0.02]
				Irbesartan	41	37 (90.2)	-0.22 (1.04)	-2.5	-0.76	-0.24	0.62	1.6	
			Week 58	Sparsentan	47	43 (91.5)	-0.83 (1.12)	-4.2	-1.45	-0.76	-0.09	2.0	-0.45 [-0.91, 0.01]
				Irbesartan	41	33 (80.5)	-0.35 (1.01)	-3.3	-0.84	-0.30	0.24	2.3	
			Week 110	Sparsentan	47	38 (80.9)	-0.52 (1.22)	-3.1	-1.16	-0.75	-0.04	3.2	-0.44 [-0.93, 0.05]
				Irbesartan	41	28 (68.3)	0.02 (1.21)	-3.4	-0.57	0.07	0.35	2.8	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Sex	Overall	Sparsentan						Interaction:	0.037	#
Male	Week 6	Sparsentan	139	135 (97.1)	-0.72 (0.12)	(-0.96, -0.47)	-0.80 (0.18)	(-1.14, -0.46)	<0.001	*
		Irbesartan	143	137 (95.8)	0.08 (0.12)	(-0.16, 0.32)				
	Week 36	Sparsentan	139	135 (97.1)	-0.76 (0.12)	(-1.01, -0.52)	-0.84 (0.18)	(-1.19, -0.50)	<0.001	*
		Irbesartan	143	132 (92.3)	0.08 (0.12)	(-0.16, 0.33)				
Week 58	Sparsentan	139	129 (92.8)	-0.63 (0.13)	(-0.88, -0.38)	-0.77 (0.18)	(-1.12, -0.41)	<0.001	*	
	Irbesartan	143	127 (88.8)	0.13 (0.13)	(-0.11, 0.38)					
Week 110	Sparsentan	139	116 (83.5)	-0.73 (0.13)	(-0.99, -0.47)	-0.91 (0.19)	(-1.29, -0.54)	<0.001	*	
	Irbesartan	143	108 (75.5)	0.18 (0.14)	(-0.09, 0.45)					
Female	Week 6	Sparsentan	63	60 (95.2)	-0.55 (0.14)	(-0.83, -0.26)	-0.27 (0.21)	(-0.68, 0.14)	0.200	
		Irbesartan	59	56 (94.9)	-0.28 (0.15)	(-0.57, 0.01)				
	Week 36	Sparsentan	63	59 (93.7)	-0.82 (0.14)	(-1.11, -0.54)	-0.60 (0.21)	(-1.02, -0.19)	0.004	*
		Irbesartan	59	55 (93.2)	-0.22 (0.15)	(-0.51, 0.08)				
Week 58	Sparsentan	63	59 (93.7)	-0.81 (0.15)	(-1.09, -0.52)	-0.50 (0.21)	(-0.92, -0.09)	0.018	*	
	Irbesartan	59	51 (86.4)	-0.30 (0.15)	(-0.61, -0.00)					
Week 110	Sparsentan	63	53 (84.1)	-0.59 (0.15)	(-0.88, -0.29)	-0.37 (0.23)	(-0.81, 0.07)	0.102		
	Irbesartan	59	42 (71.2)	-0.22 (0.17)	(-0.55, 0.11)					

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.656
<= 45 years	Week 6	Sparsentan	96	93 (96.9)	-0.77 (0.16)	(-1.08, -0.46)	-0.69 (0.22)	(-1.13, -0.26)	0.002 *
		Irbesartan	99	95 (96.0)	-0.08 (0.15)	(-0.38, 0.23)			
	Week 36	Sparsentan	96	91 (94.8)	-0.81 (0.16)	(-1.12, -0.51)	-0.62 (0.22)	(-1.05, -0.18)	0.005 *
		Irbesartan	99	91 (91.9)	-0.20 (0.16)	(-0.50, 0.11)			
	Week 58	Sparsentan	96	89 (92.7)	-0.66 (0.16)	(-0.97, -0.35)	-0.52 (0.23)	(-0.97, -0.08)	0.020 *
		Irbesartan	99	84 (84.8)	-0.13 (0.16)	(-0.45, 0.18)			
	Week 110	Sparsentan	96	76 (79.2)	-0.68 (0.17)	(-1.01, -0.35)	-0.68 (0.24)	(-1.16, -0.20)	0.006 *
		Irbesartan	99	69 (69.7)	-0.00 (0.18)	(-0.35, 0.35)			
> 45 years	Week 6	Sparsentan	106	102 (96.2)	-0.56 (0.12)	(-0.79, -0.32)	-0.57 (0.17)	(-0.91, -0.24)	<0.001 *
		Irbesartan	103	98 (95.1)	0.02 (0.12)	(-0.22, 0.26)			
	Week 36	Sparsentan	106	103 (97.2)	-0.74 (0.12)	(-0.97, -0.50)	-0.90 (0.17)	(-1.24, -0.56)	<0.001 *
		Irbesartan	103	96 (93.2)	0.16 (0.12)	(-0.08, 0.40)			
	Week 58	Sparsentan	106	99 (93.4)	-0.70 (0.12)	(-0.93, -0.46)	-0.82 (0.17)	(-1.17, -0.48)	<0.001 *
		Irbesartan	103	94 (91.3)	0.13 (0.12)	(-0.12, 0.37)			
	Week 110	Sparsentan	106	93 (87.7)	-0.68 (0.12)	(-0.93, -0.44)	-0.80 (0.18)	(-1.16, -0.45)	<0.001 *
		Irbesartan	103	81 (78.6)	0.12 (0.13)	(-0.14, 0.38)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.077
<= 18 years	Week 6	Sparsentan	9	8 (88.9)	-0.70 (0.65)	(-2.02, 0.61)	-0.40 (1.10)	(-2.62, 1.82)	0.717
		Irbesartan	5	5 (100.0)	-0.30 (0.83)	(-1.98, 1.38)			
	Week 36	Sparsentan	9	8 (88.9)	-0.69 (0.65)	(-2.00, 0.62)	-0.47 (1.09)	(-2.67, 1.73)	0.667
		Irbesartan	5	5 (100.0)	-0.22 (0.83)	(-1.90, 1.46)			
	Week 58	Sparsentan	9	7 (77.8)	-0.46 (0.68)	(-1.81, 0.90)	0.12 (1.13)	(-2.16, 2.40)	0.916
		Irbesartan	5	4 (80.0)	-0.58 (0.87)	(-2.33, 1.18)			
	Week 110	Sparsentan	9	5 (55.6)	0.46 (0.77)	(-1.08, 2.00)	1.22 (1.22)	(-1.23, 3.67)	0.321
		Irbesartan	5	4 (80.0)	-0.76 (0.92)	(-2.60, 1.07)			
> 18 to 40 years	Week 6	Sparsentan	102	100 (98.0)	-0.83 (0.14)	(-1.12, -0.55)	-0.74 (0.20)	(-1.13, -0.34)	<0.001 *
		Irbesartan	109	105 (96.3)	-0.10 (0.14)	(-0.37, 0.18)			
	Week 36	Sparsentan	102	97 (95.1)	-0.82 (0.15)	(-1.10, -0.54)	-0.64 (0.20)	(-1.04, -0.24)	0.002 *
		Irbesartan	109	100 (91.7)	-0.18 (0.14)	(-0.46, 0.10)			
	Week 58	Sparsentan	102	95 (93.1)	-0.76 (0.15)	(-1.05, -0.48)	-0.62 (0.21)	(-1.03, -0.21)	0.003 *
		Irbesartan	109	92 (84.4)	-0.14 (0.15)	(-0.43, 0.14)			
	Week 110	Sparsentan	102	82 (80.4)	-0.74 (0.16)	(-1.04, -0.43)	-0.74 (0.22)	(-1.17, -0.31)	<0.001 *
		Irbesartan	109	79 (72.5)	0.00 (0.16)	(-0.31, 0.31)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
> 40 years	Week 6	Sparsentan	91	87 (95.6)	-0.49 (0.13)	(-0.74, -0.23)	-0.57 (0.19)	(-0.94, -0.21)	0.002	*
		Irbesartan	88	83 (94.3)	0.08 (0.13)	(-0.18, 0.34)				
	Week 36	Sparsentan	91	89 (97.8)	-0.75 (0.13)	(-1.00, -0.49)	-0.96 (0.19)	(-1.33, -0.60)	<0.001	*
		Irbesartan	88	82 (93.2)	0.22 (0.13)	(-0.04, 0.48)				
	Week 58	Sparsentan	91	86 (94.5)	-0.61 (0.13)	(-0.87, -0.36)	-0.84 (0.19)	(-1.21, -0.47)	<0.001	*
		Irbesartan	88	82 (93.2)	0.22 (0.13)	(-0.04, 0.49)				
	Week 110	Sparsentan	91	82 (90.1)	-0.70 (0.13)	(-0.96, -0.44)	-0.91 (0.20)	(-1.30, -0.52)	<0.001	*
		Irbesartan	88	67 (76.1)	0.21 (0.15)	(-0.07, 0.50)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

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Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.061
North America	Week 6	Sparsentan	35	35 (100.0)	-0.63 (0.29)	(-1.21, -0.05)	-0.60 (0.39)	(-1.37, 0.17)	0.124
		Irbesartan	46	45 (97.8)	-0.03 (0.26)	(-0.53, 0.48)			
	Week 36	Sparsentan	35	33 (94.3)	-1.17 (0.30)	(-1.75, -0.58)	-1.39 (0.40)	(-2.17, -0.61)	<0.001 *
		Irbesartan	46	43 (93.5)	0.22 (0.26)	(-0.29, 0.74)			
	Week 58	Sparsentan	35	28 (80.0)	-0.77 (0.31)	(-1.39, -0.16)	-0.94 (0.41)	(-1.76, -0.13)	0.023 *
		Irbesartan	46	40 (87.0)	0.17 (0.27)	(-0.36, 0.70)			
	Week 110	Sparsentan	35	27 (77.1)	-1.06 (0.33)	(-1.70, -0.42)	-1.01 (0.45)	(-1.89, -0.14)	0.024 *
		Irbesartan	46	31 (67.4)	-0.05 (0.30)	(-0.65, 0.55)			
Europe	Week 6	Sparsentan	98	92 (93.9)	-0.74 (0.14)	(-1.01, -0.48)	-0.70 (0.18)	(-1.06, -0.34)	<0.001 *
		Irbesartan	115	109 (94.8)	-0.04 (0.13)	(-0.29, 0.20)			
	Week 36	Sparsentan	98	94 (95.9)	-0.71 (0.13)	(-0.97, -0.44)	-0.63 (0.18)	(-0.99, -0.27)	<0.001 *
		Irbesartan	115	108 (93.9)	-0.08 (0.13)	(-0.32, 0.17)			
	Week 58	Sparsentan	98	93 (94.9)	-0.61 (0.14)	(-0.88, -0.34)	-0.61 (0.19)	(-0.97, -0.24)	0.001 *
		Irbesartan	115	100 (87.0)	-0.00 (0.13)	(-0.25, 0.25)			
	Week 110	Sparsentan	98	80 (81.6)	-0.65 (0.14)	(-0.93, -0.37)	-0.74 (0.20)	(-1.12, -0.35)	<0.001 *
		Irbesartan	115	88 (76.5)	0.09 (0.14)	(-0.18, 0.36)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Asia Pacific	Week 6	Sparsentan	69	68 (98.6)	-0.61 (0.14)	(-0.88, -0.34)	-0.61 (0.22)	(-1.05, -0.17)	0.007	*
		Irbesartan	41	39 (95.1)	-0.00 (0.18)	(-0.35, 0.35)				
	Week 36	Sparsentan	69	67 (97.1)	-0.70 (0.14)	(-0.97, -0.44)	-0.60 (0.23)	(-1.04, -0.15)	0.009	*
		Irbesartan	41	36 (87.8)	-0.11 (0.18)	(-0.46, 0.25)				
	Week 58	Sparsentan	69	67 (97.1)	-0.76 (0.14)	(-1.03, -0.49)	-0.56 (0.23)	(-1.00, -0.11)	0.015	*
		Irbesartan	41	38 (92.7)	-0.20 (0.18)	(-0.56, 0.16)				
	Week 110	Sparsentan	69	62 (89.9)	-0.59 (0.14)	(-0.86, -0.31)	-0.70 (0.24)	(-1.17, -0.22)	0.004	*
		Irbesartan	41	31 (75.6)	0.11 (0.20)	(-0.28, 0.49)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

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Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline BMI	Overall	Sparsentan						Interaction:	0.050	#
< 27 kg/m**2	Week 6	Sparsentan	83	78 (94.0)	-0.56 (0.12)	(-0.80, -0.32)	-0.48 (0.17)	(-0.81, -0.15)	0.005	*
		Irbesartan	94	89 (94.7)	-0.08 (0.12)	(-0.31, 0.15)				
	Week 36	Sparsentan	83	80 (96.4)	-0.75 (0.12)	(-0.99, -0.51)	-0.47 (0.17)	(-0.80, -0.13)	0.006	*
		Irbesartan	94	85 (90.4)	-0.29 (0.12)	(-0.52, -0.06)				
	Week 58	Sparsentan	83	76 (91.6)	-0.78 (0.12)	(-1.02, -0.53)	-0.50 (0.17)	(-0.84, -0.16)	0.004	*
		Irbesartan	94	79 (84.0)	-0.28 (0.12)	(-0.52, -0.04)				
	Week 110	Sparsentan	83	70 (84.3)	-0.53 (0.13)	(-0.79, -0.28)	-0.36 (0.18)	(-0.72, 0.00)	0.052	
		Irbesartan	94	66 (70.2)	-0.17 (0.13)	(-0.43, 0.08)				
≥ 27 kg/m**2	Week 6	Sparsentan	119	117 (98.3)	-0.73 (0.14)	(-1.00, -0.45)	-0.71 (0.21)	(-1.11, -0.31)	<0.001	*
		Irbesartan	107	103 (96.3)	-0.02 (0.15)	(-0.31, 0.28)				
	Week 36	Sparsentan	119	114 (95.8)	-0.79 (0.14)	(-1.07, -0.51)	-0.98 (0.21)	(-1.39, -0.58)	<0.001	*
		Irbesartan	107	101 (94.4)	0.19 (0.15)	(-0.10, 0.49)				
	Week 58	Sparsentan	119	112 (94.1)	-0.61 (0.14)	(-0.90, -0.33)	-0.82 (0.21)	(-1.23, -0.41)	<0.001	*
		Irbesartan	107	99 (92.5)	0.20 (0.15)	(-0.09, 0.50)				
	Week 110	Sparsentan	119	99 (83.2)	-0.79 (0.15)	(-1.09, -0.50)	-1.03 (0.22)	(-1.47, -0.60)	<0.001	*
		Irbesartan	107	83 (77.6)	0.24 (0.16)	(-0.08, 0.56)				

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Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
					Change from Baseline		Treatment Difference			
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Randomization strata	Overall	Sparsentan								Interaction: 0.011 #
eGFR Low and UP High	Week 6	Sparsentan	71	68 (95.8)	-0.86 (0.20)	(-1.25, -0.47)	-0.85 (0.28)	(-1.40, -0.31)	0.002	*
		Irbesartan	74	73 (98.6)	-0.01 (0.19)	(-0.39, 0.37)				
	Week 36	Sparsentan	71	68 (95.8)	-0.95 (0.20)	(-1.34, -0.55)	-0.99 (0.28)	(-1.54, -0.44)	<0.001	*
		Irbesartan	74	68 (91.9)	0.04 (0.20)	(-0.35, 0.43)				
	Week 58	Sparsentan	71	67 (94.4)	-0.89 (0.20)	(-1.28, -0.49)	-0.93 (0.28)	(-1.49, -0.38)	0.001	*
		Irbesartan	74	64 (86.5)	0.05 (0.20)	(-0.35, 0.44)				
Week 110	Sparsentan	71	59 (83.1)	-0.75 (0.21)	(-1.17, -0.33)	-0.65 (0.31)	(-1.25, -0.05)	0.034	*	
	Irbesartan	74	54 (73.0)	-0.10 (0.22)	(-0.53, 0.33)					
eGFR Low and UP Low	Week 6	Sparsentan	55	52 (94.5)	-0.46 (0.12)	(-0.69, -0.24)	-0.27 (0.16)	(-0.59, 0.05)	0.102	
		Irbesartan	55	53 (96.4)	-0.20 (0.12)	(-0.42, 0.03)				
	Week 36	Sparsentan	55	51 (92.7)	-0.53 (0.12)	(-0.75, -0.30)	-0.50 (0.16)	(-0.82, -0.18)	0.003	*
		Irbesartan	55	51 (92.7)	-0.03 (0.12)	(-0.25, 0.20)				
	Week 58	Sparsentan	55	52 (94.5)	-0.53 (0.12)	(-0.75, -0.30)	-0.45 (0.16)	(-0.77, -0.13)	0.006	*
		Irbesartan	55	51 (92.7)	-0.08 (0.12)	(-0.30, 0.15)				
	Week 110	Sparsentan	55	46 (83.6)	-0.41 (0.12)	(-0.65, -0.17)	-0.38 (0.17)	(-0.72, -0.03)	0.031	*
		Irbesartan	55	42 (76.4)	-0.03 (0.13)	(-0.28, 0.21)				

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
eGFR High and UP High	Week 6	Sparsentan	37	36 (97.3)	-0.79 (0.30)	(-1.37, -0.21)	-0.90 (0.43)	(-1.74, -0.06)	0.036	*
		Irbesartan	36	32 (88.9)	0.11 (0.31)	(-0.50, 0.71)				
	Week 36	Sparsentan	37	37 (100.0)	-1.05 (0.29)	(-1.63, -0.47)	-1.08 (0.43)	(-1.92, -0.24)	0.012	*
		Irbesartan	36	32 (88.9)	0.03 (0.31)	(-0.58, 0.64)				
	Week 58	Sparsentan	37	32 (86.5)	-0.87 (0.30)	(-1.47, -0.28)	-0.87 (0.44)	(-1.73, -0.00)	0.049	*
		Irbesartan	36	31 (86.1)	-0.01 (0.32)	(-0.63, 0.62)				
	Week 110	Sparsentan	37	30 (81.1)	-1.26 (0.32)	(-1.89, -0.64)	-1.58 (0.46)	(-2.49, -0.67)	<0.001	*
		Irbesartan	36	27 (75.0)	0.31 (0.34)	(-0.35, 0.97)				
eGFR High and UP Low	Week 6	Sparsentan	39	39 (100.0)	-0.57 (0.11)	(-0.78, -0.35)	-0.60 (0.16)	(-0.91, -0.29)	<0.001	*
		Irbesartan	37	35 (94.6)	0.03 (0.11)	(-0.19, 0.25)				
	Week 36	Sparsentan	39	38 (97.4)	-0.65 (0.11)	(-0.86, -0.43)	-0.50 (0.16)	(-0.81, -0.19)	0.002	*
		Irbesartan	37	36 (97.3)	-0.15 (0.11)	(-0.37, 0.07)				
	Week 58	Sparsentan	39	37 (94.9)	-0.46 (0.11)	(-0.67, -0.24)	-0.47 (0.16)	(-0.79, -0.15)	0.004	*
		Irbesartan	37	32 (86.5)	0.02 (0.12)	(-0.21, 0.25)				
	Week 110	Sparsentan	39	34 (87.2)	-0.47 (0.11)	(-0.70, -0.25)	-0.76 (0.17)	(-1.09, -0.42)	<0.001	*
		Irbesartan	37	27 (73.0)	0.29 (0.13)	(0.04, 0.53)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
					Change from Baseline		Treatment Difference			
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline eGFR Group 1	Overall	Sparsentan								Interaction: 0.008 #
< 60 mL/min/1.73 m**2	Week 6	Sparsentan	127	121 (95.3)	-0.63 (0.12)	(-0.87, -0.39)	-0.57 (0.17)	(-0.91, -0.24)	<0.001	*
		Irbesartan	129	126 (97.7)	-0.06 (0.12)	(-0.30, 0.17)				
	Week 36	Sparsentan	127	120 (94.5)	-0.67 (0.12)	(-0.91, -0.43)	-0.66 (0.17)	(-1.00, -0.32)	<0.001	*
		Irbesartan	129	119 (92.2)	-0.01 (0.12)	(-0.25, 0.22)				
	Week 58	Sparsentan	127	120 (94.5)	-0.64 (0.12)	(-0.88, -0.40)	-0.65 (0.17)	(-1.00, -0.31)	<0.001	*
		Irbesartan	129	115 (89.1)	0.01 (0.12)	(-0.23, 0.26)				
Week 110	Sparsentan	127	105 (82.7)	-0.51 (0.13)	(-0.77, -0.26)	-0.50 (0.19)	(-0.86, -0.13)	0.007	*	
	Irbesartan	129	97 (75.2)	-0.02 (0.13)	(-0.28, 0.24)					
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	48 (98.0)	-0.71 (0.21)	(-1.13, -0.30)	-0.85 (0.30)	(-1.45, -0.25)	0.005	*
		Irbesartan	48	44 (91.7)	0.14 (0.22)	(-0.29, 0.56)				
	Week 36	Sparsentan	49	48 (98.0)	-0.99 (0.21)	(-1.40, -0.57)	-1.33 (0.31)	(-1.93, -0.73)	<0.001	*
		Irbesartan	48	45 (93.8)	0.34 (0.22)	(-0.09, 0.77)				
	Week 58	Sparsentan	49	43 (87.8)	-0.77 (0.22)	(-1.20, -0.34)	-1.06 (0.31)	(-1.68, -0.45)	<0.001	*
		Irbesartan	48	42 (87.5)	0.29 (0.22)	(-0.14, 0.73)				
	Week 110	Sparsentan	49	43 (87.8)	-1.07 (0.22)	(-1.50, -0.63)	-1.54 (0.32)	(-2.18, -0.91)	<0.001	*
		Irbesartan	48	36 (75.0)	0.47 (0.24)	(0.01, 0.94)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 110 are included in the model. Only key visits up to Week 110 are displayed. A first order regressive covariance structure was used.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	26 (100.0)	-0.80 (0.22)	(-1.24, -0.37)	-0.65 (0.32)	(-1.28, -0.02)	0.044	*
		Irbesartan	25	23 (92.0)	-0.16 (0.23)	(-0.61, 0.30)				
	Week 36	Sparsentan	26	26 (100.0)	-0.94 (0.22)	(-1.37, -0.51)	-0.28 (0.32)	(-0.90, 0.35)	0.387	
		Irbesartan	25	23 (92.0)	-0.66 (0.23)	(-1.12, -0.21)				
	Week 58	Sparsentan	26	25 (96.2)	-0.79 (0.22)	(-1.23, -0.35)	-0.19 (0.33)	(-0.84, 0.45)	0.558	
		Irbesartan	25	21 (84.0)	-0.60 (0.24)	(-1.07, -0.13)				
	Week 110	Sparsentan	26	21 (80.8)	-0.82 (0.24)	(-1.29, -0.36)	-0.57 (0.35)	(-1.27, 0.13)	0.109	
		Irbesartan	25	17 (68.0)	-0.25 (0.26)	(-0.77, 0.27)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.004 #
< 45 mL/min/1.73 m**2	Week 6	Sparsentan	82	78 (95.1)	-0.54 (0.15)	(-0.83, -0.25)	-0.40 (0.21)	(-0.82, 0.01)	0.056
		Irbesartan	80	78 (97.5)	-0.14 (0.15)	(-0.43, 0.16)			
	Week 36	Sparsentan	82	78 (95.1)	-0.58 (0.15)	(-0.87, -0.29)	-0.59 (0.21)	(-1.01, -0.17)	0.006 *
		Irbesartan	80	72 (90.0)	0.01 (0.15)	(-0.29, 0.30)			
	Week 58	Sparsentan	82	77 (93.9)	-0.53 (0.15)	(-0.82, -0.23)	-0.51 (0.22)	(-0.93, -0.08)	0.019 *
		Irbesartan	80	69 (86.3)	-0.02 (0.15)	(-0.33, 0.28)			
Week 110	Sparsentan	82	67 (81.7)	-0.39 (0.16)	(-0.70, -0.08)	-0.35 (0.23)	(-0.80, 0.10)	0.127	
	Irbesartan	80	59 (73.8)	-0.04 (0.17)	(-0.37, 0.28)				
45 to < 60 mL/min/1.73 m**2	Week 6	Sparsentan	45	43 (95.6)	-0.80 (0.21)	(-1.21, -0.38)	-0.86 (0.29)	(-1.44, -0.29)	0.003 *
		Irbesartan	49	48 (98.0)	0.07 (0.20)	(-0.33, 0.46)			
	Week 36	Sparsentan	45	42 (93.3)	-0.84 (0.21)	(-1.26, -0.42)	-0.81 (0.29)	(-1.39, -0.23)	0.006 *
		Irbesartan	49	47 (95.9)	-0.03 (0.20)	(-0.43, 0.36)			
	Week 58	Sparsentan	45	43 (95.6)	-0.85 (0.21)	(-1.26, -0.43)	-0.93 (0.30)	(-1.51, -0.35)	0.002 *
		Irbesartan	49	46 (93.9)	0.08 (0.21)	(-0.32, 0.48)			
Week 110	Sparsentan	45	38 (84.4)	-0.73 (0.22)	(-1.17, -0.29)	-0.76 (0.32)	(-1.38, -0.14)	0.017 *	
	Irbesartan	49	38 (77.6)	0.03 (0.22)	(-0.41, 0.46)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
					Change from Baseline		Treatment Difference			
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	48 (98.0)	-0.71 (0.21)	(-1.13, -0.30)	-0.85 (0.30)	(-1.45, -0.25)	0.005 *	
		Irbesartan	48	44 (91.7)	0.14 (0.22)	(-0.29, 0.56)				
	Week 36	Sparsentan	49	48 (98.0)	-0.99 (0.21)	(-1.40, -0.57)	-1.33 (0.31)	(-1.93, -0.73)	<0.001 *	
		Irbesartan	48	45 (93.8)	0.34 (0.22)	(-0.09, 0.77)				
	Week 58	Sparsentan	49	43 (87.8)	-0.77 (0.22)	(-1.20, -0.34)	-1.06 (0.31)	(-1.68, -0.45)	<0.001 *	
		Irbesartan	48	42 (87.5)	0.29 (0.22)	(-0.14, 0.73)				
	Week 110	Sparsentan	49	43 (87.8)	-1.07 (0.22)	(-1.50, -0.63)	-1.54 (0.32)	(-2.18, -0.91)	<0.001 *	
		Irbesartan	48	36 (75.0)	0.47 (0.24)	(0.01, 0.94)				
	>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	26 (100.0)	-0.80 (0.22)	(-1.24, -0.37)	-0.65 (0.32)	(-1.28, -0.02)	0.044 *
			Irbesartan	25	23 (92.0)	-0.16 (0.23)	(-0.61, 0.30)			
		Week 36	Sparsentan	26	26 (100.0)	-0.94 (0.22)	(-1.37, -0.51)	-0.28 (0.32)	(-0.90, 0.35)	0.387
			Irbesartan	25	23 (92.0)	-0.66 (0.23)	(-1.12, -0.21)			
Week 58		Sparsentan	26	25 (96.2)	-0.79 (0.22)	(-1.23, -0.35)	-0.19 (0.33)	(-0.84, 0.45)	0.558	
		Irbesartan	25	21 (84.0)	-0.60 (0.24)	(-1.07, -0.13)				
Week 110		Sparsentan	26	21 (80.8)	-0.82 (0.24)	(-1.29, -0.36)	-0.57 (0.35)	(-1.27, 0.13)	0.109	
		Irbesartan	25	17 (68.0)	-0.25 (0.26)	(-0.77, 0.27)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis					
					Change from Baseline		Treatment Difference			
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline urine protein excretion	Overall	Sparsentan								Interaction: <0.001 #
<= 1.75 g/day	Week 6	Sparsentan	98	95 (96.9)	-0.17 (0.10)	(-0.36, 0.03)	-0.33 (0.14)	(-0.61, -0.05)	0.020	*
		Irbesartan	93	88 (94.6)	0.16 (0.10)	(-0.04, 0.36)				
	Week 36	Sparsentan	98	94 (95.9)	-0.24 (0.10)	(-0.43, -0.04)	-0.47 (0.14)	(-0.75, -0.20)	<0.001	*
		Irbesartan	93	87 (93.5)	0.24 (0.10)	(0.04, 0.44)				
	Week 58	Sparsentan	98	93 (94.9)	-0.11 (0.10)	(-0.30, 0.09)	-0.35 (0.14)	(-0.63, -0.07)	0.016	*
		Irbesartan	93	82 (88.2)	0.24 (0.10)	(0.04, 0.45)				
	Week 110	Sparsentan	98	85 (86.7)	-0.07 (0.10)	(-0.27, 0.13)	-0.44 (0.15)	(-0.73, -0.14)	0.004	*
		Irbesartan	93	70 (75.3)	0.37 (0.11)	(0.15, 0.59)				
> 1.75 g/day	Week 6	Sparsentan	104	100 (96.2)	-1.23 (0.18)	(-1.58, -0.87)	-1.06 (0.25)	(-1.56, -0.56)	<0.001	*
		Irbesartan	109	105 (96.3)	-0.16 (0.18)	(-0.51, 0.19)				
	Week 36	Sparsentan	104	100 (96.2)	-1.37 (0.18)	(-1.73, -1.01)	-1.17 (0.26)	(-1.67, -0.66)	<0.001	*
		Irbesartan	109	100 (91.7)	-0.20 (0.18)	(-0.55, 0.15)				
	Week 58	Sparsentan	104	95 (91.3)	-1.30 (0.19)	(-1.66, -0.93)	-1.13 (0.26)	(-1.64, -0.62)	<0.001	*
		Irbesartan	109	96 (88.1)	-0.17 (0.18)	(-0.53, 0.19)				
	Week 110	Sparsentan	104	84 (80.8)	-1.33 (0.19)	(-1.71, -0.95)	-1.16 (0.28)	(-1.70, -0.62)	<0.001	*
		Irbesartan	109	80 (73.4)	-0.17 (0.20)	(-0.56, 0.22)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.413
Yes	Week 6	Sparsentan	90	86 (95.6)	-0.76 (0.16)	(-1.07, -0.45)	-0.72 (0.23)	(-1.17, -0.28)	0.001 *
		Irbesartan	88	82 (93.2)	-0.03 (0.16)	(-0.35, 0.28)			
	Week 36	Sparsentan	90	87 (96.7)	-0.93 (0.16)	(-1.24, -0.62)	-0.94 (0.23)	(-1.38, -0.49)	<0.001 *
		Irbesartan	88	82 (93.2)	0.01 (0.16)	(-0.31, 0.33)			
	Week 58	Sparsentan	90	83 (92.2)	-0.76 (0.16)	(-1.07, -0.45)	-0.73 (0.23)	(-1.18, -0.28)	0.002 *
		Irbesartan	88	80 (90.9)	-0.03 (0.16)	(-0.36, 0.29)			
	Week 110	Sparsentan	90	75 (83.3)	-0.88 (0.17)	(-1.21, -0.55)	-0.91 (0.24)	(-1.38, -0.44)	<0.001 *
		Irbesartan	88	69 (78.4)	0.03 (0.17)	(-0.31, 0.37)			
No	Week 6	Sparsentan	112	109 (97.3)	-0.58 (0.12)	(-0.82, -0.34)	-0.56 (0.17)	(-0.89, -0.22)	0.001 *
		Irbesartan	114	111 (97.4)	-0.03 (0.12)	(-0.26, 0.21)			
	Week 36	Sparsentan	112	107 (95.5)	-0.65 (0.12)	(-0.89, -0.41)	-0.62 (0.17)	(-0.96, -0.28)	<0.001 *
		Irbesartan	114	105 (92.1)	-0.03 (0.12)	(-0.27, 0.21)			
	Week 58	Sparsentan	112	105 (93.8)	-0.62 (0.12)	(-0.86, -0.38)	-0.65 (0.18)	(-0.99, -0.30)	<0.001 *
		Irbesartan	114	98 (86.0)	0.03 (0.13)	(-0.22, 0.27)			
	Week 110	Sparsentan	112	94 (83.9)	-0.52 (0.13)	(-0.78, -0.27)	-0.62 (0.19)	(-0.99, -0.25)	<0.001 *
		Irbesartan	114	81 (71.1)	0.10 (0.14)	(-0.17, 0.37)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 110 are included in the model. Only key visits up to Week 110 are displayed. A first order regressive covariance structure was used.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.955
<= 5 years	Week 6	Sparsentan	113	108 (95.6)	-0.71 (0.14)	(-0.98, -0.45)	-0.59 (0.19)	(-0.96, -0.23)	0.002 *
		Irbesartan	127	123 (96.9)	-0.12 (0.13)	(-0.37, 0.13)			
	Week 36	Sparsentan	113	107 (94.7)	-0.93 (0.14)	(-1.20, -0.66)	-0.82 (0.19)	(-1.19, -0.45)	<0.001 *
		Irbesartan	127	117 (92.1)	-0.11 (0.13)	(-0.36, 0.15)			
	Week 58	Sparsentan	113	104 (92.0)	-0.68 (0.14)	(-0.95, -0.41)	-0.61 (0.19)	(-0.98, -0.24)	0.001 *
		Irbesartan	127	116 (91.3)	-0.07 (0.13)	(-0.33, 0.18)			
	Week 110	Sparsentan	113	96 (85.0)	-0.94 (0.14)	(-1.22, -0.66)	-0.89 (0.20)	(-1.29, -0.49)	<0.001 *
		Irbesartan	127	93 (73.2)	-0.05 (0.14)	(-0.34, 0.23)			
> 5 years	Week 6	Sparsentan	89	87 (97.8)	-0.62 (0.14)	(-0.90, -0.35)	-0.76 (0.21)	(-1.17, -0.35)	<0.001 *
		Irbesartan	75	70 (93.3)	0.14 (0.15)	(-0.16, 0.44)			
	Week 36	Sparsentan	89	87 (97.8)	-0.60 (0.14)	(-0.88, -0.33)	-0.76 (0.21)	(-1.17, -0.35)	<0.001 *
		Irbesartan	75	70 (93.3)	0.16 (0.15)	(-0.14, 0.46)			
	Week 58	Sparsentan	89	84 (94.4)	-0.70 (0.14)	(-0.97, -0.42)	-0.83 (0.21)	(-1.25, -0.40)	<0.001 *
		Irbesartan	75	62 (82.7)	0.13 (0.16)	(-0.19, 0.45)			
	Week 110	Sparsentan	89	73 (82.0)	-0.36 (0.15)	(-0.65, -0.06)	-0.63 (0.23)	(-1.07, -0.19)	0.005 *
		Irbesartan	75	57 (76.0)	0.27 (0.17)	(-0.06, 0.61)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 110 are included in the model. Only key visits up to Week 110 are displayed. A first order regressive covariance structure was used.

An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2UEC\_FSCM: Change from baseline in urine protein excretion by subgroup  
Full Analysis Set

Change from baseline in urine protein excretion					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.137
Yes	Week 6	Sparsentan	155	149 (96.1)	-0.71 (0.12)	(-0.94, -0.48)	-0.65 (0.17)	(-0.97, -0.33)	<0.001 *
		Irbesartan	161	153 (95.0)	-0.06 (0.12)	(-0.28, 0.17)			
	Week 36	Sparsentan	155	149 (96.1)	-0.79 (0.12)	(-1.03, -0.56)	-0.84 (0.17)	(-1.16, -0.51)	<0.001 *
		Irbesartan	161	150 (93.2)	0.04 (0.12)	(-0.18, 0.27)			
	Week 58	Sparsentan	155	145 (93.5)	-0.67 (0.12)	(-0.90, -0.43)	-0.70 (0.17)	(-1.03, -0.37)	<0.001 *
		Irbesartan	161	145 (90.1)	0.04 (0.12)	(-0.20, 0.27)			
	Week 110	Sparsentan	155	131 (84.5)	-0.71 (0.12)	(-0.95, -0.47)	-0.83 (0.18)	(-1.18, -0.48)	<0.001 *
		Irbesartan	161	122 (75.8)	0.12 (0.13)	(-0.13, 0.37)			
No	Week 6	Sparsentan	47	46 (97.9)	-0.55 (0.16)	(-0.87, -0.24)	-0.64 (0.23)	(-1.10, -0.18)	0.007 *
		Irbesartan	41	40 (97.6)	0.09 (0.17)	(-0.25, 0.42)			
	Week 36	Sparsentan	47	45 (95.7)	-0.73 (0.16)	(-1.05, -0.41)	-0.50 (0.24)	(-0.97, -0.04)	0.034 *
		Irbesartan	41	37 (90.2)	-0.23 (0.17)	(-0.57, 0.11)			
	Week 58	Sparsentan	47	43 (91.5)	-0.74 (0.16)	(-1.06, -0.42)	-0.61 (0.24)	(-1.09, -0.14)	0.012 *
		Irbesartan	41	33 (80.5)	-0.13 (0.18)	(-0.48, 0.22)			
	Week 110	Sparsentan	47	38 (80.9)	-0.60 (0.17)	(-0.93, -0.26)	-0.42 (0.26)	(-0.93, 0.09)	0.104
		Irbesartan	41	28 (68.3)	-0.17 (0.19)	(-0.55, 0.21)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 110 are included in the model. Only key visits up to Week 110 are displayed. A first order regressive covariance structure was used.

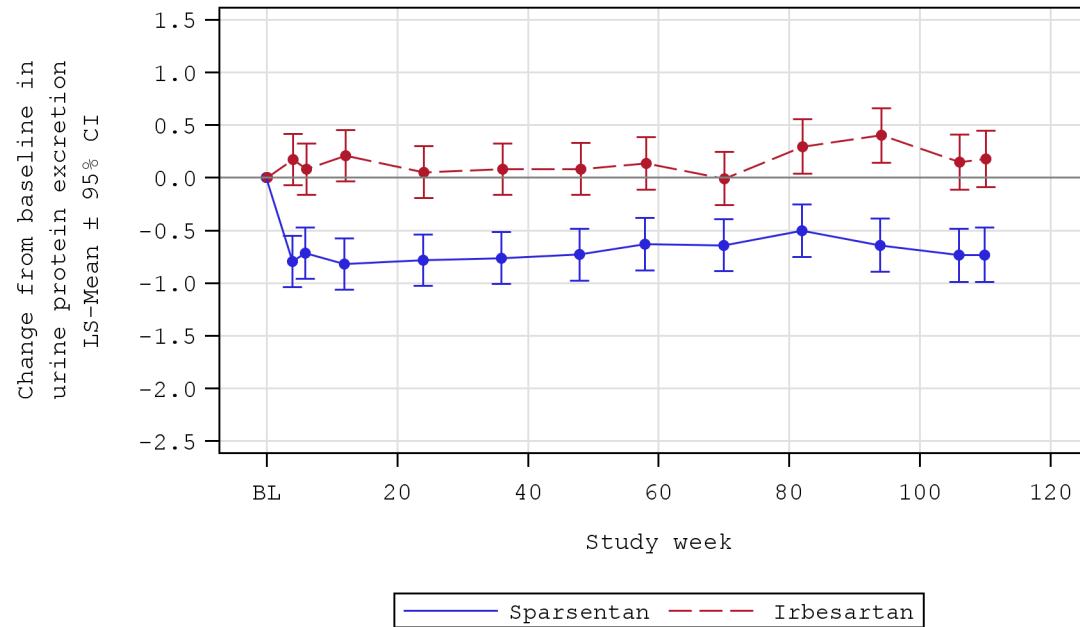
An increase reflects a worsening of the status of the patient. Results are presented in g/day.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

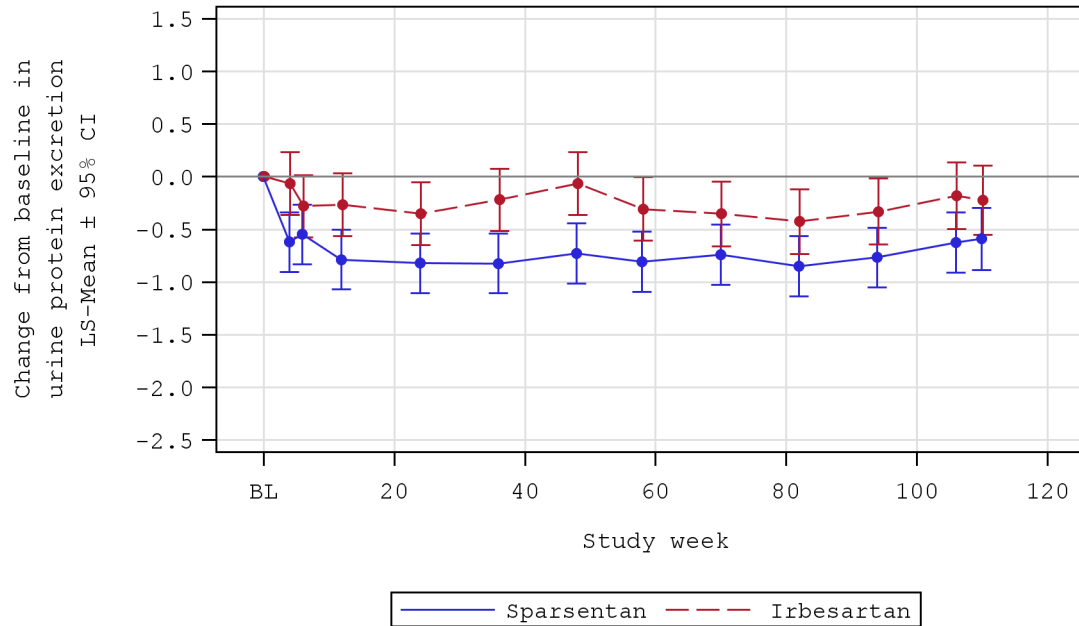
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Sex: Male



Sparsentan	135	135	129	116
Irbesartan	137	132	127	108

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

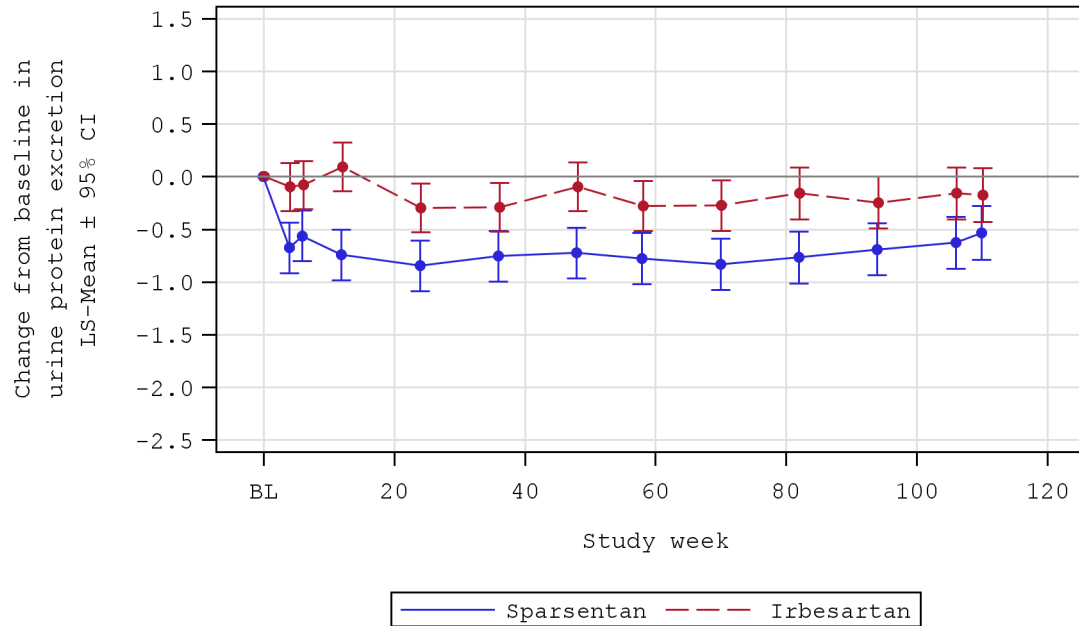
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Sex: Female



Sparsentan	60	59	59	53
Irbesartan	56	55	51	42

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

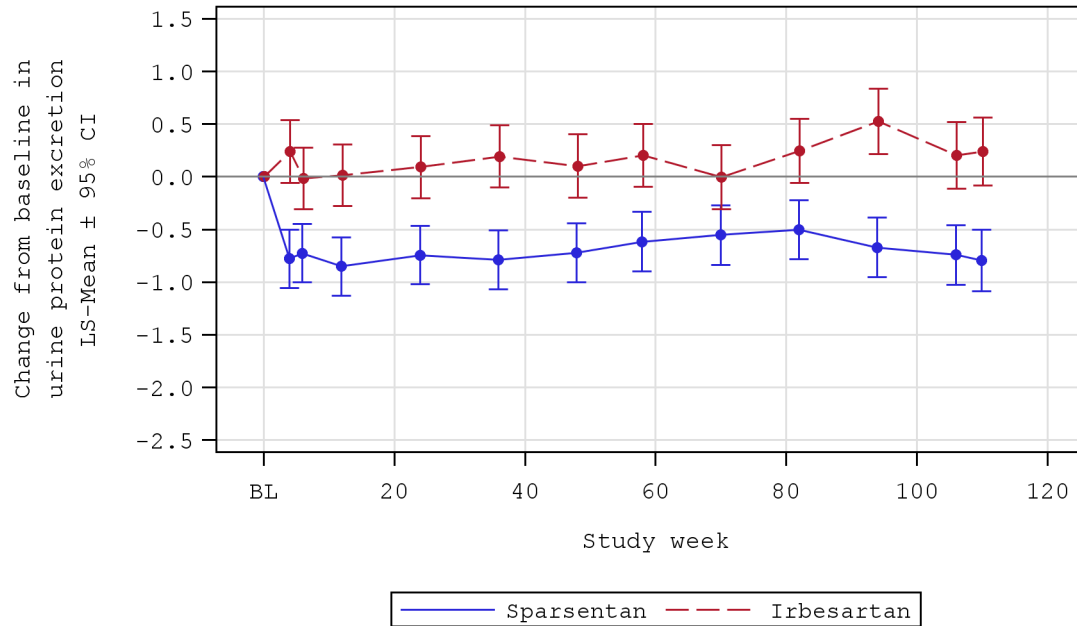
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline BMI: < 27 kg/m\*\*2



Sparsentan	78	80	76	70
Irbesartan	89	85	79	66

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

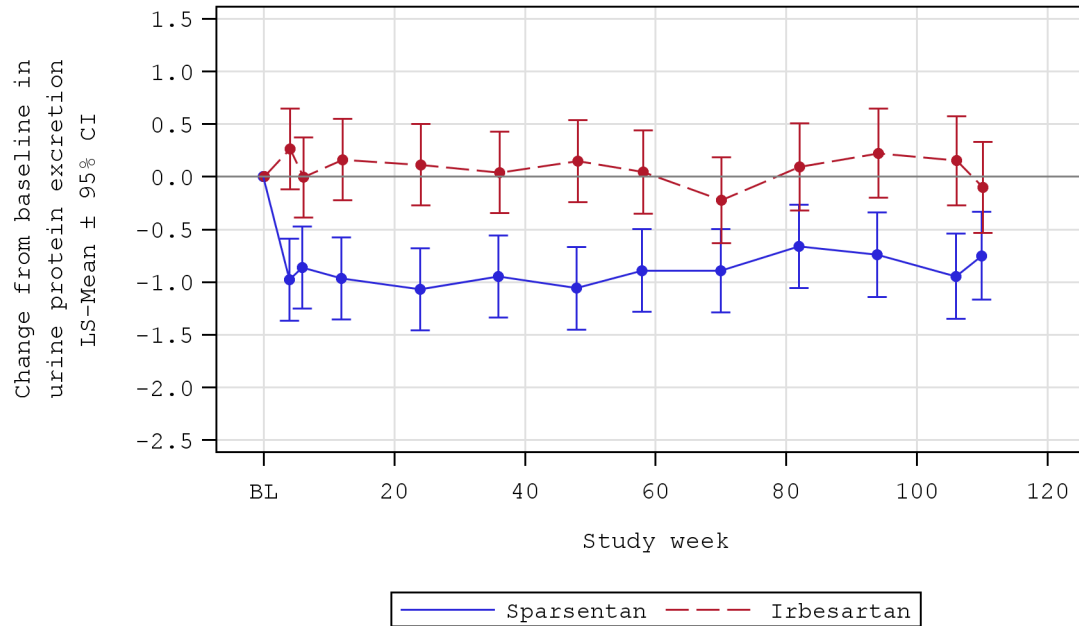
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline BMI:  $\geq 27 \text{ kg/m}^2$



Sparsentan	117	114	112	99
Irbesartan	103	101	99	83

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low =  $30 < \text{eGFR} < 60 \text{ ml/min/1.73m}^2$ , eGFR High  $\geq 60 \text{ ml/min/1.73m}^2$ , UP Low =  $\leq 1.75 \text{ g/day}$ , UP High =  $> 1.75 \text{ g/day}$ .  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

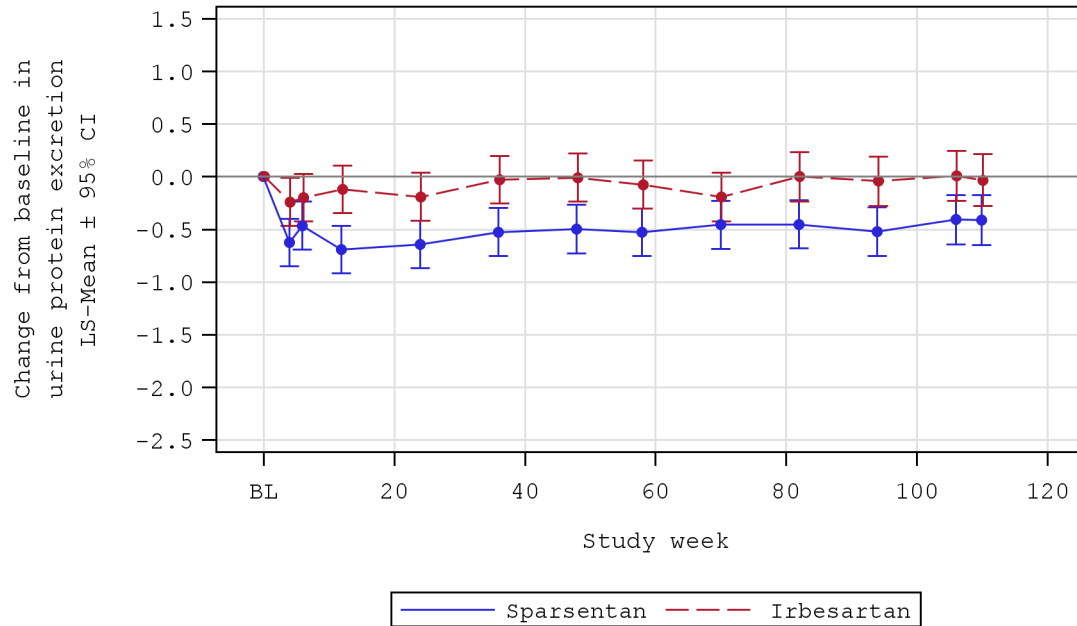
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR Low and UP High



Sparsentan	68	68	67	59
Irbesartan	73	68	64	54

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

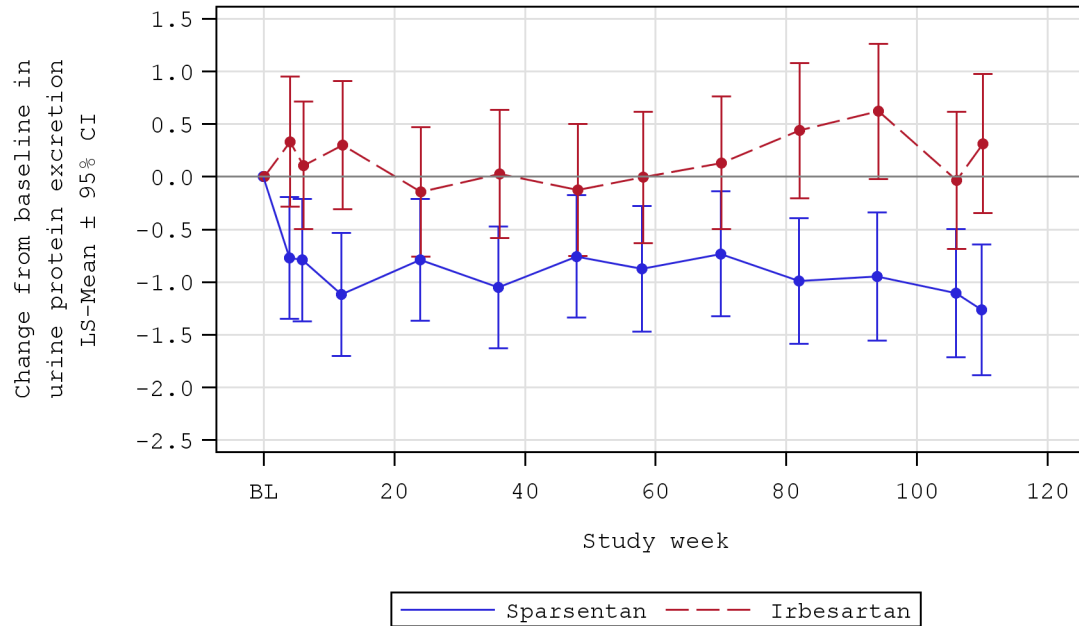
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR Low and UP Low



Sparsentan	52	51	52	46
Irbesartan	53	51	51	42

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR High and UP High

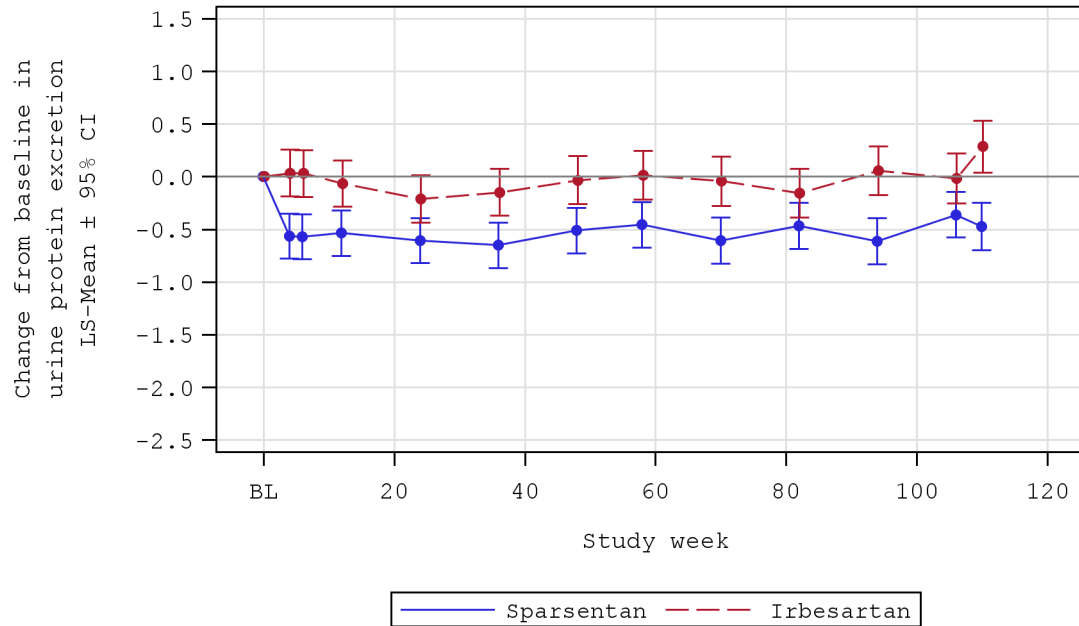


Sparsentan	36	37	32	30
Irbesartan	32	32	31	27

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM



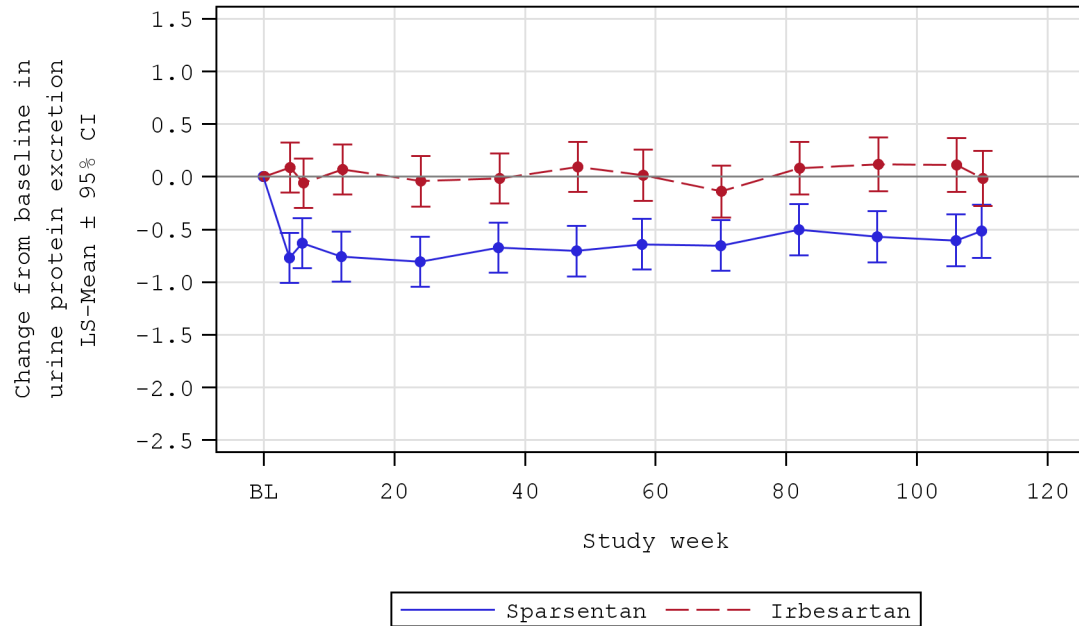
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR High and UP Low



Sparsentan	39	38	37	34
Irbesartan	35	36	32	27

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

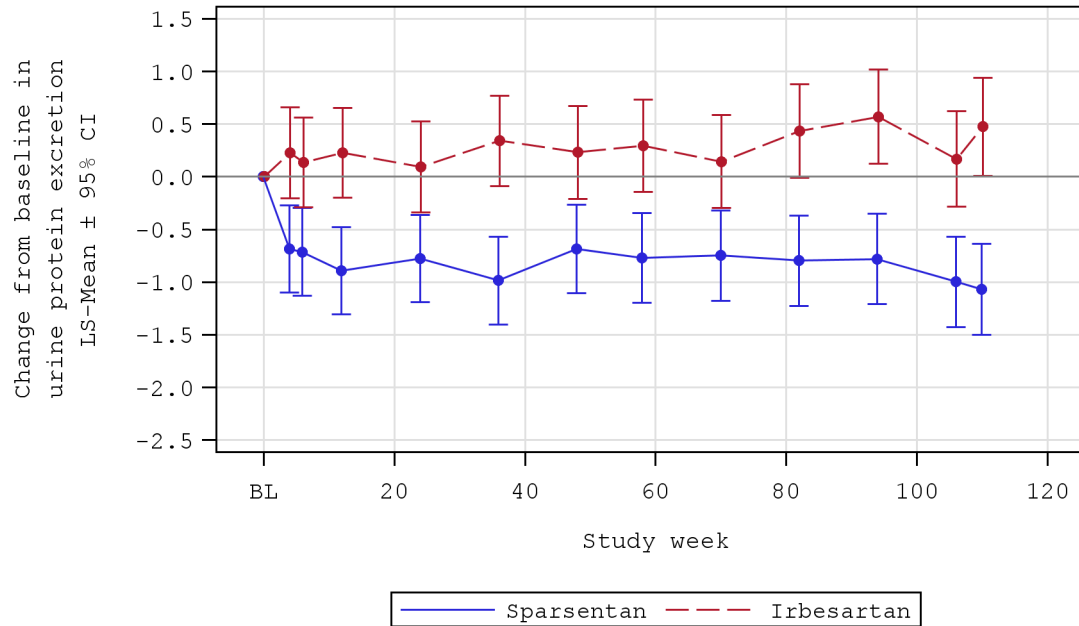
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1: < 60 mL/min/1.73 m\*\*2



Sparsentan	121	120	120	105
Irbesartan	126	119	115	97

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

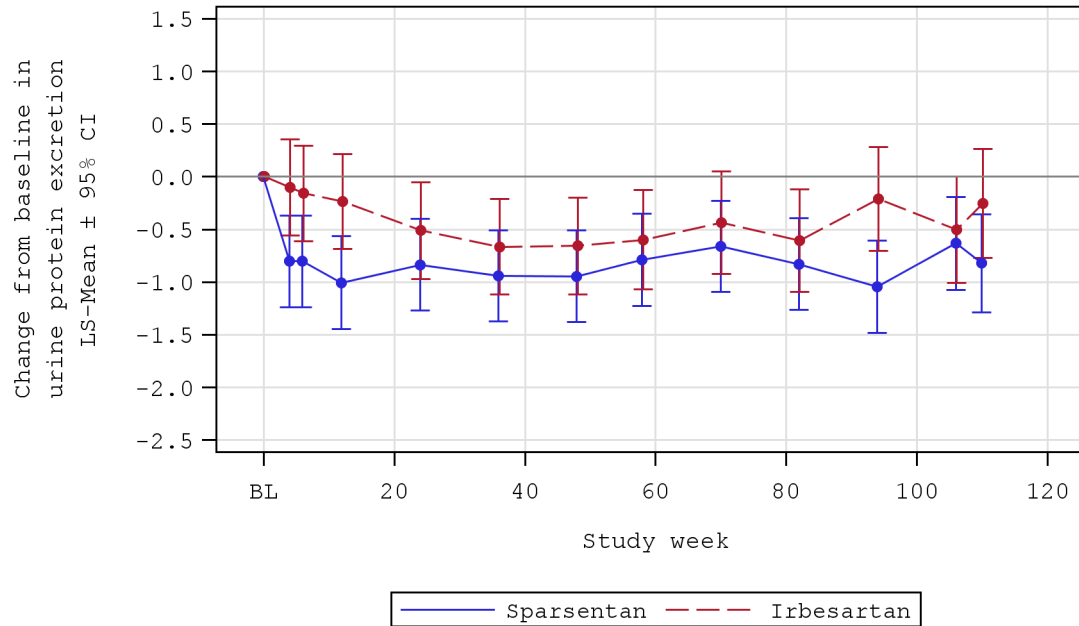
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1: 60 to < 90 mL/min/1.73 m\*\*2



Sparsentan	48	48	43	43
Irbesartan	44	45	42	36

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

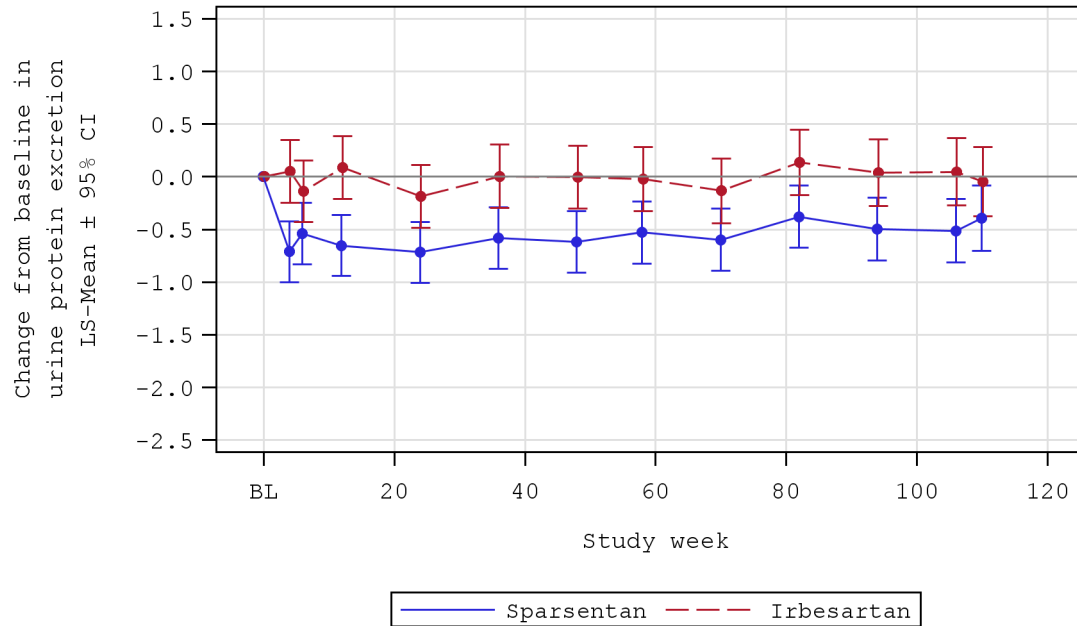
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1:  $\geq 90$  mL/min/1.73 m<sup>2</sup>



Sparsentan	26	26	25	21
Irbesartan	23	23	21	17

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High  $\geq 60$  ml/min/1.73m<sup>2</sup>, UP Low =  $\leq 1.75$  g/day, UP High =  $> 1.75$  g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

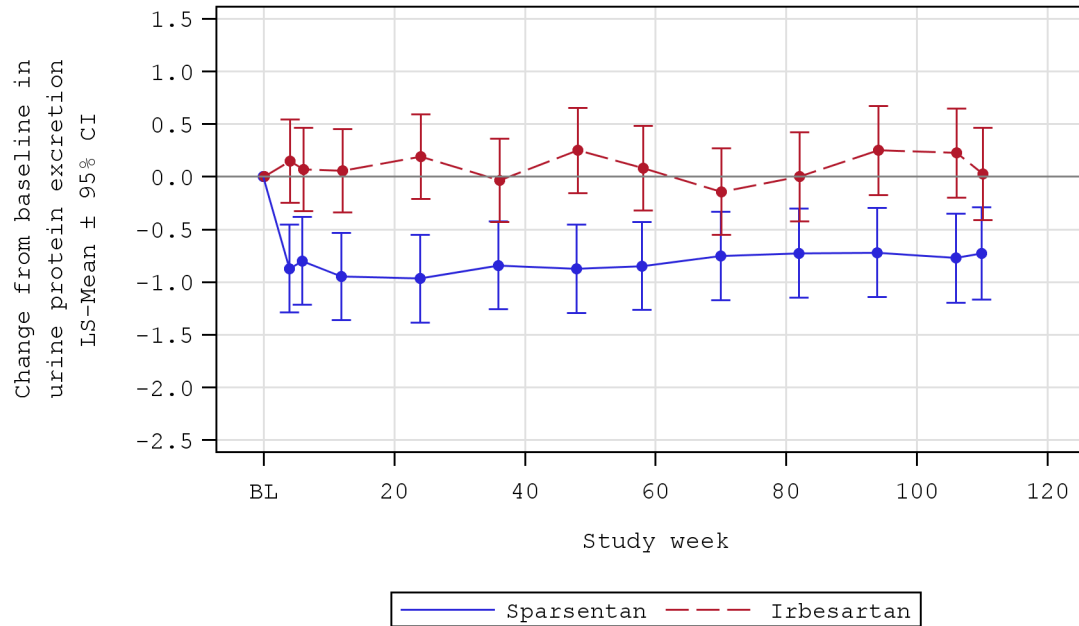
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: < 45 mL/min/1.73 m\*\*2



Sparsentan	78	78	77	67
Irbesartan	78	72	69	59

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

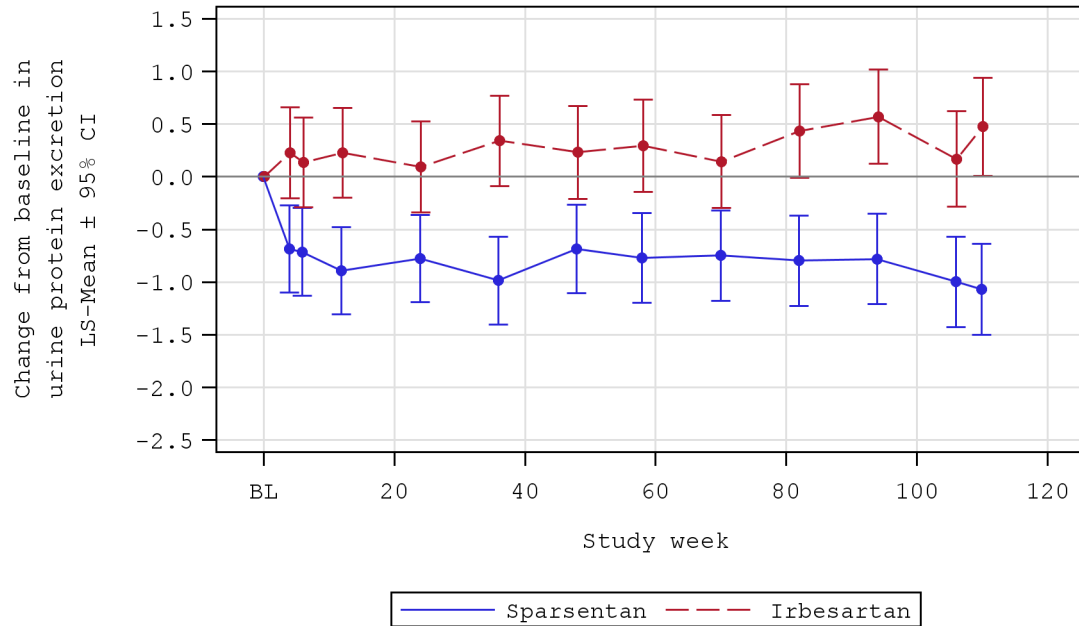
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: 45 to < 60 mL/min/1.73 m\*\*2



Sparsentan	43	42	43	38
Irbesartan	48	47	46	38

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

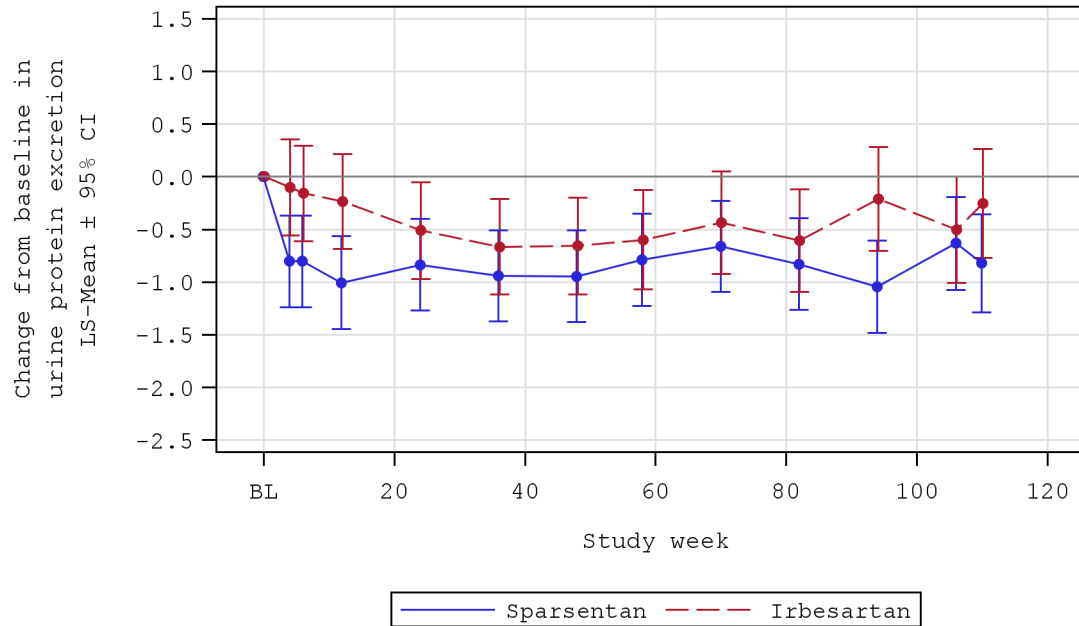
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: 60 to < 90 mL/min/1.73 m\*\*2



Sparsentan	48	48	43	43
Irbesartan	44	45	42	36

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2:  $\geq 90$  mL/min/1.73 m<sup>2</sup>

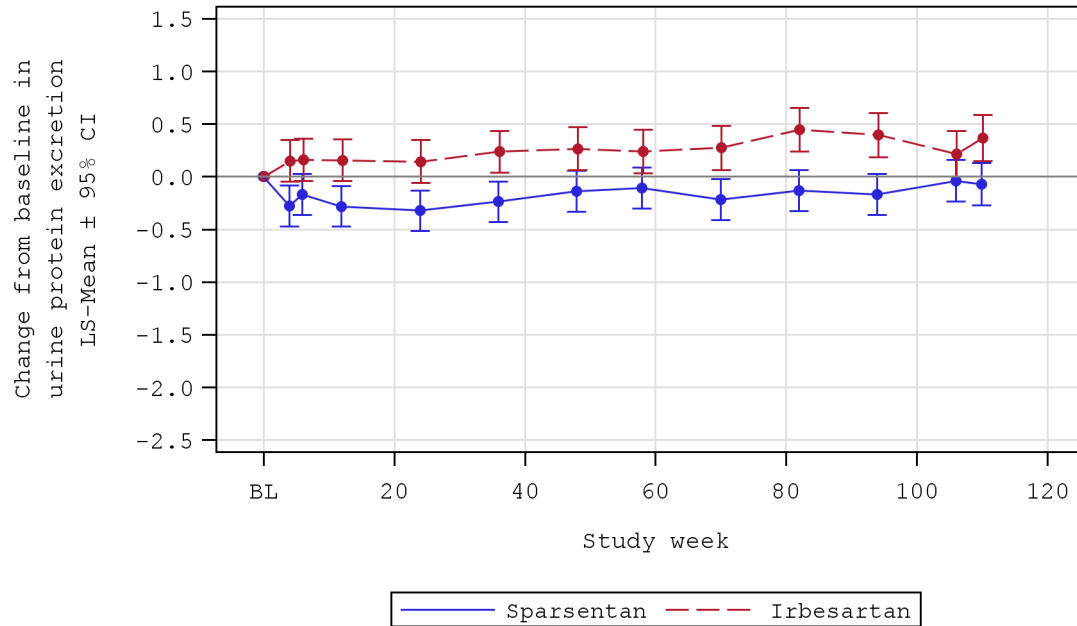


Sparsentan	26	26	25	21
Irbesartan	23	23	21	17

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High  $\geq 60$  ml/min/1.73m<sup>2</sup>, UP Low =  $\leq 1.75$  g/day, UP High =  $> 1.75$  g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM



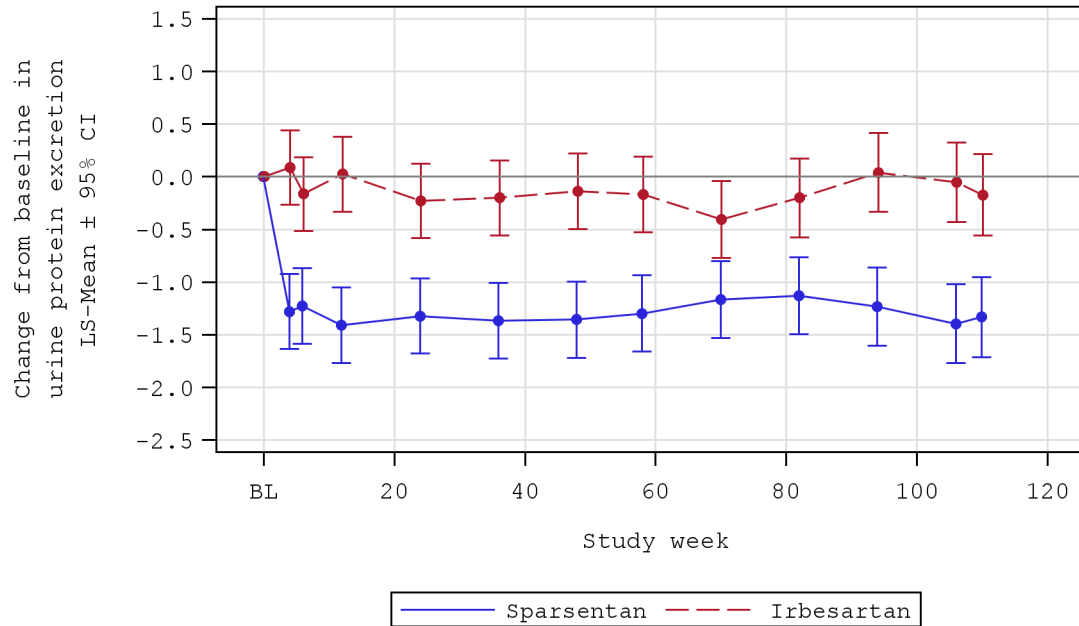
Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline urine protein excretion: <= 1.75 g/day



Sparsentan	95	94	93	85
Irbesartan	88	87	82	70

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

Figure PF2UEC\_FSGM: Change from baseline in urine protein excretion by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline urine protein excretion: > 1.75 g/day



Sparsentan	100	100	95	84
Irbesartan	105	100	96	80

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. An increase reflects a worsening of the status of the patient. Results are presented in g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2UEC\_FSCM

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Week 58	Sparsentan							Interaction test: 0.075
	Week 110	Sparsentan							Interaction test: 0.834
Male	Week 58	Sparsentan	139	129 (92.8)	58 (45.0)	2.196 [1.483, 3.251]	3.173 [1.825, 5.518]	24.5 [12.6, 36.4]	<0.001 *
		Irbesartan	143	127 (88.8)	26 (20.5)				
	Week 110	Sparsentan	139	116 (83.5)	56 (48.3)	2.172 [1.456, 3.241]	3.267 [1.826, 5.845]	26.1 [13.2, 39.0]	<0.001 *
		Irbesartan	143	108 (75.5)	24 (22.2)				
Female	Week 58	Sparsentan	63	59 (93.7)	32 (54.2)	1.317 [0.880, 1.971]	1.693 [0.794, 3.610]	13.1 [-7.3, 33.4]	0.186
		Irbesartan	59	51 (86.4)	21 (41.2)				
	Week 110	Sparsentan	63	53 (84.1)	28 (52.8)	2.017 [1.143, 3.559]	3.156 [1.317, 7.565]	26.6 [5.6, 47.7]	0.012 *
		Irbesartan	59	42 (71.2)	11 (26.2)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Partial remission means an urinary protein excretion of <1.0 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Week 58	Sparsentan							Interaction test: 0.075
	Week 110	Sparsentan							Interaction test: 0.446
<= 45 years	Week 58	Sparsentan	96	89 (92.7)	40 (44.9)	1.398 [0.950, 2.058]	1.723 [0.927, 3.203]	12.8 [-2.7, 28.3]	0.089
		Irbesartan	99	84 (84.8)	27 (32.1)				
	Week 110	Sparsentan	96	76 (79.2)	37 (48.7)	1.866 [1.179, 2.954]	2.688 [1.334, 5.417]	22.6 [5.9, 39.3]	0.006 *
		Irbesartan	99	69 (69.7)	18 (26.1)				
> 45 years	Week 58	Sparsentan	106	99 (93.4)	50 (50.5)	2.374 [1.536, 3.667]	3.776 [2.007, 7.102]	29.2 [15.3, 43.1]	<0.001 *
		Irbesartan	103	94 (91.3)	20 (21.3)				
	Week 110	Sparsentan	106	93 (87.7)	47 (50.5)	2.408 [1.508, 3.845]	3.847 [1.965, 7.530]	29.5 [14.9, 44.2]	<0.001 *
		Irbesartan	103	81 (78.6)	17 (21.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Week 58	Sparsentan							Interaction test: 0.323
	Week 110	Sparsentan							Interaction test: 0.701
<= 18 years	Week 58	Sparsentan	9	7 (77.8)	2 (28.6)	0.571 [0.124, 2.632]	0.400 [0.031, 5.151]	-21.4 [-100.0, 57.6]	0.576
		Irbesartan	5	4 (80.0)	2 (50.0)				
	Week 110	Sparsentan	9	5 (55.6)	1 (20.0)	0.800 [0.070, 9.180]	0.750 [0.032, 17.506]	-5.0 [-82.5, 72.5]	1.000
		Irbesartan	5	4 (80.0)	1 (25.0)				
> 18 to 40 years	Week 58	Sparsentan	102	95 (93.1)	49 (51.6)	1.898 [1.289, 2.795]	2.855 [1.550, 5.257]	24.4 [9.8, 39.0]	<0.001 *
		Irbesartan	109	92 (84.4)	25 (27.2)				
	Week 110	Sparsentan	102	82 (80.4)	41 (50.0)	2.079 [1.329, 3.253]	3.158 [1.610, 6.193]	25.9 [10.4, 41.5]	0.001 *
		Irbesartan	109	79 (72.5)	19 (24.1)				
> 40 years	Week 58	Sparsentan	91	86 (94.5)	39 (45.3)	1.859 [1.190, 2.905]	2.572 [1.331, 4.972]	21.0 [5.7, 36.2]	0.006 *
		Irbesartan	88	82 (93.2)	20 (24.4)				
	Week 110	Sparsentan	91	82 (90.1)	42 (51.2)	2.288 [1.397, 3.747]	3.640 [1.773, 7.473]	28.8 [12.8, 44.9]	<0.001 *
		Irbesartan	88	67 (76.1)	15 (22.4)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Week 58	Sparsentan							Interaction test: 0.576
	Week 110	Sparsentan							Interaction test: 0.933
North America	Week 58	Sparsentan	35	28 (80.0)	12 (42.9)	2.449 [1.103, 5.436]	3.536 [1.169, 10.696]	25.4 [0.5, 50.2]	0.029 *
		Irbesartan	46	40 (87.0)	7 (17.5)				
	Week 110	Sparsentan	35	27 (77.1)	12 (44.4)	1.968 [0.906, 4.278]	2.743 [0.883, 8.522]	21.9 [-5.4, 49.2]	0.097
		Irbesartan	46	31 (67.4)	7 (22.6)				
Europe	Week 58	Sparsentan	98	93 (94.9)	40 (43.0)	1.536 [1.038, 2.273]	1.941 [1.066, 3.533]	15.0 [0.6, 29.4]	0.035 *
		Irbesartan	115	100 (87.0)	28 (28.0)				
	Week 110	Sparsentan	98	80 (81.6)	43 (53.8)	2.150 [1.420, 3.256]	3.486 [1.815, 6.696]	28.8 [13.4, 44.1]	<0.001 *
		Irbesartan	115	88 (76.5)	22 (25.0)				
Asia Pacific	Week 58	Sparsentan	69	67 (97.1)	38 (56.7)	1.796 [1.076, 2.999]	2.839 [1.229, 6.560]	25.1 [4.1, 46.2]	0.015 *
		Irbesartan	41	38 (92.7)	12 (31.6)				
	Week 110	Sparsentan	69	62 (89.9)	29 (46.8)	2.417 [1.123, 5.199]	3.662 [1.319, 10.166]	27.4 [6.4, 48.5]	0.013 *
		Irbesartan	41	31 (75.6)	6 (19.4)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Week 58	Sparsentan							Interaction test: 0.916
	Week 110	Sparsentan							Interaction test: 0.671
< 27 kg/m**2	Week 58	Sparsentan	83	76 (91.6)	43 (56.6)	1.862 [1.264, 2.744]	2.986 [1.543, 5.778]	26.2 [9.8, 42.6]	0.001 *
		Irbesartan	94	79 (84.0)	24 (30.4)				
	Week 110	Sparsentan	83	70 (84.3)	39 (55.7)	2.043 [1.308, 3.191]	3.355 [1.636, 6.880]	28.4 [11.1, 45.8]	<0.001 *
		Irbesartan	94	66 (70.2)	18 (27.3)				
≥ 27 kg/m**2	Week 58	Sparsentan	119	112 (94.1)	47 (42.0)	1.806 [1.188, 2.747]	2.389 [1.313, 4.348]	18.7 [5.4, 32.0]	0.005 *
		Irbesartan	107	99 (92.5)	23 (23.2)				
	Week 110	Sparsentan	119	99 (83.2)	45 (45.5)	2.358 [1.444, 3.850]	3.490 [1.779, 6.845]	26.2 [12.1, 40.3]	<0.001 *
		Irbesartan	107	83 (77.6)	16 (19.3)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
Full Analysis Set

Partial remission										
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
Randomization strata	Week 58	Sparsentan							Interaction test:	0.360
	Week 110	Sparsentan							Interaction test:	0.282
eGFR Low and UP High	Week 58	Sparsentan	71	67 (94.4)	17 (25.4)	2.706 [1.139, 6.430]	3.287 [1.204, 8.976]	16.0 [1.8, 30.2]	0.021	*
		Irbesartan	74	64 (86.5)	6 (9.4)					
	Week 110	Sparsentan	71	59 (83.1)	19 (32.2)	2.898 [1.251, 6.716]	3.800 [1.385, 10.425]	21.1 [4.7, 37.4]	0.012	*
		Irbesartan	74	54 (73.0)	6 (11.1)					
eGFR Low and UP Low	Week 58	Sparsentan	55	52 (94.5)	32 (61.5)	1.365 [0.941, 1.978]	1.948 [0.888, 4.271]	16.4 [-4.5, 37.4]	0.116	
		Irbesartan	55	51 (92.7)	23 (45.1)					
	Week 110	Sparsentan	55	46 (83.6)	25 (54.3)	1.522 [0.937, 2.470]	2.143 [0.909, 5.050]	18.6 [-4.1, 41.3]	0.091	
		Irbesartan	55	42 (76.4)	15 (35.7)					
eGFR High and UP High	Week 58	Sparsentan	37	32 (86.5)	14 (43.8)	2.260 [0.997, 5.127]	3.241 [1.045, 10.054]	24.4 [-0.9, 49.7]	0.058	
		Irbesartan	36	31 (86.1)	6 (19.4)					
	Week 110	Sparsentan	37	30 (81.1)	17 (56.7)	3.825 [1.469, 9.961]	7.519 [2.082, 27.154]	41.9 [16.1, 67.6]	0.002	*
		Irbesartan	36	27 (75.0)	4 (14.8)					

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Partial remission means an urinary protein excretion of <1.0 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024



Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
eGFR High and UP Low	Week 58	Sparsentan	39	37 (94.9)	27 (73.0)	1.946 [1.194, 3.171]	4.500 [1.624, 12.468]	35.5 [10.5, 60.4]	0.004 *
		Irbesartan	37	32 (86.5)	12 (37.5)				
	Week 110	Sparsentan	39	34 (87.2)	23 (67.6)	1.826 [1.060, 3.147]	3.555 [1.230, 10.273]	30.6 [3.2, 58.0]	0.022 *
		Irbesartan	37	27 (73.0)	10 (37.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Week 58	Sparsentan							Interaction test: 0.139
	Week 110	Sparsentan							Interaction test: 0.384
< 60 mL/min/1.73 m**2	Week 58	Sparsentan	127	120 (94.5)	51 (42.5)	1.577 [1.094, 2.273]	2.003 [1.157, 3.466]	15.5 [2.7, 28.4]	0.014 *
		Irbesartan	129	115 (89.1)	31 (27.0)				
	Week 110	Sparsentan	127	105 (82.7)	44 (41.9)	1.936 [1.246, 3.007]	2.610 [1.405, 4.850]	20.3 [6.8, 33.7]	0.003 *
		Irbesartan	129	97 (75.2)	21 (21.6)				
60 to < 90 mL/min/1.73 m**2	Week 58	Sparsentan	49	43 (87.8)	22 (51.2)	3.581 [1.615, 7.941]	6.286 [2.198, 17.979]	36.9 [16.2, 57.5]	<0.001 *
		Irbesartan	48	42 (87.5)	6 (14.3)				
	Week 110	Sparsentan	49	43 (87.8)	26 (60.5)	3.110 [1.533, 6.309]	6.336 [2.269, 17.697]	41.0 [19.0, 63.1]	<0.001 *
		Irbesartan	48	36 (75.0)	7 (19.4)				
≥ 90 mL/min/1.73 m**2	Week 58	Sparsentan	26	25 (96.2)	17 (68.0)	1.428 [0.846, 2.409]	2.338 [0.704, 7.759]	20.4 [-12.1, 52.9]	0.231
		Irbesartan	25	21 (84.0)	10 (47.6)				
	Week 110	Sparsentan	26	21 (80.8)	14 (66.7)	1.619 [0.851, 3.082]	2.857 [0.759, 10.751]	25.5 [-10.7, 61.7]	0.190
		Irbesartan	25	17 (68.0)	7 (41.2)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Week 58	Sparsentan							Interaction test: 0.239
	Week 110	Sparsentan							Interaction test: 0.591
< 45 mL/min/1.73 m**2	Week 58	Sparsentan	82	77 (93.9)	31 (40.3)	1.462 [0.914, 2.338]	1.773 [0.883, 3.562]	12.7 [-3.9, 29.3]	0.119
		Irbesartan	80	69 (86.3)	19 (27.5)				
	Week 110	Sparsentan	82	67 (81.7)	26 (38.8)	1.908 [1.060, 3.434]	2.484 [1.114, 5.539]	18.5 [1.3, 35.6]	0.032 *
		Irbesartan	80	59 (73.8)	12 (20.3)				
45 to < 60 mL/min/1.73 m**2	Week 58	Sparsentan	45	43 (95.6)	20 (46.5)	1.783 [0.996, 3.192]	2.464 [1.012, 6.000]	20.4 [-1.4, 42.3]	0.051
		Irbesartan	49	46 (93.9)	12 (26.1)				
	Week 110	Sparsentan	45	38 (84.4)	18 (47.4)	2.000 [1.032, 3.877]	2.900 [1.086, 7.744]	23.7 [0.2, 47.2]	0.054
		Irbesartan	49	38 (77.6)	9 (23.7)				
60 to < 90 mL/min/1.73 m**2	Week 58	Sparsentan	49	43 (87.8)	22 (51.2)	3.581 [1.615, 7.941]	6.286 [2.198, 17.979]	36.9 [16.2, 57.5]	<0.001 *
		Irbesartan	48	42 (87.5)	6 (14.3)				
	Week 110	Sparsentan	49	43 (87.8)	26 (60.5)	3.110 [1.533, 6.309]	6.336 [2.269, 17.697]	41.0 [19.0, 63.1]	<0.001 *
		Irbesartan	48	36 (75.0)	7 (19.4)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
>= 90 mL/min/1.73 m**2	Week 58	Sparsentan	26	25 (96.2)	17 (68.0)	1.428 [0.846, 2.409]	2.338 [0.704, 7.759]	20.4 [-12.1, 52.9]	0.231
		Irbesartan	25	21 (84.0)	10 (47.6)				
	Week 110	Sparsentan	26	21 (80.8)	14 (66.7)	1.619 [0.851, 3.082]	2.857 [0.759, 10.751]	25.5 [-10.7, 61.7]	0.190
		Irbesartan	25	17 (68.0)	7 (41.2)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
Full Analysis Set

Partial remission										
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
Baseline urine protein excretion	Week 58	Sparsentan							Interaction test:	0.174
	Week 110	Sparsentan							Interaction test:	0.177
<= 1.75 g/day	Week 58	Sparsentan	98	93 (94.9)	59 (63.4)	1.530 [1.134, 2.065]	2.450 [1.332, 4.505]	22.0 [6.4, 37.6]	0.004	*
		Irbesartan	93	82 (88.2)	34 (41.5)					
	Week 110	Sparsentan	98	85 (86.7)	51 (60.0)	1.750 [1.211, 2.528]	2.875 [1.490, 5.547]	25.7 [9.2, 42.3]	0.002	*
		Irbesartan	93	70 (75.3)	24 (34.3)					
> 1.75 g/day	Week 58	Sparsentan	104	95 (91.3)	31 (32.6)	2.410 [1.346, 4.313]	3.093 [1.498, 6.386]	19.1 [6.4, 31.8]	0.002	*
		Irbesartan	109	96 (88.1)	13 (13.5)					
	Week 110	Sparsentan	104	84 (80.8)	33 (39.3)	2.857 [1.553, 5.257]	4.059 [1.875, 8.787]	25.5 [11.4, 39.6]	<0.001	*
		Irbesartan	109	80 (73.4)	11 (13.8)					

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Partial remission means an urinary protein excretion of <1.0 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission										
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
Baseline use of antihypertensives	Week 58	Sparsentan							Interaction test:	0.477
	Week 110	Sparsentan							Interaction test:	0.262
Yes	Week 58	Sparsentan	90	83 (92.2)	33 (39.8)	1.590 [1.001, 2.526]	1.980 [1.013, 3.871]	14.8 [-0.6, 30.2]	0.047	*
		Irbesartan	88	80 (90.9)	20 (25.0)					
	Week 110	Sparsentan	90	75 (83.3)	34 (45.3)	1.738 [1.088, 2.776]	2.350 [1.162, 4.750]	19.2 [2.5, 35.9]	0.024	*
		Irbesartan	88	69 (78.4)	18 (26.1)					
No	Week 58	Sparsentan	112	105 (93.8)	57 (54.3)	1.970 [1.367, 2.841]	3.123 [1.737, 5.613]	26.7 [12.7, 40.7]	<0.001	*
		Irbesartan	114	98 (86.0)	27 (27.6)					
	Week 110	Sparsentan	112	94 (83.9)	50 (53.2)	2.534 [1.595, 4.027]	4.278 [2.187, 8.368]	32.2 [17.6, 46.8]	<0.001	*
		Irbesartan	114	81 (71.1)	17 (21.0)					

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission										
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
Time since renal biopsy	Week 58	Sparsentan							Interaction test:	0.024 #
	Week 110	Sparsentan							Interaction test:	0.659
<= 5 years	Week 58	Sparsentan	113	104 (92.0)	45 (43.3)	1.394 [0.983, 1.977]	1.695 [0.975, 2.945]	12.2 [-1.4, 25.9]	0.069	
		Irbesartan	127	116 (91.3)	36 (31.0)					
	Week 110	Sparsentan	113	96 (85.0)	53 (55.2)	2.054 [1.404, 3.005]	3.353 [1.822, 6.169]	28.3 [13.8, 42.8]	<0.001	*
		Irbesartan	127	93 (73.2)	25 (26.9)					
> 5 years	Week 58	Sparsentan	89	84 (94.4)	45 (53.6)	3.019 [1.705, 5.349]	5.350 [2.452, 11.670]	35.8 [20.1, 51.5]	<0.001	*
		Irbesartan	75	62 (82.7)	11 (17.7)					
	Week 110	Sparsentan	89	73 (82.0)	31 (42.5)	2.421 [1.298, 4.513]	3.469 [1.520, 7.919]	24.9 [8.3, 41.5]	0.003	*
		Irbesartan	75	57 (76.0)	10 (17.5)					

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URP\_FSPM: Patients with partial remission by subgroup  
 Full Analysis Set

Partial remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Week 58	Sparsentan							Interaction test: 0.672
	Week 110	Sparsentan							Interaction test: 0.659
Yes	Week 58	Sparsentan	155	145 (93.5)	64 (44.1)	1.730 [1.240, 2.413]	2.306 [1.403, 3.790]	18.6 [7.2, 30.1]	0.001 *
		Irbesartan	161	145 (90.1)	37 (25.5)				
	Week 110	Sparsentan	155	131 (84.5)	61 (46.6)	2.029 [1.397, 2.947]	2.926 [1.698, 5.040]	23.6 [11.5, 35.7]	<0.001 *
		Irbesartan	161	122 (75.8)	28 (23.0)				
No	Week 58	Sparsentan	47	43 (91.5)	26 (60.5)	1.995 [1.127, 3.532]	3.518 [1.345, 9.203]	30.2 [6.1, 54.3]	0.011 *
		Irbesartan	41	33 (80.5)	10 (30.3)				
	Week 110	Sparsentan	47	38 (80.9)	23 (60.5)	2.421 [1.213, 4.832]	4.600 [1.571, 13.473]	35.5 [10.1, 61.0]	0.006 *
		Irbesartan	41	28 (68.3)	7 (25.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Partial remission means an urinary protein excretion of <1.0 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024



Table PT2URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.488
Male	Sparsentan	139	101 (72.7)	7.6	(6.0, 24.1)	2.730	(1.992, 3.742)	<0.001 *
	Irbesartan	143	70 (49.0)	81.1	(57.4, NE)			
Female	Sparsentan	63	56 (88.9)	6.1	(4.1, 12.3)	2.577	(1.646, 4.036)	<0.001 *
	Irbesartan	59	36 (61.0)	22.4	(6.7, 59.6)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.101
<= 45 years	Sparsentan	96	73 (76.0)	7.4	(4.7, 24.0)	2.221	(1.557, 3.168)	<0.001 *
	Irbesartan	99	56 (56.6)	47.6	(25.1, 70.0)			
> 45 years	Sparsentan	106	84 (79.2)	6.3	(4.9, 12.7)	3.108	(2.156, 4.481)	<0.001 *
	Irbesartan	103	50 (48.5)	92.1	(51.9, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: 0.888
<= 18 years	Sparsentan	9	4 (44.4)	NE		3.253	(0.280, 37.739)	0.346
	Irbesartan	5	3 (60.0)	70.0	(6.0, NE)			
> 18 to 40 years	Sparsentan	102	79 (77.5)	6.6	(4.7, 13.1)	2.522	(1.772, 3.590)	<0.001 *
	Irbesartan	109	58 (53.2)	68.6	(36.0, NE)			
> 40 years	Sparsentan	91	74 (81.3)	6.1	(4.9, 12.4)	2.699	(1.841, 3.958)	<0.001 *
	Irbesartan	88	45 (51.1)	70.9	(36.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URPT\_FSTM: Time to partial remission by subgroup  
Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan							Interaction test: 0.779
North America	Sparsentan	35	27 (77.1)	13.3	(6.3, 34.6)	3.632	(1.897, 6.954)	<0.001 *
	Irbesartan	46	16 (34.8)	NE				
Europe	Sparsentan	98	73 (74.5)	7.6	(5.0, 24.1)	2.630	(1.846, 3.747)	<0.001 *
	Irbesartan	115	62 (53.9)	68.6	(36.1, NE)			
Asia Pacific	Sparsentan	69	57 (82.6)	4.9	(4.1, 6.9)	2.356	(1.483, 3.743)	<0.001 *
	Irbesartan	41	28 (68.3)	24.7	(6.9, 92.1)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb\_tte, created on: 27FEB2024

Table PT2URPT\_FSTM: Time to partial remission by subgroup  
Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan							Interaction test: 0.996
< 27 kg/m**2	Sparsentan	83	71 (85.5)	4.6	(4.1, 8.0)	2.542	(1.757, 3.677)	<0.001 *
	Irbesartan	94	55 (58.5)	46.1	(24.0, 92.1)			
≥ 27 kg/m**2	Sparsentan	119	86 (72.3)	12.0	(6.1, 25.1)	2.900	(2.025, 4.152)	<0.001 *
	Irbesartan	107	50 (46.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb\_tte, created on: 27FEB2024

Table PT2URPT\_FSTM: Time to partial remission by subgroup  
Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan							Interaction test: 0.965
eGFR Low and UP High	Sparsentan	71	42 (59.2)	67.3	(25.1, NE)	2.454	(1.475, 4.083)	<0.001 *
	Irbesartan	74	23 (31.1)	NE				
eGFR Low and UP Low	Sparsentan	55	50 (90.9)	4.3	(4.1, 4.7)	2.378	(1.550, 3.649)	<0.001 *
	Irbesartan	55	41 (74.5)	12.1	(4.9, 35.7)			
eGFR High and UP High	Sparsentan	37	27 (73.0)	24.1	(8.0, 69.3)	3.193	(1.628, 6.263)	<0.001 *
	Irbesartan	36	13 (36.1)	NE				
eGFR High and UP Low	Sparsentan	39	38 (97.4)	4.1	(4.1, 4.4)	2.641	(1.590, 4.387)	<0.001 *
	Irbesartan	37	29 (78.4)	12.1	(6.1, 26.1)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb\_tte, created on: 27FEB2024

Table PT2URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.318
< 60 mL/min/1.73 m**2	Sparsentan	127	94 (74.0)	12.0	(6.0, 23.7)	2.531	(1.817, 3.526)	<0.001 *
	Irbesartan	129	64 (49.6)	70.9	(40.7, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	40 (81.6)	6.1	(4.6, 22.6)	3.982	(2.227, 7.119)	<0.001 *
	Irbesartan	48	22 (45.8)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	23 (88.5)	4.1	(4.1, 12.1)	1.760	(0.940, 3.294)	0.077
	Irbesartan	25	20 (80.0)	17.4	(4.6, 36.3)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: 0.173
< 45 mL/min/1.73 m**2	Sparsentan	82	56 (68.3)	12.4	(6.1, 48.6)	2.105	(1.390, 3.187)	<0.001 *
	Irbesartan	80	39 (48.8)	81.1	(36.0, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	38 (84.4)	5.4	(4.1, 12.7)	3.810	(2.126, 6.829)	<0.001 *
	Irbesartan	49	25 (51.0)	68.6	(24.1, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	40 (81.6)	6.1	(4.6, 22.6)	3.982	(2.227, 7.119)	<0.001 *
	Irbesartan	48	22 (45.8)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	23 (88.5)	4.1	(4.1, 12.1)	1.760	(0.940, 3.294)	0.077
	Irbesartan	25	20 (80.0)	17.4	(4.6, 36.3)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024



Table PT2URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.486
<= 1.75 g/day	Sparsentan	98	89 (90.8)	4.3	(4.1, 4.7)	2.165	(1.567, 2.991)	<0.001 *
	Irbesartan	93	68 (73.1)	12.1	(6.1, 26.1)			
> 1.75 g/day	Sparsentan	104	68 (65.4)	36.1	(22.6, 69.3)	3.227	(2.117, 4.918)	<0.001 *
	Irbesartan	109	38 (34.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URPT\_FSTM: Time to partial remission by subgroup  
Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.118
Yes	Sparsentan	90	69 (76.7)	7.6	(6.0, 23.7)	3.498	(2.326, 5.259)	<0.001 *
	Irbesartan	88	46 (52.3)	70.0	(47.6, NE)			
No	Sparsentan	112	88 (78.6)	6.1	(4.7, 12.7)	2.146	(1.533, 3.004)	<0.001 *
	Irbesartan	114	60 (52.6)	59.6	(26.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb\_tte, created on: 27FEB2024

Table PT2URPT\_FSTM: Time to partial remission by subgroup  
 Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.132
<= 5 years	Sparsentan	113	90 (79.6)	7.1	(5.0, 13.3)	2.387	(1.739, 3.275)	<0.001 *
	Irbesartan	127	74 (58.3)	46.1	(25.6, 70.0)			
> 5 years	Sparsentan	89	67 (75.3)	6.5	(4.7, 12.7)	3.384	(2.181, 5.250)	<0.001 *
	Irbesartan	75	32 (42.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URPT\_FSTM: Time to partial remission by subgroup  
Full Analysis Set

Time to partial remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension		Sparsentan				Interaction test: 0.550		
Yes	Sparsentan	155	122 (78.7)	7.6	(6.0, 13.1)	2.977	(2.222, 3.989)	<0.001 *
	Irbesartan	161	80 (49.7)	70.6	(47.6, NE)			
No	Sparsentan	47	35 (74.5)	4.4	(4.1, 22.6)	1.989	(1.169, 3.384)	0.011 *
	Irbesartan	41	26 (63.4)	41.6	(6.9, 92.1)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb\_tte, created on: 27FEB2024

Figure PF2URPT\_FSKM: Time to partial remission by subgroup  
Full Analysis Set

Not done. No significant subgroups.

Partial remission means an urinary protein excretion of <1.0 g/day. No event means no partial remission.  
eGFR = estimated glomerular filtration rate.  
Reference table: PT2URPT\_FSTM

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Week 58	Sparsentan							Interaction test: 0.886
	Week 110	Sparsentan							Interaction test: 0.687
Male	Week 58	Sparsentan	139	129 (92.8)	20 (15.5)	3.938 [1.525, 10.171]	4.477 [1.625, 12.335]	11.6 [3.7, 19.5]	0.003 *
		Irbesartan	143	127 (88.8)	5 (3.9)				
	Week 110	Sparsentan	139	116 (83.5)	17 (14.7)	5.276 [1.591, 17.500]	6.010 [1.709, 21.140]	11.9 [3.8, 19.9]	0.002 *
		Irbesartan	143	108 (75.5)	3 (2.8)				
Female	Week 58	Sparsentan	63	59 (93.7)	8 (13.6)	3.458 [0.769, 15.550]	3.843 [0.777, 19.003]	9.6 [-2.4, 21.7]	0.103
		Irbesartan	59	51 (86.4)	2 (3.9)				
	Week 110	Sparsentan	63	53 (84.1)	9 (17.0)	3.566 [0.814, 15.630]	4.091 [0.834, 20.078]	12.2 [-1.9, 26.3]	0.105
		Irbesartan	59	42 (71.2)	2 (4.8)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Week 58	Sparsentan							Interaction test: 0.391
	Week 110	Sparsentan							Interaction test: 0.762
<= 45 years	Week 58	Sparsentan	96	89 (92.7)	13 (14.6)	6.135 [1.427, 26.378]	7.013 [1.532, 32.100]	12.2 [3.0, 21.4]	0.005 *
		Irbesartan	99	84 (84.8)	2 (2.4)				
	Week 110	Sparsentan	96	76 (79.2)	12 (15.8)	5.447 [1.264, 23.481]	6.281 [1.352, 29.172]	12.9 [2.4, 23.4]	0.010 *
		Irbesartan	99	69 (69.7)	2 (2.9)				
> 45 years	Week 58	Sparsentan	106	99 (93.4)	15 (15.2)	2.848 [1.078, 7.529]	3.179 [1.107, 9.130]	9.8 [0.4, 19.3]	0.033 *
		Irbesartan	103	94 (91.3)	5 (5.3)				
	Week 110	Sparsentan	106	93 (87.7)	14 (15.1)	4.065 [1.211, 13.642]	4.608 [1.274, 16.665]	11.4 [1.8, 20.9]	0.019 *
		Irbesartan	103	81 (78.6)	3 (3.7)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Week 58	Sparsentan							Interaction test: 0.586
	Week 110	Sparsentan							Interaction test: 0.889
<= 18 years	Week 58	Sparsentan	9	7 (77.8)	1 (14.3)	1.875 + [0.093, 37.632]	2.077 + [0.068, 63.417]	14.3 [-31.3, 59.9]	1.000
		Irbesartan	5	4 (80.0)	0 (0.0)				
	Week 110	Sparsentan	9	5 (55.6)	1 (20.0)	2.500 + [0.128, 48.848]	3.000 + [0.095, 95.170]	20.0 [-37.6, 77.6]	1.000
		Irbesartan	5	4 (80.0)	0 (0.0)				
> 18 to 40 years	Week 58	Sparsentan	102	95 (93.1)	13 (13.7)	6.295 [1.461, 27.127]	7.134 [1.563, 32.569]	11.5 [2.9, 20.1]	0.005 *
		Irbesartan	109	92 (84.4)	2 (2.2)				
	Week 110	Sparsentan	102	82 (80.4)	11 (13.4)	5.299 [1.213, 23.152]	5.965 [1.278, 27.845]	10.9 [1.5, 20.3]	0.018 *
		Irbesartan	109	79 (72.5)	2 (2.5)				
> 40 years	Week 58	Sparsentan	91	86 (94.5)	14 (16.3)	2.670 [1.007, 7.080]	2.994 [1.027, 8.734]	10.2 [-0.4, 20.7]	0.050
		Irbesartan	88	82 (93.2)	5 (6.1)				
	Week 110	Sparsentan	91	82 (90.1)	14 (17.1)	3.813 [1.143, 12.716]	4.392 [1.206, 16.001]	12.6 [1.7, 23.5]	0.019 *
		Irbesartan	88	67 (76.1)	3 (4.5)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Complete remission means an urinary protein excretion of <0.3 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024



Table PT2URF\_FSPM: Patients with complete remission by subgroup  
Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Week 58	Sparsentan							Interaction test: 0.631
	Week 110	Sparsentan							Interaction test: 0.418
North America	Week 58	Sparsentan	35	28 (80.0)	2 (7.1)	7.069 + [0.352, 141.826]	7.642 + [0.353, 165.537]	7.1 [-5.4, 19.7]	0.166
		Irbesartan	46	40 (87.0)	0 (0.0)				
	Week 110	Sparsentan	35	27 (77.1)	4 (14.8)	10.286 + [0.579, 182.754]	12.064 + [0.619, 235.190]	14.8 [-2.0, 31.7]	0.041 *
		Irbesartan	46	31 (67.4)	0 (0.0)				
Europe	Week 58	Sparsentan	98	93 (94.9)	11 (11.8)	3.943 [1.135, 13.692]	4.337 [1.170, 16.076]	8.8 [0.4, 17.2]	0.025 *
		Irbesartan	115	100 (87.0)	3 (3.0)				
	Week 110	Sparsentan	98	80 (81.6)	11 (13.8)	2.420 [0.879, 6.663]	2.646 [0.877, 7.984]	8.1 [-2.1, 18.2]	0.113
		Irbesartan	115	88 (76.5)	5 (5.7)				
Asia Pacific	Week 58	Sparsentan	69	67 (97.1)	15 (22.4)	2.127 [0.760, 5.949]	2.452 [0.750, 8.016]	11.9 [-4.2, 27.9]	0.188
		Irbesartan	41	38 (92.7)	4 (10.5)				
	Week 110	Sparsentan	69	62 (89.9)	11 (17.7)	11.683 + [0.711, 191.976]	14.068 + [0.801, 247.107]	17.7 [5.8, 29.7]	0.014 *
		Irbesartan	41	31 (75.6)	0 (0.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Complete remission means an urinary protein excretion of <0.3 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Week 58	Sparsentan							Interaction test: 0.092
	Week 110	Sparsentan							Interaction test: 0.766
< 27 kg/m**2	Week 58	Sparsentan	83	76 (91.6)	17 (22.4)	8.836 [2.112, 36.955]	11.093 [2.466, 49.910]	19.8 [8.6, 31.1]	<0.001 *
		Irbesartan	94	79 (84.0)	2 (2.5)				
	Week 110	Sparsentan	83	70 (84.3)	13 (18.6)	4.086 [1.219, 13.695]	4.789 [1.298, 17.671]	14.0 [2.2, 25.9]	0.015 *
		Irbesartan	94	66 (70.2)	3 (4.5)				
≥ 27 kg/m**2	Week 58	Sparsentan	119	112 (94.1)	11 (9.8)	1.945 [0.700, 5.403]	2.048 [0.686, 6.113]	4.8 [-3.2, 12.7]	0.297
		Irbesartan	107	99 (92.5)	5 (5.1)				
	Week 110	Sparsentan	119	99 (83.2)	13 (13.1)	5.449 [1.266, 23.462]	6.122 [1.340, 27.972]	10.7 [2.2, 19.3]	0.013 *
		Irbesartan	107	83 (77.6)	2 (2.4)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Complete remission means an urinary protein excretion of <0.3 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Week 58	Sparsentan							Interaction test: 0.688
	Week 110	Sparsentan							Interaction test: 0.547
eGFR Low and UP High	Week 58	Sparsentan	71	67 (94.4)	2 (3.0)	1.910 [0.178, 20.557]	1.938 [0.171, 21.916]	1.4 [-5.2, 8.0]	1.000
		Irbesartan	74	64 (86.5)	1 (1.6)				
	Week 110	Sparsentan	71	59 (83.1)	4 (6.8)	1.831 [0.349, 9.596]	1.891 [0.332, 10.765]	3.1 [-6.9, 13.0]	0.681
		Irbesartan	74	54 (73.0)	2 (3.7)				
eGFR Low and UP Low	Week 58	Sparsentan	55	52 (94.5)	10 (19.2)	2.452 [0.822, 7.317]	2.798 [0.816, 9.590]	11.4 [-3.6, 26.3]	0.149
		Irbesartan	55	51 (92.7)	4 (7.8)				
	Week 110	Sparsentan	55	46 (83.6)	9 (19.6)	2.739 [0.794, 9.445]	3.162 [0.794, 12.593]	12.4 [-3.7, 28.6]	0.123
		Irbesartan	55	42 (76.4)	3 (7.1)				
eGFR High and UP High	Week 58	Sparsentan	37	32 (86.5)	6 (18.8)	12.606 + [0.740, 214.695]	15.453 + [0.831, 287.224]	18.8 [2.1, 35.4]	0.024 *
		Irbesartan	36	31 (86.1)	0 (0.0)				
	Week 110	Sparsentan	37	30 (81.1)	5 (16.7)	9.935 + [0.575, 171.695]	11.863 + [0.624, 225.468]	16.7 [-0.2, 33.5]	0.053
		Irbesartan	36	27 (75.0)	0 (0.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Complete remission means an urinary protein excretion of <0.3 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission										
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
eGFR High and UP Low	Week 58	Sparsentan	39	37 (94.9)	10 (27.0)	4.324 [1.022, 18.299]	5.556 [1.116, 27.648]	20.8 [1.3, 40.3]	0.028	*
		Irbesartan	37	32 (86.5)	2 (6.3)					
	Week 110	Sparsentan	39	34 (87.2)	8 (23.5)	13.600 + [0.820, 225.566]	17.642 + [0.969, 321.145]	23.5 [5.9, 41.1]	0.007	*
		Irbesartan	37	27 (73.0)	0 (0.0)					

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Week 58	Sparsentan							Interaction test: 0.361
	Week 110	Sparsentan							Interaction test: 0.320
< 60 mL/min/1.73 m**2	Week 58	Sparsentan	127	120 (94.5)	12 (10.0)	2.300 [0.837, 6.324]	2.444 [0.833, 7.173]	5.7 [-1.7, 13.0]	0.130
		Irbesartan	129	115 (89.1)	5 (4.3)				
	Week 110	Sparsentan	127	105 (82.7)	12 (11.4)	2.217 [0.811, 6.064]	2.374 [0.804, 7.008]	6.3 [-2.2, 14.8]	0.132
		Irbesartan	129	97 (75.2)	5 (5.2)				
60 to < 90 mL/min/1.73 m**2	Week 58	Sparsentan	49	43 (87.8)	12 (27.9)	11.721 [1.594, 86.191]	15.871 [1.958, 128.656]	25.5 [9.0, 42.1]	0.002 *
		Irbesartan	48	42 (87.5)	1 (2.4)				
	Week 110	Sparsentan	49	43 (87.8)	9 (20.9)	15.977 + [0.962, 265.384]	20.101 + [1.127, 358.682]	20.9 [6.2, 35.6]	0.003 *
		Irbesartan	48	36 (75.0)	0 (0.0)				
≥ 90 mL/min/1.73 m**2	Week 58	Sparsentan	26	25 (96.2)	4 (16.0)	3.360 [0.406, 27.800]	3.810 [0.392, 37.068]	11.2 [-10.2, 32.6]	0.357
		Irbesartan	25	21 (84.0)	1 (4.8)				
	Week 110	Sparsentan	26	21 (80.8)	5 (23.8)	9.000 + [0.533, 152.090]	11.667 + [0.597, 227.896]	23.8 [0.3, 47.3]	0.053
		Irbesartan	25	17 (68.0)	0 (0.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Complete remission means an urinary protein excretion of <0.3 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Week 58	Sparsentan							Interaction test: 0.356
	Week 110	Sparsentan							Interaction test: NE
< 45 mL/min/1.73 m**2	Week 58	Sparsentan	82	77 (93.9)	7 (9.1)	1.568 [0.480, 5.128]	1.625 [0.455, 5.810]	3.3 [-6.5, 13.1]	0.540
		Irbesartan	80	69 (86.3)	4 (5.8)				
	Week 110	Sparsentan	82	67 (81.7)	7 (10.4)	all n<10			NE
		Irbesartan	80	59 (73.8)	2 (3.4)				
45 to < 60 mL/min/1.73 m**2	Week 58	Sparsentan	45	43 (95.6)	5 (11.6)	5.349 [0.651, 43.960]	5.921 [0.663, 52.907]	9.5 [-3.3, 22.2]	0.103
		Irbesartan	49	46 (93.9)	1 (2.2)				
	Week 110	Sparsentan	45	38 (84.4)	5 (13.2)	all n<10			NE
		Irbesartan	49	38 (77.6)	3 (7.9)				
60 to < 90 mL/min/1.73 m**2	Week 58	Sparsentan	49	43 (87.8)	12 (27.9)	11.721 [1.594, 86.191]	15.871 [1.958, 128.656]	25.5 [9.0, 42.1]	0.002 *
		Irbesartan	48	42 (87.5)	1 (2.4)				
	Week 110	Sparsentan	49	43 (87.8)	9 (20.9)	all n<10			NE
		Irbesartan	48	36 (75.0)	0 (0.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Complete remission means an urinary protein excretion of <0.3 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
>= 90 mL/min/1.73 m**2	Week 58	Sparsentan	26	25 (96.2)	4 (16.0)	3.360 [0.406, 27.800]	3.810 [0.392, 37.068]	11.2 [-10.2, 32.6]	0.357
		Irbesartan	25	21 (84.0)	1 (4.8)				
	Week 110	Sparsentan	26	21 (80.8)	5 (23.8)	all n<10			NE
		Irbesartan	25	17 (68.0)	0 (0.0)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Week 58	Sparsentan							Interaction test: 0.469
	Week 110	Sparsentan							Interaction test: 0.930
<= 1.75 g/day	Week 58	Sparsentan	98	93 (94.9)	21 (22.6)	3.086 [1.309, 7.273]	3.694 [1.411, 9.676]	15.3 [3.9, 26.6]	0.006 *
		Irbesartan	93	82 (88.2)	6 (7.3)				
	Week 110	Sparsentan	98	85 (86.7)	17 (20.0)	4.667 [1.426, 15.277]	5.583 [1.563, 19.939]	15.7 [4.7, 26.8]	0.004 *
		Irbesartan	93	70 (75.3)	3 (4.3)				
> 1.75 g/day	Week 58	Sparsentan	104	95 (91.3)	7 (7.4)	7.074 [0.887, 56.395]	7.557 [0.911, 62.659]	6.3 [-0.4, 13.0]	0.035 *
		Irbesartan	109	96 (88.1)	1 (1.0)				
	Week 110	Sparsentan	104	84 (80.8)	9 (10.7)	4.286 [0.955, 19.232]	4.680 [0.979, 22.374]	8.2 [-0.5, 16.9]	0.057
		Irbesartan	109	80 (73.4)	2 (2.5)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024



Table PT2URF\_FSPM: Patients with complete remission by subgroup  
Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Week 58	Sparsentan							Interaction test: 0.048 #
	Week 110	Sparsentan							Interaction test: 0.405
Yes	Week 58	Sparsentan	90	83 (92.2)	8 (9.6)	1.542 [0.527, 4.515]	1.600 [0.500, 5.116]	3.4 [-6.1, 12.9]	0.566
		Irbesartan	88	80 (90.9)	5 (6.3)				
	Week 110	Sparsentan	90	75 (83.3)	10 (13.3)	3.067 [0.880, 10.683]	3.385 [0.891, 12.860]	9.0 [-1.5, 19.5]	0.081
		Irbesartan	88	69 (78.4)	3 (4.3)				
No	Week 58	Sparsentan	112	105 (93.8)	20 (19.0)	9.333 [2.240, 38.894]	11.294 [2.564, 49.742]	17.0 [8.0, 26.0]	<0.001 *
		Irbesartan	114	98 (86.0)	2 (2.0)				
	Week 110	Sparsentan	112	94 (83.9)	16 (17.0)	6.894 [1.634, 29.085]	8.103 [1.803, 36.418]	14.6 [5.1, 24.0]	0.002 *
		Irbesartan	114	81 (71.1)	2 (2.5)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
Complete remission means an urinary protein excretion of <0.3 g/day.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Week 58	Sparsentan							Interaction test: 0.402
	Week 110	Sparsentan							Interaction test: 0.548
<= 5 years	Week 58	Sparsentan	113	104 (92.0)	17 (16.3)	3.160 [1.295, 7.713]	3.582 [1.355, 9.471]	11.2 [2.1, 20.3]	0.008 *
		Irbesartan	127	116 (91.3)	6 (5.2)				
	Week 110	Sparsentan	113	96 (85.0)	16 (16.7)	3.875 [1.345, 11.161]	4.450 [1.428, 13.866]	12.4 [2.8, 21.9]	0.008 *
		Irbesartan	127	93 (73.2)	4 (4.3)				
> 5 years	Week 58	Sparsentan	89	84 (94.4)	11 (13.1)	8.119 [1.076, 61.243]	9.192 [1.154, 73.220]	11.5 [2.2, 20.8]	0.014 *
		Irbesartan	75	62 (82.7)	1 (1.6)				
	Week 110	Sparsentan	89	73 (82.0)	10 (13.7)	7.808 [1.029, 59.228]	8.889 [1.103, 71.644]	11.9 [1.8, 22.1]	0.023 *
		Irbesartan	75	57 (76.0)	1 (1.8)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URF\_FSPM: Patients with complete remission by subgroup  
 Full Analysis Set

Complete remission									
Subgroup	Time	Treatment	N	nval (%)	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Week 58	Sparsentan							Interaction test: 0.152
	Week 110	Sparsentan							Interaction test: 0.555
Yes	Week 58	Sparsentan	155	145 (93.5)	16 (11.0)	2.286 [0.969, 5.390]	2.445 [0.974, 6.136]	6.2 [-0.7, 13.1]	0.080
		Irbesartan	161	145 (90.1)	7 (4.8)				
	Week 110	Sparsentan	155	131 (84.5)	16 (12.2)	3.725 [1.281, 10.834]	4.104 [1.332, 12.647]	8.9 [1.7, 16.2]	0.010 *
		Irbesartan	161	122 (75.8)	4 (3.3)				
No	Week 58	Sparsentan	47	43 (91.5)	12 (27.9)	19.318 + [1.185, 314.849]	26.587 + [1.510, 468.101]	27.9 [11.8, 44.0]	<0.001 *
		Irbesartan	41	33 (80.5)	0 (0.0)				
	Week 110	Sparsentan	47	38 (80.9)	10 (26.3)	7.368 [1.000, 54.273]	9.643 [1.155, 80.538]	22.7 [4.0, 41.4]	0.018 *
		Irbesartan	41	28 (68.3)	1 (3.6)				

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method). Percentages are based on number of evaluable values.  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 Complete remission means an urinary protein excretion of <0.3 g/day.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb, created on: 19FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.358
Male	Sparsentan	139	40 (28.8)	NE		3.837	(2.080, 7.077)	<0.001 *
	Irbesartan	143	14 (9.8)	NE				
Female	Sparsentan	63	22 (34.9)	111.4	(109.9, NE)	2.677	(1.213, 5.905)	0.015 *
	Irbesartan	59	9 (15.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test: 0.890	
<= 45 years	Sparsentan	96	34 (35.4)	NE		3.268	(1.718, 6.218)	<0.001 *
	Irbesartan	99	13 (13.1)	NE				
> 45 years	Sparsentan	106	28 (26.4)	112.3	(112.3, NE)	3.147	(1.526, 6.491)	0.002 *
	Irbesartan	103	10 (9.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb\_tte, created on: 27FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis		Sparsentan				Interaction test:		0.636
<= 18 years	Sparsentan	9	1 (11.1)	NE		NE		NE
	Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Sparsentan	102	35 (34.3)	112.3	(111.4, NE)	3.545	(1.855, 6.774)	<0.001 *
	Irbesartan	109	13 (11.9)	NE				
> 40 years	Sparsentan	91	26 (28.6)	NE		3.159	(1.474, 6.770)	0.003 *
	Irbesartan	88	9 (10.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb\_tte, created on: 27FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan							Interaction test: 0.393
North America	Sparsentan	35	9 (25.7)	NE		3.676	(0.984, 13.731)	0.053
	Irbesartan	46	3 (6.5)	NE				
Europe	Sparsentan	98	30 (30.6)	112.3	(111.4, 112.3)	4.387	(2.208, 8.719)	<0.001 *
	Irbesartan	115	12 (10.4)	NE				
Asia Pacific	Sparsentan	69	23 (33.3)	NE		2.081	(0.926, 4.674)	0.076
	Irbesartan	41	8 (19.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.755
< 27 kg/m**2	Sparsentan	83	31 (37.3)	112.3	(104.4, NE)	3.309	(1.688, 6.488)	<0.001 *
	Irbesartan	94	12 (12.8)	NE				
≥ 27 kg/m**2	Sparsentan	119	31 (26.1)	NE		3.126	(1.528, 6.395)	0.002 *
	Irbesartan	107	10 (9.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024



Table PT2URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan							Interaction test: 0.644
eGFR Low and UP High	Sparsentan	71	10 (14.1)	112.3	(112.3, NE)	2.676	(0.830, 8.620)	0.099
	Irbesartan	74	4 (5.4)	NE				
eGFR Low and UP Low	Sparsentan	55	22 (40.0)	111.4	(82.1, NE)	2.777	(1.313, 5.873)	0.008 *
	Irbesartan	55	10 (18.2)	NE				
eGFR High and UP High	Sparsentan	37	9 (24.3)	NE		9.656	(1.213, 76.871)	0.032 *
	Irbesartan	36	1 (2.8)	NE				
eGFR High and UP Low	Sparsentan	39	21 (53.8)	70.1	(47.7, NE)	2.848	(1.252, 6.478)	0.013 *
	Irbesartan	37	8 (21.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.030 #
< 60 mL/min/1.73 m**2	Sparsentan	127	31 (24.4)	112.3	(112.3, NE)	2.506	(1.328, 4.728)	0.005 *
	Irbesartan	129	14 (10.9)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	19 (38.8)	NE		13.934	(2.615, 74.264)	0.002 *
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	12 (46.2)	NE		1.718	(0.674, 4.383)	0.257
	Irbesartan	25	7 (28.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb\_tte, created on: 27FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: 0.033 #
< 45 mL/min/1.73 m**2	Sparsentan	82	17 (20.7)	112.3	(111.4, NE)	1.891	(0.842, 4.247)	0.123
	Irbesartan	80	9 (11.3)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	14 (31.1)	NE		3.980	(1.407, 11.260)	0.009 *
	Irbesartan	49	5 (10.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	19 (38.8)	NE		13.934	(2.615, 74.264)	0.002 *
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	12 (46.2)	NE		1.718	(0.674, 4.383)	0.257
	Irbesartan	25	7 (28.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.560
<= 1.75 g/day	Sparsentan	98	44 (44.9)	111.4	(70.1, NE)	2.891	(1.669, 5.006)	<0.001 *
	Irbesartan	93	18 (19.4)	NE				
> 1.75 g/day	Sparsentan	104	18 (17.3)	112.3	(112.3, NE)	4.367	(1.590, 11.997)	0.004 *
	Irbesartan	109	5 (4.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: alb\_tte, created on: 27FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.279
Yes	Sparsentan	90	21 (23.3)	NE		2.284	(1.069, 4.877)	0.033 *
	Irbesartan	88	10 (11.4)	NE				
No	Sparsentan	112	41 (36.6)	112.3	(111.4, NE)	3.867	(2.048, 7.303)	<0.001 *
	Irbesartan	114	13 (11.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.512
<= 5 years	Sparsentan	113	39 (34.5)	NE		2.985	(1.703, 5.232)	<0.001 *
	Irbesartan	127	18 (14.2)	NE				
> 5 years	Sparsentan	89	23 (25.8)	112.3	(111.4, 112.3)	4.838	(1.767, 13.248)	0.002 *
	Irbesartan	75	5 (6.7)	NE				

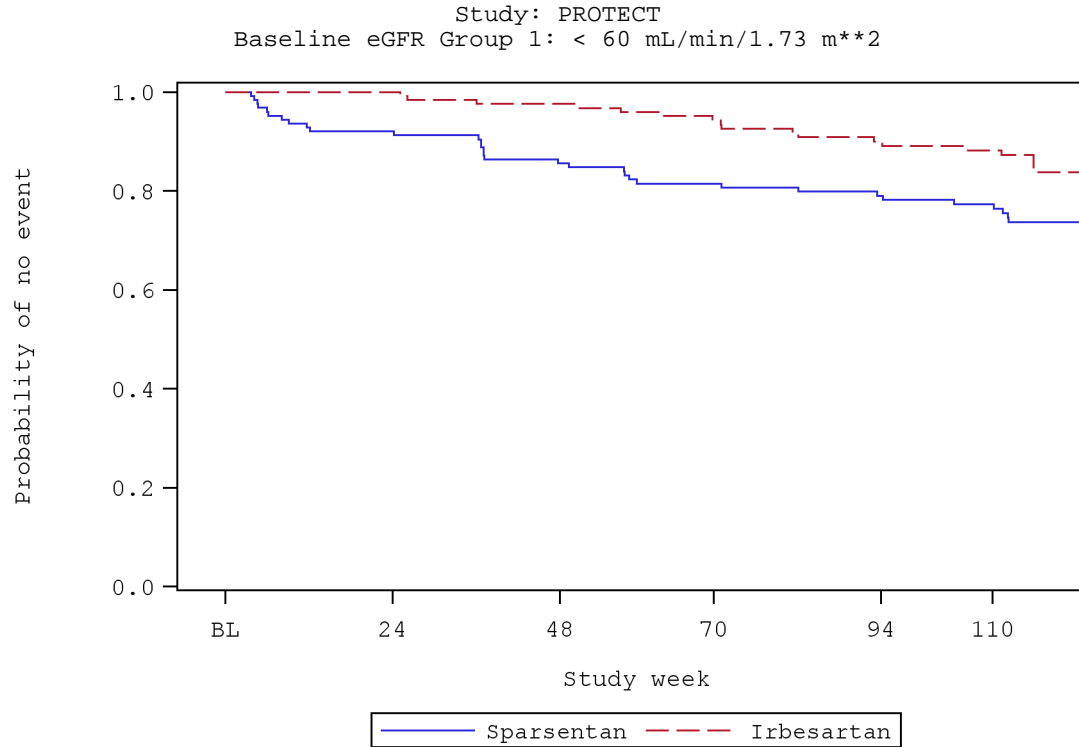
N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Table PT2URFT\_FSTM: Time to complete remission by subgroup  
 Full Analysis Set

Time to complete remission				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension		Sparsentan				Interaction test: 0.429		
Yes	Sparsentan	155	44 (28.4)	NE		3.676	(2.013, 6.713)	<0.001 *
	Irbesartan	161	14 (8.7)	NE				
No	Sparsentan	47	18 (38.3)	111.4	(60.0, NE)	2.255	(0.997, 5.100)	0.051
	Irbesartan	41	9 (22.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate. Baseline was omitted for subgroup Baseline protein excretion.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: alb\_tte, created on: 27FEB2024

Figure PF2URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set

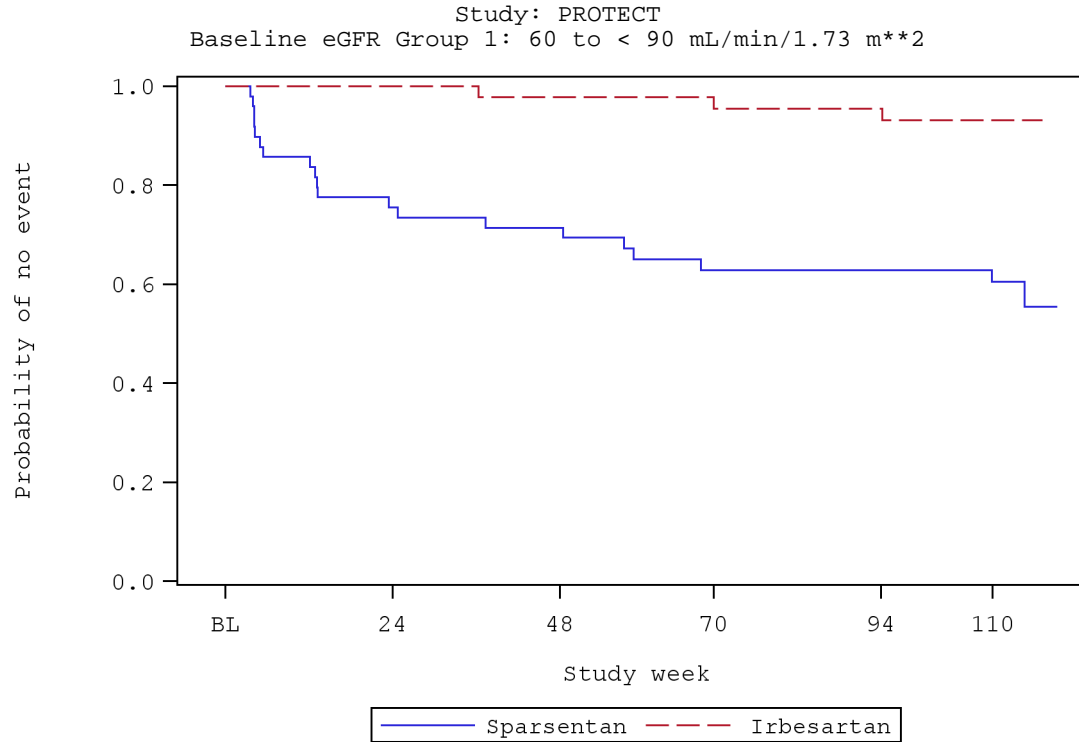


Sparsentan	127	115	105	100	93	88
Irbesartan	129	128	119	111	101	96

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT2URFT\_FSTM



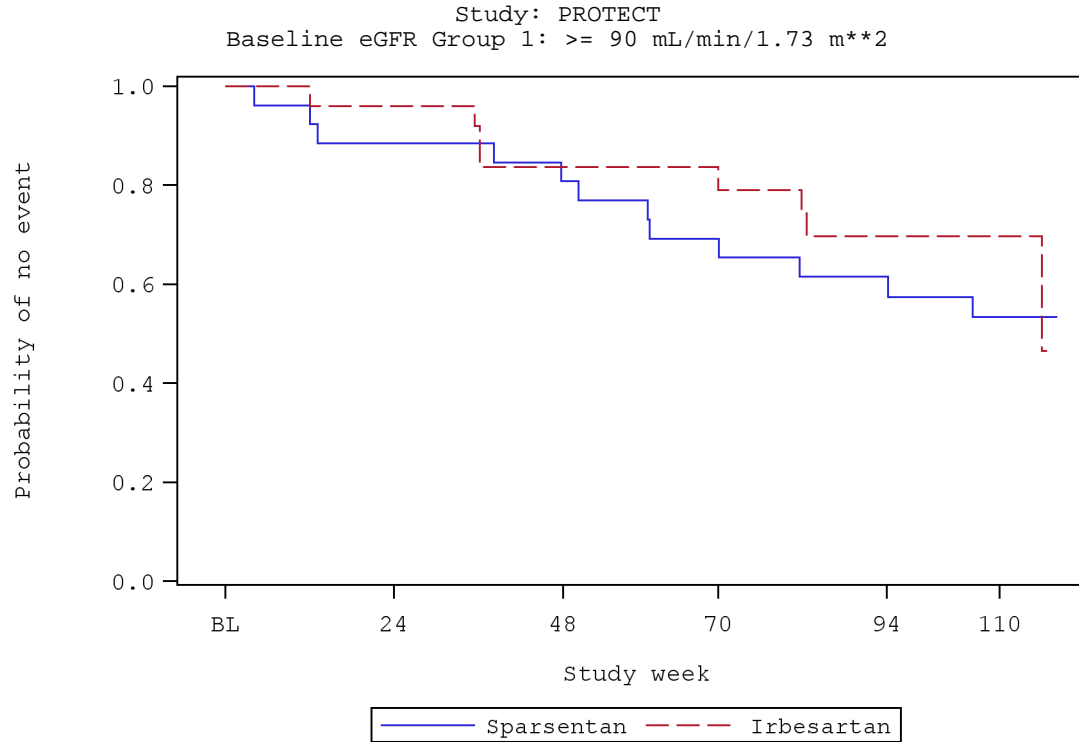
Figure PF2URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	49	37	35	29	29	26
Irbesartan	48	46	44	43	40	37

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT2URFT\_FSTM

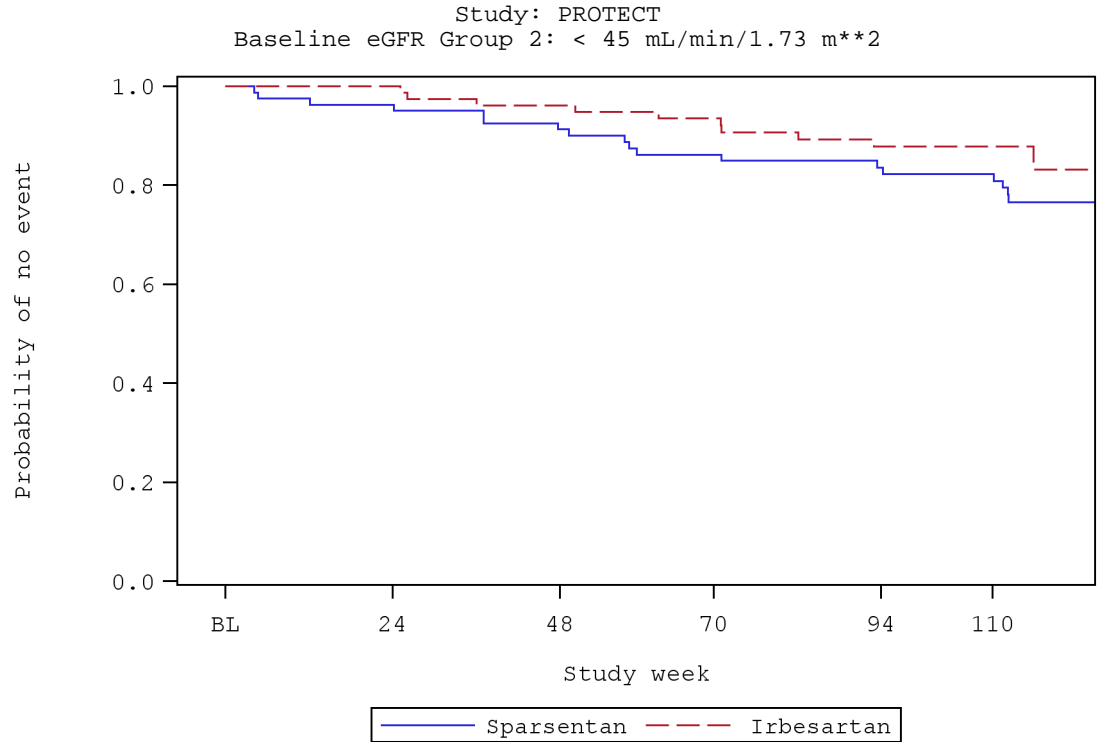
Figure PF2URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	26	23	21	18	15	13
Irbesartan	25	24	18	18	15	15

Complete remission means an urinary protein excretion of  $<0.3$  g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT2URFT\_FSTM

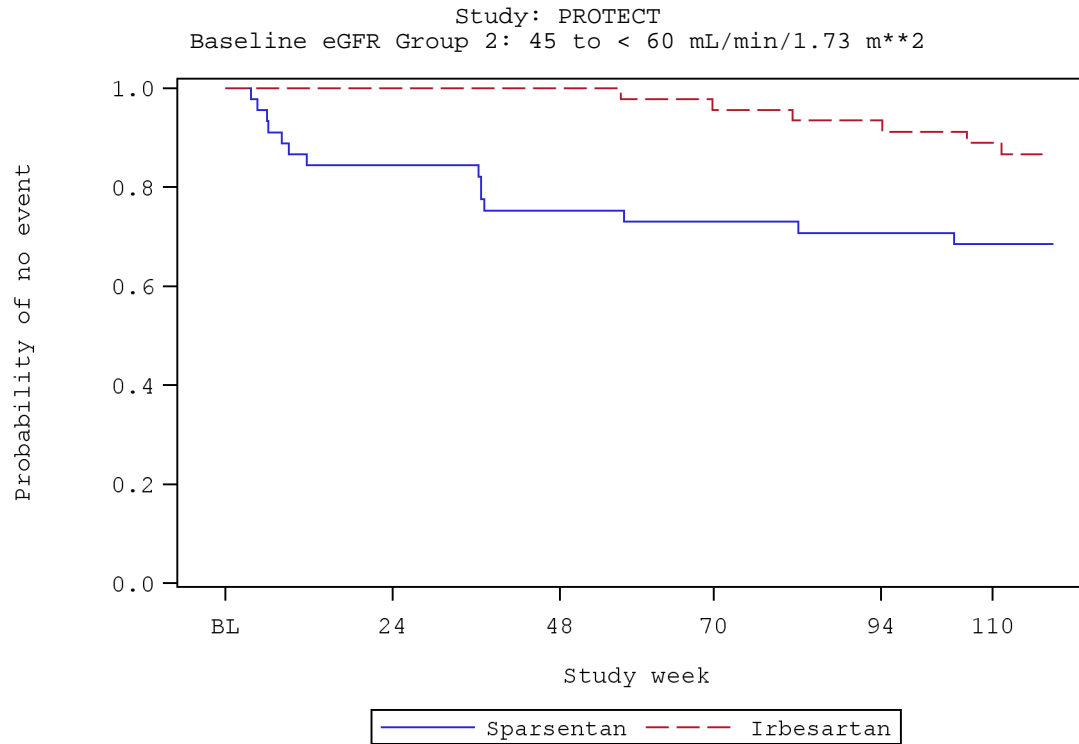
Figure PF2URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	82	77	72	68	62	60
Irbesartan	80	79	72	67	59	57

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT2URFT\_FSTM

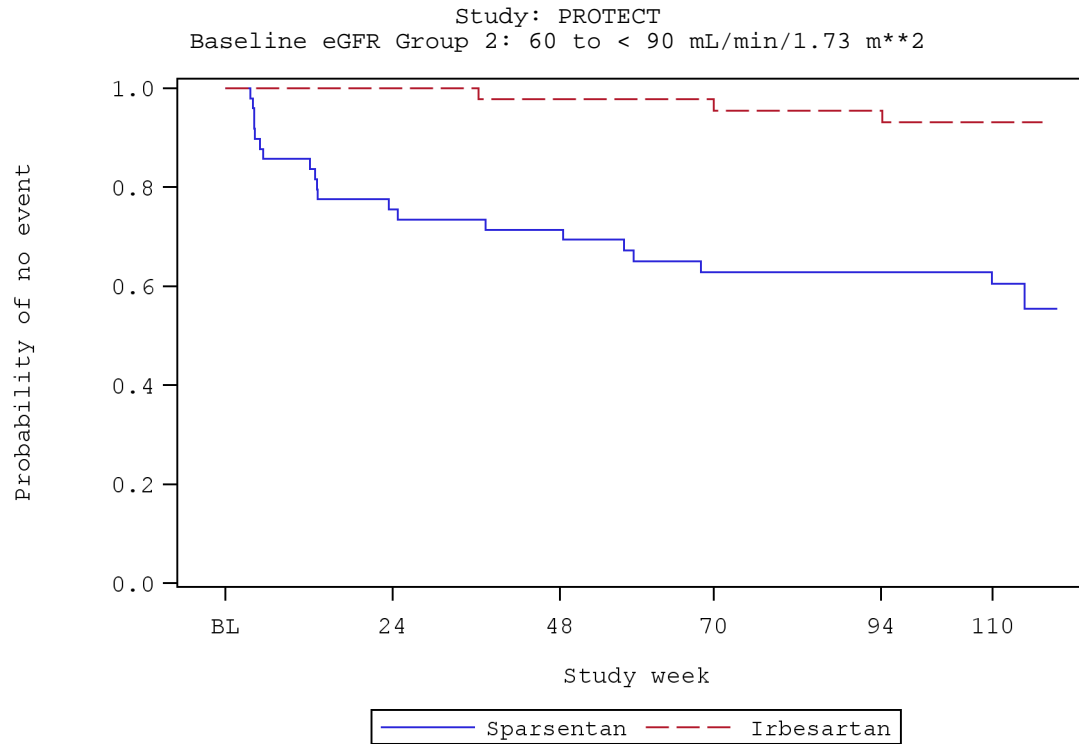
Figure PF2URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	45	38	33	32	31	28
Irbesartan	49	49	47	44	42	39

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT2URFT\_FSTM

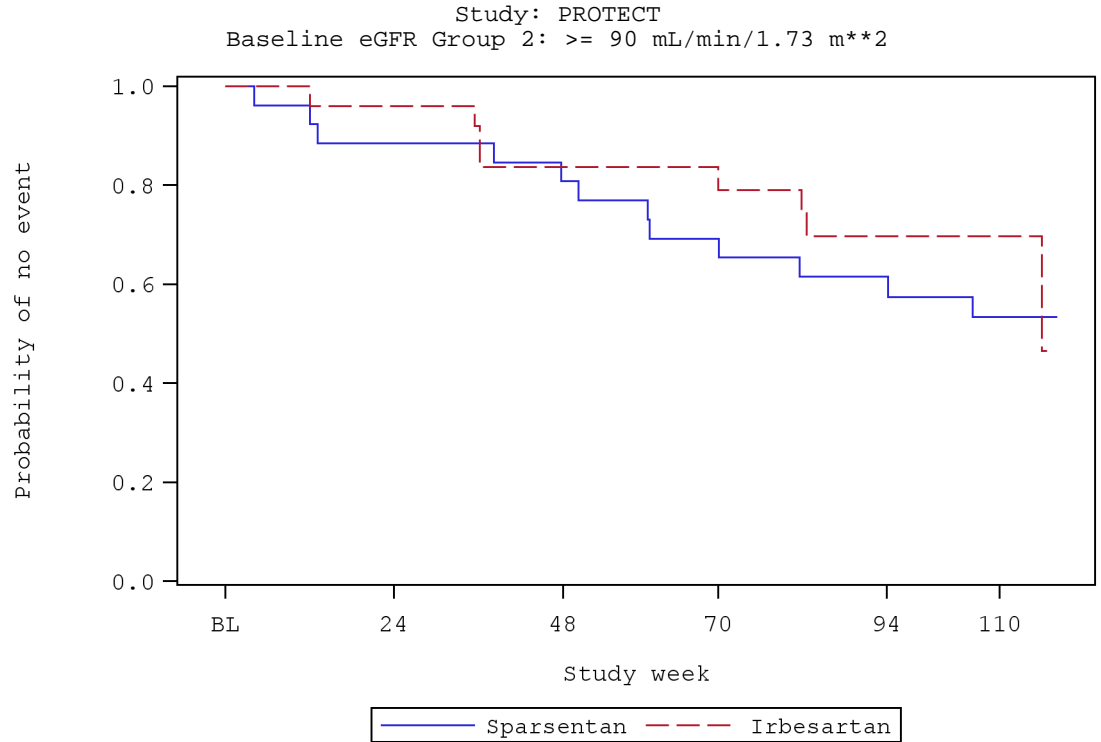
Figure PF2URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	49	37	35	29	29	26
Irbesartan	48	46	44	43	40	37

Complete remission means an urinary protein excretion of <0.3 g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT2URFT\_FSTM

Figure PF2URFT\_FSKM: Time to complete remission by subgroup  
 Full Analysis Set



Sparsentan	26	23	21	18	15	13
Irbesartan	25	24	18	18	15	15

Complete remission means an urinary protein excretion of  $<0.3$  g/day. No event means no complete remission.  
 eGFR = estimated glomerular filtration rate.  
 Reference table: PT2URFT\_FSTM

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Sex														
Male	eGFR	Baseline	Sparsentan	139	139 (100.0)	53.42 (22.51)	25.0	36.00	45.00	66.00	126.0			
			Irbesartan	143	143 (100.0)	57.02 (23.53)	26.0	38.00	50.00	70.00	123.0			
		Week 6	Sparsentan	139	134 (96.4)	52.42 (22.36)	21.0	35.00	47.50	65.00	121.0			
			Irbesartan	143	139 (97.2)	55.72 (23.29)	21.0	37.00	50.00	70.00	120.0			
		Week 36	Sparsentan	139	135 (97.1)	50.21 (21.52)	18.0	34.00	46.00	63.00	123.0			
			Irbesartan	143	130 (90.9)	53.07 (24.76)	7.0	34.00	48.00	66.00	125.0			
		Week 58	Sparsentan	139	127 (91.4)	50.13 (23.00)	13.0	33.00	44.00	64.00	127.0			
			Irbesartan	143	124 (86.7)	50.98 (23.81)	11.0	33.00	46.00	69.00	122.0			
		Week 110	Sparsentan	139	117 (84.2)	47.26 (23.76)	14.0	30.00	42.00	60.00	126.0			
			Irbesartan	143	111 (77.6)	44.68 (24.71)	11.0	26.00	38.00	61.00	123.0			
			Change from baseline in eGFR	Week 6	Sparsentan	139	134 (96.4)	-0.88 (6.68)	-27.0	-4.00	-1.00	2.00	20.0	0.05 [-0.19, 0.28]
					Irbesartan	143	139 (97.2)	-1.18 (6.62)	-31.0	-5.00	-1.00	2.00	20.0	
		Week 36		Sparsentan	139	135 (97.1)	-3.28 (7.75)	-30.0	-8.00	-3.00	1.00	26.0	0.08 [-0.16, 0.32]	
				Irbesartan	143	130 (90.9)	-3.98 (8.96)	-32.0	-8.00	-4.00	1.00	33.0		
		Week 58		Sparsentan	139	127 (91.4)	-3.25 (8.60)	-33.0	-8.00	-3.00	1.00	38.0	0.32 [0.07, 0.57]	
				Irbesartan	143	124 (86.7)	-6.05 (8.72)	-32.0	-11.00	-7.00	0.00	20.0		
		Week 110		Sparsentan	139	117 (84.2)	-5.89 (10.66)	-34.0	-12.00	-6.00	0.00	45.0	0.44 [0.18, 0.71]	
				Irbesartan	143	111 (77.6)	-10.65 (10.87)	-59.0	-16.00	-10.00	-4.00	13.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Female	eGFR	Baseline	Sparsentan	63	63 (100.0)	64.21 (26.63)	24.0	41.00	59.00	86.00	127.0			
			Irbesartan	59	59 (100.0)	57.19 (23.89)	27.0	39.00	50.00	70.00	123.0			
		Week 6	Sparsentan	63	60 (95.2)	63.28 (24.51)	26.0	44.00	60.00	82.00	120.0			
			Irbesartan	59	58 (98.3)	54.81 (23.52)	19.0	39.00	47.00	67.00	116.0			
		Week 36	Sparsentan	63	56 (88.9)	62.55 (23.00)	25.0	45.00	63.00	81.50	114.0			
			Irbesartan	59	56 (94.9)	53.34 (24.33)	15.0	37.00	44.50	66.00	114.0			
		Week 58	Sparsentan	63	61 (96.8)	60.33 (25.40)	18.0	43.00	57.00	84.00	132.0			
			Irbesartan	59	49 (83.1)	52.22 (24.30)	13.0	34.00	45.00	64.00	116.0			
		Week 110	Sparsentan	63	54 (85.7)	60.57 (25.04)	13.0	41.00	57.50	76.00	113.0			
			Irbesartan	59	44 (74.6)	49.95 (23.93)	23.0	32.50	43.00	56.50	113.0			
		Change from baseline in eGFR	Change from baseline in eGFR	Week 6	Sparsentan	63	60 (95.2)	-1.68 (9.58)	-43.0	-4.00	-0.50	3.00	14.0	0.12 [-0.24, 0.48]
					Irbesartan	59	58 (98.3)	-2.67 (6.44)	-19.0	-7.00	-3.00	3.00	9.0	
				Week 36	Sparsentan	63	56 (88.9)	-4.09 (9.54)	-39.0	-7.50	-2.00	0.50	17.0	0.11 [-0.26, 0.48]
					Irbesartan	59	56 (94.9)	-5.00 (6.33)	-24.0	-9.00	-4.50	-0.50	11.0	
Week 58	Sparsentan			63	61 (96.8)	-4.80 (9.88)	-34.0	-10.00	-6.00	1.00	24.0	-0.03 [-0.40, 0.35]		
	Irbesartan			59	49 (83.1)	-4.55 (8.97)	-24.0	-9.00	-3.00	0.00	23.0			
Week 110	Sparsentan			63	54 (85.7)	-4.11 (8.82)	-30.0	-10.00	-3.50	1.00	17.0	0.28 [-0.12, 0.68]		
	Irbesartan			59	44 (74.6)	-6.64 (9.17)	-26.0	-13.00	-6.50	0.00	15.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age														
<= 45 years	eGFR	Baseline	Sparsentan	96	96 (100.0)	63.15 (27.41)	25.0	41.00	55.50	83.00	127.0			
			Irbesartan	99	99 (100.0)	65.52 (26.04)	26.0	42.00	63.00	80.00	123.0			
		Week 6	Sparsentan	96	94 (97.9)	60.94 (26.38)	23.0	38.00	56.00	82.00	121.0			
			Irbesartan	99	98 (99.0)	63.04 (25.32)	23.0	43.00	57.00	80.00	120.0			
		Week 36	Sparsentan	96	91 (94.8)	58.91 (25.37)	18.0	37.00	51.00	78.00	123.0			
			Irbesartan	99	91 (91.9)	61.60 (28.00)	7.0	40.00	59.00	80.00	125.0			
		Week 58	Sparsentan	96	89 (92.7)	59.52 (27.26)	13.0	40.00	53.00	77.00	132.0			
			Irbesartan	99	82 (82.8)	59.61 (26.81)	13.0	38.00	56.00	77.00	122.0			
		Week 110	Sparsentan	96	77 (80.2)	57.58 (28.02)	18.0	34.00	50.00	75.00	126.0			
			Irbesartan	99	69 (69.7)	54.87 (28.25)	11.0	34.00	52.00	66.00	123.0			
			Change from baseline in eGFR	Week 6	Sparsentan	96	94 (97.9)	-2.27 (8.80)	-43.0	-5.00	-1.00	1.00	15.0	0.03 [-0.26, 0.31]
					Irbesartan	99	98 (99.0)	-2.47 (6.30)	-17.0	-7.00	-2.00	2.00	14.0	
		Week 36		Sparsentan	96	91 (94.8)	-5.02 (9.21)	-39.0	-10.00	-3.00	0.00	12.0	-0.00 [-0.29, 0.29]	
				Irbesartan	99	91 (91.9)	-5.01 (9.85)	-32.0	-12.00	-5.00	1.00	33.0		
		Week 58		Sparsentan	96	89 (92.7)	-4.64 (9.68)	-34.0	-8.00	-4.00	1.00	24.0	0.16 [-0.14, 0.46]	
				Irbesartan	99	82 (82.8)	-6.21 (9.90)	-32.0	-13.00	-7.00	0.00	23.0		
		Week 110		Sparsentan	96	77 (80.2)	-5.30 (9.11)	-30.0	-11.00	-4.00	0.00	15.0	0.47 [0.14, 0.80]	
				Irbesartan	99	69 (69.7)	-10.59 (13.34)	-59.0	-17.00	-11.00	-1.00	15.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 45 years	eGFR	Baseline	Sparsentan	106	106 (100.0)	51.02 (19.56)	24.0	36.00	45.00	64.00	103.0			
			Irbesartan	103	103 (100.0)	48.95 (17.55)	27.0	36.00	44.00	56.00	104.0			
		Week 6	Sparsentan	106	100 (94.3)	50.93 (19.40)	21.0	34.50	49.00	64.00	104.0			
			Irbesartan	103	99 (96.1)	47.94 (18.36)	19.0	35.00	43.00	59.00	102.0			
		Week 36	Sparsentan	106	100 (94.3)	49.21 (18.74)	23.0	34.00	46.00	62.50	97.0			
			Irbesartan	103	95 (92.2)	45.05 (17.36)	21.0	32.00	40.00	54.00	98.0			
		Week 58	Sparsentan	106	99 (93.4)	47.97 (19.71)	18.0	32.00	44.00	61.00	97.0			
			Irbesartan	103	91 (88.3)	43.88 (18.02)	11.0	30.00	38.00	55.00	95.0			
		Week 110	Sparsentan	106	94 (88.7)	46.46 (20.83)	13.0	30.00	43.00	62.00	95.0			
			Irbesartan	103	86 (83.5)	39.21 (18.47)	12.0	26.00	34.50	47.00	92.0			
			Change from baseline in eGFR	Week 6	Sparsentan	106	100 (94.3)	-0.06 (6.30)	-21.0	-3.00	0.00	3.00	20.0	0.11 [-0.17, 0.39]
					Irbesartan	103	99 (96.1)	-0.78 (6.78)	-31.0	-4.00	0.00	3.00	20.0	
		Week 36		Sparsentan	106	100 (94.3)	-2.15 (7.14)	-22.0	-6.00	-2.00	1.00	26.0	0.21 [-0.07, 0.50]	
				Irbesartan	103	95 (92.2)	-3.60 (6.34)	-25.0	-6.00	-4.00	-1.00	18.0		
Week 58	Sparsentan	106		99 (93.4)	-2.96 (8.40)	-19.0	-8.00	-3.00	1.00	38.0	0.27 [-0.02, 0.55]			
	Irbesartan	103		91 (88.3)	-5.10 (7.67)	-30.0	-9.00	-5.00	-1.00	20.0				
Week 110	Sparsentan	106		94 (88.7)	-5.35 (10.93)	-34.0	-11.00	-6.00	0.00	45.0	0.35 [0.05, 0.64]			
	Irbesartan	103		86 (83.5)	-8.64 (7.57)	-30.0	-13.00	-9.00	-4.00	13.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age at IgAN diagnosis														
<= 18 years	eGFR	Baseline	Sparsentan	9	9 (100.0)	57.33 (29.45)	32.0	35.00	41.00	70.00	119.0			
			Irbesartan	5	5 (100.0)	66.60 (34.46)	35.0	39.00	66.00	72.00	121.0			
		Week 6	Sparsentan	9	9 (100.0)	50.00 (20.99)	26.0	36.00	38.00	62.00	86.0			
			Irbesartan	5	5 (100.0)	64.80 (32.64)	34.0	37.00	61.00	79.00	113.0			
		Week 36	Sparsentan	9	8 (88.9)	51.75 (20.76)	34.0	35.00	46.00	66.50	85.0			
			Irbesartan	5	5 (100.0)	61.40 (38.80)	26.0	31.00	59.00	68.00	123.0			
		Week 58	Sparsentan	9	7 (77.8)	49.43 (24.58)	27.0	33.00	39.00	69.00	96.0			
			Irbesartan	5	4 (80.0)	67.00 (35.47)	26.0	40.50	66.00	93.50	110.0			
		Week 110	Sparsentan	9	5 (55.6)	56.60 (37.85)	23.0	29.00	41.00	77.00	113.0			
			Irbesartan	5	4 (80.0)	49.25 (39.76)	22.0	23.00	34.00	75.50	107.0			
		Change from baseline in eGFR		Week 6	Sparsentan	9	9 (100.0)	-7.33 (10.75)	-33.0	-8.00	-4.00	-1.00	1.0	-0.59 [-1.71, 0.52]
					Irbesartan	5	5 (100.0)	-1.80 (5.63)	-8.0	-5.00	-2.00	-1.00	7.0	
				Week 36	Sparsentan	9	8 (88.9)	-8.75 (12.10)	-34.0	-14.50	-3.00	-0.50	0.0	-0.35 [-1.48, 0.77]
					Irbesartan	5	5 (100.0)	-5.20 (4.44)	-9.0	-8.00	-7.00	-4.00	2.0	
				Week 58	Sparsentan	9	7 (77.8)	-11.00 (9.63)	-24.0	-23.00	-7.00	-2.00	-2.0	-0.38 [-1.62, 0.86]
					Irbesartan	5	4 (80.0)	-7.50 (8.39)	-13.0	-12.00	-11.00	-3.00	5.0	
				Week 110	Sparsentan	9	5 (55.6)	-6.00 (3.94)	-11.0	-7.00	-6.00	-6.00	0.0	2.34 [0.64, 4.04]
					Irbesartan	5	4 (80.0)	-16.00 (4.69)	-22.0	-19.50	-15.50	-12.50	-11.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 18 to 40 years	eGFR	Baseline	Sparsentan	102	102 (100.0)	63.25 (26.67)	25.0	42.00	56.50	85.00	127.0			
			Irbesartan	109	109 (100.0)	60.08 (25.53)	26.0	40.00	52.00	71.00	123.0			
		Week 6	Sparsentan	102	99 (97.1)	61.66 (26.00)	23.0	40.00	57.00	82.00	121.0			
			Irbesartan	109	106 (97.2)	58.44 (24.43)	23.0	40.00	54.00	72.00	120.0			
		Week 36	Sparsentan	102	97 (95.1)	59.29 (24.88)	18.0	39.00	55.00	77.00	123.0			
			Irbesartan	109	101 (92.7)	56.11 (26.73)	7.0	36.00	49.00	67.00	125.0			
		Week 58	Sparsentan	102	95 (93.1)	59.43 (26.68)	13.0	40.00	53.00	77.00	132.0			
			Irbesartan	109	90 (82.6)	55.01 (26.50)	13.0	35.00	49.50	71.00	122.0			
		Week 110	Sparsentan	102	83 (81.4)	57.30 (26.58)	23.0	35.00	52.00	74.00	126.0			
			Irbesartan	109	80 (73.4)	49.45 (26.84)	11.0	31.00	42.00	63.00	123.0			
		Change from baseline in eGFR		Week 6	Sparsentan	102	99 (97.1)	-1.43 (8.26)	-43.0	-4.00	-1.00	2.00	15.0	0.08 [-0.19, 0.36]
					Irbesartan	109	106 (97.2)	-2.04 (6.04)	-17.0	-5.00	-1.50	2.00	14.0	
	Week 36			Sparsentan	102	97 (95.1)	-4.58 (8.46)	-39.0	-10.00	-3.00	1.00	10.0	0.00 [-0.27, 0.28]	
				Irbesartan	109	101 (92.7)	-4.61 (9.19)	-32.0	-10.00	-3.00	1.00	33.0		
	Week 58			Sparsentan	102	95 (93.1)	-4.20 (9.13)	-34.0	-8.00	-4.00	1.00	24.0	0.14 [-0.15, 0.43]	
				Irbesartan	109	90 (82.6)	-5.44 (9.12)	-32.0	-10.00	-5.00	0.00	23.0		
	Week 110	Sparsentan	102	83 (81.4)	-5.72 (9.64)	-30.0	-12.00	-4.00	1.00	15.0	0.40 [0.09, 0.71]			
		Irbesartan	109	80 (73.4)	-10.14 (12.16)	-59.0	-15.00	-9.00	-3.00	15.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 40 years	eGFR	Baseline	Sparsentan	91	91 (100.0)	49.47 (18.57)	24.0	35.00	45.00	64.00	101.0		
			Irbesartan	88	88 (100.0)	52.80 (19.63)	27.0	38.00	47.50	65.50	106.0		
		Week 6	Sparsentan	91	86 (94.5)	49.62 (18.80)	21.0	34.00	46.00	62.00	99.0		
			Irbesartan	88	86 (97.7)	51.22 (20.75)	19.0	36.00	45.00	69.00	108.0		
		Week 36	Sparsentan	91	86 (94.5)	47.87 (18.40)	22.0	33.00	45.00	62.00	97.0		
			Irbesartan	88	80 (90.9)	48.90 (19.96)	15.0	33.00	42.50	61.50	109.0		
		Week 58	Sparsentan	91	86 (94.5)	47.14 (19.47)	18.0	31.00	43.50	60.00	97.0		
			Irbesartan	88	79 (89.8)	46.35 (18.83)	11.0	32.00	42.00	59.00	95.0		
		Week 110	Sparsentan	91	83 (91.2)	45.33 (20.80)	13.0	29.00	43.00	59.00	95.0		
			Irbesartan	88	71 (80.7)	42.32 (20.38)	12.0	26.00	37.00	57.00	92.0		
		Change from baseline in eGFR	Week 6	Sparsentan	91	86 (94.5)	-0.13 (6.24)	-21.0	-3.00	0.00	3.00	20.0	0.14 [-0.16, 0.44]
				Irbesartan	88	86 (97.7)	-1.09 (7.28)	-31.0	-5.00	-2.00	3.00	20.0	
	Week 36		Sparsentan	91	86 (94.5)	-1.84 (7.37)	-22.0	-5.00	-2.00	1.00	26.0	0.27 [-0.03, 0.58]	
			Irbesartan	88	80 (90.9)	-3.83 (7.16)	-25.0	-7.00	-4.00	0.00	18.0		
Week 58	Sparsentan		91	86 (94.5)	-2.67 (8.68)	-19.0	-8.00	-3.00	0.00	38.0	0.36 [0.05, 0.66]		
	Irbesartan		88	79 (89.8)	-5.73 (8.52)	-30.0	-10.00	-6.00	0.00	20.0			
Week 110	Sparsentan	91	83 (91.2)	-4.89 (10.88)	-34.0	-11.00	-6.00	-1.00	45.0	0.36 [0.04, 0.68]			
	Irbesartan	88	71 (80.7)	-8.44 (8.53)	-30.0	-14.00	-9.00	-3.00	13.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Geographic region												
North America	eGFR	Baseline	Sparsentan	35	35 (100.0)	56.54 (25.93)	25.0	38.00	50.00	67.00	121.0	
			Irbesartan	46	46 (100.0)	55.93 (23.44)	26.0	36.00	50.50	71.00	123.0	
	Week 6	Sparsentan	35	35 (100.0)	54.14 (25.21)	21.0	33.00	50.00	67.00	120.0		
		Irbesartan	46	45 (97.8)	53.33 (22.58)	21.0	36.00	46.00	68.00	107.0		
	Week 36	Sparsentan	35	34 (97.1)	53.41 (22.67)	23.0	34.00	48.00	65.00	104.0		
		Irbesartan	46	44 (95.7)	51.77 (24.21)	21.0	33.00	44.50	65.50	123.0		
	Week 58	Sparsentan	35	31 (88.6)	53.74 (24.97)	21.0	32.00	52.00	72.00	119.0		
		Irbesartan	46	39 (84.8)	49.72 (22.95)	16.0	31.00	43.00	67.00	114.0		
	Week 110	Sparsentan	35	26 (74.3)	49.88 (24.98)	16.0	28.00	45.00	71.00	113.0		
		Irbesartan	46	32 (69.6)	44.00 (21.59)	14.0	26.00	43.00	57.50	113.0		
	Change from baseline in eGFR	Week 6	Sparsentan	35	35 (100.0)	-2.40 (8.17)	-33.0	-4.00	-1.00	1.00	9.0	-0.04 [-0.48, 0.41]
			Irbesartan	46	45 (97.8)	-2.11 (7.92)	-17.0	-7.00	-3.00	3.00	20.0	
		Week 36	Sparsentan	35	34 (97.1)	-3.47 (7.99)	-34.0	-5.00	-2.00	0.00	9.0	0.09 [-0.35, 0.54]
			Irbesartan	46	44 (95.7)	-4.25 (8.37)	-19.0	-10.50	-4.00	0.50	18.0	
		Week 58	Sparsentan	35	31 (88.6)	-4.42 (10.28)	-33.0	-10.00	-4.00	1.00	24.0	-0.05 [-0.52, 0.43]
			Irbesartan	46	39 (84.8)	-3.97 (9.26)	-20.0	-10.00	-4.00	1.00	20.0	
		Week 110	Sparsentan	35	26 (74.3)	-5.19 (8.29)	-27.0	-12.00	-3.50	1.00	11.0	0.29 [-0.23, 0.81]
			Irbesartan	46	32 (69.6)	-8.13 (11.41)	-46.0	-14.50	-9.50	-1.00	15.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Europe	eGFR	Baseline	Sparsentan	98	98 (100.0)	55.84 (24.05)	24.0	36.00	47.50	71.00	127.0		
			Irbesartan	115	115 (100.0)	56.87 (24.29)	27.0	38.00	50.00	69.00	123.0		
		Week 6	Sparsentan	98	92 (93.9)	55.27 (23.93)	25.0	34.00	50.00	70.00	117.0		
			Irbesartan	115	112 (97.4)	55.64 (24.10)	19.0	37.00	48.00	69.00	120.0		
		Week 36	Sparsentan	98	91 (92.9)	53.55 (23.54)	18.0	36.00	47.00	68.00	123.0		
			Irbesartan	115	106 (92.2)	53.99 (25.91)	7.0	34.00	47.50	66.00	125.0		
		Week 58	Sparsentan	98	93 (94.9)	52.45 (24.38)	13.0	36.00	45.00	64.00	132.0		
			Irbesartan	115	98 (85.2)	51.84 (25.29)	11.0	33.00	46.50	65.00	122.0		
		Week 110	Sparsentan	98	83 (84.7)	49.95 (22.85)	14.0	33.00	45.00	64.00	126.0		
			Irbesartan	115	92 (80.0)	47.49 (26.59)	11.0	27.50	40.00	62.50	123.0		
		Change from baseline in eGFR	Week 6	Sparsentan	98	92 (93.9)	-0.50 (6.31)	-17.0	-4.00	0.00	3.00	15.0	0.16 [-0.12, 0.44]
				Irbesartan	115	112 (97.4)	-1.52 (6.33)	-31.0	-5.00	-1.50	2.00	14.0	
	Week 36		Sparsentan	98	91 (92.9)	-3.00 (7.62)	-30.0	-8.00	-2.00	2.00	17.0	0.10 [-0.18, 0.38]	
			Irbesartan	115	106 (92.2)	-3.76 (8.21)	-32.0	-7.00	-4.00	0.00	33.0		
	Week 58		Sparsentan	98	93 (94.9)	-3.23 (7.25)	-24.0	-8.00	-2.00	1.00	14.0	0.32 [0.04, 0.61]	
			Irbesartan	115	98 (85.2)	-5.76 (8.29)	-32.0	-10.00	-5.50	0.00	11.0		
Week 110	Sparsentan	98	83 (84.7)	-5.47 (7.74)	-30.0	-11.00	-5.00	0.00	17.0	0.43 [0.13, 0.73]			
	Irbesartan	115	92 (80.0)	-9.47 (10.35)	-59.0	-15.00	-9.00	-3.00	13.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Asia Pacific	eGFR	Baseline	Sparsentan	69	69 (100.0)	58.25 (24.19)	26.0	40.00	51.00	74.00	121.0			
			Irbesartan	41	41 (100.0)	58.90 (22.11)	31.0	41.00	52.00	69.00	112.0			
		Week 6	Sparsentan	69	67 (97.1)	57.33 (22.30)	26.0	38.00	54.00	70.00	121.0			
			Irbesartan	41	40 (97.6)	57.30 (22.20)	23.0	41.50	53.50	70.00	119.0			
		Week 36	Sparsentan	69	66 (95.7)	54.44 (21.60)	19.0	34.00	50.50	67.00	108.0			
			Irbesartan	41	36 (87.8)	52.36 (21.19)	15.0	37.50	47.50	70.00	101.0			
		Week 58	Sparsentan	69	64 (92.8)	54.72 (23.93)	14.0	37.00	50.00	73.00	121.0			
			Irbesartan	41	36 (87.8)	51.72 (21.33)	13.0	35.00	47.50	67.50	98.0			
		Week 110	Sparsentan	69	62 (89.9)	54.16 (27.49)	13.0	31.00	49.50	69.00	123.0			
			Irbesartan	41	31 (75.6)	44.55 (21.20)	11.0	29.00	38.00	59.00	104.0			
			Change from baseline in eGFR	Week 6	Sparsentan	69	67 (97.1)	-1.33 (9.03)	-43.0	-4.00	-1.00	3.00	20.0	0.00 [-0.39, 0.39]
					Irbesartan	41	40 (97.6)	-1.35 (5.71)	-19.0	-4.00	-1.50	2.00	9.0	
		Week 36		Sparsentan	69	66 (95.7)	-4.26 (9.35)	-39.0	-8.00	-4.00	-1.00	26.0	0.18 [-0.23, 0.59]	
				Irbesartan	41	36 (87.8)	-5.89 (8.28)	-24.0	-10.50	-5.50	-0.50	11.0		
Week 58	Sparsentan	69		64 (92.8)	-4.20 (10.71)	-34.0	-8.00	-5.00	0.00	38.0	0.28 [-0.13, 0.69]			
	Irbesartan	41		36 (87.8)	-7.06 (9.54)	-24.0	-13.00	-8.00	0.00	23.0				
Week 110	Sparsentan	69		62 (89.9)	-5.19 (13.30)	-34.0	-12.00	-6.50	0.00	45.0	0.47 [0.04, 0.91]			
	Irbesartan	41		31 (75.6)	-11.06 (10.34)	-34.0	-18.00	-11.00	-4.00	11.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline BMI														
< 27 kg/m**2	eGFR	Baseline	Sparsentan	83	83 (100.0)	58.66 (24.61)	24.0	39.00	53.00	72.00	126.0			
			Irbesartan	94	94 (100.0)	58.84 (26.11)	26.0	39.00	50.00	70.00	123.0			
		Week 6	Sparsentan	83	78 (94.0)	58.38 (22.75)	26.0	38.00	55.00	74.00	117.0			
			Irbesartan	94	91 (96.8)	56.63 (25.44)	19.0	37.00	50.00	71.00	120.0			
		Week 36	Sparsentan	83	77 (92.8)	56.36 (23.52)	19.0	34.00	51.00	72.00	123.0			
			Irbesartan	94	86 (91.5)	54.64 (28.46)	7.0	35.00	46.00	69.00	125.0			
		Week 58	Sparsentan	83	77 (92.8)	55.97 (24.36)	14.0	37.00	53.00	73.00	127.0			
			Irbesartan	94	75 (79.8)	53.33 (25.75)	13.0	34.00	49.00	67.00	122.0			
		Week 110	Sparsentan	83	70 (84.3)	53.83 (25.28)	13.0	33.00	49.50	68.00	126.0			
			Irbesartan	94	69 (73.4)	50.14 (27.37)	12.0	32.00	41.00	63.00	123.0			
			Change from baseline in eGFR	Week 6	Sparsentan	83	78 (94.0)	-1.12 (8.99)	-43.0	-4.00	0.00	3.00	20.0	0.15 [-0.15, 0.45]
					Irbesartan	94	91 (96.8)	-2.23 (5.56)	-16.0	-5.00	-2.00	2.00	12.0	
		Week 36		Sparsentan	83	77 (92.8)	-3.62 (9.89)	-39.0	-8.00	-2.00	0.00	26.0	0.19 [-0.12, 0.50]	
				Irbesartan	94	86 (91.5)	-5.41 (8.95)	-32.0	-11.00	-4.50	-1.00	33.0		
Week 58	Sparsentan	83		77 (92.8)	-3.39 (10.66)	-34.0	-8.00	-3.00	0.00	38.0	0.32 [-0.00, 0.64]			
	Irbesartan	94		75 (79.8)	-6.40 (8.16)	-32.0	-11.00	-7.00	-2.00	16.0				
Week 110	Sparsentan	83		70 (84.3)	-5.13 (11.91)	-34.0	-11.00	-6.00	0.00	45.0	0.36 [0.03, 0.70]			
	Irbesartan	94		69 (73.4)	-9.07 (9.87)	-36.0	-15.00	-9.00	-3.00	15.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
>= 27 kg/m**2	eGFR	Baseline	Sparsentan	119	119 (100.0)	55.47 (24.15)	25.0	38.00	49.00	69.00	127.0			
			Irbesartan	107	107 (100.0)	55.66 (21.19)	27.0	38.00	50.00	70.00	114.0			
		Week 6	Sparsentan	119	116 (97.5)	54.03 (23.97)	21.0	35.00	49.00	66.50	121.0			
			Irbesartan	107	105 (98.1)	54.59 (21.42)	21.0	39.00	48.00	69.00	119.0			
		Week 36	Sparsentan	119	114 (95.8)	52.12 (21.92)	18.0	36.00	47.00	65.00	114.0			
			Irbesartan	107	100 (93.5)	51.87 (20.70)	20.0	34.00	48.00	65.00	125.0			
		Week 58	Sparsentan	119	111 (93.3)	51.68 (24.08)	13.0	33.00	47.00	64.00	132.0			
			Irbesartan	107	98 (91.6)	49.81 (22.37)	11.0	32.00	45.50	66.00	121.0			
		Week 110	Sparsentan	119	101 (84.9)	49.83 (24.59)	14.0	31.00	45.00	63.00	123.0			
			Irbesartan	107	85 (79.4)	43.15 (21.71)	11.0	26.00	40.00	56.00	106.0			
		Change from baseline in eGFR		Week 6	Sparsentan	119	116 (97.5)	-1.14 (6.69)	-27.0	-4.00	-1.00	2.00	15.0	-0.01 [-0.27, 0.26]
					Irbesartan	107	105 (98.1)	-1.09 (7.37)	-31.0	-5.00	-1.00	3.00	20.0	
				Week 36	Sparsentan	119	114 (95.8)	-3.45 (7.07)	-26.0	-8.00	-3.00	1.00	12.0	-0.02 [-0.28, 0.25]
					Irbesartan	107	100 (93.5)	-3.33 (7.52)	-25.0	-6.50	-4.00	1.00	18.0	
				Week 58	Sparsentan	119	111 (93.3)	-4.01 (7.77)	-33.0	-8.00	-4.00	1.00	13.0	0.12 [-0.15, 0.39]
					Irbesartan	107	98 (91.6)	-5.03 (9.25)	-30.0	-10.00	-4.00	0.00	23.0	
				Week 110	Sparsentan	119	101 (84.9)	-5.47 (8.74)	-30.0	-11.00	-4.00	0.00	15.0	0.44 [0.15, 0.73]
					Irbesartan	107	85 (79.4)	-9.85 (11.16)	-59.0	-15.00	-11.00	-4.00	13.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Randomization strata														
eGFR Low and UP High	eGFR	Baseline	Sparsentan	71	71 (100.0)	40.28 (9.79)	25.0	31.00	39.00	47.00	67.0			
		Week 6	Irbesartan	74	74 (100.0)	41.78 (9.97)	26.0	34.00	39.00	50.00	66.0			
			Sparsentan	71	69 (97.2)	39.55 (10.77)	21.0	32.00	38.00	49.00	64.0			
		Week 36	Irbesartan	74	72 (97.3)	39.76 (9.92)	21.0	32.00	39.00	46.50	64.0			
			Sparsentan	71	68 (95.8)	38.09 (10.70)	18.0	31.50	36.50	46.00	65.0			
		Week 58	Irbesartan	74	66 (89.2)	36.35 (12.05)	7.0	29.00	35.50	42.00	88.0			
			Sparsentan	71	67 (94.4)	36.61 (10.86)	13.0	28.00	36.00	45.00	61.0			
		Week 110	Irbesartan	74	62 (83.8)	34.76 (12.05)	11.0	27.00	34.00	43.00	66.0			
			Sparsentan	71	59 (83.1)	33.95 (10.76)	14.0	27.00	33.00	42.00	56.0			
		Week 110	Irbesartan	74	57 (77.0)	30.81 (11.67)	11.0	24.00	29.00	40.00	63.0			
			Sparsentan	71	69 (97.2)	-0.54 (5.72)	-27.0	-3.00	-1.00	2.00	15.0	0.24 [-0.09, 0.57]		
		Change from baseline in eGFR	Change from baseline in eGFR	Week 6	Irbesartan	74	72 (97.3)	-1.82 (4.80)	-14.0	-5.00	-2.00	2.00	9.0	
					Sparsentan	71	68 (95.8)	-2.37 (5.54)	-15.0	-5.50	-2.50	1.00	10.0	0.47 [0.13, 0.82]
				Week 36	Irbesartan	74	66 (89.2)	-5.85 (8.85)	-32.0	-9.00	-5.00	-2.00	33.0	
Sparsentan	71				67 (94.4)	-4.01 (5.91)	-18.0	-8.00	-4.00	0.00	10.0	0.42 [0.07, 0.77]		
Week 58	Irbesartan			74	62 (83.8)	-6.97 (8.18)	-32.0	-10.00	-7.00	-1.00	11.0			
	Sparsentan			71	59 (83.1)	-6.64 (7.84)	-31.0	-12.00	-7.00	0.00	8.0	0.58 [0.21, 0.95]		
Week 110	Irbesartan			74	57 (77.0)	-11.23 (7.89)	-36.0	-16.00	-11.00	-5.00	7.0			
	Sparsentan			71	69 (97.2)	-0.54 (5.72)	-27.0	-3.00	-1.00	2.00	15.0	0.24 [-0.09, 0.57]		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR Low and UP Low	eGFR	Baseline	Sparsentan	55	55 (100.0)	42.13 (9.23)	24.0	35.00	42.00	50.00	62.0			
			Irbesartan	55	55 (100.0)	44.25 (9.36)	27.0	38.00	43.00	50.00	71.0			
		Week 6	Sparsentan	55	51 (92.7)	42.25 (10.06)	26.0	34.00	40.00	50.00	70.0			
			Irbesartan	55	54 (98.2)	43.24 (9.93)	19.0	36.00	43.00	48.00	68.0			
		Week 36	Sparsentan	55	49 (89.1)	42.27 (11.81)	25.0	34.00	41.00	48.00	76.0			
			Irbesartan	55	52 (94.5)	41.65 (9.85)	28.0	33.50	40.00	48.50	70.0			
		Week 58	Sparsentan	55	51 (92.7)	41.92 (13.53)	18.0	33.00	41.00	48.00	88.0			
			Irbesartan	55	49 (89.1)	41.51 (12.39)	22.0	33.00	39.00	49.00	82.0			
		Week 110	Sparsentan	55	47 (85.5)	39.96 (15.15)	13.0	29.00	35.00	48.00	95.0			
			Irbesartan	55	44 (80.0)	36.02 (11.13)	11.0	27.50	35.00	41.00	66.0			
		Change from baseline in eGFR		Week 6	Sparsentan	55	51 (92.7)	0.41 (6.69)	-17.0	-3.00	0.00	3.00	20.0	0.22 [-0.16, 0.61]
					Irbesartan	55	54 (98.2)	-1.09 (6.78)	-31.0	-4.00	0.00	3.00	16.0	
				Week 36	Sparsentan	55	49 (89.1)	0.20 (7.08)	-11.0	-4.00	-1.00	3.00	26.0	0.47 [0.08, 0.87]
	Irbesartan				55	52 (94.5)	-2.63 (4.79)	-12.0	-5.50	-3.00	0.50	13.0		
Week 58	Sparsentan			55	51 (92.7)	-0.49 (8.78)	-11.0	-6.00	-2.00	3.00	38.0	0.36 [-0.04, 0.75]		
	Irbesartan			55	49 (89.1)	-3.39 (7.42)	-23.0	-8.00	-4.00	1.00	23.0			
Week 110	Sparsentan	55	47 (85.5)	-2.72 (11.20)	-15.0	-9.00	-4.00	0.00	45.0	0.49 [0.07, 0.90]				
	Irbesartan	55	44 (80.0)	-7.50 (8.08)	-34.0	-12.00	-7.00	-1.50	11.0					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP High	eGFR	Baseline	Sparsentan	37	37 (100.0)	82.43 (20.55)	51.0	66.00	76.00	91.00	127.0			
			Irbesartan	36	36 (100.0)	79.81 (14.96)	57.0	69.00	74.00	91.50	109.0			
		Week 6	Sparsentan	37	35 (94.6)	78.86 (18.69)	56.0	62.00	74.00	90.00	121.0			
			Irbesartan	36	34 (94.4)	77.09 (14.76)	56.0	63.00	76.00	85.00	108.0			
		Week 36	Sparsentan	37	37 (100.0)	74.35 (18.89)	34.0	62.00	71.00	85.00	118.0			
			Irbesartan	36	33 (91.7)	73.76 (16.97)	40.0	62.00	68.00	86.00	123.0			
		Week 58	Sparsentan	37	34 (91.9)	78.12 (21.35)	46.0	62.00	74.00	88.00	132.0			
			Irbesartan	36	30 (83.3)	71.13 (15.67)	37.0	59.00	71.50	83.00	101.0			
		Week 110	Sparsentan	37	30 (81.1)	72.97 (21.22)	43.0	59.00	64.50	86.00	123.0			
			Irbesartan	36	27 (75.0)	64.15 (20.28)	14.0	53.00	63.00	74.00	106.0			
			Change from baseline in eGFR	Week 6	Sparsentan	37	35 (94.6)	-3.91 (11.54)	-43.0	-8.00	-1.00	4.00	12.0	-0.09 [-0.56, 0.38]
		Irbesartan			36	34 (94.4)	-3.09 (6.45)	-17.0	-6.00	-3.00	2.00	12.0		
		Week 36		Sparsentan	37	37 (100.0)	-8.08 (11.79)	-39.0	-13.00	-5.00	0.00	8.0	-0.16 [-0.63, 0.31]	
				Irbesartan	36	33 (91.7)	-6.42 (9.11)	-25.0	-12.00	-6.00	-1.00	14.0		
		Week 58		Sparsentan	37	34 (91.9)	-5.29 (12.83)	-34.0	-14.00	-5.50	5.00	24.0	0.21 [-0.28, 0.70]	
				Irbesartan	36	30 (83.3)	-7.60 (8.63)	-23.0	-14.00	-8.00	-2.00	10.0		
		Week 110		Sparsentan	37	30 (81.1)	-6.10 (11.65)	-34.0	-12.00	-5.00	1.00	15.0	0.63 [0.10, 1.16]	
				Irbesartan	36	27 (75.0)	-14.48 (14.85)	-59.0	-22.00	-12.00	-5.00	13.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP Low	eGFR	Baseline	Sparsentan	39	39 (100.0)	83.15 (16.11)	57.0	70.00	80.00	96.00	126.0			
			Irbesartan	37	37 (100.0)	84.57 (22.97)	50.0	66.00	79.00	104.00	123.0			
		Week 6	Sparsentan	39	39 (100.0)	81.46 (16.34)	53.0	67.00	78.00	98.00	113.0			
			Irbesartan	37	37 (100.0)	83.92 (20.66)	43.0	69.00	76.00	101.00	120.0			
		Week 36	Sparsentan	39	37 (94.9)	77.57 (16.79)	47.0	65.00	73.00	91.00	123.0			
			Irbesartan	37	35 (94.6)	82.49 (23.25)	41.0	64.00	74.00	102.00	125.0			
		Week 58	Sparsentan	39	36 (92.3)	77.75 (16.89)	53.0	63.50	74.50	90.00	127.0			
			Irbesartan	37	32 (86.5)	79.94 (22.04)	37.0	64.50	74.50	95.00	122.0			
		Week 110	Sparsentan	39	35 (89.7)	78.03 (18.80)	43.0	64.00	76.00	88.00	126.0			
			Irbesartan	37	27 (73.0)	77.22 (25.01)	33.0	61.00	69.00	107.00	123.0			
		Change from baseline in eGFR	Change from baseline in eGFR	Week 6	Sparsentan	39	39 (100.0)	-1.69 (7.14)	-20.0	-7.00	-1.00	3.00	14.0	-0.13 [-0.58, 0.32]
					Irbesartan	37	37 (100.0)	-0.65 (9.02)	-19.0	-6.00	0.00	6.00	20.0	
				Week 36	Sparsentan	39	37 (94.9)	-6.00 (7.15)	-26.0	-12.00	-4.00	-2.00	12.0	-0.50 [-0.97, -0.03]
					Irbesartan	37	35 (94.6)	-1.80 (9.42)	-24.0	-10.00	-3.00	5.00	18.0	
				Week 58	Sparsentan	39	36 (92.3)	-6.44 (8.93)	-33.0	-10.50	-6.00	-1.50	13.0	-0.18 [-0.66, 0.29]
					Irbesartan	37	32 (86.5)	-4.59 (11.22)	-24.0	-12.50	-2.00	2.00	20.0	
				Week 110	Sparsentan	39	35 (89.7)	-5.94 (10.43)	-30.0	-13.00	-5.00	3.00	12.0	-0.16 [-0.67, 0.34]
					Irbesartan	37	27 (73.0)	-4.19 (11.28)	-36.0	-13.00	-3.00	5.00	15.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m<sup>2</sup>.

eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High >= 60 ml/min/1.73m<sup>2</sup>, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 1														
< 60 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	127	127 (100.0)	41.06 (9.32)	24.0	33.00	41.00	49.00	59.0			
			Irbesartan	129	129 (100.0)	42.27 (8.60)	26.0	35.00	41.00	50.00	59.0			
		Week 6	Sparsentan	127	121 (95.3)	40.95 (10.86)	21.0	33.00	38.00	50.00	73.0			
			Irbesartan	129	127 (98.4)	41.72 (10.87)	19.0	34.00	41.00	47.00	76.0			
		Week 36	Sparsentan	127	118 (92.9)	39.92 (11.22)	18.0	32.00	39.00	47.00	76.0			
			Irbesartan	129	118 (91.5)	39.16 (12.20)	7.0	32.00	37.00	45.00	88.0			
		Week 58	Sparsentan	127	118 (92.9)	38.87 (12.23)	13.0	30.00	38.50	46.00	88.0			
			Irbesartan	129	111 (86.0)	38.30 (13.43)	11.0	29.00	36.00	45.00	82.0			
		Week 110	Sparsentan	127	107 (84.3)	36.90 (13.28)	13.0	28.00	34.00	45.00	95.0			
			Irbesartan	129	102 (79.1)	33.95 (12.97)	11.0	25.00	32.00	41.00	72.0			
			Change from baseline in eGFR	Week 6	Sparsentan	127	121 (95.3)	0.15 (6.28)	-27.0	-3.00	0.00	2.00	20.0	0.13 [-0.12, 0.38]
					Irbesartan	129	127 (98.4)	-0.65 (6.12)	-31.0	-4.00	0.00	3.00	20.0	
		Week 36		Sparsentan	127	118 (92.9)	-1.18 (6.28)	-15.0	-5.00	-2.00	2.00	26.0	0.31 [0.06, 0.57]	
				Irbesartan	129	118 (91.5)	-3.34 (7.51)	-32.0	-6.00	-3.00	0.00	33.0		
Week 58	Sparsentan	127		118 (92.9)	-2.36 (7.43)	-18.0	-7.00	-2.50	1.00	38.0	0.25 [-0.01, 0.51]			
	Irbesartan	129		111 (86.0)	-4.26 (8.02)	-30.0	-9.00	-5.00	1.00	23.0				
Week 110	Sparsentan	127		107 (84.3)	-4.58 (9.41)	-31.0	-10.00	-6.00	0.00	45.0	0.42 [0.15, 0.70]			
	Irbesartan	129		102 (79.1)	-8.32 (8.27)	-34.0	-14.00	-9.00	-2.00	13.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
60 to < 90 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	49	49 (100.0)	72.49 (8.30)	60.0	66.00	71.00	78.00	89.0			
			Irbesartan	48	48 (100.0)	71.58 (7.62)	60.0	66.00	70.00	76.00	89.0			
		Week 6	Sparsentan	49	47 (95.9)	69.62 (8.73)	56.0	62.00	69.00	75.00	90.0			
			Irbesartan	48	45 (93.8)	68.64 (9.54)	49.0	61.00	69.00	74.00	85.0			
		Week 36	Sparsentan	49	47 (95.9)	66.15 (10.57)	34.0	62.00	65.00	72.00	88.0			
			Irbesartan	48	44 (91.7)	64.77 (11.13)	36.0	58.00	64.00	70.00	87.0			
		Week 58	Sparsentan	49	44 (89.8)	67.16 (9.86)	46.0	59.50	65.00	74.00	91.0			
			Irbesartan	48	41 (85.4)	62.78 (11.30)	34.0	55.00	64.00	72.00	83.0			
		Week 110	Sparsentan	49	43 (87.8)	65.33 (12.16)	36.0	59.00	64.00	71.00	100.0			
			Irbesartan	48	37 (77.1)	57.08 (14.58)	14.0	52.00	57.00	67.00	86.0			
		Change from baseline in eGFR		Week 6	Sparsentan	49	47 (95.9)	-2.70 (7.27)	-21.0	-7.00	-2.00	3.00	12.0	0.04 [-0.37, 0.45]
					Irbesartan	48	45 (93.8)	-2.98 (7.62)	-19.0	-7.00	-3.00	3.00	14.0	
				Week 36	Sparsentan	49	47 (95.9)	-6.21 (8.66)	-30.0	-12.00	-5.00	-1.00	12.0	0.05 [-0.36, 0.46]
					Irbesartan	48	44 (91.7)	-6.64 (8.51)	-30.0	-12.00	-6.00	-1.50	12.0	
				Week 58	Sparsentan	49	44 (89.8)	-5.18 (9.56)	-24.0	-11.00	-7.00	-1.50	24.0	0.39 [-0.04, 0.82]
					Irbesartan	48	41 (85.4)	-8.95 (9.92)	-32.0	-16.00	-9.00	-2.00	10.0	
				Week 110	Sparsentan	49	43 (87.8)	-7.30 (10.41)	-34.0	-13.00	-5.00	0.00	14.0	0.57 [0.12, 1.01]
					Irbesartan	48	37 (77.1)	-14.14 (13.78)	-59.0	-21.00	-12.00	-5.00	6.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
>= 90 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	26	26 (100.0)	104.00 (11.43)	90.0	95.00	102.00	111.00	127.0			
			Irbesartan	25	25 (100.0)	105.56 (9.98)	91.0	98.00	105.00	112.00	123.0			
		Week 6	Sparsentan	26	26 (100.0)	99.77 (11.42)	77.0	92.00	99.00	104.00	121.0			
			Irbesartan	25	25 (100.0)	101.48 (10.76)	80.0	95.00	101.00	108.00	120.0			
		Week 36	Sparsentan	26	26 (100.0)	94.73 (12.40)	63.0	88.00	93.50	101.00	123.0			
			Irbesartan	25	24 (96.0)	100.63 (15.82)	73.0	89.00	99.50	112.50	125.0			
		Week 58	Sparsentan	26	26 (100.0)	96.31 (16.63)	61.0	85.00	94.00	105.00	132.0			
			Irbesartan	25	21 (84.0)	97.90 (13.03)	77.0	88.00	96.00	110.00	122.0			
		Week 110	Sparsentan	26	21 (80.8)	97.33 (16.09)	60.0	87.00	96.00	108.00	126.0			
			Irbesartan	25	16 (64.0)	98.94 (15.96)	68.0	86.00	105.00	112.00	123.0			
		Change from baseline in eGFR		Week 6	Sparsentan	26	26 (100.0)	-4.23 (12.06)	-43.0	-5.00	-1.00	1.00	13.0	-0.02 [-0.56, 0.53]
					Irbesartan	25	25 (100.0)	-4.08 (6.00)	-16.0	-8.00	-3.00	0.00	7.0	
				Week 36	Sparsentan	26	26 (100.0)	-9.27 (11.24)	-39.0	-13.00	-6.50	-2.00	7.0	-0.42 [-0.98, 0.14]
					Irbesartan	25	24 (96.0)	-4.67 (10.49)	-24.0	-12.00	-7.00	3.50	15.0	
				Week 58	Sparsentan	26	26 (100.0)	-7.69 (12.92)	-34.0	-17.00	-6.00	1.00	13.0	-0.12 [-0.70, 0.46]
					Irbesartan	25	21 (84.0)	-6.33 (8.96)	-24.0	-11.00	-8.00	0.00	16.0	
				Week 110	Sparsentan	26	21 (80.8)	-5.10 (12.77)	-30.0	-17.00	-6.00	5.00	15.0	0.10 [-0.55, 0.75]
					Irbesartan	25	16 (64.0)	-6.38 (12.33)	-30.0	-15.50	-6.00	0.50	15.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 2														
< 45 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	82	82 (100.0)	35.37 (5.59)	24.0	31.00	35.00	41.00	44.0			
			Irbesartan	80	80 (100.0)	36.58 (4.61)	26.0	33.00	37.00	40.00	44.0			
		Week 6	Sparsentan	82	79 (96.3)	36.08 (8.13)	21.0	31.00	34.00	40.00	58.0			
			Irbesartan	80	78 (97.5)	35.69 (6.41)	19.0	31.00	36.50	40.00	50.0			
		Week 36	Sparsentan	82	76 (92.7)	34.32 (8.11)	18.0	29.00	34.00	38.50	57.0			
			Irbesartan	80	71 (88.8)	32.23 (6.84)	7.0	29.00	33.00	37.00	47.0			
		Week 58	Sparsentan	82	76 (92.7)	32.68 (8.68)	13.0	27.00	33.00	38.00	63.0			
			Irbesartan	80	66 (82.5)	30.65 (7.82)	11.0	27.00	31.00	35.00	52.0			
		Week 110	Sparsentan	82	68 (82.9)	30.76 (9.53)	13.0	26.00	30.00	34.50	75.0			
			Irbesartan	80	63 (78.8)	27.60 (8.30)	11.0	23.00	26.00	32.00	55.0			
			Change from baseline in eGFR	Week 6	Sparsentan	82	79 (96.3)	0.85 (5.00)	-10.0	-2.00	0.00	3.00	15.0	0.37 [0.05, 0.68]
					Irbesartan	80	78 (97.5)	-0.90 (4.49)	-16.0	-4.00	-1.00	2.00	10.0	
		Week 36		Sparsentan	82	76 (92.7)	-1.05 (5.87)	-15.0	-4.00	-1.00	2.50	21.0	0.54 [0.22, 0.87]	
				Irbesartan	80	71 (88.8)	-4.32 (6.15)	-32.0	-6.00	-3.00	0.00	5.0		
Week 58	Sparsentan	82		76 (92.7)	-2.82 (6.34)	-18.0	-6.00	-3.00	0.00	27.0	0.46 [0.13, 0.80]			
	Irbesartan	80		66 (82.5)	-5.77 (6.46)	-30.0	-9.00	-6.00	-1.00	8.0				
Week 110	Sparsentan	82		68 (82.9)	-4.82 (7.97)	-21.0	-10.00	-6.00	-1.00	39.0	0.57 [0.22, 0.92]			
	Irbesartan	80		63 (78.8)	-9.08 (6.90)	-28.0	-14.00	-10.00	-3.00	11.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
45 to < 60 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	45	45 (100.0)	51.42 (4.57)	45.0	48.00	51.00	55.00	59.0			
			Irbesartan	49	49 (100.0)	51.57 (4.47)	45.0	48.00	51.00	55.00	59.0			
		Week 6	Sparsentan	45	42 (93.3)	50.12 (9.36)	23.0	45.00	50.00	54.00	73.0			
			Irbesartan	49	49 (100.0)	51.31 (9.57)	21.0	46.00	50.00	57.00	76.0			
		Week 36	Sparsentan	45	42 (93.3)	50.05 (8.69)	34.0	45.00	48.00	56.00	76.0			
			Irbesartan	49	47 (95.9)	49.64 (11.00)	33.0	42.00	48.00	54.00	88.0			
		Week 58	Sparsentan	45	42 (93.3)	50.07 (9.46)	38.0	44.00	47.50	53.00	88.0			
			Irbesartan	49	45 (91.8)	49.51 (12.00)	27.0	42.00	49.00	56.00	82.0			
		Week 110	Sparsentan	45	39 (86.7)	47.59 (12.14)	27.0	40.00	47.00	52.00	95.0			
			Irbesartan	49	39 (79.6)	44.21 (12.64)	11.0	36.00	42.00	55.00	72.0			
		Change from baseline in eGFR		Week 6	Sparsentan	45	42 (93.3)	-1.17 (8.07)	-27.0	-4.00	-0.50	2.00	20.0	-0.11 [-0.52, 0.30]
					Irbesartan	49	49 (100.0)	-0.27 (8.11)	-31.0	-4.00	0.00	4.00	20.0	
				Week 36	Sparsentan	45	42 (93.3)	-1.40 (7.04)	-12.0	-5.00	-2.00	2.00	26.0	0.05 [-0.36, 0.47]
					Irbesartan	49	47 (95.9)	-1.85 (9.07)	-19.0	-7.00	-3.00	2.00	33.0	
				Week 58	Sparsentan	45	42 (93.3)	-1.52 (9.10)	-15.0	-8.00	-2.00	2.00	38.0	0.06 [-0.36, 0.48]
					Irbesartan	49	45 (91.8)	-2.04 (9.53)	-20.0	-8.00	-3.00	4.00	23.0	
				Week 110	Sparsentan	45	39 (86.7)	-4.15 (11.60)	-31.0	-11.00	-5.00	0.00	45.0	0.27 [-0.17, 0.72]
					Irbesartan	49	39 (79.6)	-7.10 (10.08)	-34.0	-15.00	-7.00	1.00	13.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
60 to < 90 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	49	49 (100.0)	72.49 (8.30)	60.0	66.00	71.00	78.00	89.0			
			Irbesartan	48	48 (100.0)	71.58 (7.62)	60.0	66.00	70.00	76.00	89.0			
		Week 6	Sparsentan	49	47 (95.9)	69.62 (8.73)	56.0	62.00	69.00	75.00	90.0			
			Irbesartan	48	45 (93.8)	68.64 (9.54)	49.0	61.00	69.00	74.00	85.0			
		Week 36	Sparsentan	49	47 (95.9)	66.15 (10.57)	34.0	62.00	65.00	72.00	88.0			
			Irbesartan	48	44 (91.7)	64.77 (11.13)	36.0	58.00	64.00	70.00	87.0			
		Week 58	Sparsentan	49	44 (89.8)	67.16 (9.86)	46.0	59.50	65.00	74.00	91.0			
			Irbesartan	48	41 (85.4)	62.78 (11.30)	34.0	55.00	64.00	72.00	83.0			
		Week 110	Sparsentan	49	43 (87.8)	65.33 (12.16)	36.0	59.00	64.00	71.00	100.0			
			Irbesartan	48	37 (77.1)	57.08 (14.58)	14.0	52.00	57.00	67.00	86.0			
		Change from baseline in eGFR		Week 6	Sparsentan	49	47 (95.9)	-2.70 (7.27)	-21.0	-7.00	-2.00	3.00	12.0	0.04 [-0.37, 0.45]
					Irbesartan	48	45 (93.8)	-2.98 (7.62)	-19.0	-7.00	-3.00	3.00	14.0	
				Week 36	Sparsentan	49	47 (95.9)	-6.21 (8.66)	-30.0	-12.00	-5.00	-1.00	12.0	0.05 [-0.36, 0.46]
					Irbesartan	48	44 (91.7)	-6.64 (8.51)	-30.0	-12.00	-6.00	-1.50	12.0	
				Week 58	Sparsentan	49	44 (89.8)	-5.18 (9.56)	-24.0	-11.00	-7.00	-1.50	24.0	0.39 [-0.04, 0.82]
					Irbesartan	48	41 (85.4)	-8.95 (9.92)	-32.0	-16.00	-9.00	-2.00	10.0	
				Week 110	Sparsentan	49	43 (87.8)	-7.30 (10.41)	-34.0	-13.00	-5.00	0.00	14.0	0.57 [0.12, 1.01]
					Irbesartan	48	37 (77.1)	-14.14 (13.78)	-59.0	-21.00	-12.00	-5.00	6.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	eGFR	Baseline	Sparsentan	26	26 (100.0)	104.00 (11.43)	90.0	95.00	102.00	111.00	127.0		
			Irbesartan	25	25 (100.0)	105.56 (9.98)	91.0	98.00	105.00	112.00	123.0		
		Week 6	Sparsentan	26	26 (100.0)	99.77 (11.42)	77.0	92.00	99.00	104.00	121.0		
			Irbesartan	25	25 (100.0)	101.48 (10.76)	80.0	95.00	101.00	108.00	120.0		
		Week 36	Sparsentan	26	26 (100.0)	94.73 (12.40)	63.0	88.00	93.50	101.00	123.0		
			Irbesartan	25	24 (96.0)	100.63 (15.82)	73.0	89.00	99.50	112.50	125.0		
		Week 58	Sparsentan	26	26 (100.0)	96.31 (16.63)	61.0	85.00	94.00	105.00	132.0		
			Irbesartan	25	21 (84.0)	97.90 (13.03)	77.0	88.00	96.00	110.00	122.0		
		Week 110	Sparsentan	26	21 (80.8)	97.33 (16.09)	60.0	87.00	96.00	108.00	126.0		
		Irbesartan	25	16 (64.0)	98.94 (15.96)	68.0	86.00	105.00	112.00	123.0			
		Change from baseline in eGFR	Week 6	Sparsentan	26	26 (100.0)	-4.23 (12.06)	-43.0	-5.00	-1.00	1.00	13.0	-0.02 [-0.56, 0.53]
			Irbesartan	25	25 (100.0)	-4.08 (6.00)	-16.0	-8.00	-3.00	0.00	7.0		
	Week 36		Sparsentan	26	26 (100.0)	-9.27 (11.24)	-39.0	-13.00	-6.50	-2.00	7.0	-0.42 [-0.98, 0.14]	
			Irbesartan	25	24 (96.0)	-4.67 (10.49)	-24.0	-12.00	-7.00	3.50	15.0		
	Week 58		Sparsentan	26	26 (100.0)	-7.69 (12.92)	-34.0	-17.00	-6.00	1.00	13.0	-0.12 [-0.70, 0.46]	
			Irbesartan	25	21 (84.0)	-6.33 (8.96)	-24.0	-11.00	-8.00	0.00	16.0		
	Week 110		Sparsentan	26	21 (80.8)	-5.10 (12.77)	-30.0	-17.00	-6.00	5.00	15.0	0.10 [-0.55, 0.75]	
			Irbesartan	25	16 (64.0)	-6.38 (12.33)	-30.0	-15.50	-6.00	0.50	15.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline urine protein excretion														
<= 1.75 g/day	eGFR	Baseline	Sparsentan	98	98 (100.0)	59.05 (24.14)	24.0	40.00	53.50	77.00	126.0			
			Irbesartan	93	93 (100.0)	59.77 (25.25)	27.0	40.00	52.00	71.00	123.0			
		Week 6	Sparsentan	98	94 (95.9)	59.35 (23.61)	21.0	38.00	54.00	73.00	113.0			
			Irbesartan	93	91 (97.8)	58.98 (24.75)	19.0	41.00	53.00	72.00	120.0			
		Week 36	Sparsentan	98	91 (92.9)	57.85 (22.67)	23.0	38.00	55.00	73.00	123.0			
			Irbesartan	93	86 (92.5)	57.51 (26.15)	20.0	37.00	49.50	70.00	125.0			
		Week 58	Sparsentan	98	92 (93.9)	56.70 (23.26)	18.0	38.50	52.50	72.00	127.0			
			Irbesartan	93	79 (84.9)	56.34 (25.14)	13.0	36.00	52.00	72.00	122.0			
		Week 110	Sparsentan	98	87 (88.8)	55.64 (25.71)	13.0	33.00	52.00	75.00	126.0			
			Irbesartan	93	71 (76.3)	49.86 (26.35)	11.0	31.00	42.00	64.00	123.0			
			Change from baseline in eGFR	Week 6	Sparsentan	98	94 (95.9)	-0.27 (6.80)	-17.0	-4.00	0.00	3.00	20.0	0.09 [-0.20, 0.38]
					Irbesartan	93	91 (97.8)	-0.91 (7.95)	-31.0	-5.00	0.00	4.00	20.0	
				Week 36	Sparsentan	98	91 (92.9)	-1.77 (7.25)	-18.0	-5.00	-2.00	2.00	26.0	0.06 [-0.23, 0.36]
					Irbesartan	93	86 (92.5)	-2.24 (7.44)	-24.0	-7.00	-3.00	2.00	18.0	
				Week 58	Sparsentan	98	92 (93.9)	-2.72 (9.36)	-33.0	-7.00	-2.50	1.50	38.0	0.10 [-0.20, 0.40]
					Irbesartan	93	79 (84.9)	-3.67 (9.21)	-24.0	-10.00	-3.00	1.00	23.0	
				Week 110	Sparsentan	98	87 (88.8)	-3.47 (10.48)	-27.0	-10.00	-4.00	0.00	45.0	0.36 [0.04, 0.67]
					Irbesartan	93	71 (76.3)	-7.27 (10.87)	-59.0	-12.00	-6.00	-1.00	15.0	

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 1.75 g/day	eGFR	Baseline	Sparsentan	104	104 (100.0)	54.64 (24.43)	25.0	37.00	48.00	66.00	127.0			
			Irbesartan	109	109 (100.0)	54.76 (21.90)	26.0	38.00	48.00	68.00	116.0			
		Week 6	Sparsentan	104	100 (96.2)	52.42 (23.07)	23.0	34.50	47.50	63.00	121.0			
			Irbesartan	109	106 (97.2)	52.42 (21.65)	21.0	37.00	46.00	64.00	116.0			
		Week 36	Sparsentan	104	100 (96.2)	50.18 (22.05)	18.0	34.00	46.00	63.50	118.0			
			Irbesartan	109	100 (91.7)	49.40 (22.58)	7.0	33.00	42.50	62.50	123.0			
		Week 58	Sparsentan	104	96 (92.3)	50.31 (24.83)	13.0	32.00	44.50	61.50	132.0			
			Irbesartan	109	94 (86.2)	47.13 (22.04)	11.0	31.00	40.50	60.00	116.0			
		Week 110	Sparsentan	104	84 (80.8)	47.14 (23.36)	14.0	30.00	42.50	56.00	123.0			
			Irbesartan	109	84 (77.1)	43.07 (22.56)	11.0	26.00	38.50	57.00	113.0			
			Change from baseline in eGFR	Week 6	Sparsentan	104	100 (96.2)	-1.94 (8.37)	-43.0	-4.00	-1.00	1.50	15.0	0.04 [-0.23, 0.31]
		Irbesartan			109	106 (97.2)	-2.23 (5.09)	-17.0	-5.00	-2.00	1.00	12.0		
		Week 36		Sparsentan	104	100 (96.2)	-5.11 (8.88)	-39.0	-9.50	-3.50	0.00	10.0	0.11 [-0.17, 0.39]	
				Irbesartan	109	100 (91.7)	-6.05 (8.55)	-32.0	-10.00	-5.00	-2.00	33.0		
Week 58	Sparsentan	104		96 (92.3)	-4.75 (8.65)	-34.0	-10.00	-5.00	0.00	13.0	0.30 [0.01, 0.59]			
	Irbesartan	109		94 (86.2)	-7.27 (8.11)	-32.0	-12.00	-7.00	-2.00	11.0				
Week 110	Sparsentan	104		84 (80.8)	-7.25 (9.42)	-34.0	-12.00	-7.00	-1.00	15.0	0.43 [0.12, 0.74]			
	Irbesartan	109		84 (77.1)	-11.40 (9.93)	-46.0	-17.50	-11.00	-4.50	13.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Baseline use of antihypertensives												
Yes	eGFR	Baseline	Sparsentan	90	90 (100.0)	50.91 (21.73)	25.0	35.00	44.00	65.00	127.0	
			Irbesartan	88	88 (100.0)	50.57 (18.19)	26.0	37.00	45.50	59.50	108.0	
	Week 6	Sparsentan	90	86 (95.6)	50.20 (21.77)	23.0	33.00	44.00	60.00	121.0		
		Irbesartan	88	84 (95.5)	48.64 (18.17)	19.0	35.00	44.00	57.50	108.0		
	Week 36	Sparsentan	90	86 (95.6)	48.28 (20.70)	18.0	33.00	45.00	57.00	114.0		
		Irbesartan	88	80 (90.9)	47.14 (19.70)	7.0	32.50	40.50	57.00	109.0		
	Week 58	Sparsentan	90	82 (91.1)	48.35 (22.56)	13.0	32.00	43.50	59.00	132.0		
		Irbesartan	88	77 (87.5)	43.65 (17.50)	11.0	31.00	38.00	54.00	97.0		
	Week 110	Sparsentan	90	75 (83.3)	46.47 (21.90)	14.0	31.00	40.00	56.00	123.0		
		Irbesartan	88	72 (81.8)	38.81 (17.80)	11.0	26.00	34.00	47.00	86.0		
	Change from baseline in eGFR	Week 6	Sparsentan	90	86 (95.6)	-0.83 (6.90)	-27.0	-4.00	-1.00	3.00	13.0	0.15 [-0.15, 0.45]
			Irbesartan	88	84 (95.5)	-1.87 (6.95)	-31.0	-4.50	-1.00	2.00	13.0	
		Week 36	Sparsentan	90	86 (95.6)	-2.74 (7.08)	-30.0	-5.00	-2.00	1.00	21.0	0.09 [-0.22, 0.39]
			Irbesartan	88	80 (90.9)	-3.40 (8.29)	-32.0	-6.00	-4.00	0.00	33.0	
Week 58		Sparsentan	90	82 (91.1)	-3.27 (7.19)	-18.0	-8.00	-4.00	0.00	27.0	0.39 [0.07, 0.70]	
		Irbesartan	88	77 (87.5)	-6.43 (9.12)	-32.0	-11.00	-7.00	1.00	12.0		
Week 110		Sparsentan	90	75 (83.3)	-4.88 (9.41)	-34.0	-11.00	-6.00	0.00	39.0	0.56 [0.23, 0.89]	
		Irbesartan	88	72 (81.8)	-10.60 (11.03)	-59.0	-16.50	-10.00	-3.00	13.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
No	eGFR	Baseline	Sparsentan	112	112 (100.0)	61.50 (25.35)	24.0	41.00	57.00	80.00	126.0		
			Irbesartan	114	114 (100.0)	62.09 (25.99)	28.0	41.00	55.50	76.00	123.0		
		Week 6	Sparsentan	112	108 (96.4)	60.22 (24.02)	21.0	38.00	57.00	74.00	120.0		
			Irbesartan	114	113 (99.1)	60.51 (25.40)	23.0	40.00	54.00	76.00	120.0		
		Week 36	Sparsentan	112	105 (93.8)	58.38 (23.20)	19.0	39.00	56.00	76.00	123.0		
			Irbesartan	114	106 (93.0)	57.69 (26.88)	15.0	37.00	53.50	70.00	125.0		
		Week 58	Sparsentan	112	106 (94.6)	57.37 (24.82)	14.0	39.00	53.00	74.00	127.0		
			Irbesartan	114	96 (84.2)	57.50 (26.49)	13.0	35.50	54.50	76.00	122.0		
		Week 110	Sparsentan	112	96 (85.7)	55.38 (26.44)	13.0	32.50	51.00	69.50	126.0		
			Irbesartan	114	83 (72.8)	52.58 (27.68)	11.0	32.00	46.00	65.00	123.0		
		Change from baseline in eGFR	Week 6	Sparsentan	112	108 (96.4)	-1.37 (8.26)	-43.0	-4.00	-0.50	2.00	20.0	0.01 [-0.26, 0.27]
				Irbesartan	114	113 (99.1)	-1.43 (6.33)	-17.0	-5.00	-2.00	3.00	20.0	
	Week 36		Sparsentan	112	105 (93.8)	-4.15 (9.16)	-39.0	-8.00	-3.00	1.00	26.0	0.09 [-0.18, 0.36]	
			Irbesartan	114	106 (93.0)	-4.96 (8.20)	-25.0	-11.00	-4.00	0.00	18.0		
	Week 58		Sparsentan	112	106 (94.6)	-4.13 (10.26)	-34.0	-8.00	-3.50	1.00	38.0	0.09 [-0.19, 0.37]	
			Irbesartan	114	96 (84.2)	-4.98 (8.51)	-24.0	-10.00	-5.00	0.00	23.0		
	Week 110	Sparsentan	112	96 (85.7)	-5.68 (10.68)	-31.0	-11.50	-5.50	0.00	45.0	0.28 [-0.02, 0.57]		
		Irbesartan	114	83 (72.8)	-8.57 (10.07)	-46.0	-14.00	-9.00	-3.00	15.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Time since renal biopsy												
<= 5 years	eGFR	Baseline	Sparsentan	113	113 (100.0)	59.08 (25.60)	24.0	39.00	52.00	74.00	126.0	
			Irbesartan	127	127 (100.0)	61.36 (24.93)	28.0	41.00	56.00	73.00	123.0	
	Week 6	Sparsentan	113	107 (94.7)	58.21 (25.55)	21.0	37.00	53.00	75.00	121.0		
		Irbesartan	127	124 (97.6)	59.52 (25.06)	21.0	40.00	54.50	74.00	120.0		
	Week 36	Sparsentan	113	106 (93.8)	56.29 (24.64)	18.0	36.00	50.00	72.00	123.0		
		Irbesartan	127	116 (91.3)	58.09 (26.68)	7.0	38.00	50.00	70.50	125.0		
	Week 58	Sparsentan	113	105 (92.9)	56.21 (25.76)	13.0	38.00	53.00	72.00	127.0		
		Irbesartan	127	113 (89.0)	55.10 (25.08)	11.0	36.00	50.00	71.00	122.0		
	Week 110	Sparsentan	113	96 (85.0)	54.02 (26.70)	13.0	32.50	50.00	69.00	126.0		
		Irbesartan	127	95 (74.8)	50.85 (25.83)	11.0	32.00	44.00	64.00	123.0		
	Change from baseline in eGFR	Week 6	Sparsentan	113	107 (94.7)	-1.30 (8.36)	-43.0	-4.00	-1.00	3.00	20.0	0.04 [-0.22, 0.30]
			Irbesartan	127	124 (97.6)	-1.61 (7.22)	-31.0	-5.00	-2.00	3.00	20.0	
		Week 36	Sparsentan	113	106 (93.8)	-3.72 (9.04)	-39.0	-8.00	-2.00	0.00	26.0	0.04 [-0.23, 0.30]
			Irbesartan	127	116 (91.3)	-4.07 (9.51)	-32.0	-9.50	-4.00	1.00	33.0	
Week 58		Sparsentan	113	105 (92.9)	-3.85 (9.76)	-34.0	-8.00	-4.00	0.00	38.0	0.17 [-0.09, 0.44]	
		Irbesartan	127	113 (89.0)	-5.58 (10.19)	-32.0	-12.00	-7.00	1.00	23.0		
Week 110		Sparsentan	113	96 (85.0)	-5.44 (11.62)	-34.0	-12.00	-6.50	0.00	45.0	0.33 [0.04, 0.61]	
		Irbesartan	127	95 (74.8)	-9.35 (12.19)	-59.0	-16.00	-9.00	-1.00	15.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 5 years	eGFR	Baseline	Sparsentan	89	89 (100.0)	53.87 (22.42)	28.0	36.00	44.00	66.00	127.0			
			Irbesartan	75	75 (100.0)	49.80 (19.15)	26.0	36.00	43.00	64.00	108.0			
		Week 6	Sparsentan	89	87 (97.8)	52.79 (20.53)	26.0	36.00	46.00	64.00	114.0			
			Irbesartan	75	73 (97.3)	48.55 (18.13)	19.0	35.00	43.00	59.00	95.0			
		Week 36	Sparsentan	89	85 (95.5)	50.76 (19.51)	25.0	34.00	47.00	64.00	114.0			
			Irbesartan	75	70 (93.3)	44.97 (17.95)	23.0	32.00	38.00	57.00	93.0			
		Week 58	Sparsentan	89	83 (93.3)	49.93 (21.76)	21.0	33.00	43.00	61.00	132.0			
			Irbesartan	75	60 (80.0)	44.25 (19.77)	18.0	29.00	38.50	55.00	97.0			
		Week 110	Sparsentan	89	75 (84.3)	48.20 (22.09)	14.0	31.00	41.00	62.00	113.0			
			Irbesartan	75	60 (80.0)	38.78 (20.41)	11.0	24.50	34.00	46.50	106.0			
			Change from baseline in eGFR	Week 6	Sparsentan	89	87 (97.8)	-0.92 (6.77)	-33.0	-4.00	0.00	2.00	14.0	0.12 [-0.20, 0.43]
					Irbesartan	75	73 (97.3)	-1.63 (5.38)	-16.0	-5.00	-2.00	2.00	13.0	
		Week 36		Sparsentan	89	85 (95.5)	-3.27 (7.32)	-34.0	-6.00	-3.00	1.00	10.0	0.21 [-0.11, 0.53]	
				Irbesartan	75	70 (93.3)	-4.66 (5.62)	-20.0	-8.00	-4.00	-1.00	10.0		
Week 58	Sparsentan	89		83 (93.3)	-3.64 (8.10)	-29.0	-8.00	-3.00	1.00	24.0	0.29 [-0.04, 0.63]			
	Irbesartan	75		60 (80.0)	-5.72 (5.30)	-18.0	-9.00	-5.50	-2.00	6.0				
Week 110	Sparsentan	89		75 (84.3)	-5.19 (7.89)	-30.0	-10.00	-4.00	0.00	14.0	0.60 [0.25, 0.95]			
	Irbesartan	75		60 (80.0)	-9.77 (7.31)	-28.0	-14.50	-10.00	-5.00	13.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: History of hypertension													
Yes	eGFR	Baseline	Sparsentan	155	155 (100.0)	53.36 (22.75)	24.0	36.00	46.00	66.00	127.0		
			Irbesartan	161	161 (100.0)	54.63 (21.55)	26.0	38.00	50.00	67.00	123.0		
	Week 6	Sparsentan	155	147 (94.8)	52.64 (22.43)	21.0	35.00	48.00	67.00	121.0			
		Irbesartan	161	157 (97.5)	53.21 (21.71)	19.0	37.00	47.00	64.00	120.0			
	Week 36	Sparsentan	155	146 (94.2)	50.86 (21.98)	18.0	34.00	46.00	63.00	118.0			
		Irbesartan	161	148 (91.9)	51.60 (23.87)	7.0	33.00	45.00	65.00	125.0			
	Week 58	Sparsentan	155	147 (94.8)	50.44 (23.26)	13.0	33.00	45.00	61.00	132.0			
		Irbesartan	161	140 (87.0)	49.19 (22.79)	11.0	32.50	44.50	63.50	122.0			
	Week 110	Sparsentan	155	133 (85.8)	48.52 (23.98)	13.0	30.00	44.00	61.00	123.0			
		Irbesartan	161	125 (77.6)	43.63 (22.71)	11.0	26.00	38.00	56.00	123.0			
	Change from baseline in eGFR		Week 6	Sparsentan	155	147 (94.8)	-0.70 (7.11)	-33.0	-4.00	0.00	3.00	20.0	0.10 [-0.12, 0.33]
				Irbesartan	161	157 (97.5)	-1.41 (6.83)	-31.0	-5.00	-1.00	2.00	20.0	
			Week 36	Sparsentan	155	146 (94.2)	-2.79 (7.63)	-34.0	-6.00	-2.00	1.00	26.0	0.07 [-0.16, 0.30]
				Irbesartan	161	148 (91.9)	-3.33 (8.04)	-32.0	-7.00	-4.00	1.00	33.0	
Week 58			Sparsentan	155	147 (94.8)	-3.40 (8.09)	-24.0	-8.00	-3.00	1.00	38.0	0.21 [-0.03, 0.44]	
			Irbesartan	161	140 (87.0)	-5.14 (8.71)	-30.0	-10.50	-5.00	0.00	23.0		
Week 110			Sparsentan	155	133 (85.8)	-5.07 (9.41)	-34.0	-11.00	-5.00	0.00	45.0	0.46 [0.22, 0.71]	
			Irbesartan	161	125 (77.6)	-9.65 (10.34)	-59.0	-15.00	-9.00	-3.00	13.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSHM: Course of eGFR by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
No	eGFR	Baseline	Sparsentan	47	47 (100.0)	68.06 (26.16)	29.0	45.00	69.00	89.00	126.0			
			Irbesartan	41	41 (100.0)	66.66 (28.59)	31.0	41.00	63.00	91.00	123.0			
		Week 6	Sparsentan	47	47 (100.0)	65.60 (24.40)	26.0	46.00	62.00	86.00	120.0			
			Irbesartan	41	40 (97.6)	64.25 (27.27)	23.0	42.00	55.00	84.00	119.0			
		Week 36	Sparsentan	47	45 (95.7)	63.49 (22.17)	27.0	46.00	64.00	77.00	123.0			
			Irbesartan	41	38 (92.7)	59.18 (26.58)	17.0	37.00	54.50	77.00	123.0			
		Week 58	Sparsentan	47	41 (87.2)	64.20 (24.82)	26.0	42.00	63.00	82.00	127.0			
			Irbesartan	41	33 (80.5)	60.42 (26.56)	13.0	35.00	52.00	82.00	114.0			
		Week 110	Sparsentan	47	38 (80.9)	61.79 (25.54)	26.0	40.00	59.50	81.00	126.0			
			Irbesartan	41	30 (73.2)	56.80 (29.06)	24.0	31.00	47.50	73.00	113.0			
		Change from baseline in eGFR	Change from baseline in eGFR	Week 6	Sparsentan	47	47 (100.0)	-2.47 (9.17)	-43.0	-5.00	-1.00	1.00	12.0	-0.01 [-0.43, 0.42]
					Irbesartan	41	40 (97.6)	-2.43 (5.54)	-16.0	-6.00	-2.50	1.50	7.0	
				Week 36	Sparsentan	47	45 (95.7)	-5.89 (9.90)	-39.0	-12.00	-4.00	-1.00	21.0	0.23 [-0.20, 0.67]
					Irbesartan	41	38 (92.7)	-8.03 (8.12)	-30.0	-12.00	-5.50	-3.00	4.0	
Week 58	Sparsentan			47	41 (87.2)	-5.02 (11.86)	-34.0	-10.00	-5.00	1.00	27.0	0.25 [-0.21, 0.71]		
	Irbesartan			41	33 (80.5)	-7.70 (8.97)	-32.0	-10.00	-7.00	-5.00	16.0			
Week 110	Sparsentan			47	38 (80.9)	-6.24 (12.42)	-30.0	-12.00	-7.50	0.00	39.0	0.22 [-0.26, 0.70]		
	Irbesartan			41	30 (73.2)	-8.93 (11.52)	-36.0	-14.00	-11.00	-3.00	15.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. Only key visits up to Week 110 are displayed.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
 Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.818
Male	Week 6	Sparsentan	139	134 (96.4)	-0.97 (0.73)	(-2.39, 0.46)	0.17 (1.02)	(-1.84, 2.17)	0.872
		Irbesartan	143	139 (97.2)	-1.13 (0.72)	(-2.54, 0.28)			
	Week 36	Sparsentan	139	135 (97.1)	-3.41 (0.73)	(-4.84, -1.98)	0.50 (1.03)	(-1.52, 2.52)	0.629
		Irbesartan	143	130 (90.9)	-3.91 (0.73)	(-5.34, -2.48)			
Week 58	Sparsentan	139	127 (91.4)	-3.41 (0.74)	(-4.86, -1.97)	2.53 (1.05)	(0.48, 4.59)	0.016 *	
	Irbesartan	143	124 (86.7)	-5.95 (0.74)	(-7.40, -4.49)				
Week 110	Sparsentan	139	117 (84.2)	-6.43 (0.77)	(-7.93, -4.93)	4.28 (1.10)	(2.13, 6.44)	<0.001 *	
	Irbesartan	143	111 (77.6)	-10.71 (0.79)	(-12.25, -9.17)				
Female	Week 6	Sparsentan	63	60 (95.2)	-1.75 (1.03)	(-3.79, 0.28)	0.94 (1.49)	(-1.98, 3.86)	0.529
		Irbesartan	59	58 (98.3)	-2.69 (1.06)	(-4.78, -0.60)			
	Week 36	Sparsentan	63	56 (88.9)	-4.09 (1.05)	(-6.15, -2.02)	1.37 (1.51)	(-1.59, 4.33)	0.364
		Irbesartan	59	56 (94.9)	-5.46 (1.07)	(-7.56, -3.35)			
	Week 58	Sparsentan	63	61 (96.8)	-4.91 (1.04)	(-6.94, -2.87)	0.57 (1.52)	(-2.42, 3.56)	0.709
		Irbesartan	59	49 (83.1)	-5.48 (1.11)	(-7.66, -3.29)			
	Week 110	Sparsentan	63	54 (85.7)	-4.24 (1.08)	(-6.36, -2.12)	3.40 (1.61)	(0.23, 6.56)	0.035 *
		Irbesartan	59	44 (74.6)	-7.64 (1.19)	(-9.98, -5.30)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.879
<= 45 years	Week 6	Sparsentan	96	94 (97.9)	-2.40 (0.98)	(-4.32, -0.49)	0.09 (1.37)	(-2.59, 2.78)	0.945
		Irbesartan	99	98 (99.0)	-2.50 (0.96)	(-4.38, -0.61)			
	Week 36	Sparsentan	96	91 (94.8)	-5.25 (0.98)	(-7.18, -3.32)	-0.13 (1.38)	(-2.85, 2.59)	0.926
		Irbesartan	99	91 (91.9)	-5.12 (0.97)	(-7.03, -3.21)			
Week 58	Sparsentan	96	89 (92.7)	-5.01 (0.99)	(-6.95, -3.06)	1.68 (1.41)	(-1.10, 4.45)	0.235	
	Irbesartan	99	82 (82.8)	-6.69 (1.01)	(-8.66, -4.71)				
Week 110	Sparsentan	96	77 (80.2)	-6.32 (1.05)	(-8.38, -4.26)	4.81 (1.52)	(1.82, 7.79)	0.002 *	
	Irbesartan	99	69 (69.7)	-11.12 (1.10)	(-13.29, -8.96)				
> 45 years	Week 6	Sparsentan	106	100 (94.3)	0.01 (0.71)	(-1.37, 1.40)	0.82 (1.01)	(-1.15, 2.79)	0.416
		Irbesartan	103	99 (96.1)	-0.81 (0.72)	(-2.21, 0.60)			
	Week 36	Sparsentan	106	100 (94.3)	-2.01 (0.71)	(-3.39, -0.62)	1.74 (1.01)	(-0.25, 3.73)	0.086
		Irbesartan	103	95 (92.2)	-3.74 (0.72)	(-5.17, -2.32)			
Week 58	Sparsentan	106	99 (93.4)	-2.77 (0.71)	(-4.17, -1.38)	2.29 (1.02)	(0.28, 4.30)	0.025 *	
	Irbesartan	103	91 (88.3)	-5.06 (0.74)	(-6.51, -3.62)				
Week 110	Sparsentan	106	94 (88.7)	-5.07 (0.73)	(-6.50, -3.65)	3.67 (1.05)	(1.61, 5.74)	<0.001 *	
	Irbesartan	103	86 (83.5)	-8.74 (0.76)	(-10.23, -7.25)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 110 are included in the model. Only key visits up to Week 110 are displayed. A first order regressive covariance structure was used.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.100
<= 18 years	Week 6	Sparsentan	9	9 (100.0)	-7.77 (3.72)	(-15.18, -0.35)	-6.56 (6.34)	(-19.22, 6.10)	0.305
		Irbesartan	5	5 (100.0)	-1.21 (5.06)	(-11.31, 8.89)			
	Week 36	Sparsentan	9	8 (88.9)	-9.58 (3.90)	(-17.34, -1.82)	-4.97 (6.46)	(-17.84, 7.91)	0.444
		Irbesartan	5	5 (100.0)	-4.61 (5.06)	(-14.71, 5.49)			
Week 58	Sparsentan	9	7 (77.8)	-11.83 (4.17)	(-20.13, -3.53)	-4.40 (6.94)	(-18.21, 9.42)	0.528	
	Irbesartan	5	4 (80.0)	-7.43 (5.44)	(-18.24, 3.38)				
Week 110	Sparsentan	9	5 (55.6)	-11.92 (4.88)	(-21.62, -2.22)	4.29 (7.50)	(-10.64, 19.23)	0.569	
	Irbesartan	5	4 (80.0)	-16.21 (5.53)	(-27.22, -5.21)				
> 18 to 40 years	Week 6	Sparsentan	102	99 (97.1)	-1.44 (0.87)	(-3.15, 0.28)	0.73 (1.22)	(-1.66, 3.12)	0.547
		Irbesartan	109	106 (97.2)	-2.17 (0.85)	(-3.83, -0.51)			
	Week 36	Sparsentan	102	97 (95.1)	-4.59 (0.88)	(-6.32, -2.87)	0.29 (1.23)	(-2.12, 2.70)	0.814
		Irbesartan	109	101 (92.7)	-4.88 (0.86)	(-6.56, -3.20)			
Week 58	Sparsentan	102	95 (93.1)	-4.14 (0.88)	(-5.88, -2.41)	1.82 (1.25)	(-0.63, 4.28)	0.146	
	Irbesartan	109	90 (82.6)	-5.97 (0.89)	(-7.71, -4.23)				
Week 110	Sparsentan	102	83 (81.4)	-5.81 (0.93)	(-7.64, -3.97)	4.97 (1.33)	(2.35, 7.59)	<0.001 *	
	Irbesartan	109	80 (73.4)	-10.78 (0.95)	(-12.65, -8.91)				

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 110 are included in the model. Only key visits up to Week 110 are displayed. A first order regressive covariance structure was used.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
 Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 40 years	Week 6	Sparsentan	91	86 (94.5)	-0.29 (0.79)	(-1.85, 1.27)	0.53 (1.13)	(-1.69, 2.75)	0.639
		Irbesartan	88	86 (97.7)	-0.82 (0.80)	(-2.40, 0.75)			
	Week 36	Sparsentan	91	86 (94.5)	-1.88 (0.80)	(-3.44, -0.31)	1.77 (1.14)	(-0.47, 4.02)	0.120
		Irbesartan	88	80 (90.9)	-3.65 (0.82)	(-5.26, -2.05)			
	Week 58	Sparsentan	91	86 (94.5)	-2.87 (0.80)	(-4.43, -1.30)	2.51 (1.15)	(0.26, 4.76)	0.029 *
		Irbesartan	88	79 (89.8)	-5.38 (0.83)	(-7.00, -3.75)			
	Week 110	Sparsentan	91	83 (91.2)	-4.93 (0.81)	(-6.52, -3.35)	3.36 (1.18)	(1.03, 5.68)	0.005 *
		Irbesartan	88	71 (80.7)	-8.29 (0.87)	(-9.99, -6.59)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
 Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.631
North America	Week 6	Sparsentan	35	35 (100.0)	-2.55 (1.39)	(-5.29, 0.19)	-0.82 (1.86)	(-4.47, 2.83)	0.660
		Irbesartan	46	45 (97.8)	-1.73 (1.22)	(-4.14, 0.67)			
	Week 36	Sparsentan	35	34 (97.1)	-3.71 (1.41)	(-6.47, -0.94)	0.37 (1.87)	(-3.31, 4.05)	0.844
		Irbesartan	46	44 (95.7)	-4.07 (1.23)	(-6.50, -1.65)			
	Week 58	Sparsentan	35	31 (88.6)	-4.65 (1.44)	(-7.49, -1.82)	-0.58 (1.93)	(-4.38, 3.21)	0.763
		Irbesartan	46	39 (84.8)	-4.07 (1.28)	(-6.58, -1.55)			
Week 110	Sparsentan	35	26 (74.3)	-5.24 (1.55)	(-8.29, -2.19)	4.05 (2.10)	(-0.07, 8.18)	0.054	
	Irbesartan	46	32 (69.6)	-9.29 (1.41)	(-12.07, -6.52)				
Europe	Week 6	Sparsentan	98	92 (93.9)	-0.59 (0.82)	(-2.20, 1.01)	1.00 (1.11)	(-1.18, 3.18)	0.370
		Irbesartan	115	112 (97.4)	-1.59 (0.75)	(-3.07, -0.12)			
	Week 36	Sparsentan	98	91 (92.9)	-3.06 (0.82)	(-4.67, -1.44)	0.99 (1.12)	(-1.21, 3.19)	0.376
		Irbesartan	115	106 (92.2)	-4.05 (0.76)	(-5.54, -2.55)			
	Week 58	Sparsentan	98	93 (94.9)	-3.30 (0.82)	(-4.91, -1.69)	2.73 (1.13)	(0.50, 4.95)	0.016 *
		Irbesartan	115	98 (85.2)	-6.03 (0.78)	(-7.56, -4.50)			
Week 110	Sparsentan	98	83 (84.7)	-6.30 (0.86)	(-7.98, -4.62)	3.42 (1.18)	(1.10, 5.74)	0.004 *	
	Irbesartan	115	92 (80.0)	-9.72 (0.81)	(-11.32, -8.12)				

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Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
 Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Asia Pacific	Week 6	Sparsentan	69	67 (97.1)	-1.22 (1.11)	(-3.40, 0.95)	0.25 (1.81)	(-3.31, 3.81)	0.891
		Irbesartan	41	40 (97.6)	-1.47 (1.43)	(-4.29, 1.34)			
	Week 36	Sparsentan	69	66 (95.7)	-4.21 (1.11)	(-6.40, -2.03)	1.44 (1.84)	(-2.18, 5.06)	0.436
		Irbesartan	41	36 (87.8)	-5.65 (1.47)	(-8.53, -2.77)			
	Week 58	Sparsentan	69	64 (92.8)	-4.23 (1.12)	(-6.43, -2.03)	3.14 (1.86)	(-0.51, 6.80)	0.092
		Irbesartan	41	36 (87.8)	-7.37 (1.48)	(-10.29, -4.46)			
	Week 110	Sparsentan	69	62 (89.9)	-5.00 (1.14)	(-7.24, -2.76)	5.86 (1.96)	(2.02, 9.71)	0.003 *
		Irbesartan	41	31 (75.6)	-10.86 (1.59)	(-13.98, -7.74)			

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Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.187
< 27 kg/m**2	Week 6	Sparsentan	83	78 (94.0)	-1.29 (0.97)	(-3.19, 0.61)	1.04 (1.33)	(-1.56, 3.64)	0.432
		Irbesartan	94	91 (96.8)	-2.33 (0.90)	(-4.10, -0.56)			
	Week 36	Sparsentan	83	77 (92.8)	-3.61 (0.97)	(-5.52, -1.71)	2.03 (1.34)	(-0.59, 4.65)	0.129
		Irbesartan	94	86 (91.5)	-5.64 (0.92)	(-7.44, -3.85)			
	Week 58	Sparsentan	83	77 (92.8)	-3.51 (0.97)	(-5.42, -1.60)	3.49 (1.37)	(0.81, 6.17)	0.011 *
		Irbesartan	94	75 (79.8)	-7.00 (0.95)	(-8.87, -5.13)			
Week 110	Sparsentan	83	70 (84.3)	-4.96 (1.01)	(-6.95, -2.97)	4.72 (1.44)	(1.91, 7.54)	0.001 *	
	Irbesartan	94	69 (73.4)	-9.68 (1.01)	(-11.67, -7.69)				
≥ 27 kg/m**2	Week 6	Sparsentan	119	116 (97.5)	-1.03 (0.75)	(-2.51, 0.45)	0.02 (1.10)	(-2.13, 2.17)	0.984
		Irbesartan	107	105 (98.1)	-1.05 (0.79)	(-2.61, 0.51)			
	Week 36	Sparsentan	119	114 (95.8)	-3.48 (0.76)	(-4.97, -1.99)	-0.09 (1.11)	(-2.26, 2.08)	0.934
		Irbesartan	107	100 (93.5)	-3.39 (0.80)	(-4.97, -1.81)			
	Week 58	Sparsentan	119	111 (93.3)	-4.03 (0.76)	(-5.53, -2.53)	0.91 (1.12)	(-1.28, 3.10)	0.414
		Irbesartan	107	98 (91.6)	-4.95 (0.81)	(-6.54, -3.35)			
	Week 110	Sparsentan	119	101 (84.9)	-6.16 (0.79)	(-7.72, -4.61)	3.92 (1.17)	(1.62, 6.22)	<0.001 *
		Irbesartan	107	85 (79.4)	-10.08 (0.86)	(-11.77, -8.39)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 110 are included in the model. Only key visits up to Week 110 are displayed. A first order regressive covariance structure was used.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan						Interaction:	0.013 #
eGFR Low and UP High	Week 6	Sparsentan	71	69 (97.2)	-0.71 (0.83)	(-2.34, 0.92)	1.19 (1.16)	(-1.09, 3.47)	0.307
		Irbesartan	74	72 (97.3)	-1.89 (0.81)	(-3.49, -0.30)			
	Week 36	Sparsentan	71	68 (95.8)	-2.68 (0.83)	(-4.31, -1.04)	3.43 (1.17)	(1.13, 5.73)	0.004 *
		Irbesartan	74	66 (89.2)	-6.11 (0.82)	(-7.73, -4.49)			
	Week 58	Sparsentan	71	67 (94.4)	-4.43 (0.84)	(-6.08, -2.79)	3.38 (1.19)	(1.05, 5.71)	0.005 *
		Irbesartan	74	62 (83.8)	-7.81 (0.84)	(-9.47, -6.16)			
	Week 110	Sparsentan	71	59 (83.1)	-7.49 (0.88)	(-9.22, -5.77)	4.60 (1.26)	(2.12, 7.07)	<0.001 *
		Irbesartan	74	57 (77.0)	-12.09 (0.90)	(-13.86, -10.32)			
eGFR Low and UP Low	Week 6	Sparsentan	55	51 (92.7)	0.26 (0.99)	(-1.69, 2.20)	1.30 (1.40)	(-1.45, 4.05)	0.353
		Irbesartan	55	54 (98.2)	-1.04 (0.98)	(-2.98, 0.89)			
	Week 36	Sparsentan	55	49 (89.1)	0.01 (1.00)	(-1.95, 1.98)	2.62 (1.41)	(-0.15, 5.39)	0.064
		Irbesartan	55	52 (94.5)	-2.61 (0.99)	(-4.55, -0.66)			
	Week 58	Sparsentan	55	51 (92.7)	-0.50 (1.00)	(-2.46, 1.46)	2.87 (1.42)	(0.09, 5.65)	0.043 *
		Irbesartan	55	49 (89.1)	-3.37 (1.00)	(-5.34, -1.40)			
	Week 110	Sparsentan	55	47 (85.5)	-2.77 (1.02)	(-4.78, -0.75)	4.86 (1.47)	(1.98, 7.75)	0.001 *
		Irbesartan	55	44 (80.0)	-7.63 (1.05)	(-9.70, -5.56)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
eGFR High and UP High	Week 6	Sparsentan	37	35 (94.6)	-3.60 (1.84)	(-7.23, 0.02)	-0.52 (2.63)	(-5.69, 4.65)	0.844	
		Irbesartan	36	34 (94.4)	-3.08 (1.87)	(-6.77, 0.60)				
	Week 36	Sparsentan	37	37 (100.0)	-7.95 (1.82)	(-11.54, -4.37)	-1.54 (2.64)	(-6.74, 3.65)	0.560	
		Irbesartan	36	33 (91.7)	-6.41 (1.91)	(-10.17, -2.65)				
	Week 58	Sparsentan	37	34 (91.9)	-4.97 (1.86)	(-8.63, -1.30)	2.12 (2.72)	(-3.24, 7.47)	0.437	
		Irbesartan	36	30 (83.3)	-7.09 (1.98)	(-10.99, -3.19)				
	Week 110	Sparsentan	37	30 (81.1)	-6.70 (1.97)	(-10.58, -2.83)	7.80 (2.87)	(2.16, 13.44)	0.007 *	
		Irbesartan	36	27 (75.0)	-14.50 (2.09)	(-18.60, -10.40)				
eGFR High and UP Low	Week 6	Sparsentan	39	39 (100.0)	-1.74 (1.48)	(-4.65, 1.16)	-1.19 (2.12)	(-5.35, 2.97)	0.575	
		Irbesartan	37	37 (100.0)	-0.56 (1.52)	(-3.54, 2.43)				
	Week 36	Sparsentan	39	37 (94.9)	-5.85 (1.49)	(-8.78, -2.91)	-4.09 (2.14)	(-8.30, 0.13)	0.057	
		Irbesartan	37	35 (94.6)	-1.76 (1.54)	(-4.78, 1.26)				
	Week 58	Sparsentan	39	36 (92.3)	-6.50 (1.50)	(-9.45, -3.55)	-1.85 (2.19)	(-6.14, 2.45)	0.398	
		Irbesartan	37	32 (86.5)	-4.65 (1.59)	(-7.77, -1.53)				
	Week 110	Sparsentan	39	35 (89.7)	-5.73 (1.53)	(-8.73, -2.72)	-1.01 (2.28)	(-5.49, 3.47)	0.658	
		Irbesartan	37	27 (73.0)	-4.72 (1.69)	(-8.04, -1.39)				

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LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
 Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.056
< 60 mL/min/1.73 m**2	Week 6	Sparsentan	127	121 (95.3)	0.17 (0.63)	(-1.07, 1.41)	0.92 (0.89)	(-0.82, 2.66)	0.298
		Irbesartan	129	127 (98.4)	-0.75 (0.62)	(-1.97, 0.47)			
	Week 36	Sparsentan	127	118 (92.9)	-1.23 (0.63)	(-2.47, 0.01)	2.54 (0.89)	(0.78, 4.29)	0.005 *
		Irbesartan	129	118 (91.5)	-3.77 (0.63)	(-5.01, -2.53)			
	Week 58	Sparsentan	127	118 (92.9)	-2.45 (0.64)	(-3.70, -1.20)	2.54 (0.90)	(0.76, 4.31)	0.005 *
		Irbesartan	129	111 (86.0)	-4.99 (0.64)	(-6.25, -3.73)			
Week 110	Sparsentan	127	107 (84.3)	-4.97 (0.66)	(-6.27, -3.68)	4.14 (0.95)	(2.29, 6.00)	<0.001 *	
	Irbesartan	129	102 (79.1)	-9.12 (0.68)	(-10.45, -7.79)				
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	47 (95.9)	-2.86 (1.43)	(-5.67, -0.05)	-0.01 (2.04)	(-4.02, 4.01)	0.997
		Irbesartan	48	45 (93.8)	-2.85 (1.45)	(-5.71, 0.00)			
	Week 36	Sparsentan	49	47 (95.9)	-6.39 (1.43)	(-9.19, -3.58)	-0.17 (2.06)	(-4.21, 3.88)	0.935
		Irbesartan	48	44 (91.7)	-6.22 (1.48)	(-9.12, -3.32)			
	Week 58	Sparsentan	49	44 (89.8)	-5.39 (1.45)	(-8.24, -2.53)	2.98 (2.10)	(-1.15, 7.10)	0.156
		Irbesartan	48	41 (85.4)	-8.37 (1.51)	(-11.33, -5.40)			
	Week 110	Sparsentan	49	43 (87.8)	-8.29 (1.49)	(-11.22, -5.37)	5.49 (2.17)	(1.21, 9.76)	0.012 *
		Irbesartan	48	37 (77.1)	-13.78 (1.58)	(-16.89, -10.67)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
 Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	26 (100.0)	-4.21 (2.08)	(-8.31, -0.12)	-0.11 (2.97)	(-5.96, 5.73)	0.969
		Irbesartan	25	25 (100.0)	-4.10 (2.12)	(-8.27, 0.08)			
	Week 36	Sparsentan	26	26 (100.0)	-9.25 (2.08)	(-13.35, -5.16)	-4.61 (2.99)	(-10.50, 1.27)	0.124
		Irbesartan	25	24 (96.0)	-4.64 (2.15)	(-8.87, -0.41)			
	Week 58	Sparsentan	26	26 (100.0)	-7.67 (2.08)	(-11.77, -3.58)	-1.69 (3.08)	(-7.74, 4.37)	0.584
		Irbesartan	25	21 (84.0)	-5.99 (2.26)	(-10.45, -1.53)			
	Week 110	Sparsentan	26	21 (80.8)	-4.12 (2.24)	(-8.52, 0.29)	2.47 (3.37)	(-4.17, 9.10)	0.465
		Irbesartan	25	16 (64.0)	-6.58 (2.52)	(-11.55, -1.62)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.098
< 45 mL/min/1.73 m**2	Week 6	Sparsentan	82	79 (96.3)	0.79 (0.69)	(-0.57, 2.14)	1.66 (0.98)	(-0.27, 3.59)	0.092
		Irbesartan	80	78 (97.5)	-0.87 (0.70)	(-2.24, 0.50)			
	Week 36	Sparsentan	82	76 (92.7)	-1.27 (0.69)	(-2.64, 0.09)	3.56 (0.99)	(1.61, 5.50)	<0.001 *
		Irbesartan	80	71 (88.8)	-4.83 (0.71)	(-6.22, -3.44)			
	Week 58	Sparsentan	82	76 (92.7)	-3.03 (0.70)	(-4.40, -1.66)	3.43 (1.00)	(1.46, 5.40)	<0.001 *
		Irbesartan	80	66 (82.5)	-6.46 (0.72)	(-7.87, -5.05)			
	Week 110	Sparsentan	82	68 (82.9)	-5.44 (0.72)	(-6.86, -4.02)	4.26 (1.05)	(2.20, 6.32)	<0.001 *
		Irbesartan	80	63 (78.8)	-9.70 (0.76)	(-11.19, -8.21)			
45 to < 60 mL/min/1.73 m**2	Week 6	Sparsentan	45	42 (93.3)	-1.00 (1.26)	(-3.48, 1.48)	-0.46 (1.74)	(-3.88, 2.95)	0.790
		Irbesartan	49	49 (100.0)	-0.54 (1.19)	(-2.88, 1.81)			
	Week 36	Sparsentan	45	42 (93.3)	-1.20 (1.26)	(-3.68, 1.28)	0.89 (1.75)	(-2.54, 4.32)	0.612
		Irbesartan	49	47 (95.9)	-2.08 (1.20)	(-4.44, 0.28)			
	Week 58	Sparsentan	45	42 (93.3)	-1.47 (1.27)	(-3.96, 1.02)	1.31 (1.76)	(-2.15, 4.78)	0.456
		Irbesartan	49	45 (91.8)	-2.79 (1.22)	(-5.19, -0.39)			
	Week 110	Sparsentan	45	39 (86.7)	-4.29 (1.31)	(-6.85, -1.73)	4.00 (1.84)	(0.38, 7.62)	0.031 *
		Irbesartan	49	39 (79.6)	-8.29 (1.30)	(-10.84, -5.74)			

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Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
60 to < 90 mL/min/1.73 m**2	Week 6	Sparsentan	49	47 (95.9)	-2.86 (1.43)	(-5.67, -0.05)	-0.01 (2.04)	(-4.02, 4.01)	0.997	
		Irbesartan	48	45 (93.8)	-2.85 (1.45)	(-5.71, 0.00)				
	Week 36	Sparsentan	49	47 (95.9)	-6.39 (1.43)	(-9.19, -3.58)	-0.17 (2.06)	(-4.21, 3.88)	0.935	
		Irbesartan	48	44 (91.7)	-6.22 (1.48)	(-9.12, -3.32)				
	Week 58	Sparsentan	49	44 (89.8)	-5.39 (1.45)	(-8.24, -2.53)	2.98 (2.10)	(-1.15, 7.10)	0.156	
		Irbesartan	48	41 (85.4)	-8.37 (1.51)	(-11.33, -5.40)				
	Week 110	Sparsentan	49	43 (87.8)	-8.29 (1.49)	(-11.22, -5.37)	5.49 (2.17)	(1.21, 9.76)	0.012 *	
		Irbesartan	48	37 (77.1)	-13.78 (1.58)	(-16.89, -10.67)				
>= 90 mL/min/1.73 m**2	Week 6	Sparsentan	26	26 (100.0)	-4.21 (2.08)	(-8.31, -0.12)	-0.11 (2.97)	(-5.96, 5.73)	0.969	
		Irbesartan	25	25 (100.0)	-4.10 (2.12)	(-8.27, 0.08)				
	Week 36	Sparsentan	26	26 (100.0)	-9.25 (2.08)	(-13.35, -5.16)	-4.61 (2.99)	(-10.50, 1.27)	0.124	
		Irbesartan	25	24 (96.0)	-4.64 (2.15)	(-8.87, -0.41)				
	Week 58	Sparsentan	26	26 (100.0)	-7.67 (2.08)	(-11.77, -3.58)	-1.69 (3.08)	(-7.74, 4.37)	0.584	
		Irbesartan	25	21 (84.0)	-5.99 (2.26)	(-10.45, -1.53)				
	Week 110	Sparsentan	26	21 (80.8)	-4.12 (2.24)	(-8.52, 0.29)	2.47 (3.37)	(-4.17, 9.10)	0.465	
		Irbesartan	25	16 (64.0)	-6.58 (2.52)	(-11.55, -1.62)				

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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline urine protein excretion	Overall		Sparsentan							Interaction: 0.801
<= 1.75 g/day	Week 6	Sparsentan	98	94 (95.9)	-0.35 (0.86)	(-2.04, 1.33)	0.52 (1.23)	(-1.89, 2.93)	0.672	
		Irbesartan	93	91 (97.8)	-0.87 (0.88)	(-2.60, 0.85)				
	Week 36	Sparsentan	98	91 (92.9)	-1.77 (0.86)	(-3.47, -0.08)	0.47 (1.24)	(-1.97, 2.90)	0.706	
		Irbesartan	93	86 (92.5)	-2.24 (0.89)	(-3.99, -0.49)				
	Week 58	Sparsentan	98	92 (93.9)	-2.79 (0.86)	(-4.49, -1.10)	0.93 (1.26)	(-1.54, 3.39)	0.461	
		Irbesartan	93	79 (84.9)	-3.72 (0.91)	(-5.51, -1.93)				
	Week 110	Sparsentan	98	87 (88.8)	-3.31 (0.88)	(-5.04, -1.58)	4.17 (1.31)	(1.60, 6.74)	0.001 *	
		Irbesartan	93	71 (76.3)	-7.48 (0.97)	(-9.38, -5.59)				
> 1.75 g/day	Week 6	Sparsentan	104	100 (96.2)	-1.93 (0.81)	(-3.52, -0.33)	0.36 (1.14)	(-1.88, 2.59)	0.755	
		Irbesartan	109	106 (97.2)	-2.28 (0.79)	(-3.84, -0.73)				
	Week 36	Sparsentan	104	100 (96.2)	-5.24 (0.82)	(-6.84, -3.64)	1.02 (1.15)	(-1.23, 3.27)	0.372	
		Irbesartan	109	100 (91.7)	-6.26 (0.80)	(-7.84, -4.68)				
	Week 58	Sparsentan	104	96 (92.3)	-4.84 (0.83)	(-6.46, -3.22)	2.85 (1.17)	(0.56, 5.14)	0.015 *	
		Irbesartan	109	94 (86.2)	-7.69 (0.82)	(-9.30, -6.07)				
	Week 110	Sparsentan	104	84 (80.8)	-8.06 (0.87)	(-9.77, -6.35)	3.90 (1.24)	(1.48, 6.33)	0.002 *	
		Irbesartan	109	84 (77.1)	-11.96 (0.88)	(-13.68, -10.24)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values up to Week 110 are included in the model. Only key visits up to Week 110 are displayed. A first order regressive covariance structure was used.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline use of antihypertensives		Overall	Sparsentan							Interaction: 0.201
Yes	Week 6	Sparsentan	90	86 (95.6)	-0.71 (0.83)	(-2.35, 0.92)	1.37 (1.19)	(-0.96, 3.70)	0.249	
		Irbesartan	88	84 (95.5)	-2.08 (0.85)	(-3.74, -0.42)				
	Week 36	Sparsentan	90	86 (95.6)	-2.75 (0.83)	(-4.38, -1.11)	0.76 (1.20)	(-1.58, 3.11)	0.524	
		Irbesartan	88	80 (90.9)	-3.51 (0.86)	(-5.19, -1.83)				
Week 58	Sparsentan	90	82 (91.1)	-3.33 (0.84)	(-4.98, -1.67)	3.25 (1.21)	(0.87, 5.63)	0.007 *		
	Irbesartan	88	77 (87.5)	-6.58 (0.87)	(-8.29, -4.87)					
Week 110	Sparsentan	90	75 (83.3)	-4.96 (0.88)	(-6.69, -3.24)	5.93 (1.26)	(3.46, 8.41)	<0.001 *		
	Irbesartan	88	72 (81.8)	-10.90 (0.91)	(-12.67, -9.12)					
No	Week 6	Sparsentan	112	108 (96.4)	-1.57 (0.84)	(-3.21, 0.07)	-0.30 (1.18)	(-2.61, 2.00)	0.795	
		Irbesartan	114	113 (99.1)	-1.26 (0.82)	(-2.88, 0.35)				
	Week 36	Sparsentan	112	105 (93.8)	-4.26 (0.84)	(-5.91, -2.61)	0.81 (1.19)	(-1.52, 3.14)	0.495	
		Irbesartan	114	106 (93.0)	-5.07 (0.84)	(-6.71, -3.43)				
	Week 58	Sparsentan	112	106 (94.6)	-4.29 (0.84)	(-5.95, -2.64)	0.97 (1.21)	(-1.40, 3.34)	0.422	
		Irbesartan	114	96 (84.2)	-5.26 (0.86)	(-6.95, -3.57)				
	Week 110	Sparsentan	112	96 (85.7)	-6.26 (0.88)	(-7.98, -4.54)	2.72 (1.28)	(0.21, 5.23)	0.034 *	
		Irbesartan	114	83 (72.8)	-8.98 (0.93)	(-10.81, -7.16)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.949
<= 5 years	Week 6	Sparsentan	113	107 (94.7)	-1.45 (0.88)	(-3.17, 0.27)	0.15 (1.20)	(-2.21, 2.51)	0.901
		Irbesartan	127	124 (97.6)	-1.60 (0.82)	(-3.21, 0.01)			
	Week 36	Sparsentan	113	106 (93.8)	-3.75 (0.88)	(-5.47, -2.02)	0.36 (1.21)	(-2.02, 2.74)	0.765
		Irbesartan	127	116 (91.3)	-4.11 (0.83)	(-5.75, -2.47)			
	Week 58	Sparsentan	113	105 (92.9)	-4.02 (0.88)	(-5.75, -2.29)	1.80 (1.22)	(-0.60, 4.20)	0.141
		Irbesartan	127	113 (89.0)	-5.82 (0.85)	(-7.48, -4.16)			
Week 110	Sparsentan	113	96 (85.0)	-5.55 (0.92)	(-7.35, -3.76)	4.25 (1.29)	(1.72, 6.78)	0.001 *	
	Irbesartan	127	95 (74.8)	-9.80 (0.91)	(-11.58, -8.01)				
> 5 years	Week 6	Sparsentan	89	87 (97.8)	-0.79 (0.74)	(-2.25, 0.67)	0.92 (1.10)	(-1.24, 3.08)	0.404
		Irbesartan	75	73 (97.3)	-1.71 (0.81)	(-3.31, -0.11)			
	Week 36	Sparsentan	89	85 (95.5)	-3.30 (0.75)	(-4.77, -1.83)	1.64 (1.11)	(-0.54, 3.82)	0.140
		Irbesartan	75	70 (93.3)	-4.94 (0.82)	(-6.55, -3.33)			
	Week 58	Sparsentan	89	83 (93.3)	-3.60 (0.76)	(-5.08, -2.11)	2.35 (1.15)	(0.09, 4.60)	0.041 *
		Irbesartan	75	60 (80.0)	-5.94 (0.86)	(-7.64, -4.25)			
Week 110	Sparsentan	89	75 (84.3)	-5.77 (0.79)	(-7.32, -4.22)	4.34 (1.19)	(2.00, 6.67)	<0.001 *	
	Irbesartan	75	60 (80.0)	-10.11 (0.89)	(-11.85, -8.36)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGC\_FSCM: Change from baseline in eGFR by subgroup  
Full Analysis Set

Change from baseline in eGFR					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.738
Yes	Week 6	Sparsentan	155	147 (94.8)	-0.79 (0.66)	(-2.09, 0.50)	0.53 (0.92)	(-1.28, 2.34)	0.563
		Irbesartan	161	157 (97.5)	-1.33 (0.65)	(-2.59, -0.06)			
	Week 36	Sparsentan	155	146 (94.2)	-2.80 (0.66)	(-4.10, -1.50)	0.52 (0.93)	(-1.31, 2.34)	0.577
		Irbesartan	161	148 (91.9)	-3.32 (0.65)	(-4.60, -2.04)			
	Week 58	Sparsentan	155	147 (94.8)	-3.50 (0.66)	(-4.80, -2.20)	1.68 (0.94)	(-0.16, 3.53)	0.073
		Irbesartan	161	140 (87.0)	-5.18 (0.67)	(-6.49, -3.87)			
	Week 110	Sparsentan	155	133 (85.8)	-5.43 (0.69)	(-6.78, -4.09)	4.46 (0.98)	(2.53, 6.39)	<0.001 *
		Irbesartan	161	125 (77.6)	-9.89 (0.71)	(-11.28, -8.51)			
No	Week 6	Sparsentan	47	47 (100.0)	-2.43 (1.35)	(-5.09, 0.23)	0.39 (1.99)	(-3.53, 4.32)	0.843
		Irbesartan	41	40 (97.6)	-2.82 (1.45)	(-5.68, 0.03)			
	Week 36	Sparsentan	47	45 (95.7)	-6.14 (1.37)	(-8.83, -3.45)	2.56 (2.02)	(-1.41, 6.53)	0.205
		Irbesartan	41	38 (92.7)	-8.70 (1.47)	(-11.60, -5.81)			
	Week 58	Sparsentan	47	41 (87.2)	-5.07 (1.41)	(-7.83, -2.31)	3.64 (2.08)	(-0.45, 7.73)	0.081
		Irbesartan	41	33 (80.5)	-8.71 (1.52)	(-11.70, -5.71)			
	Week 110	Sparsentan	47	38 (80.9)	-6.62 (1.47)	(-9.51, -3.73)	3.39 (2.21)	(-0.95, 7.73)	0.126
		Irbesartan	41	30 (73.2)	-10.01 (1.64)	(-13.23, -6.79)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures.

LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.

A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

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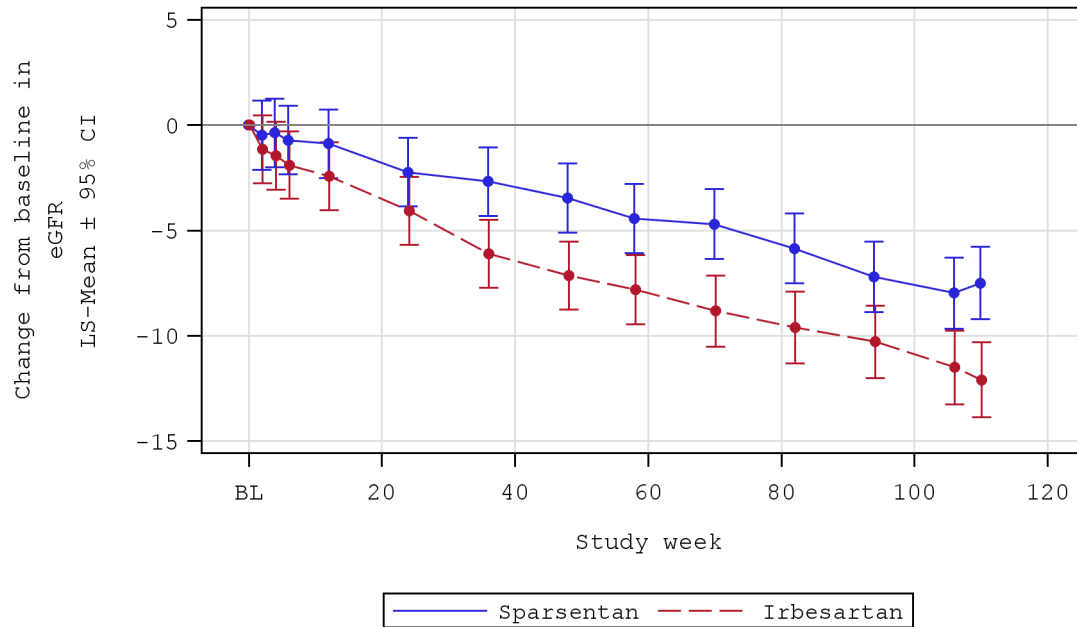
A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m<sup>2</sup>.

eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High >= 60 ml/min/1.73m<sup>2</sup>, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

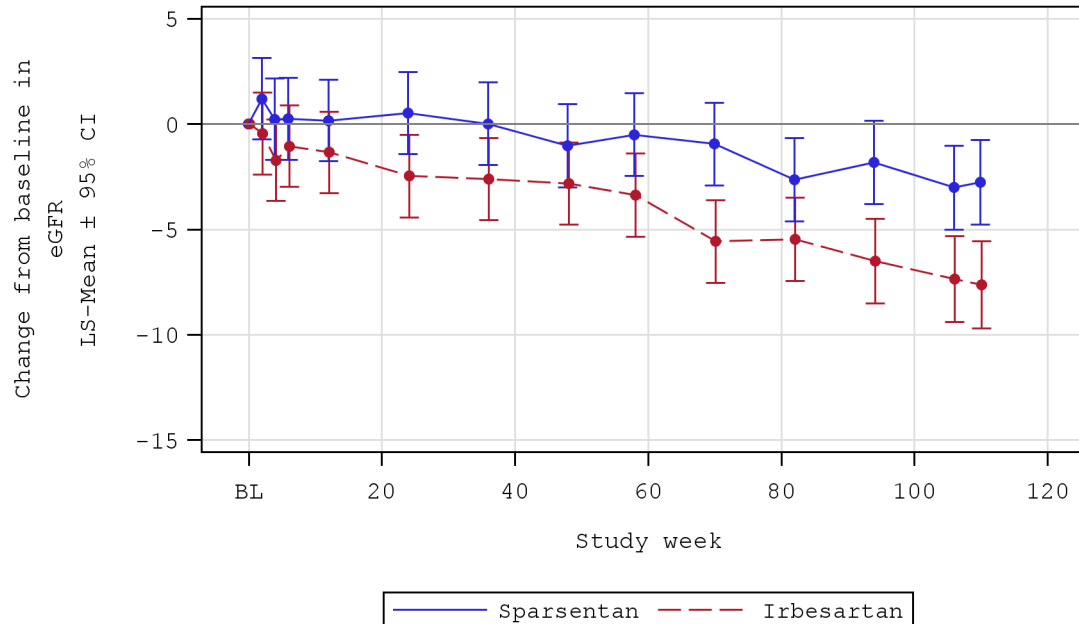
Figure PF2GGC\_FSGM: Change from baseline in eGFR by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR Low and UP High



Sparsentan	69	68	67	59
Irbesartan	72	66	62	57

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m<sup>2</sup>. eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High >= 60 ml/min/1.73m<sup>2</sup>, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. Reference table: PT2GGC\_FSCM.

Figure PF2GGC\_FSGM: Change from baseline in eGFR by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR Low and UP Low

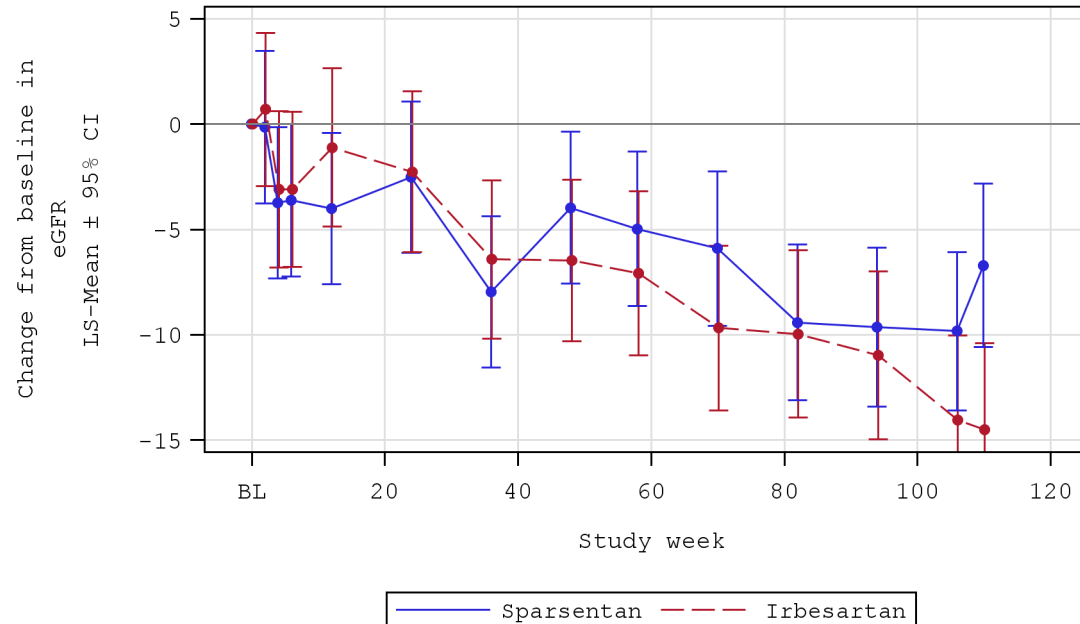


Sparsentan	51	49	51	47
Irbesartan	54	52	49	44

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m<sup>2</sup>. eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High >= 60 ml/min/1.73m<sup>2</sup>, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. Reference table: PT2GGC\_FSCM.



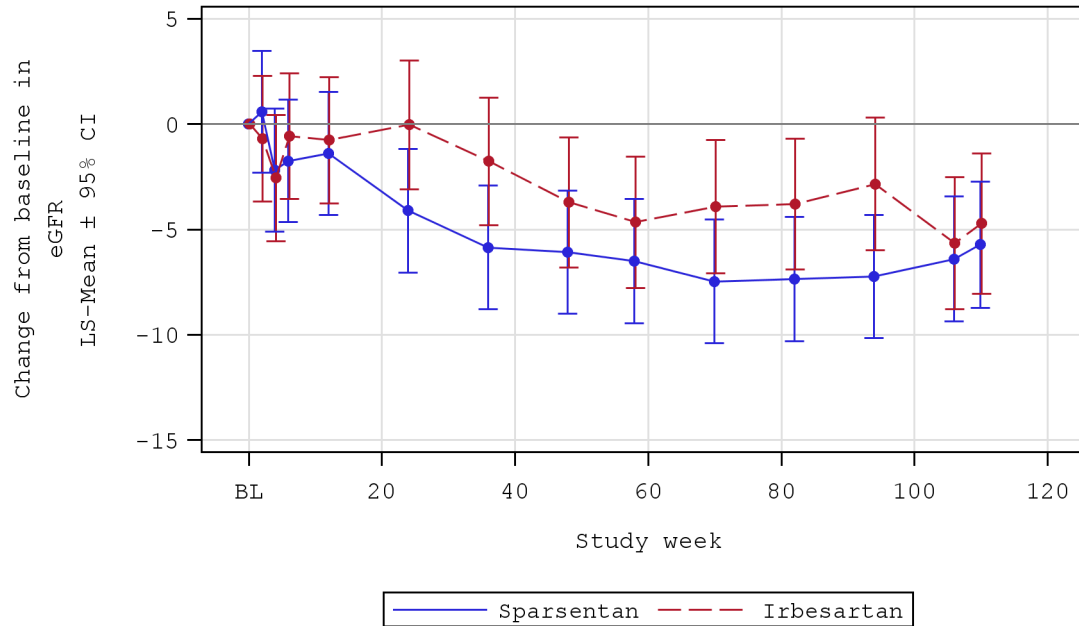
Figure PF2GGC\_FSGM: Change from baseline in eGFR by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR High and UP High



Sparsentan	35	37	34	30
Irbesartan	34	33	30	27

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m<sup>2</sup>. eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High >= 60 ml/min/1.73m<sup>2</sup>, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. Reference table: PT2GGC\_FSCM.

Figure PF2GGC\_FSGM: Change from baseline in eGFR by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Randomization strata: eGFR High and UP Low



Sparsentan	39	37	36	35
Irbesartan	37	35	32	27

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m<sup>2</sup>. eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High >= 60 ml/min/1.73m<sup>2</sup>, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. Reference table: PT2GGC\_FSCM.

Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan					Interaction:	0.015 #
Male	Baseline to Week 58	Sparsentan	139	-3.00 (0.81)	(-4.60, -1.39)	1.69 (1.15)	(-0.58, 3.96)	0.144
		Irbesartan	143	-4.68 (0.82)	(-6.29, -3.08)			
Female	Baseline to Week 58	Sparsentan	63	-4.41 (1.32)	(-7.01, -1.81)	-0.23 (1.90)	(-3.99, 3.53)	0.904
		Irbesartan	59	-4.18 (1.38)	(-6.90, -1.46)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan					Interaction:	0.490
<= 45 years	Baseline to Week 58	Sparsentan	96	-4.18 (1.16)	(-6.46, -1.89)	0.74 (1.64)	(-2.48, 3.97)	0.651
		Irbesartan	99	-4.92 (1.16)	(-7.20, -2.64)			
> 45 years	Baseline to Week 58	Sparsentan	106	-2.77 (0.80)	(-4.34, -1.20)	1.39 (1.15)	(-0.87, 3.64)	0.227
		Irbesartan	103	-4.15 (0.82)	(-5.77, -2.53)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan					Interaction:	<0.001 #
<= 18 years	Baseline to Week 58	Sparsentan	9	-5.49 (3.64)	(-13.03, 2.06)	1.51 (5.92)	(-10.74, 13.76)	0.801
		Irbesartan	5	-7.00 (4.67)	(-16.64, 2.65)			
> 18 to 40 years	Baseline to Week 58	Sparsentan	102	-3.88 (1.04)	(-5.92, -1.84)	0.33 (1.45)	(-2.53, 3.19)	0.821
		Irbesartan	109	-4.21 (1.02)	(-6.22, -2.20)			
> 40 years	Baseline to Week 58	Sparsentan	91	-2.63 (0.91)	(-4.43, -0.82)	2.13 (1.32)	(-0.46, 4.73)	0.106
		Irbesartan	88	-4.76 (0.95)	(-6.62, -2.90)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan					Interaction:	0.121
North America	Baseline to Week 58	Sparsentan	35	-3.14 (1.66)	(-6.43, 0.15)	0.47 (2.21)	(-3.91, 4.85)	0.833
		Irbesartan	46	-3.61 (1.46)	(-6.51, -0.71)			
Europe	Baseline to Week 58	Sparsentan	98	-2.96 (0.93)	(-4.80, -1.12)	1.60 (1.28)	(-0.92, 4.13)	0.213
		Irbesartan	115	-4.56 (0.88)	(-6.29, -2.83)			
Asia Pacific	Baseline to Week 58	Sparsentan	69	-4.24 (1.29)	(-6.78, -1.69)	1.10 (2.13)	(-3.11, 5.32)	0.605
		Irbesartan	41	-5.34 (1.70)	(-8.70, -1.98)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan					Interaction:	0.078
< 27 kg/m**2	Baseline to Week 58	Sparsentan	83	-3.07 (1.19)	(-5.40, -0.73)	2.17 (1.64)	(-1.06, 5.41)	0.187
		Irbesartan	94	-5.24 (1.14)	(-7.49, -3.00)			
>= 27 kg/m**2	Baseline to Week 58	Sparsentan	119	-3.66 (0.83)	(-5.29, -2.04)	0.29 (1.21)	(-2.08, 2.67)	0.808
		Irbesartan	107	-3.96 (0.88)	(-5.69, -2.23)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		p-value
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Randomization strata	Overall	Sparsentan					Interaction:	0.005 #
eGFR Low and UP High	Baseline to Week 58	Sparsentan	71	-3.99 (1.05)	(-6.05, -1.92)	2.82 (1.47)	(-0.09, 5.72)	0.058
		Irbesartan	74	-6.80 (1.04)	(-8.85, -4.76)			
eGFR Low and UP Low	Baseline to Week 58	Sparsentan	55	-1.17 (1.11)	(-3.37, 1.03)	1.09 (1.57)	(-2.02, 4.19)	0.491
		Irbesartan	55	-2.26 (1.11)	(-4.45, -0.06)			
eGFR High and UP High	Baseline to Week 58	Sparsentan	37	-3.02 (2.09)	(-7.18, 1.13)	2.50 (3.05)	(-3.55, 8.56)	0.414
		Irbesartan	36	-5.53 (2.22)	(-9.93, -1.12)			
eGFR High and UP Low	Baseline to Week 58	Sparsentan	39	-6.08 (1.68)	(-9.41, -2.76)	-3.20 (2.43)	(-8.02, 1.61)	0.190
		Irbesartan	37	-2.88 (1.76)	(-6.36, 0.60)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan					Interaction:	0.057
< 60 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	127	-2.84 (0.75)	(-4.31, -1.37)	1.59 (1.06)	(-0.49, 3.67)	0.134
		Irbesartan	129	-4.43 (0.75)	(-5.90, -2.96)			
60 to < 90 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	49	-3.69 (1.56)	(-6.78, -0.61)	2.67 (2.26)	(-1.79, 7.13)	0.239
		Irbesartan	48	-6.36 (1.63)	(-9.58, -3.14)			
>= 90 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	26	-5.88 (2.49)	(-10.85, -0.91)	-4.29 (3.62)	(-11.51, 2.93)	0.240
		Irbesartan	25	-1.59 (2.63)	(-6.82, 3.64)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

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Source Data: alb, created on: 19FEB2024

Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		p-value
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline eGFR Group 2	Overall	Sparsentan					Interaction:	0.043 #
< 45 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	82	-3.66 (0.88)	(-5.39, -1.92)	2.36 (1.26)	(-0.13, 4.84)	0.063
		Irbesartan	80	-6.01 (0.90)	(-7.79, -4.24)			
45 to < 60 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	45	-1.40 (1.40)	(-4.18, 1.37)	0.70 (1.94)	(-3.13, 4.54)	0.717
		Irbesartan	49	-2.11 (1.34)	(-4.76, 0.54)			
60 to < 90 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	49	-3.69 (1.56)	(-6.78, -0.61)	2.67 (2.26)	(-1.79, 7.13)	0.239
		Irbesartan	48	-6.36 (1.63)	(-9.58, -3.14)			
>= 90 mL/min/1.73 m**2	Baseline to Week 58	Sparsentan	26	-5.88 (2.49)	(-10.85, -0.91)	-4.29 (3.62)	(-11.51, 2.93)	0.240
		Irbesartan	25	-1.59 (2.63)	(-6.82, 3.64)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model					
				Annualized Slope			Slope Difference		
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline urine protein excretion	Overall	Sparsentan					Interaction:	0.620	
<= 1.75 g/day	Baseline to Week 58	Sparsentan	98	-2.88 (0.99)	(-4.82, -0.93)	-0.12 (1.43)	(-2.93, 2.70)	0.935	
		Irbesartan	93	-2.76 (1.03)	(-4.79, -0.73)				
> 1.75 g/day	Baseline to Week 58	Sparsentan	104	-3.98 (0.95)	(-5.85, -2.11)	2.07 (1.34)	(-0.56, 4.70)	0.123	
		Irbesartan	109	-6.05 (0.94)	(-7.90, -4.20)				

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan					Interaction:	0.614
Yes	Baseline to Week 58	Sparsentan	90	-2.84 (0.94)	(-4.70, -0.98)	1.80 (1.35)	(-0.86, 4.47)	0.184
		Irbesartan	88	-4.65 (0.97)	(-6.56, -2.73)			
No	Baseline to Week 58	Sparsentan	112	-3.87 (0.99)	(-5.82, -1.93)	0.57 (1.40)	(-2.19, 3.32)	0.687
		Irbesartan	114	-4.44 (0.99)	(-6.39, -2.48)			

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan					Interaction:	0.393
<= 5 years	Baseline to Week 58	Sparsentan	113	-3.28 (1.03)	(-5.31, -1.25)	0.82 (1.42)	(-1.98, 3.62)	0.566
		Irbesartan	127	-4.10 (0.98)	(-6.03, -2.17)			
> 5 years	Baseline to Week 58	Sparsentan	89	-3.60 (0.83)	(-5.24, -1.96)	1.66 (1.25)	(-0.80, 4.12)	0.186
		Irbesartan	75	-5.25 (0.93)	(-7.09, -3.42)			

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Table PT2GGA\_FSRM: Rate of change in eGFR (total slope) - Year 1 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan					Interaction:	0.348
Yes	Baseline to Week 58	Sparsentan	155	-2.98 (0.75)	(-4.46, -1.51)	1.12 (1.06)	(-0.96, 3.21)	0.289
		Irbesartan	161	-4.11 (0.75)	(-5.58, -2.64)			
No	Baseline to Week 58	Sparsentan	47	-5.01 (1.66)	(-8.30, -1.73)	1.15 (2.44)	(-3.68, 5.98)	0.639
		Irbesartan	41	-6.16 (1.79)	(-9.70, -2.62)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan					Interaction:	0.023 #
Male	Baseline to Week 110	Sparsentan	139	-3.12 (0.51)	(-4.12, -2.12)	1.43 (0.72)	(0.01, 2.85)	0.049 *
		Irbesartan	143	-4.55 (0.51)	(-5.56, -3.54)			
Female	Baseline to Week 110	Sparsentan	63	-2.88 (0.72)	(-4.30, -1.46)	0.53 (1.05)	(-1.54, 2.60)	0.615
		Irbesartan	59	-3.41 (0.76)	(-4.92, -1.90)			

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Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan					Interaction:	0.147
<= 45 years	Baseline to Week 110	Sparsentan	96	-3.32 (0.71)	(-4.71, -1.93)	1.72 (1.00)	(-0.26, 3.70)	0.089
		Irbesartan	99	-5.03 (0.71)	(-6.44, -3.63)			
> 45 years	Baseline to Week 110	Sparsentan	106	-2.81 (0.47)	(-3.73, -1.89)	0.68 (0.67)	(-0.65, 2.01)	0.313
		Irbesartan	103	-3.49 (0.48)	(-4.45, -2.54)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Age at IgAN diagnosis	Overall	Sparsentan					Interaction:	0.001 #	
<= 18 years	Baseline to Week 110	Sparsentan	9	-7.59 (2.91)	(-13.94, -1.23)	-0.98 (4.58)	(-11.07, 9.10)	0.834	
		Irbesartan	5	-6.60 (3.54)	(-14.43, 1.23)				
> 18 to 40 years	Baseline to Week 110	Sparsentan	102	-2.94 (0.62)	(-4.16, -1.73)	1.80 (0.87)	(0.09, 3.51)	0.040 *	
		Irbesartan	109	-4.74 (0.61)	(-5.95, -3.53)				
> 40 years	Baseline to Week 110	Sparsentan	91	-2.77 (0.53)	(-3.81, -1.72)	0.72 (0.77)	(-0.80, 2.23)	0.351	
		Irbesartan	88	-3.49 (0.56)	(-4.58, -2.39)				

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Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Geographic region	Overall	Sparsentan					Interaction:	0.053	
North America	Baseline to Week 110	Sparsentan	35	-2.74 (1.01)	(-4.74, -0.74)	1.63 (1.35)	(-1.05, 4.31)	0.230	
		Irbesartan	46	-4.37 (0.90)	(-6.16, -2.59)				
Europe	Baseline to Week 110	Sparsentan	98	-3.02 (0.56)	(-4.12, -1.91)	1.32 (0.77)	(-0.19, 2.84)	0.086	
		Irbesartan	115	-4.34 (0.53)	(-5.38, -3.30)				
Asia Pacific	Baseline to Week 110	Sparsentan	69	-3.26 (0.78)	(-4.80, -1.71)	0.48 (1.31)	(-2.10, 3.07)	0.711	
		Irbesartan	41	-3.74 (1.05)	(-5.82, -1.67)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		p-value
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline BMI	Overall	Sparsentan					Interaction:	0.019 #
< 27 kg/m**2	Baseline to Week 110	Sparsentan	83	-2.93 (0.69)	(-4.30, -1.57)	1.48 (0.96)	(-0.42, 3.38)	0.127
		Irbesartan	94	-4.41 (0.67)	(-5.73, -3.09)			
>= 27 kg/m**2	Baseline to Week 110	Sparsentan	119	-3.13 (0.52)	(-4.14, -2.11)	0.98 (0.76)	(-0.51, 2.46)	0.198
		Irbesartan	107	-4.10 (0.55)	(-5.19, -3.02)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan					Interaction:	<0.001 #
eGFR Low and UP High	Baseline to Week 110	Sparsentan	71	-4.11 (0.67)	(-5.44, -2.78)	2.00 (0.95)	(0.12, 3.87)	0.037 *
		Irbesartan	74	-6.11 (0.67)	(-7.43, -4.78)			
eGFR Low and UP Low	Baseline to Week 110	Sparsentan	55	-1.74 (0.68)	(-3.09, -0.39)	1.23 (0.96)	(-0.68, 3.14)	0.206
		Irbesartan	55	-2.96 (0.68)	(-4.32, -1.61)			
eGFR High and UP High	Baseline to Week 110	Sparsentan	37	-3.31 (1.21)	(-5.71, -0.91)	2.31 (1.76)	(-1.20, 5.82)	0.194
		Irbesartan	36	-5.62 (1.29)	(-8.18, -3.07)			
eGFR High and UP Low	Baseline to Week 110	Sparsentan	39	-2.95 (0.96)	(-4.85, -1.04)	-1.14 (1.40)	(-3.92, 1.64)	0.417
		Irbesartan	37	-1.81 (1.02)	(-3.83, 0.22)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model				
				Annualized Slope			Slope Difference	
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan					Interaction:	0.008 #
< 60 mL/min/1.73 m**2	Baseline to Week 110	Sparsentan	127	-3.09 (0.47)	(-4.02, -2.16)	1.26 (0.67)	(-0.06, 2.58)	0.061
		Irbesartan	129	-4.35 (0.48)	(-5.29, -3.42)			
60 to < 90 mL/min/1.73 m**2	Baseline to Week 110	Sparsentan	49	-3.22 (0.91)	(-5.02, -1.43)	2.69 (1.31)	(0.10, 5.28)	0.042 *
		Irbesartan	48	-5.91 (0.95)	(-7.78, -4.04)			
>= 90 mL/min/1.73 m**2	Baseline to Week 110	Sparsentan	26	-2.78 (1.37)	(-5.52, -0.05)	-2.04 (2.02)	(-6.09, 2.01)	0.317
		Irbesartan	25	-0.74 (1.49)	(-3.72, 2.24)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed post-baseline values up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. Source Data: alb, created on: 19FEB2024

Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		p-value
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline eGFR Group 2	Overall	Sparsentan					Interaction:	0.003 #
< 45 mL/min/1.73 m**2	Baseline to Week 110	Sparsentan	82	-3.69 (0.59)	(-4.85, -2.53)	1.41 (0.84)	(-0.26, 3.07)	0.097
		Irbesartan	80	-5.09 (0.60)	(-6.28, -3.90)			
45 to < 60 mL/min/1.73 m**2	Baseline to Week 110	Sparsentan	45	-2.17 (0.87)	(-3.89, -0.46)	1.25 (1.20)	(-1.14, 3.63)	0.302
		Irbesartan	49	-3.42 (0.84)	(-5.08, -1.76)			
60 to < 90 mL/min/1.73 m**2	Baseline to Week 110	Sparsentan	49	-3.22 (0.91)	(-5.02, -1.43)	2.69 (1.31)	(0.10, 5.28)	0.042 *
		Irbesartan	48	-5.91 (0.95)	(-7.78, -4.04)			
>= 90 mL/min/1.73 m**2	Baseline to Week 110	Sparsentan	26	-2.78 (1.37)	(-5.52, -0.05)	-2.04 (2.02)	(-6.09, 2.01)	0.317
		Irbesartan	25	-0.74 (1.49)	(-3.72, 2.24)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline urine protein excretion	Overall	Sparsentan					Interaction:	0.718	
<= 1.75 g/day	Baseline to Week 110	Sparsentan	98	-2.18 (0.55)	(-3.26, -1.09)	0.96 (0.80)	(-0.62, 2.53)	0.233	
		Irbesartan	93	-3.13 (0.58)	(-4.28, -1.99)				
> 1.75 g/day	Baseline to Week 110	Sparsentan	104	-3.94 (0.61)	(-5.15, -2.74)	1.28 (0.86)	(-0.42, 2.98)	0.139	
		Irbesartan	109	-5.23 (0.61)	(-6.42, -4.03)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
 Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline use of antihypertensives	Overall	Sparsentan					Interaction:	0.430	
Yes	Baseline to Week 110	Sparsentan	90	-2.73 (0.56)	(-3.83, -1.63)	1.68 (0.81)	(0.09, 3.27)	0.038 *	
		Irbesartan	88	-4.41 (0.58)	(-5.56, -3.27)				
No	Baseline to Week 110	Sparsentan	112	-3.30 (0.60)	(-4.47, -2.12)	0.81 (0.85)	(-0.86, 2.48)	0.342	
		Irbesartan	114	-4.11 (0.61)	(-5.30, -2.92)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan					Interaction:	0.102
<= 5 years	Baseline to Week 110	Sparsentan	113	-2.69 (0.62)	(-3.91, -1.47)	1.52 (0.86)	(-0.17, 3.21)	0.078
		Irbesartan	127	-4.21 (0.60)	(-5.38, -3.03)			
> 5 years	Baseline to Week 110	Sparsentan	89	-3.51 (0.50)	(-4.50, -2.51)	0.77 (0.76)	(-0.73, 2.26)	0.312
		Irbesartan	75	-4.27 (0.56)	(-5.39, -3.16)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGB\_FSRM: Rate of change in eGFR (total slope) - Year 2 - by subgroup  
Full Analysis Set

Rate of change in eGFR (total slope) - year 2				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope			Slope Difference		
				LS-Mean (STE)	95% CI		LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.205
Yes	Baseline to Week 110	Sparsentan	155	-2.95 (0.45)	(-3.84, -2.05)	1.36 (0.65)	(0.09, 2.63)		0.036 *
		Irbesartan	161	-4.30 (0.46)	(-5.21, -3.40)				
No	Baseline to Week 110	Sparsentan	47	-3.43 (0.98)	(-5.37, -1.48)	0.49 (1.44)	(-2.37, 3.35)		0.735
		Irbesartan	41	-3.91 (1.06)	(-6.01, -1.82)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Sex	Overall	Sparsentan					Interaction:	0.018 #	
Male	Week 6 to Week 58	Sparsentan	139	-2.73 (0.86)	(-4.43, -1.04)	2.37 (1.22)	(-0.03, 4.77)	0.053	
		Irbesartan	143	-5.10 (0.87)	(-6.80, -3.40)				
Female	Week 6 to Week 58	Sparsentan	63	-3.92 (1.40)	(-6.69, -1.15)	-0.64 (2.03)	(-4.65, 3.37)	0.754	
		Irbesartan	59	-3.28 (1.47)	(-6.19, -0.37)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan					Interaction:	0.378
<= 45 years	Week 6 to Week 58	Sparsentan	96	-3.66 (1.22)	(-6.06, -1.25)	1.26 (1.73)	(-2.14, 4.66)	0.466
		Irbesartan	99	-4.92 (1.22)	(-7.33, -2.51)			
> 45 years	Week 6 to Week 58	Sparsentan	106	-2.61 (0.86)	(-4.29, -0.92)	1.66 (1.23)	(-0.76, 4.08)	0.177
		Irbesartan	103	-4.27 (0.88)	(-6.00, -2.53)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		p-value
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Age at IgAN diagnosis	Overall	Sparsentan					Interaction:	0.005 #
<= 18 years	Week 6 to Week 58	Sparsentan	9	-5.27 (3.88)	(-13.36, 2.83)	3.07 (6.30)	(-10.05, 16.18)	0.632
		Irbesartan	5	-8.33 (4.95)	(-18.65, 1.99)			
> 18 to 40 years	Week 6 to Week 58	Sparsentan	102	-3.50 (1.10)	(-5.66, -1.34)	0.92 (1.54)	(-2.11, 3.96)	0.549
		Irbesartan	109	-4.42 (1.08)	(-6.56, -2.29)			
> 40 years	Week 6 to Week 58	Sparsentan	91	-2.39 (0.98)	(-4.32, -0.47)	2.14 (1.41)	(-0.63, 4.91)	0.129
		Irbesartan	88	-4.53 (1.01)	(-6.52, -2.55)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan					Interaction:	0.648
North America	Week 6 to Week 58	Sparsentan	35	-2.81 (1.78)	(-6.32, 0.71)	0.79 (2.37)	(-3.90, 5.48)	0.740
		Irbesartan	46	-3.59 (1.57)	(-6.69, -0.49)			
Europe	Week 6 to Week 58	Sparsentan	98	-2.88 (0.99)	(-4.84, -0.93)	1.67 (1.36)	(-1.01, 4.36)	0.221
		Irbesartan	115	-4.55 (0.93)	(-6.39, -2.72)			
Asia Pacific	Week 6 to Week 58	Sparsentan	69	-3.57 (1.35)	(-6.24, -0.90)	2.12 (2.25)	(-2.31, 6.55)	0.347
		Irbesartan	41	-5.69 (1.79)	(-9.23, -2.15)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline BMI	Overall	Sparsentan					Interaction:	0.035 #	
< 27 kg/m**2	Week 6 to Week 58	Sparsentan	83	-2.82 (1.25)	(-5.28, -0.36)	2.15 (1.74)	(-1.27, 5.56)	0.217	
		Irbesartan	94	-4.97 (1.20)	(-7.34, -2.60)				
>= 27 kg/m**2	Week 6 to Week 58	Sparsentan	119	-3.27 (0.89)	(-5.02, -1.52)	1.00 (1.30)	(-1.55, 3.55)	0.442	
		Irbesartan	107	-4.27 (0.95)	(-6.13, -2.41)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan					Interaction:	0.002 #
eGFR Low and UP High	Week 6 to Week 58	Sparsentan	71	-4.00 (1.09)	(-6.15, -1.85)	2.72 (1.54)	(-0.31, 5.75)	0.079
		Irbesartan	74	-6.72 (1.08)	(-8.85, -4.58)			
eGFR Low and UP Low	Week 6 to Week 58	Sparsentan	55	-1.05 (1.19)	(-3.39, 1.30)	1.16 (1.68)	(-2.15, 4.47)	0.492
		Irbesartan	55	-2.20 (1.19)	(-4.54, 0.13)			
eGFR High and UP High	Week 6 to Week 58	Sparsentan	37	-1.79 (2.24)	(-6.22, 2.63)	3.00 (3.26)	(-3.46, 9.45)	0.360
		Irbesartan	36	-4.79 (2.37)	(-9.49, -0.09)			
eGFR High and UP Low	Week 6 to Week 58	Sparsentan	39	-5.67 (1.80)	(-9.23, -2.11)	-1.63 (2.62)	(-6.79, 3.54)	0.534
		Irbesartan	37	-4.04 (1.89)	(-7.78, -0.30)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. Source Data: alb, created on: 19FEB2024



Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
				Annualized Slope		Slope Difference		
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan					Interaction:	0.010 #
< 60 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	127	-2.80 (0.79)	(-4.35, -1.25)	1.65 (1.12)	(-0.54, 3.84)	0.140
		Irbesartan	129	-4.45 (0.79)	(-6.00, -2.90)			
60 to < 90 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	49	-3.13 (1.65)	(-6.38, 0.11)	2.80 (2.38)	(-1.89, 7.49)	0.240
		Irbesartan	48	-5.94 (1.71)	(-9.32, -2.56)			
>= 90 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	26	-4.56 (2.70)	(-9.93, 0.80)	-1.83 (3.95)	(-9.67, 6.00)	0.643
		Irbesartan	25	-2.73 (2.88)	(-8.44, 2.98)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
				Annualized Slope		Slope Difference		
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan					Interaction:	0.013 #
< 45 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	82	-3.91 (0.91)	(-5.70, -2.12)	2.43 (1.30)	(-0.13, 4.99)	0.063
		Irbesartan	80	-6.34 (0.93)	(-8.18, -4.51)			
45 to < 60 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	45	-0.87 (1.52)	(-3.87, 2.12)	0.84 (2.10)	(-3.30, 4.98)	0.690
		Irbesartan	49	-1.71 (1.45)	(-4.57, 1.14)			
60 to < 90 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	49	-3.13 (1.65)	(-6.38, 0.11)	2.80 (2.38)	(-1.89, 7.49)	0.240
		Irbesartan	48	-5.94 (1.71)	(-9.32, -2.56)			
>= 90 mL/min/1.73 m**2	Week 6 to Week 58	Sparsentan	26	-4.56 (2.70)	(-9.93, 0.80)	-1.83 (3.95)	(-9.67, 6.00)	0.643
		Irbesartan	25	-2.73 (2.88)	(-8.44, 2.98)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model. All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. Source Data: alb, created on: 19FEB2024

Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline urine protein excretion	Overall	Sparsentan					Interaction:	0.817	
<= 1.75 g/day	Week 6 to Week 58	Sparsentan	98	-2.84 (1.05)	(-4.90, -0.79)	0.17 (1.51)	(-2.80, 3.15)	0.908	
		Irbesartan	93	-3.02 (1.10)	(-5.17, -0.86)				
> 1.75 g/day	Week 6 to Week 58	Sparsentan	104	-3.35 (1.02)	(-5.35, -1.35)	2.55 (1.43)	(-0.27, 5.36)	0.076	
		Irbesartan	109	-5.90 (1.01)	(-7.88, -3.92)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model					
				Annualized Slope			Slope Difference		
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline use of antihypertensives	Overall	Sparsentan					Interaction:	0.178	
Yes	Week 6 to Week 58	Sparsentan	90	-3.03 (1.00)	(-5.01, -1.06)	1.63 (1.44)	(-1.21, 4.46)	0.260	
		Irbesartan	88	-4.66 (1.03)	(-6.69, -2.63)				
No	Week 6 to Week 58	Sparsentan	112	-3.13 (1.05)	(-5.19, -1.07)	1.37 (1.49)	(-1.56, 4.29)	0.359	
		Irbesartan	114	-4.49 (1.06)	(-6.57, -2.42)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan					Interaction:	0.366
<= 5 years	Week 6 to Week 58	Sparsentan	113	-2.87 (1.09)	(-5.01, -0.74)	1.26 (1.50)	(-1.69, 4.22)	0.400
		Irbesartan	127	-4.14 (1.04)	(-6.18, -2.10)			
> 5 years	Week 6 to Week 58	Sparsentan	89	-3.34 (0.90)	(-5.10, -1.58)	1.97 (1.34)	(-0.67, 4.61)	0.144
		Irbesartan	75	-5.31 (1.00)	(-7.28, -3.34)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 58 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLA\_FSRM: Rate of change in eGFR (chronic slope) - Year 1 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 1				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan					Interaction:	0.768
Yes	Week 6 to Week 58	Sparsentan	155	-2.97 (0.80)	(-4.53, -1.40)	1.30 (1.13)	(-0.92, 3.51)	0.251
		Irbesartan	161	-4.26 (0.80)	(-5.83, -2.70)			
No	Week 6 to Week 58	Sparsentan	47	-3.60 (1.74)	(-7.04, -0.16)	2.19 (2.57)	(-2.88, 7.27)	0.394
		Irbesartan	41	-5.79 (1.88)	(-9.52, -2.07)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Sex	Overall	Sparsentan					Interaction:	0.037	#
Male	Week 6 to Week 110	Sparsentan	139	-3.07 (0.52)	(-4.09, -2.05)	1.61 (0.74)	(0.16, 3.07)	0.030	*
		Irbesartan	143	-4.68 (0.53)	(-5.71, -3.65)				
Female	Week 6 to Week 110	Sparsentan	63	-2.52 (0.74)	(-3.98, -1.06)	0.49 (1.08)	(-1.65, 2.62)	0.653	
		Irbesartan	59	-3.01 (0.79)	(-4.57, -1.45)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan					Interaction:	0.089
<= 45 years	Week 6 to Week 110	Sparsentan	96	-3.05 (0.72)	(-4.47, -1.62)	2.02 (1.03)	(-0.00, 4.05)	0.050
		Irbesartan	99	-5.07 (0.73)	(-6.52, -3.62)			
> 45 years	Week 6 to Week 110	Sparsentan	106	-2.77 (0.48)	(-3.71, -1.83)	0.67 (0.69)	(-0.69, 2.03)	0.332
		Irbesartan	103	-3.44 (0.50)	(-4.42, -2.46)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Age at IgAN diagnosis	Overall	Sparsentan					Interaction:	0.005 #	
<= 18 years	Week 6 to Week 110	Sparsentan	9	-8.80 (3.38)	(-16.29, -1.30)	-2.00 (5.35)	(-14.03, 10.02)	0.716	
		Irbesartan	5	-6.79 (4.16)	(-16.20, 2.62)				
> 18 to 40 years	Week 6 to Week 110	Sparsentan	102	-2.70 (0.63)	(-3.94, -1.45)	2.21 (0.89)	(0.45, 3.97)	0.014 *	
		Irbesartan	109	-4.91 (0.63)	(-6.15, -3.66)				
> 40 years	Week 6 to Week 110	Sparsentan	91	-2.71 (0.54)	(-3.78, -1.64)	0.52 (0.79)	(-1.03, 2.07)	0.506	
		Irbesartan	88	-3.24 (0.57)	(-4.36, -2.11)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan					Interaction:	0.305
North America	Week 6 to Week 110	Sparsentan	35	-2.58 (1.02)	(-4.62, -0.55)	1.85 (1.38)	(-0.88, 4.58)	0.182
		Irbesartan	46	-4.43 (0.92)	(-6.26, -2.61)			
Europe	Week 6 to Week 110	Sparsentan	98	-3.02 (0.58)	(-4.16, -1.89)	1.31 (0.79)	(-0.24, 2.87)	0.098
		Irbesartan	115	-4.34 (0.54)	(-5.40, -3.27)			
Asia Pacific	Week 6 to Week 110	Sparsentan	69	-2.91 (0.80)	(-4.49, -1.33)	0.71 (1.34)	(-1.94, 3.36)	0.596
		Irbesartan	41	-3.62 (1.08)	(-5.75, -1.49)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

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UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

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Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan					Interaction:	0.008 #
< 27 kg/m**2	Week 6 to Week 110	Sparsentan	83	-2.83 (0.71)	(-4.22, -1.43)	1.39 (0.99)	(-0.56, 3.33)	0.161
		Irbesartan	94	-4.21 (0.69)	(-5.57, -2.86)			
>= 27 kg/m**2	Week 6 to Week 110	Sparsentan	119	-2.95 (0.53)	(-4.00, -1.91)	1.27 (0.78)	(-0.26, 2.80)	0.102
		Irbesartan	107	-4.23 (0.57)	(-5.35, -3.11)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan					Interaction:	<0.001 #
eGFR Low and UP High	Week 6 to Week 110	Sparsentan	71	-4.16 (0.68)	(-5.50, -2.82)	1.87 (0.96)	(-0.03, 3.77)	0.054
		Irbesartan	74	-6.03 (0.68)	(-7.37, -4.68)			
eGFR Low and UP Low	Week 6 to Week 110	Sparsentan	55	-1.79 (0.70)	(-3.16, -0.41)	1.25 (0.99)	(-0.70, 3.20)	0.207
		Irbesartan	55	-3.04 (0.70)	(-4.42, -1.65)			
eGFR High and UP High	Week 6 to Week 110	Sparsentan	37	-2.94 (1.26)	(-5.46, -0.43)	2.41 (1.85)	(-1.25, 6.08)	0.194
		Irbesartan	36	-5.36 (1.35)	(-8.03, -2.68)			
eGFR High and UP Low	Week 6 to Week 110	Sparsentan	39	-2.39 (0.99)	(-4.35, -0.42)	-0.37 (1.45)	(-3.23, 2.50)	0.801
		Irbesartan	37	-2.02 (1.05)	(-4.11, 0.07)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline eGFR Group 1	Overall	Sparsentan					Interaction:	<0.001 #	
< 60 mL/min/1.73 m**2	Week 6 to Week 110	Sparsentan	127	-3.13 (0.48)	(-4.07, -2.18)	1.25 (0.68)	(-0.09, 2.59)	0.068	
		Irbesartan	129	-4.37 (0.48)	(-5.32, -3.42)				
60 to < 90 mL/min/1.73 m**2	Week 6 to Week 110	Sparsentan	49	-3.00 (0.94)	(-4.86, -1.15)	2.69 (1.36)	(0.01, 5.37)	0.049 *	
		Irbesartan	48	-5.69 (0.98)	(-7.63, -3.76)				
≥ 90 mL/min/1.73 m**2	Week 6 to Week 110	Sparsentan	26	-1.89 (1.42)	(-4.73, 0.95)	-0.89 (2.11)	(-5.11, 3.32)	0.674	
		Irbesartan	25	-1.00 (1.56)	(-4.11, 2.12)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan					Interaction:	<0.001 #
< 45 mL/min/1.73 m**2	Week 6 to Week 110	Sparsentan	82	-3.78 (0.59)	(-4.95, -2.61)	1.36 (0.85)	(-0.32, 3.03)	0.112
		Irbesartan	80	-5.13 (0.61)	(-6.33, -3.93)			
45 to < 60 mL/min/1.73 m**2	Week 6 to Week 110	Sparsentan	45	-2.14 (0.89)	(-3.90, -0.39)	1.31 (1.23)	(-1.13, 3.76)	0.290
		Irbesartan	49	-3.46 (0.86)	(-5.16, -1.75)			
60 to < 90 mL/min/1.73 m**2	Week 6 to Week 110	Sparsentan	49	-3.00 (0.94)	(-4.86, -1.15)	2.69 (1.36)	(0.01, 5.37)	0.049 *
		Irbesartan	48	-5.69 (0.98)	(-7.63, -3.76)			
>= 90 mL/min/1.73 m**2	Week 6 to Week 110	Sparsentan	26	-1.89 (1.42)	(-4.73, 0.95)	-0.89 (2.11)	(-5.11, 3.32)	0.674
		Irbesartan	25	-1.00 (1.56)	(-4.11, 2.12)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model					
				Annualized Slope			Slope Difference		
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline urine protein excretion	Overall	Sparsentan					Interaction:	0.447	
<= 1.75 g/day	Week 6 to Week 110	Sparsentan	98	-2.06 (0.57)	(-3.18, -0.95)	1.20 (0.83)	(-0.43, 2.83)	0.147	
		Irbesartan	93	-3.26 (0.60)	(-4.45, -2.08)				
> 1.75 g/day	Week 6 to Week 110	Sparsentan	104	-3.76 (0.63)	(-5.00, -2.52)	1.32 (0.89)	(-0.43, 3.07)	0.138	
		Irbesartan	109	-5.08 (0.63)	(-6.32, -3.85)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
 Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model					
				Annualized Slope			Slope Difference		
Subgroup	Time	Treatment	N	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
Baseline use of antihypertensives	Overall	Sparsentan					Interaction:	0.130	
Yes	Week 6 to Week 110	Sparsentan	90	-2.80 (0.57)	(-3.93, -1.67)	1.61 (0.82)	(-0.02, 3.23)	0.053	
		Irbesartan	88	-4.41 (0.59)	(-5.58, -3.24)				
No	Week 6 to Week 110	Sparsentan	112	-2.97 (0.61)	(-4.18, -1.77)	1.10 (0.87)	(-0.62, 2.82)	0.208	
		Irbesartan	114	-4.07 (0.62)	(-5.30, -2.85)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024



Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model				
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference		
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan					Interaction:	0.079
<= 5 years	Week 6 to Week 110	Sparsentan	113	-2.49 (0.63)	(-3.72, -1.25)	1.75 (0.87)	(0.02, 3.47)	0.047 *
		Irbesartan	127	-4.23 (0.61)	(-5.43, -3.04)			
> 5 years	Week 6 to Week 110	Sparsentan	89	-3.42 (0.52)	(-4.44, -2.40)	0.75 (0.78)	(-0.79, 2.29)	0.338
		Irbesartan	75	-4.17 (0.58)	(-5.32, -3.02)			

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GGLB\_FSRM: Rate of change in eGFR (chronic slope) - Year 2 by subgroup  
Full Analysis Set

Rate of change in eGFR (chronic slope) - year 2				Mixed Random Coefficient Model					
Subgroup	Time	Treatment	N	Annualized Slope		Slope Difference			
				LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
History of hypertension	Overall	Sparsentan					Interaction:	0.445	
Yes	Week 6 to Week 110	Sparsentan	155	-2.94 (0.46)	(-3.86, -2.03)	1.44 (0.66)	(0.14, 2.74)	0.030 *	
		Irbesartan	161	-4.39 (0.47)	(-5.31, -3.46)				
No	Week 6 to Week 110	Sparsentan	47	-2.75 (1.00)	(-4.74, -0.75)	0.76 (1.48)	(-2.18, 3.70)	0.609	
		Irbesartan	41	-3.51 (1.09)	(-5.67, -1.35)				

N = number of patients in analysis set. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction. MRCM = Mixed Random Coefficient Model. An MRCM was applied including treatment, time, randomization strata, baseline (as covariate), and interaction term treatment-by-time, random intercept and random slope per subject. For interaction test, subgroup and interaction subgroup\*treatment were added to the model.

All observed post-baseline values from Week 6 up to Week 110 are included in the model. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model.

A decrease reflects a worsening of the status of the patient. Results are presented in ml/min/1.73 m\*\*2. eGFR = estimated glomerular filtration rate. A first order autoregressive covariance structure was used.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: alb, created on: 19FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.508
Male	Double-blind period	Sparsentan	139	15 (10.8)	0.772 [0.412, 1.445]	0.744 [0.364, 1.520]	-3.2 [-11.6, 5.2]	0.472
		Irbesartan	143	20 (14.0)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	0.468 [0.123, 1.788]	0.442 [0.105, 1.854]	-5.4 [-16.4, 5.6]	0.312
		Irbesartan	59	6 (10.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.553
<= 45 years	Double-blind period	Sparsentan	96	10 (10.4)	0.607 [0.293, 1.257]	0.561 [0.243, 1.296]	-6.8 [-17.4, 3.9]	0.215
		Irbesartan	99	17 (17.2)				
> 45 years	Double-blind period	Sparsentan	106	8 (7.5)	0.864 [0.347, 2.152]	0.853 [0.316, 2.303]	-1.2 [-9.6, 7.2]	0.804
		Irbesartan	103	9 (8.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.407
<= 18 years	Double-blind period	Sparsentan	9	3 (33.3)	1.667 [0.230, 12.091]	2.000 [0.150, 26.734]	13.3 [-48.9, 75.6]	1.000
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	7 (6.9)	0.468 [0.201, 1.090]	0.428 [0.168, 1.089]	-7.8 [-17.0, 1.4]	0.079
		Irbesartan	109	16 (14.7)				
> 40 years	Double-blind period	Sparsentan	91	8 (8.8)	0.860 [0.347, 2.127]	0.846 [0.311, 2.302]	-1.4 [-11.2, 8.3]	0.803
		Irbesartan	88	9 (10.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.728
North America	Double-blind period	Sparsentan	35	3 (8.6)	0.657 [0.177, 2.446]	0.625 [0.145, 2.696]	-4.5 [-20.4, 11.5]	0.725
		Irbesartan	46	6 (13.0)				
Europe	Double-blind period	Sparsentan	98	8 (8.2)	0.587 [0.262, 1.312]	0.550 [0.225, 1.347]	-5.7 [-15.0, 3.5]	0.201
		Irbesartan	115	16 (13.9)				
Asia Pacific	Double-blind period	Sparsentan	69	7 (10.1)	1.040 [0.324, 3.337]	1.044 [0.286, 3.810]	0.4 [-13.1, 13.9]	1.000
		Irbesartan	41	4 (9.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
ESRD = End-stage renal disease.  
Source Data: aeFF, created on: 20FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.246
< 27 kg/m**2	Double-blind period	Sparsentan	83	5 (6.0)	0.436 [0.162, 1.170]	0.399 [0.136, 1.173]	-7.8 [-17.6, 2.0]	0.133
		Irbesartan	94	13 (13.8)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	13 (10.9)	0.899 [0.436, 1.853]	0.887 [0.392, 2.008]	-1.2 [-10.5, 8.0]	0.836
		Irbesartan	107	13 (12.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.956
eGFR Low and UP High	Double-blind period	Sparsentan	71	13 (18.3)	0.713 [0.381, 1.333]	0.649 [0.293, 1.438]	-7.4 [-22.2, 7.4]	0.321
		Irbesartan	74	19 (25.7)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)	0.500 [0.095, 2.618]	0.481 [0.084, 2.742]	-3.6 [-13.9, 6.6]	0.679
		Irbesartan	55	4 (7.3)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4)	0.973 [0.145, 6.541]	0.971 [0.129, 7.294]	-0.2 [-13.3, 13.0]	1.000
		Irbesartan	36	2 (5.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)	0.949 [0.062, 14.620]	0.947 [0.057, 15.721]	-0.1 [-10.0, 9.7]	1.000
		Irbesartan	37	1 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeFF, created on: 20FEB2024



Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.846
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	14 (11.0)	0.646 [0.346, 1.206]	0.603 [0.293, 1.238]	-6.0 [-15.3, 3.2]	0.208
		Irbesartan	129	22 (17.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.980 [0.260, 3.695]	0.978 [0.230, 4.155]	-0.2 [-13.2, 12.8]	1.000
		Irbesartan	48	4 (8.3)				
>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
ESRD = End-stage renal disease.  
Source Data: aeфф, created on: 20FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.893
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	12 (14.6)	0.689 [0.352, 1.348]	0.635 [0.282, 1.433]	-6.6 [-19.6, 6.4]	0.310
		Irbesartan	80	17 (21.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	2 (4.4)	0.436 [0.089, 2.134]	0.409 [0.075, 2.224]	-5.8 [-18.3, 6.8]	0.438
		Irbesartan	49	5 (10.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.980 [0.260, 3.695]	0.978 [0.230, 4.155]	-0.2 [-13.2, 12.8]	1.000
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.506
<= 1.75 g/day	Double-blind period	Sparsentan	98	3 (3.1)	0.474 [0.122, 1.842]	0.458 [0.111, 1.887]	-3.4 [-10.5, 3.7]	0.321
		Irbesartan	93	6 (6.5)				
> 1.75 g/day	Double-blind period	Sparsentan	104	15 (14.4)	0.786 [0.426, 1.451]	0.750 [0.361, 1.558]	-3.9 [-14.8, 6.9]	0.465
		Irbesartan	109	20 (18.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.223
Yes	Double-blind period	Sparsentan	90	11 (12.2)	0.978 [0.447, 2.138]	0.975 [0.399, 2.380]	-0.3 [-11.1, 10.5]	1.000
		Irbesartan	88	11 (12.5)				
No	Double-blind period	Sparsentan	112	7 (6.3)	0.475 [0.201, 1.121]	0.440 [0.172, 1.124]	-6.9 [-15.4, 1.6]	0.115
		Irbesartan	114	15 (13.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.503
<= 5 years	Double-blind period	Sparsentan	113	13 (11.5)	0.812 [0.417, 1.581]	0.787 [0.367, 1.689]	-2.7 [-12.0, 6.6]	0.569
		Irbesartan	127	18 (14.2)				
> 5 years	Double-blind period	Sparsentan	89	5 (5.6)	0.527 [0.180, 1.542]	0.499 [0.156, 1.594]	-5.0 [-14.7, 4.6]	0.259
		Irbesartan	75	8 (10.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2GED\_FSIM: Patients with confirmed 40% reduction in eGFR, ESRD or death by subgroup  
Full Analysis Set

<u>Confirmed 40% reduction in eGFR, ESRD, or death</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.159
Yes	Double-blind period	Sparsentan	155	17 (11.0)	0.841 [0.461, 1.532]	0.821 [0.416, 1.623]	-2.1 [-9.9, 5.7]	0.607
		Irbesartan	161	21 (13.0)				
No	Double-blind period	Sparsentan	47	1 (2.1)	0.174 [0.021, 1.433]	0.157 [0.018, 1.400]	-10.1 [-23.2, 3.0]	0.093
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
ESRD = End-stage renal disease.  
Source Data: aeff, created on: 20FEB2024

Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.801
Male	Sparsentan	139	15 (10.8)	NE		0.645	(0.328, 1.270)	0.205
	Irbesartan	143	20 (14.0)	117.0	(117.0, NE)			
Female	Sparsentan	63	3 (4.8)	NE		0.625	(0.156, 2.506)	0.507
	Irbesartan	59	6 (10.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.319
<= 45 years	Sparsentan	96	10 (10.4)	NE		0.482	(0.220, 1.058)	0.069
	Irbesartan	99	17 (17.2)	NE				
> 45 years	Sparsentan	106	8 (7.5)	NE		0.865	(0.328, 2.282)	0.770
	Irbesartan	103	9 (8.7)	117.0	(117.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024



Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: 0.419
<= 18 years	Sparsentan	9	3 (33.3)	NE		NE		NE
	Irbesartan	5	1 (20.0)	111.4	(NE, NE)			
> 18 to 40 years	Sparsentan	102	7 (6.9)	NE		0.399	(0.163, 0.973)	0.043 *
	Irbesartan	109	16 (14.7)	NE				
> 40 years	Sparsentan	91	8 (8.8)	NE		0.839	(0.318, 2.214)	0.724
	Irbesartan	88	9 (10.2)	117.0	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.964
North America	Sparsentan	35	3 (8.6)	NE		0.497	(0.124, 1.996)	0.324
	Irbesartan	46	6 (13.0)	NE				
Europe	Sparsentan	98	8 (8.2)	NE		0.608	(0.257, 1.437)	0.257
	Irbesartan	115	16 (13.9)	117.0	(117.0, NE)			
Asia Pacific	Sparsentan	69	7 (10.1)	NE		0.624	(0.178, 2.192)	0.462
	Irbesartan	41	4 (9.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.355
< 27 kg/m**2	Sparsentan	83	5 (6.0)	NE		0.427	(0.152, 1.203)	0.107
	Irbesartan	94	13 (13.8)	NE				
≥ 27 kg/m**2	Sparsentan	119	13 (10.9)	NE		0.815	(0.371, 1.792)	0.611
	Irbesartan	107	13 (12.1)	117.0	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.959
eGFR Low and UP High	Sparsentan	71	13 (18.3)	NE		0.627	(0.307, 1.281)	0.200
	Irbesartan	74	19 (25.7)	117.0	(117.0, NE)			
eGFR Low and UP Low	Sparsentan	55	2 (3.6)	NE		0.479	(0.088, 2.616)	0.395
	Irbesartan	55	4 (7.3)	NE				
eGFR High and UP High	Sparsentan	37	2 (5.4)	NE		0.867	(0.122, 6.157)	0.886
	Irbesartan	36	2 (5.6)	NE				
eGFR High and UP Low	Sparsentan	39	1 (2.6)	NE		0.781	(0.049, 12.483)	0.861
	Irbesartan	37	1 (2.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	NE
< 60 mL/min/1.73 m**2	Sparsentan	127	14 (11.0)	NE		0.566	(0.288, 1.111)	0.098
	Irbesartan	129	22 (17.1)	117.0	(117.0, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	4 (8.2)	NE		0.919	(0.227, 3.721)	0.906
	Irbesartan	48	4 (8.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: NE
< 45 mL/min/1.73 m**2	Sparsentan	82	12 (14.6)	NE		0.637	(0.301, 1.346)	0.237
	Irbesartan	80	17 (21.3)	117.0	(117.0, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	2 (4.4)	NE		0.350	(0.068, 1.805)	0.210
	Irbesartan	49	5 (10.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	4 (8.2)	NE		0.919	(0.227, 3.721)	0.906
	Irbesartan	48	4 (8.3)	NE				
>= 90 mL/min/1.73 m**2	Sparsentan	26	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.628
<= 1.75 g/day	Sparsentan	98	3 (3.1)	NE		0.441	(0.110, 1.769)	0.248
	Irbesartan	93	6 (6.5)	NE				
> 1.75 g/day	Sparsentan	104	15 (14.4)	NE		0.679	(0.344, 1.341)	0.265
	Irbesartan	109	20 (18.3)	117.0	(117.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.201
Yes	Sparsentan	90	11 (12.2)	NE		0.888	(0.384, 2.051)	0.781
	Irbesartan	88	11 (12.5)	NE				
No	Sparsentan	112	7 (6.3)	NE		0.422	(0.170, 1.048)	0.063
	Irbesartan	114	15 (13.2)	117.0	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF\_tte, created on: 28FEB2024



Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.873
<= 5 years	Sparsentan	113	13 (11.5)	NE		0.655	(0.320, 1.338)	0.245
	Irbesartan	127	18 (14.2)	NE				
> 5 years	Sparsentan	89	5 (5.6)	NE		0.609	(0.192, 1.926)	0.398
	Irbesartan	75	8 (10.7)	117.0	(117.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GEDT\_FSTM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR, ESRD, or death				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension		Sparsentan				Interaction test:		0.091
Yes	Sparsentan	155	17 (11.0)	NE		0.763	(0.401, 1.454)	0.412
	Irbesartan	161	21 (13.0)	117.0	(117.0, NE)			
No	Sparsentan	47	1 (2.1)	NE		0.142	(0.016, 1.237)	0.077
	Irbesartan	41	5 (12.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Figure PF2GEDT\_FSKM: Time to confirmed 40% reduction in eGFR, ESRD or death by subgroup  
Full Analysis Set

Not done. No significant subgroups.

ESRD = End-stage renal disease. eGFR = Estimated glomerular filtration rate.  
Reference table: PT2GEDT\_FSTM

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.806
Male	Double-blind period	Sparsentan	139	15 (10.8)	0.857 [0.450, 1.633]	0.840 [0.405, 1.741]	-1.8 [-10.0, 6.4]	0.713
		Irbesartan	143	18 (12.6)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	0.702 [0.164, 3.007]	0.688 [0.147, 3.210]	-2.0 [-12.0, 7.9]	0.711
		Irbesartan	59	4 (6.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Confirmed 40% reduction in eGFR								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.654
<= 45 years	Double-blind period	Sparsentan	96	10 (10.4)	0.737 [0.344, 1.577]	0.706 [0.297, 1.677]	-3.7 [-13.9, 6.5]	0.515
		Irbesartan	99	14 (14.1)				
> 45 years	Double-blind period	Sparsentan	106	8 (7.5)	0.972 [0.379, 2.492]	0.969 [0.350, 2.688]	-0.2 [-8.4, 7.9]	1.000
		Irbesartan	103	8 (7.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 20FEB2024

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Confirmed 40% reduction in eGFR								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.542
<= 18 years	Double-blind period	Sparsentan	9	3 (33.3)	1.667 [0.230, 12.091]	2.000 [0.150, 26.734]	13.3 [-48.9, 75.6]	1.000
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	7 (6.9)	0.575 [0.239, 1.385]	0.544 [0.208, 1.423]	-5.1 [-13.8, 3.7]	0.245
		Irbesartan	109	13 (11.9)				
> 40 years	Double-blind period	Sparsentan	91	8 (8.8)	0.967 [0.380, 2.464]	0.964 [0.345, 2.692]	-0.3 [-9.8, 9.2]	1.000
		Irbesartan	88	8 (9.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

Confirmed 40% reduction in eGFR								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.799
North America	Double-blind period	Sparsentan	35	3 (8.6)	0.986 [0.236, 4.123]	0.984 [0.206, 4.713]	-0.1 [-15.0, 14.7]	1.000
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	8 (8.2)	0.671 [0.294, 1.531]	0.641 [0.257, 1.599]	-4.0 [-13.0, 5.0]	0.374
		Irbesartan	115	14 (12.2)				
Asia Pacific	Double-blind period	Sparsentan	69	7 (10.1)	1.040 [0.324, 3.337]	1.044 [0.286, 3.810]	0.4 [-13.1, 13.9]	1.000
		Irbesartan	41	4 (9.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 20FEB2024

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

<u>Confirmed 40% reduction in eGFR</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.402
< 27 kg/m**2	Double-blind period	Sparsentan	83	5 (6.0)	0.566 [0.202, 1.590]	0.538 [0.176, 1.645]	-4.6 [-13.8, 4.6]	0.295
		Irbesartan	94	10 (10.6)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	13 (10.9)	0.974 [0.465, 2.041]	0.971 [0.423, 2.231]	-0.3 [-9.4, 8.8]	1.000
		Irbesartan	107	12 (11.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeфф, created on: 20FEB2024



Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Confirmed 40% reduction in eGFR								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.929
eGFR Low and UP High	Double-blind period	Sparsentan	71	13 (18.3)	0.903 [0.463, 1.761]	0.882 [0.386, 2.015]	-2.0 [-16.2, 12.3]	0.835
		Irbesartan	74	15 (20.3)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)	0.500 [0.095, 2.618]	0.481 [0.084, 2.742]	-3.6 [-13.9, 6.6]	0.679
		Irbesartan	55	4 (7.3)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4)	0.973 [0.145, 6.541]	0.971 [0.129, 7.294]	-0.2 [-13.3, 13.0]	1.000
		Irbesartan	36	2 (5.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)	0.949 [0.062, 14.620]	0.947 [0.057, 15.721]	-0.1 [-10.0, 9.7]	1.000
		Irbesartan	37	1 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

Confirmed 40% reduction in eGFR								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.957
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	14 (11.0)	0.790 [0.411, 1.519]	0.764 [0.362, 1.611]	-2.9 [-11.8, 5.9]	0.572
		Irbesartan	129	18 (14.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.980 [0.260, 3.695]	0.978 [0.230, 4.155]	-0.2 [-13.2, 12.8]	1.000
		Irbesartan	48	4 (8.3)				
>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 20FEB2024

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Confirmed 40% reduction in eGFR								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.862
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	12 (14.6)	0.901 [0.438, 1.853]	0.884 [0.376, 2.074]	-1.6 [-14.0, 10.7]	0.830
		Irbesartan	80	13 (16.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	2 (4.4)	0.436 [0.089, 2.134]	0.409 [0.075, 2.224]	-5.8 [-18.3, 6.8]	0.438
		Irbesartan	49	5 (10.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.980 [0.260, 3.695]	0.978 [0.230, 4.155]	-0.2 [-13.2, 12.8]	1.000
		Irbesartan	48	4 (8.3)				
>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

Confirmed 40% reduction in eGFR								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.538
<= 1.75 g/day	Double-blind period	Sparsentan	98	3 (3.1)	0.569 [0.140, 2.316]	0.556 [0.129, 2.394]	-2.3 [-9.1, 4.4]	0.489
		Irbesartan	93	5 (5.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	15 (14.4)	0.925 [0.488, 1.754]	0.912 [0.430, 1.937]	-1.2 [-11.7, 9.4]	0.850
		Irbesartan	109	17 (15.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeфф, created on: 20FEB2024

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Confirmed 40% reduction in eGFR								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.333
Yes	Double-blind period	Sparsentan	90	11 (12.2)	1.076 [0.481, 2.404]	1.086 [0.436, 2.703]	0.9 [-9.7, 11.5]	1.000
		Irbesartan	88	10 (11.4)				
No	Double-blind period	Sparsentan	112	7 (6.3)	0.594 [0.243, 1.453]	0.567 [0.215, 1.497]	-4.3 [-12.4, 3.8]	0.338
		Irbesartan	114	12 (10.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 20FEB2024

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Confirmed 40% reduction in eGFR								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.471
<= 5 years	Double-blind period	Sparsentan	113	13 (11.5)	0.974 [0.485, 1.958]	0.971 [0.440, 2.139]	-0.3 [-9.3, 8.7]	1.000
		Irbesartan	127	15 (11.8)				
> 5 years	Double-blind period	Sparsentan	89	5 (5.6)	0.602 [0.199, 1.819]	0.578 [0.176, 1.903]	-3.7 [-13.1, 5.7]	0.385
		Irbesartan	75	7 (9.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2G\_FSIM: Patients with confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

<u>Confirmed 40% reduction in eGFR</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.324
Yes	Double-blind period	Sparsentan	155	17 (11.0)	0.929 [0.502, 1.721]	0.921 [0.459, 1.845]	-0.8 [-8.5, 6.8]	0.861
		Irbesartan	161	19 (11.8)				
No	Double-blind period	Sparsentan	47	1 (2.1)	0.291 [0.031, 2.688]	0.275 [0.028, 2.756]	-5.2 [-16.4, 6.1]	0.335
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.927
Male	Sparsentan	139	15 (10.8)	NE		0.719	(0.359, 1.438)	0.351
	Irbesartan	143	18 (12.6)	117.0	(117.0, NE)			
Female	Sparsentan	63	3 (4.8)	NE		0.902	(0.202, 4.039)	0.893
	Irbesartan	59	4 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024



Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.384
<= 45 years	Sparsentan	96	10 (10.4)	NE		0.576	(0.254, 1.304)	0.186
	Irbesartan	99	14 (14.1)	NE				
> 45 years	Sparsentan	106	8 (7.5)	NE		0.979	(0.360, 2.660)	0.967
	Irbesartan	103	8 (7.8)	117.0	(117.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: 0.538
<= 18 years	Sparsentan	9	3 (33.3)	NE		NE		NE
	Irbesartan	5	1 (20.0)	111.4	(NE, NE)			
> 18 to 40 years	Sparsentan	102	7 (6.9)	NE		0.485	(0.193, 1.222)	0.125
	Irbesartan	109	13 (11.9)	NE				
> 40 years	Sparsentan	91	8 (8.8)	NE		0.951	(0.350, 2.586)	0.922
	Irbesartan	88	8 (9.1)	117.0	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.986
North America	Sparsentan	35	3 (8.6)	NE		0.726	(0.162, 3.256)	0.675
	Irbesartan	46	4 (8.7)	NE				
Europe	Sparsentan	98	8 (8.2)	NE		0.697	(0.289, 1.685)	0.423
	Irbesartan	115	14 (12.2)	117.0	(117.0, NE)			
Asia Pacific	Sparsentan	69	7 (10.1)	NE		0.624	(0.178, 2.192)	0.462
	Irbesartan	41	4 (9.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan							Interaction test: 0.509
< 27 kg/m**2	Sparsentan	83	5 (6.0)	NE		0.535	(0.182, 1.572)	0.255
	Irbesartan	94	10 (10.6)	NE				
≥ 27 kg/m**2	Sparsentan	119	13 (10.9)	NE		0.892	(0.398, 1.999)	0.782
	Irbesartan	107	12 (11.2)	117.0	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.946
eGFR Low and UP High	Sparsentan	71	13 (18.3)	NE		0.794	(0.373, 1.691)	0.550
	Irbesartan	74	15 (20.3)	117.0	(117.0, NE)			
eGFR Low and UP Low	Sparsentan	55	2 (3.6)	NE		0.479	(0.088, 2.616)	0.395
	Irbesartan	55	4 (7.3)	NE				
eGFR High and UP High	Sparsentan	37	2 (5.4)	NE		0.867	(0.122, 6.157)	0.886
	Irbesartan	36	2 (5.6)	NE				
eGFR High and UP Low	Sparsentan	39	1 (2.6)	NE		0.781	(0.049, 12.483)	0.861
	Irbesartan	37	1 (2.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	NE
< 60 mL/min/1.73 m**2	Sparsentan	127	14 (11.0)	NE		0.689	(0.340, 1.395)	0.300
	Irbesartan	129	18 (14.0)	117.0	(117.0, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	4 (8.2)	NE		0.919	(0.227, 3.721)	0.906
	Irbesartan	48	4 (8.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	NE
< 45 mL/min/1.73 m**2	Sparsentan	82	12 (14.6)	NE		0.831	(0.374, 1.844)	0.648
	Irbesartan	80	13 (16.3)	117.0	(117.0, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	2 (4.4)	NE		0.350	(0.068, 1.805)	0.210
	Irbesartan	49	5 (10.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	4 (8.2)	NE		0.919	(0.227, 3.721)	0.906
	Irbesartan	48	4 (8.3)	NE				
>= 90 mL/min/1.73 m**2	Sparsentan	26	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeuff\_tte, created on: 28FEB2024

Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.641
<= 1.75 g/day	Sparsentan	98	3 (3.1)	NE		0.520	(0.124, 2.185)	0.372
	Irbesartan	93	5 (5.4)	NE				
> 1.75 g/day	Sparsentan	104	15 (14.4)	NE		0.799	(0.394, 1.623)	0.535
	Irbesartan	109	17 (15.6)	117.0	(117.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024



Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
 Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.294
Yes	Sparsentan	90	11 (12.2)	NE		0.975	(0.414, 2.299)	0.954
	Irbesartan	88	10 (11.4)	NE				
No	Sparsentan	112	7 (6.3)	NE		0.525	(0.203, 1.359)	0.184
	Irbesartan	114	12 (10.5)	117.0	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment  
 was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.844
<= 5 years	Sparsentan	113	13 (11.5)	NE		0.779	(0.370, 1.641)	0.512
	Irbesartan	127	15 (11.8)	NE				
> 5 years	Sparsentan	89	5 (5.6)	NE		0.709	(0.215, 2.337)	0.572
	Irbesartan	75	7 (9.3)	117.0	(117.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2GT\_FSTM: Time to confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

Time to confirmed 40% reduction in eGFR				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension		Sparsentan				Interaction test:		0.235
Yes	Sparsentan	155	17 (11.0)	NE		0.843	(0.436, 1.631)	0.612
	Irbesartan	161	19 (11.8)	117.0	(117.0, NE)			
No	Sparsentan	47	1 (2.1)	NE		0.227	(0.023, 2.228)	0.203
	Irbesartan	41	3 (7.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Figure PF2GT\_FSKM: Time to confirmed 40% reduction in eGFR by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_  
eGFR = estimated glomerular filtration rate.  
Reference table: PT2GT\_FSTM

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
 Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.487
Male	Double-blind period	Sparsentan	139	6 (4.3)	0.686 [0.251, 1.876]	0.672 [0.233, 1.940]	-2.0 [-7.9, 4.0]	0.598
		Irbesartan	143	9 (6.3)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	1.405 [0.243, 8.113]	1.425 [0.230, 8.844]	1.4 [-7.3, 10.0]	1.000
		Irbesartan	59	2 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
 Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.748
<= 45 years	Double-blind period	Sparsentan	96	7 (7.3)	0.902 [0.340, 2.392]	0.895 [0.311, 2.571]	-0.8 [-9.3, 7.7]	1.000
		Irbesartan	99	8 (8.1)				
> 45 years	Double-blind period	Sparsentan	106	2 (1.9)	0.648 [0.111, 3.798]	0.641 [0.105, 3.917]	-1.0 [-6.1, 4.1]	0.680
		Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
 Full Analysis Set

Reaching ESRD					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.293
<= 18 years	Double-blind period	Sparsentan	9	3 (33.3)	4.200 + [0.259, 68.038]	5.923 + [0.248, 141.482]	33.3 [-13.0, 79.7]	0.258
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	3 (2.9)	0.401 [0.109, 1.469]	0.383 [0.099, 1.484]	-4.4 [-11.2, 2.4]	0.217
		Irbesartan	109	8 (7.3)				
> 40 years	Double-blind period	Sparsentan	91	3 (3.3)	0.967 [0.201, 4.663]	0.966 [0.190, 4.919]	-0.1 [-6.5, 6.3]	1.000
		Irbesartan	88	3 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.223
North America	Double-blind period	Sparsentan	35	0 (0.0)	0.145 + [0.008, 2.608]	0.133 + [0.007, 2.556]	-8.7 [-19.4, 2.0]	0.130
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	5 (5.1)	0.838 [0.275, 2.558]	0.829 [0.255, 2.701]	-1.0 [-8.1, 6.1]	1.000
		Irbesartan	115	7 (6.1)				
Asia Pacific	Double-blind period	Sparsentan	69	4 (5.8)	5.400 + [0.298, 97.806]	5.702 + [0.299, 108.676]	5.8 [-1.7, 13.3]	0.295
		Irbesartan	41	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
ESRD = End-stage renal disease.  
Source Data: aeFF, created on: 20FEB2024



Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
 Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.221
< 27 kg/m**2	Double-blind period	Sparsentan	83	2 (2.4)	0.378 [0.078, 1.820]	0.362 [0.071, 1.846]	-4.0 [-11.0, 3.1]	0.286
		Irbesartan	94	6 (6.4)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	7 (5.9)	1.259 [0.412, 3.849]	1.275 [0.392, 4.143]	1.2 [-5.5, 7.9]	0.772
		Irbesartan	107	5 (4.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.637
eGFR Low and UP High	Double-blind period	Sparsentan	71	6 (8.5)	0.625 [0.240, 1.630]	0.591 [0.203, 1.721]	-5.1 [-16.6, 6.4]	0.429
		Irbesartan	74	10 (13.5)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)	5.000 + [0.246, 101.812]	5.187 + [0.243, 110.569]	3.6 [-3.1, 10.4]	0.495
		Irbesartan	55	0 (0.0)				
eGFR High and UP High	Double-blind period	Sparsentan	37	1 (2.7)	0.973 [0.063, 14.972]	0.972 [0.058, 16.158]	-0.1 [-10.3, 10.2]	1.000
		Irbesartan	36	1 (2.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0)	0.950 + [0.019, 46.684]	0.949 + [0.018, 49.075]	0.0 [NE, NE]	NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
ESRD = End-stage renal disease.  
Source Data: aeFF, created on: 20FEB2024

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
 Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.989
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	8 (6.3)	0.813 [0.331, 1.992]	0.800 [0.305, 2.097]	-1.5 [-8.5, 5.6]	0.808
		Irbesartan	129	10 (7.8)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	1 (2.0)	0.980 [0.063, 15.218]	0.979 [0.059, 16.116]	-0.0 [-7.8, 7.7]	1.000
		Irbesartan	48	1 (2.1)				
>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
 Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.962
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	8 (9.8)	0.867 [0.352, 2.136]	0.853 [0.312, 2.333]	-1.5 [-12.2, 9.2]	0.802
		Irbesartan	80	9 (11.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	0 (0.0)	0.362 + [0.015, 8.673]	0.355 + [0.014, 8.947]	-2.0 [-8.1, 4.0]	1.000
		Irbesartan	49	1 (2.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	1 (2.0)	0.980 [0.063, 15.218]	0.979 [0.059, 16.116]	-0.0 [-7.8, 7.7]	1.000
		Irbesartan	48	1 (2.1)				
>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
 Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.604
<= 1.75 g/day	Double-blind period	Sparsentan	98	1 (1.0)	0.474 [0.044, 5.145]	0.469 [0.042, 5.262]	-1.1 [-5.7, 3.5]	0.613
		Irbesartan	93	2 (2.2)				
> 1.75 g/day	Double-blind period	Sparsentan	104	8 (7.7)	0.932 [0.374, 2.323]	0.926 [0.343, 2.499]	-0.6 [-8.8, 7.6]	1.000
		Irbesartan	109	9 (8.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
 Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.189
Yes	Double-blind period	Sparsentan	90	6 (6.7)	1.467 [0.428, 5.021]	1.500 [0.408, 5.508]	2.1 [-5.7, 10.0]	0.747
		Irbesartan	88	4 (4.5)				
No	Double-blind period	Sparsentan	112	3 (2.7)	0.436 [0.116, 1.645]	0.421 [0.106, 1.670]	-3.5 [-9.7, 2.7]	0.333
		Irbesartan	114	7 (6.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 1.000
<= 5 years	Double-blind period	Sparsentan	113	6 (5.3)	0.843 [0.302, 2.356]	0.834 [0.280, 2.481]	-1.0 [-7.7, 5.8]	0.790
		Irbesartan	127	8 (6.3)				
> 5 years	Double-blind period	Sparsentan	89	3 (3.4)	0.843 [0.175, 4.053]	0.837 [0.164, 4.276]	-0.6 [-7.7, 6.4]	1.000
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
ESRD = End-stage renal disease.  
Source Data: aeфф, created on: 20FEB2024

Table PT2E\_FSIM: Patients reaching ESRD by subgroup  
 Full Analysis Set

Reaching ESRD								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.563
Yes	Double-blind period	Sparsentan	155	8 (5.2)	0.923 [0.366, 2.332]	0.919 [0.345, 2.446]	-0.4 [-6.0, 5.2]	1.000
		Irbesartan	161	9 (5.6)				
No	Double-blind period	Sparsentan	47	1 (2.1)	0.436 [0.041, 4.636]	0.424 [0.037, 4.854]	-2.8 [-12.8, 7.3]	0.596
		Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 ESRD = End-stage renal disease.  
 Source Data: aeff, created on: 20FEB2024



Table PT2ET\_FSTM: Time to ESRD by subgroup  
 Full Analysis Set

Time to ESRD				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.287
Male	Sparsentan	139	6 (4.3)	NE		0.568	(0.202, 1.599)	0.284
	Irbesartan	143	9 (6.3)	NE				
Female	Sparsentan	63	3 (4.8)	NE		1.913	(0.318, 11.487)	0.478
	Irbesartan	59	2 (3.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2ET\_FSTM: Time to ESRD by subgroup  
 Full Analysis Set

Time to ESRD Subgroup	Treatment	N	nev (%)	Kaplan-Meier analysis		Cox regression		
				Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.889
<= 45 years	Sparsentan	96	7 (7.3)	NE		0.750	(0.271, 2.077)	0.580
	Irbesartan	99	8 (8.1)	NE				
> 45 years	Sparsentan	106	2 (1.9)	NE		0.641	(0.107, 3.841)	0.626
	Irbesartan	103	3 (2.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2ET\_FSTM: Time to ESRD by subgroup  
 Full Analysis Set

Time to ESRD Subgroup	Treatment	N	nev (%)	Kaplan-Meier analysis		Cox regression		
				Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	3 (33.3) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	3 (2.9)	NE		0.362	(0.096, 1.366)	0.134
	Irbesartan	109	8 (7.3)	NE				
> 40 years	Sparsentan	91	3 (3.3)	NE		0.899	(0.181, 4.460)	0.896
	Irbesartan	88	3 (3.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2ET\_FSTM: Time to ESRD by subgroup  
Full Analysis Set

Time to ESRD	Treatment	N	nev (%)	Kaplan-Meier analysis		Cox regression		
				Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	NE
North America	Sparsentan	35	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	46	4 (8.7)	NE				
Europe	Sparsentan	98	5 (5.1)	NE		0.818	(0.259, 2.580)	0.731
	Irbesartan	115	7 (6.1)	NE				
Asia Pacific	Sparsentan	69	4 (5.8) No events in 1 group	NE		NE		NE
	Irbesartan	41	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2ET\_FSTM: Time to ESRD by subgroup  
 Full Analysis Set

Time to ESRD Subgroup	Treatment	N	nev (%)	Kaplan-Meier analysis		Cox regression		
				Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.311
< 27 kg/m**2	Sparsentan	83	2 (2.4)	NE		0.426	(0.086, 2.114)	0.297
	Irbesartan	94	6 (6.4)	NE				
≥ 27 kg/m**2	Sparsentan	119	7 (5.9)	NE		1.093	(0.345, 3.466)	0.880
	Irbesartan	107	5 (4.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2ET\_FSTM: Time to ESRD by subgroup  
Full Analysis Set

Time to ESRD Subgroup	Treatment	N	nev (%)	Kaplan-Meier analysis		Cox regression		
				Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	NE
eGFR Low and UP High	Sparsentan	71	6 (8.5)	NE		0.540	(0.196, 1.488)	0.234
	Irbesartan	74	10 (13.5)	NE				
eGFR Low and UP Low	Sparsentan	55	2 (3.6) No events in 1 group	NE		NE		NE
	Irbesartan	55	0 (0.0)	NE				
eGFR High and UP High	Sparsentan	37	1 (2.7)	NE		0.980	(0.061, 15.698)	0.988
	Irbesartan	36	1 (2.8)	NE				
eGFR High and UP Low	Sparsentan	39	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	37	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2ET\_FSTM: Time to ESRD by subgroup  
 Full Analysis Set

Time to ESRD				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	NE
< 60 mL/min/1.73 m**2	Sparsentan	127	8 (6.3)	NE		0.696	(0.275, 1.766)	0.446
	Irbesartan	129	10 (7.8)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	1 (2.0)	NE		0.900	(0.056, 14.389)	0.941
	Irbesartan	48	1 (2.1)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2ET\_FSTM: Time to ESRD by subgroup  
Full Analysis Set

Time to ESRD				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	NE
< 45 mL/min/1.73 m**2	Sparsentan	82	8 (9.8)	NE		0.762	(0.293, 1.978)	0.576
	Irbesartan	80	9 (11.3)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	49	1 (2.0)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	1 (2.0)	NE		0.900	(0.056, 14.389)	0.941
	Irbesartan	48	1 (2.1)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024



Table PT2ET\_FSTM: Time to ESRD by subgroup  
 Full Analysis Set

Time to ESRD				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.706
<= 1.75 g/day	Sparsentan	98	1 (1.0)	NE		0.438	(0.039, 4.889)	0.502
	Irbesartan	93	2 (2.2)	NE				
> 1.75 g/day	Sparsentan	104	8 (7.7)	NE		0.792	(0.305, 2.056)	0.632
	Irbesartan	109	9 (8.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2ET\_FSTM: Time to ESRD by subgroup  
 Full Analysis Set

Time to ESRD	Subgroup	Treatment	N	nev (%)	Kaplan-Meier analysis		Cox regression		
					Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives		Sparsentan						Interaction test:	0.184
Yes		Sparsentan	90	6 (6.7)	NE		1.350	(0.381, 4.790)	0.642
		Irbesartan	88	4 (4.5)	NE				
No		Sparsentan	112	3 (2.7)	NE		0.372	(0.096, 1.444)	0.153
		Irbesartan	114	7 (6.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2ET\_FSTM: Time to ESRD by subgroup  
 Full Analysis Set

Time to ESRD				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.832
<= 5 years	Sparsentan	113	6 (5.3)	NE		0.722	(0.250, 2.084)	0.547
	Irbesartan	127	8 (6.3)	NE				
> 5 years	Sparsentan	89	3 (3.4)	NE		0.841	(0.170, 4.169)	0.832
	Irbesartan	75	3 (4.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2ET\_FSTM: Time to ESRD by subgroup  
 Full Analysis Set

Time to ESRD				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan						Interaction test:	0.496
Yes	Sparsentan	155	8 (5.2)	NE		0.839	(0.323, 2.175)	0.718
	Irbesartan	161	9 (5.6)	NE				
No	Sparsentan	47	1 (2.1)	NE		0.431	(0.038, 4.843)	0.495
	Irbesartan	41	2 (4.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeuff\_tte, created on: 28FEB2024

CSL Vifor  
Value Dossier Analysis: 021IGAN17001  
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Program Name: PF2aeff\_tte\_KM.sas  
Run Date: 26MAR2024:17:23:32

Figure PF2ET\_FSKM: Time to ESRD by subgroup  
Full Analysis Set

Not done. No significant subgroups.

ESRD = End-stage renal disease.  
Reference table: PT2ET\_FSTM

Table PT2D\_FSIM: Patients with death by subgroup  
Full Analysis Set

Subgroup analysis will not be performed as only one death occurred.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeфф, created on: 20FEB2024

Table PT2DT\_FSTM: Time to death by subgroup  
Full Analysis Set

Not done. Less than 10 patients with events in main analysis.

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF\_tte, created on: 28FEB2024

Figure PF2DT\_FSKM: Time to death by subgroup  
Full Analysis Set

Not done. Less than 10 patients with events in main analysis.

Reference table: PT2DT\_FSTM



Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.074
Male	Double-blind period	Sparsentan	139	40 (28.8)	0.857 [0.605, 1.215]	0.800 [0.482, 1.325]	-4.8 [-16.3, 6.7]	0.441
		Irbesartan	143	48 (33.6)				
Female	Double-blind period	Sparsentan	63	7 (11.1)	0.386 [0.172, 0.863]	0.309 [0.117, 0.812]	-17.7 [-33.3, -2.1]	0.022 *
		Irbesartan	59	17 (28.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 CKD = Chronic kidney disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.042 #
<= 45 years	Double-blind period	Sparsentan	96	23 (24.0)	1.078 [0.646, 1.800]	1.103 [0.566, 2.147]	1.7 [-11.1, 14.6]	0.865
		Irbesartan	99	22 (22.2)				
> 45 years	Double-blind period	Sparsentan	106	24 (22.6)	0.542 [0.357, 0.825]	0.408 [0.224, 0.744]	-19.1 [-32.5, -5.7]	0.005 *
		Irbesartan	103	43 (41.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 CKD = Chronic kidney disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.287
<= 18 years	Double-blind period	Sparsentan	9	6 (66.7)	1.667 [0.518, 5.363]	3.000 [0.312, 28.841]	26.7 [-41.7, 95.1]	0.580
		Irbesartan	5	2 (40.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	21 (20.6)	0.724 [0.446, 1.174]	0.652 [0.346, 1.231]	-7.9 [-20.3, 4.6]	0.204
		Irbesartan	109	31 (28.4)				
> 40 years	Double-blind period	Sparsentan	91	20 (22.0)	0.604 [0.376, 0.973]	0.493 [0.255, 0.953]	-14.4 [-28.7, -0.1]	0.048 *
		Irbesartan	88	32 (36.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 CKD = Chronic kidney disease.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.495
North America	Double-blind period	Sparsentan	35	6 (17.1)	0.526 [0.227, 1.216]	0.428 [0.146, 1.251]	-15.5 [-36.4, 5.5]	0.133
		Irbesartan	46	15 (32.6)				
Europe	Double-blind period	Sparsentan	98	23 (23.5)	0.692 [0.446, 1.074]	0.598 [0.326, 1.096]	-10.4 [-23.4, 2.6]	0.099
		Irbesartan	115	39 (33.9)				
Asia Pacific	Double-blind period	Sparsentan	69	18 (26.1)	0.972 [0.511, 1.849]	0.963 [0.401, 2.310]	-0.7 [-19.8, 18.3]	1.000
		Irbesartan	41	11 (26.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
CKD = Chronic kidney disease.  
Source Data: aeFF, created on: 20FEB2024

Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.796
< 27 kg/m**2	Double-blind period	Sparsentan	83	16 (19.3)	0.671 [0.390, 1.156]	0.593 [0.293, 1.199]	-9.4 [-23.1, 4.2]	0.163
		Irbesartan	94	27 (28.7)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	31 (26.1)	0.734 [0.494, 1.090]	0.640 [0.362, 1.131]	-9.5 [-22.4, 3.4]	0.148
		Irbesartan	107	38 (35.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 CKD = Chronic kidney disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.957
eGFR Low and UP High	Double-blind period	Sparsentan	71	31 (43.7)	0.702 [0.511, 0.966]	0.472 [0.243, 0.916]	-18.5 [-35.9, -1.1]	0.031 *
		Irbesartan	74	46 (62.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	14 (25.5)	0.824 [0.452, 1.501]	0.763 [0.332, 1.757]	-5.5 [-24.1, 13.1]	0.672
		Irbesartan	55	17 (30.9)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4)	0.973 [0.145, 6.541]	0.971 [0.129, 7.294]	-0.2 [-13.3, 13.0]	1.000
		Irbesartan	36	2 (5.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0)	0.950 + [0.019, 46.684]	0.949 + [0.018, 49.075]	0.0 [NE, NE]	NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
CKD = Chronic kidney disease.  
Source Data: aeFF, created on: 20FEB2024

Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.982
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	45 (35.4)	0.737 [0.549, 0.991]	0.593 [0.359, 0.979]	-12.6 [-25.4, 0.1]	0.044 *
		Irbesartan	129	62 (48.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.653 [0.114, 3.737]	0.638 [0.102, 4.000]	-2.2 [-13.0, 8.7]	0.678
		Irbesartan	48	3 (6.3)				
>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 CKD = Chronic kidney disease.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.997
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	40 (48.8)	0.723 [0.552, 0.946]	0.459 [0.242, 0.867]	-18.7 [-34.9, -2.6]	0.018 *
		Irbesartan	80	54 (67.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	5 (11.1)	0.681 [0.240, 1.928]	0.641 [0.193, 2.125]	-5.2 [-21.2, 10.8]	0.557
		Irbesartan	49	8 (16.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.653 [0.114, 3.737]	0.638 [0.102, 4.000]	-2.2 [-13.0, 8.7]	0.678
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
CKD = Chronic kidney disease.  
Source Data: aeFF, created on: 20FEB2024



Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.584
<= 1.75 g/day	Double-blind period	Sparsentan	98	16 (16.3)	0.844 [0.458, 1.554]	0.813 [0.387, 1.709]	-3.0 [-14.9, 8.9]	0.706
		Irbesartan	93	18 (19.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	31 (29.8)	0.691 [0.480, 0.996]	0.560 [0.318, 0.986]	-13.3 [-27.0, 0.4]	0.048 *
		Irbesartan	109	47 (43.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 CKD = Chronic kidney disease.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.638
Yes	Double-blind period	Sparsentan	90	29 (32.2)	0.766 [0.520, 1.129]	0.655 [0.355, 1.208]	-9.8 [-25.1, 5.4]	0.215
		Irbesartan	88	37 (42.0)				
No	Double-blind period	Sparsentan	112	18 (16.1)	0.654 [0.385, 1.113]	0.588 [0.304, 1.138]	-8.5 [-19.8, 2.8]	0.137
		Irbesartan	114	28 (24.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 CKD = Chronic kidney disease.  
 Source Data: aeff, created on: 20FEB2024

Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.769
<= 5 years	Double-blind period	Sparsentan	113	23 (20.4)	0.680 [0.433, 1.068]	0.599 [0.330, 1.085]	-9.6 [-21.3, 2.2]	0.103
		Irbesartan	127	38 (29.9)				
> 5 years	Double-blind period	Sparsentan	89	24 (27.0)	0.749 [0.475, 1.182]	0.656 [0.338, 1.276]	-9.0 [-24.5, 6.4]	0.238
		Irbesartan	75	27 (36.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 CKD = Chronic kidney disease.  
 Source Data: aeff, created on: 20FEB2024

Table PT2CKD\_FSIM: Patients entering CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Entering CKD stage 4 or 5								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.261
Yes	Double-blind period	Sparsentan	155	42 (27.1)	0.793 [0.567, 1.110]	0.716 [0.443, 1.159]	-7.1 [-17.8, 3.7]	0.182
		Irbesartan	161	55 (34.2)				
No	Double-blind period	Sparsentan	47	5 (10.6)	0.436 [0.162, 1.172]	0.369 [0.115, 1.189]	-13.8 [-31.9, 4.4]	0.098
		Irbesartan	41	10 (24.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 CKD = Chronic kidney disease.  
 Source Data: aeff, created on: 20FEB2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.400
Male	Sparsentan	139	40 (28.8)	NE		0.713	(0.468, 1.086)	0.115
	Irbesartan	143	48 (33.6)	NE				
Female	Sparsentan	63	7 (11.1)	NE		0.497	(0.206, 1.202)	0.121
	Irbesartan	59	17 (28.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 03JUN2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.060
<= 45 years	Sparsentan	96	23 (24.0)	NE		1.027	(0.572, 1.846)	0.928
	Irbesartan	99	22 (22.2)	NE				
> 45 years	Sparsentan	106	24 (22.6)	NE		0.500	(0.303, 0.824)	0.007 *
	Irbesartan	103	43 (41.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 03JUN2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	0.525
<= 18 years	Sparsentan	9	6 (66.7)	83.4	(3.9, NE)	1.293	(0.243, 6.880)	0.763
	Irbesartan	5	2 (40.0)	NE				
> 18 to 40 years	Sparsentan	102	21 (20.6)	NE		0.749	(0.430, 1.305)	0.308
	Irbesartan	109	31 (28.4)	NE				
> 40 years	Sparsentan	91	20 (22.0)	NE		0.493	(0.282, 0.863)	0.013 *
	Irbesartan	88	32 (36.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 03JUN2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.513
North America	Sparsentan	35	6 (17.1)	NE		0.434	(0.167, 1.126)	0.086
	Irbesartan	46	15 (32.6)	NE				
Europe	Sparsentan	98	23 (23.5)	NE		0.683	(0.408, 1.146)	0.149
	Irbesartan	115	39 (33.9)	NE				
Asia Pacific	Sparsentan	69	18 (26.1)	NE		0.967	(0.452, 2.069)	0.932
	Irbesartan	41	11 (26.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 03JUN2024



Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.948
< 27 kg/m**2	Sparsentan	83	16 (19.3)	NE		0.673	(0.362, 1.250)	0.210
	Irbesartan	94	27 (28.7)	NE				
≥ 27 kg/m**2	Sparsentan	119	31 (26.1)	NE		0.641	(0.398, 1.031)	0.066
	Irbesartan	107	38 (35.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF\_tte, created on: 03JUN2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	NE
eGFR Low and UP High	Sparsentan	71	31 (43.7)	NE		0.623	(0.394, 0.983)	0.042 *
	Irbesartan	74	46 (62.2)	84.0	(51.0, 109.0)			
eGFR Low and UP Low	Sparsentan	55	14 (25.5)	NE		0.763	(0.376, 1.548)	0.454
	Irbesartan	55	17 (30.9)	NE				
eGFR High and UP High	Sparsentan	37	2 (5.4)	NE		0.937	(0.132, 6.651)	0.948
	Irbesartan	36	2 (5.6)	NE				
eGFR High and UP Low	Sparsentan	39	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	37	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 03JUN2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: NE
< 60 mL/min/1.73 m**2	Sparsentan	127	45 (35.4)	NE		0.638	(0.434, 0.937)	0.022 *
	Irbesartan	129	62 (48.1)	110.1	(93.3, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	2 (4.1)	NE		0.735	(0.121, 4.459)	0.738
	Irbesartan	48	3 (6.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 03JUN2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: NE
< 45 mL/min/1.73 m**2	Sparsentan	82	40 (48.8)	109.1	(82.1, NE)	0.611	(0.405, 0.921)	0.019 *
	Irbesartan	80	54 (67.5)	70.1	(45.4, 106.3)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	5 (11.1)	NE		0.563	(0.184, 1.722)	0.314
	Irbesartan	49	8 (16.3)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	2 (4.1)	NE		0.735	(0.121, 4.459)	0.738
	Irbesartan	48	3 (6.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	25	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeuff\_tte, created on: 03JUN2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.382
<= 1.75 g/day	Sparsentan	98	16 (16.3)	NE		0.872	(0.443, 1.714)	0.691
	Irbesartan	93	18 (19.4)	NE				
> 1.75 g/day	Sparsentan	104	31 (29.8)	NE		0.601	(0.381, 0.949)	0.029 *
	Irbesartan	109	47 (43.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф\_tte, created on: 03JUN2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.658
Yes	Sparsentan	90	29 (32.2)	NE		0.706	(0.434, 1.148)	0.161
	Irbesartan	88	37 (42.0)	NE				
No	Sparsentan	112	18 (16.1)	NE		0.595	(0.328, 1.077)	0.086
	Irbesartan	114	28 (24.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF\_tte, created on: 03JUN2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.348
<= 5 years	Sparsentan	113	23 (20.4)	NE		0.557	(0.331, 0.936)	0.027 *
	Irbesartan	127	38 (29.9)	NE				
> 5 years	Sparsentan	89	24 (27.0)	NE		0.813	(0.469, 1.410)	0.461
	Irbesartan	75	27 (36.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 03JUN2024

Table PT2CKDT\_FSTM: Time to CKD stage 4 or 5 by subgroup  
 Full Analysis Set

Time to CKD stage 4 or 5				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.358
Yes	Sparsentan	155	42 (27.1)	NE		0.710	(0.475, 1.062)	0.095
	Irbesartan	161	55 (34.2)	NE				
No	Sparsentan	47	5 (10.6)	NE		0.333	(0.110, 1.006)	0.051
	Irbesartan	41	10 (24.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 03JUN2024



CSL Vifor  
Value Dossier Analysis: 021IGAN17001  
Cutoff Date (Final): 07/Sep/2023

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Program Name: PF2aeff\_tte\_KM.sas  
Run Date: 03JUN2024:17:12:05

Figure PF2CKDT\_FSKM: Time to CKD stage 4 or 5 by subgroup  
Full Analysis Set

Not done. No significant subgroups.

CKD = Chronic kidney disease.  
Reference table: PT2CKDT\_FSTM

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.951
Male	Double-blind period	Sparsentan	139	23 (16.5)	0.876 [0.529, 1.452]	0.852 [0.462, 1.572]	-2.3 [-11.9, 7.3]	0.642
		Irbesartan	143	27 (18.9)				
Female	Double-blind period	Sparsentan	63	10 (15.9)	0.851 [0.391, 1.856]	0.823 [0.321, 2.110]	-2.8 [-17.8, 12.3]	0.811
		Irbesartan	59	11 (18.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeфф, created on: 20FEB2024

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.056
<= 45 years	Double-blind period	Sparsentan	96	22 (22.9)	1.260 [0.723, 2.198]	1.338 [0.666, 2.689]	4.7 [-7.6, 17.1]	0.479
		Irbesartan	99	18 (18.2)				
> 45 years	Double-blind period	Sparsentan	106	11 (10.4)	0.534 [0.270, 1.059]	0.481 [0.218, 1.061]	-9.0 [-19.6, 1.5]	0.080
		Irbesartan	103	20 (19.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.139
<= 18 years	Double-blind period	Sparsentan	9	3 (33.3)	1.667 [0.230, 12.091]	2.000 [0.150, 26.734]	13.3 [-48.9, 75.6]	1.000
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	21 (20.6)	1.181 [0.676, 2.065]	1.228 [0.616, 2.447]	3.2 [-8.4, 14.7]	0.601
		Irbesartan	109	19 (17.4)				
> 40 years	Double-blind period	Sparsentan	91	9 (9.9)	0.484 [0.230, 1.018]	0.427 [0.180, 1.010]	-10.6 [-22.1, 1.0]	0.060
		Irbesartan	88	18 (20.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 20FEB2024

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.275
North America	Double-blind period	Sparsentan	35	9 (25.7)	0.623 [0.322, 1.205]	0.492 [0.189, 1.283]	-15.6 [-38.4, 7.2]	0.164
		Irbesartan	46	19 (41.3)				
Europe	Double-blind period	Sparsentan	98	16 (16.3)	1.341 [0.690, 2.607]	1.408 [0.649, 3.053]	4.2 [-6.2, 14.5]	0.432
		Irbesartan	115	14 (12.2)				
Asia Pacific	Double-blind period	Sparsentan	69	8 (11.6)	0.951 [0.333, 2.712]	0.944 [0.287, 3.107]	-0.6 [-15.1, 13.9]	1.000
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 20FEB2024

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.295
< 27 kg/m**2	Double-blind period	Sparsentan	83	14 (16.9)	1.133 [0.574, 2.234]	1.159 [0.517, 2.600]	2.0 [-10.0, 13.9]	0.837
		Irbesartan	94	14 (14.9)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	19 (16.0)	0.712 [0.414, 1.224]	0.657 [0.337, 1.282]	-6.5 [-17.6, 4.7]	0.238
		Irbesartan	107	24 (22.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.431
eGFR Low and UP High	Double-blind period	Sparsentan	71	10 (14.1)	0.579 [0.287, 1.167]	0.510 [0.217, 1.198]	-10.2 [-24.3, 3.8]	0.143
		Irbesartan	74	18 (24.3)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	8 (14.5)	1.143 [0.445, 2.935]	1.167 [0.392, 3.476]	1.8 [-12.8, 16.5]	1.000
		Irbesartan	55	7 (12.7)				
eGFR High and UP High	Double-blind period	Sparsentan	37	7 (18.9)	0.851 [0.345, 2.104]	0.817 [0.262, 2.547]	-3.3 [-24.6, 18.0]	0.778
		Irbesartan	36	8 (22.2)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	8 (20.5)	1.518 [0.546, 4.221]	1.652 [0.487, 5.604]	7.0 [-12.4, 26.4]	0.546
		Irbesartan	37	5 (13.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.154
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	19 (15.0)	0.742 [0.433, 1.272]	0.697 [0.364, 1.335]	-5.2 [-15.3, 4.9]	0.325
		Irbesartan	129	26 (20.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	0.653 [0.252, 1.694]	0.605 [0.197, 1.854]	-6.5 [-22.9, 9.9]	0.413
		Irbesartan	48	9 (18.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	8 (30.8)	2.564 [0.766, 8.581]	3.259 [0.753, 14.116]	18.8 [-7.0, 44.5]	0.173
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024



Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.259
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	10 (12.2)	0.887 [0.399, 1.972]	0.871 [0.348, 2.181]	-1.6 [-13.1, 10.0]	0.818
		Irbesartan	80	11 (13.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	9 (20.0)	0.653 [0.318, 1.343]	0.567 [0.219, 1.465]	-10.6 [-30.2, 8.9]	0.344
		Irbesartan	49	15 (30.6)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	0.653 [0.252, 1.694]	0.605 [0.197, 1.854]	-6.5 [-22.9, 9.9]	0.413
		Irbesartan	48	9 (18.8)				
>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	8 (30.8)	2.564 [0.766, 8.581]	3.259 [0.753, 14.116]	18.8 [-7.0, 44.5]	0.173
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 20FEB2024

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.174
<= 1.75 g/day	Double-blind period	Sparsentan	98	16 (16.3)	1.265 [0.633, 2.529]	1.317 [0.586, 2.958]	3.4 [-7.6, 14.5]	0.545
		Irbesartan	93	12 (12.9)				
> 1.75 g/day	Double-blind period	Sparsentan	104	17 (16.3)	0.685 [0.396, 1.187]	0.624 [0.316, 1.233]	-7.5 [-19.1, 4.1]	0.232
		Irbesartan	109	26 (23.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.382
Yes	Double-blind period	Sparsentan	90	12 (13.3)	0.690 [0.350, 1.360]	0.643 [0.287, 1.438]	-6.0 [-17.9, 6.0]	0.315
		Irbesartan	88	17 (19.3)				
No	Double-blind period	Sparsentan	112	21 (18.8)	1.018 [0.590, 1.757]	1.022 [0.523, 1.998]	0.3 [-10.7, 11.4]	1.000
		Irbesartan	114	21 (18.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.046 #
<= 5 years	Double-blind period	Sparsentan	113	17 (15.0)	0.637 [0.372, 1.091]	0.573 [0.296, 1.106]	-8.6 [-19.3, 2.2]	0.105
		Irbesartan	127	30 (23.6)				
> 5 years	Double-blind period	Sparsentan	89	16 (18.0)	1.685 [0.764, 3.718]	1.836 [0.738, 4.566]	7.3 [-4.5, 19.1]	0.267
		Irbesartan	75	8 (10.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2MIS\_FSIM: Patients with systemic immunosuppressive medication by subgroup  
 Full Analysis Set

<u>Systemic immunosuppressive medication</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.994
Yes	Double-blind period	Sparsentan	155	27 (17.4)	0.876 [0.552, 1.391]	0.850 [0.482, 1.500]	-2.5 [-11.7, 6.8]	0.665
		Irbesartan	161	32 (19.9)				
No	Double-blind period	Sparsentan	47	6 (12.8)	0.872 [0.305, 2.496]	0.854 [0.252, 2.886]	-1.9 [-18.6, 14.8]	1.000
		Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.973
Male	Sparsentan	139	23 (16.5)	NE		0.799	(0.457, 1.396)	0.431
	Irbesartan	143	27 (18.9)	NE				
Female	Sparsentan	63	10 (15.9)	NE		0.828	(0.348, 1.967)	0.669
	Irbesartan	59	11 (18.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.070
<= 45 years	Sparsentan	96	22 (22.9)	NE		1.171	(0.627, 2.188)	0.620
	Irbesartan	99	18 (18.2)	NE				
> 45 years	Sparsentan	106	11 (10.4)	NE		0.496	(0.237, 1.036)	0.062
	Irbesartan	103	20 (19.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							
							Interaction test:	0.164
<= 18 years	Sparsentan	9	3 (33.3)	NE		1.493	(0.152, 14.626)	0.731
	Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Sparsentan	102	21 (20.6)	NE		1.130	(0.604, 2.113)	0.702
	Irbesartan	109	19 (17.4)	NE				
> 40 years	Sparsentan	91	9 (9.9)	NE		0.449	(0.201, 1.003)	0.051
	Irbesartan	88	18 (20.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024



Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
Full Analysis Set

Time to systemic immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.193
North America	Sparsentan	35	9 (25.7)	NE		0.520	(0.234, 1.154)	0.108
	Irbesartan	46	19 (41.3)	NE				
Europe	Sparsentan	98	16 (16.3)	NE		1.359	(0.663, 2.790)	0.402
	Irbesartan	115	14 (12.2)	NE				
Asia Pacific	Sparsentan	69	8 (11.6)	NE		0.883	(0.285, 2.741)	0.830
	Irbesartan	41	5 (12.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.345
< 27 kg/m**2	Sparsentan	83	14 (16.9)	NE		1.010	(0.480, 2.126)	0.978
	Irbesartan	94	14 (14.9)	NE				
≥ 27 kg/m**2	Sparsentan	119	19 (16.0)	NE		0.644	(0.352, 1.178)	0.154
	Irbesartan	107	24 (22.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan							Interaction test: 0.298
eGFR Low and UP High	Sparsentan	71	10 (14.1)	NE		0.481	(0.222, 1.044)	0.064
	Irbesartan	74	18 (24.3)	NE				
eGFR Low and UP Low	Sparsentan	55	8 (14.5)	NE		1.178	(0.427, 3.248)	0.752
	Irbesartan	55	7 (12.7)	NE				
eGFR High and UP High	Sparsentan	37	7 (18.9)	NE		0.812	(0.294, 2.239)	0.687
	Irbesartan	36	8 (22.2)	NE				
eGFR High and UP Low	Sparsentan	39	8 (20.5)	NE		1.511	(0.494, 4.621)	0.469
	Irbesartan	37	5 (13.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.135
< 60 mL/min/1.73 m**2	Sparsentan	127	19 (15.0)	NE		0.700	(0.385, 1.271)	0.241
	Irbesartan	129	26 (20.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	6 (12.2)	NE		0.642	(0.227, 1.816)	0.403
	Irbesartan	48	9 (18.8)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	8 (30.8)	NE		2.560	(0.679, 9.657)	0.165
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.237
< 45 mL/min/1.73 m**2	Sparsentan	82	10 (12.2)	NE		0.809	(0.343, 1.906)	0.628
	Irbesartan	80	11 (13.8)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	9 (20.0)	NE		0.608	(0.263, 1.406)	0.245
	Irbesartan	49	15 (30.6)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	6 (12.2)	NE		0.642	(0.227, 1.816)	0.403
	Irbesartan	48	9 (18.8)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	8 (30.8)	NE		2.560	(0.679, 9.657)	0.165
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.117
<= 1.75 g/day	Sparsentan	98	16 (16.3)	NE		1.274	(0.603, 2.694)	0.526
	Irbesartan	93	12 (12.9)	NE				
> 1.75 g/day	Sparsentan	104	17 (16.3)	NE		0.595	(0.322, 1.099)	0.097
	Irbesartan	109	26 (23.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.448
Yes	Sparsentan	90	12 (13.3)	NE		0.645	(0.308, 1.354)	0.246
	Irbesartan	88	17 (19.3)	NE				
No	Sparsentan	112	21 (18.8)	NE		0.934	(0.510, 1.712)	0.826
	Irbesartan	114	21 (18.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.032 #
<= 5 years	Sparsentan	113	17 (15.0)	NE		0.549	(0.302, 0.997)	0.049 *
	Irbesartan	127	30 (23.6)	NE				
> 5 years	Sparsentan	89	16 (18.0)	NE		1.606	(0.684, 3.768)	0.276
	Irbesartan	75	8 (10.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeuff\_tte, created on: 28FEB2024



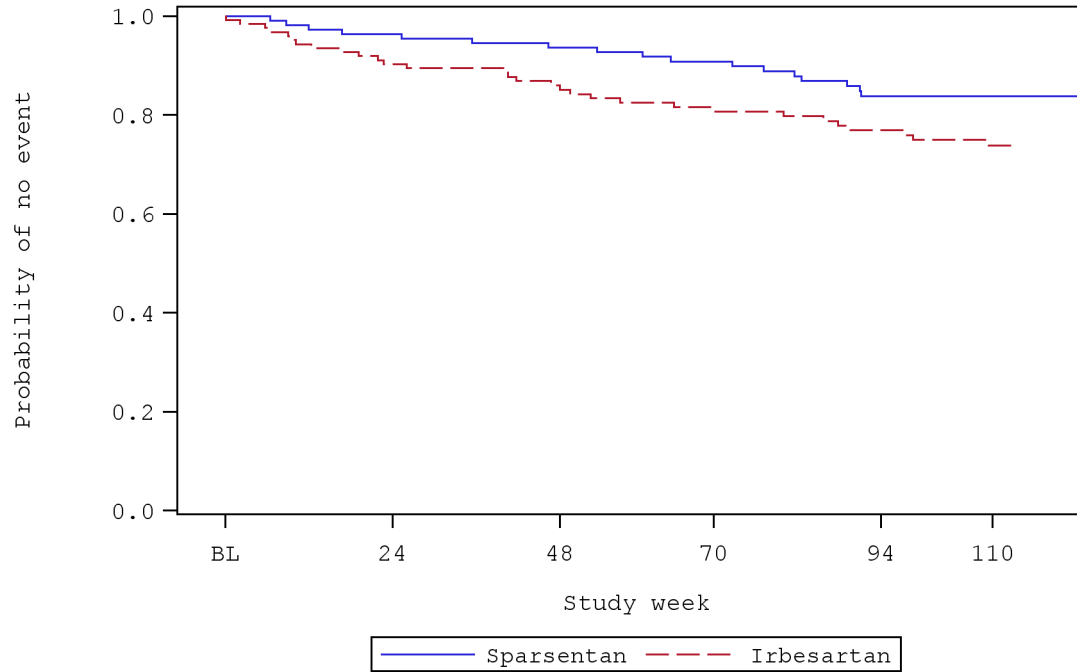
Table PT2MIST\_FSTM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

Time to systemic immunosuppressive medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension		Sparsentan				Interaction test:		0.858
Yes	Sparsentan	155	27 (17.4)	NE		0.841	(0.504, 1.405)	0.508
	Irbesartan	161	32 (19.9)	NE				
No	Sparsentan	47	6 (12.8)	NE		0.672	(0.212, 2.131)	0.500
	Irbesartan	41	6 (14.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Figure PF2MIST\_FSKM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set

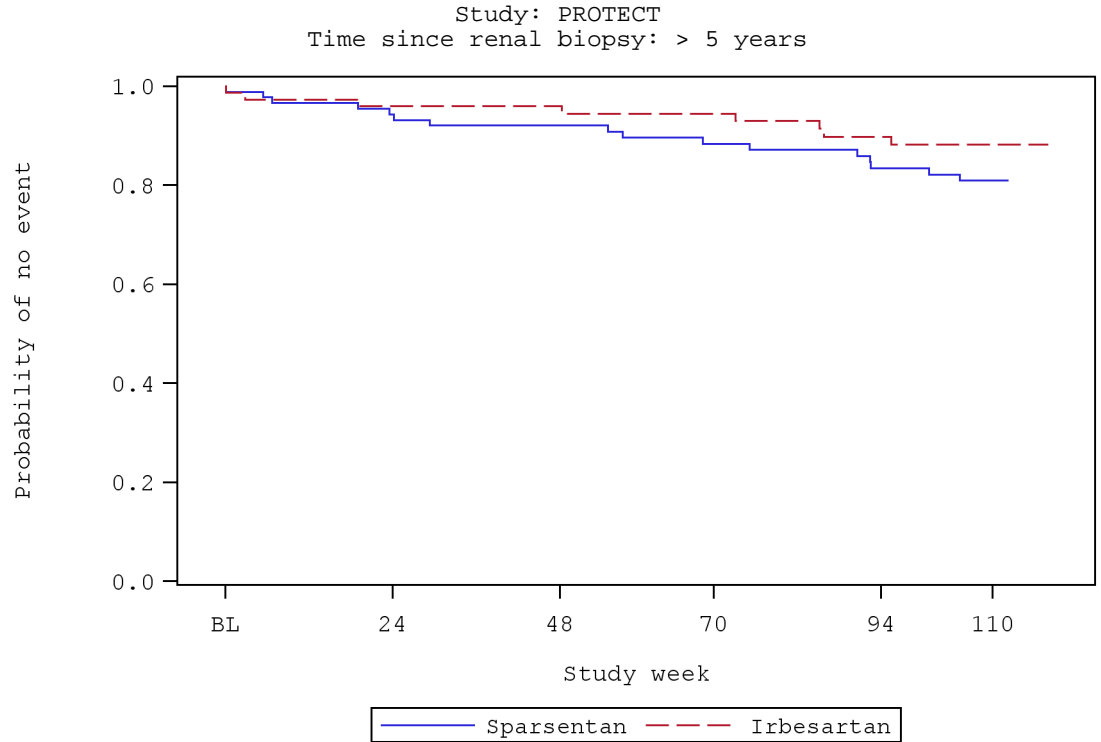
Study: PROTECT  
 Time since renal biopsy: <= 5 years



Sparsentan	113	106	101	96	83	54
Irbesartan	127	111	98	90	81	56

Reference table: PT2MIST\_FSTM

Figure PF2MIST\_FSKM: Time to systemic immunosuppressive medication by subgroup  
 Full Analysis Set



Sparsentan	89	83	77	72	67	50
Irbesartan	75	70	65	62	57	42

Reference table: PT2MIST\_FSTM

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.850
Male	Double-blind period	Sparsentan	139	5 (3.6)	0.396 [0.145, 1.081]	0.373 [0.129, 1.076]	-5.5 [-11.8, 0.9]	0.086
		Irbesartan	143	13 (9.1)				
Female	Double-blind period	Sparsentan	63	1 (1.6)	0.312 [0.033, 2.918]	0.301 [0.030, 2.979]	-3.5 [-11.5, 4.5]	0.353
		Irbesartan	59	3 (5.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.610
<= 45 years	Double-blind period	Sparsentan	96	4 (4.2)	0.458 [0.146, 1.439]	0.435 [0.129, 1.463]	-4.9 [-12.9, 3.0]	0.251
		Irbesartan	99	9 (9.1)				
> 45 years	Double-blind period	Sparsentan	106	2 (1.9)	0.278 [0.059, 1.305]	0.264 [0.053, 1.301]	-4.9 [-11.4, 1.6]	0.098
		Irbesartan	103	7 (6.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication  
 (renal indication)

Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.112
<= 18 years	Double-blind period	Sparsentan	9	3 (33.3)	4.200 + [0.259, 68.038]	5.923 + [0.248, 141.482]	33.3 [-13.0, 79.7]	0.258
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	1 (1.0)	0.107 [0.014, 0.820]	0.098 [0.012, 0.780]	-8.2 [-14.9, -1.5]	0.010 *
		Irbesartan	109	10 (9.2)				
> 40 years	Double-blind period	Sparsentan	91	2 (2.2)	0.322 [0.067, 1.554]	0.307 [0.060, 1.565]	-4.6 [-11.8, 2.6]	0.164
		Irbesartan	88	6 (6.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.276
North America	Double-blind period	Sparsentan	35	3 (8.6)	0.986 [0.236, 4.123]	0.984 [0.206, 4.713]	-0.1 [-15.0, 14.7]	1.000
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	1 (1.0)	0.130 [0.017, 1.011]	0.121 [0.015, 0.976]	-6.8 [-13.0, -0.6]	0.022 *
		Irbesartan	115	9 (7.8)				
Asia Pacific	Double-blind period	Sparsentan	69	2 (2.9)	0.396 [0.069, 2.273]	0.378 [0.060, 2.364]	-4.4 [-15.3, 6.4]	0.359
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.805
< 27 kg/m**2	Double-blind period	Sparsentan	83	3 (3.6)	0.425 [0.116, 1.548]	0.403 [0.103, 1.573]	-4.9 [-13.0, 3.2]	0.222
		Irbesartan	94	8 (8.5)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	3 (2.5)	0.337 [0.092, 1.238]	0.320 [0.083, 1.239]	-5.0 [-11.6, 1.7]	0.121
		Irbesartan	107	8 (7.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024



Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Systemic immunosuppressive renal medication  
(renal indication)

Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.273
eGFR Low and UP High	Double-blind period	Sparsentan	71	2 (2.8)	0.174 [0.040, 0.749]	0.150 [0.032, 0.696]	-13.4 [-24.0, -2.8]	0.009 *
		Irbesartan	74	12 (16.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	0 (0.0)	0.333 + [0.014, 8.008]	0.327 + [0.013, 8.212]	-1.8 [-7.2, 3.5]	1.000
		Irbesartan	55	1 (1.8)				
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	1.297 [0.312, 5.393]	1.333 [0.277, 6.427]	2.5 [-13.7, 18.7]	1.000
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0)	0.950 + [0.019, 46.684]	0.949 + [0.018, 49.075]	0.0 [NE, NE]	NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication  
 (renal indication)

Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.243
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	2 (1.6)	0.185 [0.042, 0.817]	0.172 [0.037, 0.791]	-7.0 [-13.0, -0.9]	0.019 *
		Irbesartan	129	11 (8.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	0.490 [0.094, 2.550]	0.468 [0.082, 2.684]	-4.3 [-15.9, 7.4]	0.436
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)	1.923 [0.186, 19.901]	2.000 [0.170, 23.556]	3.7 [-13.0, 20.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	1 (1.2)				NE
		Irbesartan	80	6 (7.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	1 (2.2)				NE
		Irbesartan	49	5 (10.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)				NE
		Irbesartan	48	4 (8.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication  
 (renal indication)

Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.667
<= 1.75 g/day	Double-blind period	Sparsentan	98	0 (0.0)	0.949 + [0.019, 47.367]	0.949 + [0.019, 48.329]	0.0 [NE, NE]	NE
		Irbesartan	93	0 (0.0)				
> 1.75 g/day	Double-blind period	Sparsentan	104	6 (5.8)	0.393 [0.160, 0.966]	0.356 [0.134, 0.948]	-8.9 [-17.9, 0.0]	0.042 *
		Irbesartan	109	16 (14.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.505
Yes	Double-blind period	Sparsentan	90	4 (4.4)	0.489 [0.153, 1.565]	0.465 [0.135, 1.604]	-4.6 [-13.1, 3.8]	0.246
		Irbesartan	88	8 (9.1)				
No	Double-blind period	Sparsentan	112	2 (1.8)	0.254 [0.055, 1.172]	0.241 [0.050, 1.161]	-5.2 [-11.4, 0.9]	0.102
		Irbesartan	114	8 (7.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 20FEB2024

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.601
<= 5 years	Double-blind period	Sparsentan	113	3 (2.7)	0.307 [0.088, 1.071]	0.288 [0.078, 1.058]	-6.0 [-12.6, 0.5]	0.056
		Irbesartan	127	11 (8.7)				
> 5 years	Double-blind period	Sparsentan	89	3 (3.4)	0.506 [0.125, 2.046]	0.488 [0.113, 2.115]	-3.3 [-11.3, 4.7]	0.471
		Irbesartan	75	5 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2MIK\_FSIM: Patients with systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Systemic immunosuppressive renal medication (renal indication)								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.799
Yes	Double-blind period	Sparsentan	155	5 (3.2)	0.400 [0.146, 1.094]	0.379 [0.132, 1.091]	-4.8 [-10.5, 0.8]	0.088
		Irbesartan	161	13 (8.1)				
No	Double-blind period	Sparsentan	47	1 (2.1)	0.291 [0.031, 2.688]	0.275 [0.028, 2.756]	-5.2 [-16.4, 6.1]	0.335
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.978
Male	Sparsentan	139	5 (3.6)	NE		0.329	(0.117, 0.926)	0.035 *
	Irbesartan	143	13 (9.1)	NE				
Female	Sparsentan	63	1 (1.6)	NE		0.376	(0.038, 3.690)	0.401
	Irbesartan	59	3 (5.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024



Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.677
<= 45 years	Sparsentan	96	4 (4.2)	NE		0.393	(0.121, 1.281)	0.121
	Irbesartan	99	9 (9.1)	NE				
> 45 years	Sparsentan	106	2 (1.9)	NE		0.253	(0.052, 1.224)	0.087
	Irbesartan	103	7 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	3 (33.3) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	1 (1.0)	NE		0.093	(0.012, 0.731)	0.024 *
	Irbesartan	109	10 (9.2)	NE				
> 40 years	Sparsentan	91	2 (2.2)	NE		0.291	(0.059, 1.445)	0.131
	Irbesartan	88	6 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.215
North America	Sparsentan	35	3 (8.6)	NE		0.890	(0.198, 3.992)	0.879
	Irbesartan	46	4 (8.7)	NE				
Europe	Sparsentan	98	1 (1.0)	NE		0.115	(0.015, 0.911)	0.041 *
	Irbesartan	115	9 (7.8)	NE				
Asia Pacific	Sparsentan	69	2 (2.9)	NE		0.341	(0.055, 2.124)	0.249
	Irbesartan	41	3 (7.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.889
< 27 kg/m**2	Sparsentan	83	3 (3.6)	NE		0.330	(0.087, 1.260)	0.105
	Irbesartan	94	8 (8.5)	NE				
≥ 27 kg/m**2	Sparsentan	119	3 (2.5)	NE		0.281	(0.074, 1.066)	0.062
	Irbesartan	107	8 (7.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	NE
eGFR Low and UP High	Sparsentan	71	2 (2.8)	NE		0.140	(0.031, 0.628)	0.010 *
	Irbesartan	74	12 (16.2)	NE				
eGFR Low and UP Low	Sparsentan	55	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	55	1 (1.8)	NE				
eGFR High and UP High	Sparsentan	37	4 (10.8)	NE		1.155	(0.259, 5.164)	0.850
	Irbesartan	36	3 (8.3)	NE				
eGFR High and UP Low	Sparsentan	39	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	37	0 (0.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.155
< 60 mL/min/1.73 m**2	Sparsentan	127	2 (1.6)	NE		0.152	(0.034, 0.685)	0.014 *
	Irbesartan	129	11 (8.5)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	2 (4.1)	NE		0.539	(0.096, 3.036)	0.483
	Irbesartan	48	4 (8.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	2 (7.7)	NE		1.687	(0.152, 18.691)	0.670
	Irbesartan	25	1 (4.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	NE
< 45 mL/min/1.73 m**2	Sparsentan	82	1 (1.2) all n<10	NE		NE		NE
	Irbesartan	80	6 (7.5)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	1 (2.2) all n<10	NE		NE		NE
	Irbesartan	49	5 (10.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	2 (4.1) all n<10	NE		NE		NE
	Irbesartan	48	4 (8.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	2 (7.7) all n<10	NE		NE		NE
	Irbesartan	25	1 (4.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: NE
<= 1.75 g/day	Sparsentan	98	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	93	0 (0.0)	NE				
> 1.75 g/day	Sparsentan	104	6 (5.8)	NE		0.321	(0.125, 0.821)	0.018 *
	Irbesartan	109	16 (14.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024



Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.585
Yes	Sparsentan	90	4 (4.4)	NE		0.385	(0.116, 1.281)	0.120
	Irbesartan	88	8 (9.1)	NE				
No	Sparsentan	112	2 (1.8)	NE		0.246	(0.052, 1.159)	0.076
	Irbesartan	114	8 (7.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.435
<= 5 years	Sparsentan	113	3 (2.7)	NE		0.238	(0.066, 0.854)	0.028 *
	Irbesartan	127	11 (8.7)	NE				
> 5 years	Sparsentan	89	3 (3.4)	NE		0.511	(0.122, 2.141)	0.358
	Irbesartan	75	5 (6.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.

95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).

A Cox proportional model was applied with factors treatment, randomization strata.

\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF\_tte, created on: 28FEB2024

Table PT2MIKT\_FSTM: Time to systemic immunosuppressive renal medication by subgroup  
 Full Analysis Set

Time to systemic Immunosuppressive renal medication				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.639
Yes	Sparsentan	155	5 (3.2)	NE		0.365	(0.130, 1.024)	0.056
	Irbesartan	161	13 (8.1)	NE				
No	Sparsentan	47	1 (2.1)	NE		0.188	(0.019, 1.826)	0.149
	Irbesartan	41	3 (7.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF\_tte, created on: 28FEB2024

Figure PF2MIKT\_FSKM: Time to systemic immunosuppressive renal medication by subgroup  
Full Analysis Set

Not done. No significant subgroups.

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.713
Male	Double-blind period	Sparsentan	139	28 (20.1)	1.108 [0.686, 1.791]	1.135 [0.627, 2.055]	2.0 [-7.9, 11.9]	0.762
		Irbesartan	143	26 (18.2)				
Female	Double-blind period	Sparsentan	63	11 (17.5)	0.937 [0.439, 1.996]	0.923 [0.367, 2.324]	-1.2 [-16.5, 14.1]	1.000
		Irbesartan	59	11 (18.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.365
<= 45 years	Double-blind period	Sparsentan	96	21 (21.9)	1.274 [0.717, 2.263]	1.351 [0.663, 2.753]	4.7 [-7.4, 16.8]	0.471
		Irbesartan	99	17 (17.2)				
> 45 years	Double-blind period	Sparsentan	106	18 (17.0)	0.875 [0.492, 1.556]	0.849 [0.420, 1.716]	-2.4 [-13.9, 9.0]	0.721
		Irbesartan	103	20 (19.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.330
<= 18 years	Double-blind period	Sparsentan	9	5 (55.6)	2.778 [0.438, 17.629]	5.000 [0.388, 64.387]	35.6 [-27.8, 98.9]	0.301
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	16 (15.7)	1.221 [0.628, 2.374]	1.262 [0.582, 2.738]	2.8 [-7.6, 13.2]	0.562
		Irbesartan	109	14 (12.8)				
> 40 years	Double-blind period	Sparsentan	91	18 (19.8)	0.791 [0.457, 1.371]	0.740 [0.365, 1.499]	-5.2 [-18.5, 8.1]	0.474
		Irbesartan	88	22 (25.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.805
North America	Double-blind period	Sparsentan	35	6 (17.1)	0.789 [0.317, 1.962]	0.745 [0.242, 2.292]	-4.6 [-24.4, 15.2]	0.779
		Irbesartan	46	10 (21.7)				
Europe	Double-blind period	Sparsentan	98	18 (18.4)	1.112 [0.619, 1.997]	1.137 [0.559, 2.312]	1.8 [-9.3, 13.0]	0.721
		Irbesartan	115	19 (16.5)				
Asia Pacific	Double-blind period	Sparsentan	69	15 (21.7)	1.114 [0.518, 2.397]	1.146 [0.438, 2.996]	2.2 [-15.3, 19.7]	1.000
		Irbesartan	41	8 (19.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024



Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.851
< 27 kg/m**2	Double-blind period	Sparsentan	83	16 (19.3)	1.007 [0.550, 1.844]	1.008 [0.477, 2.133]	0.1 [-12.6, 12.9]	1.000
		Irbesartan	94	18 (19.1)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	23 (19.3)	1.088 [0.629, 1.884]	1.110 [0.566, 2.175]	1.6 [-9.5, 12.6]	0.864
		Irbesartan	107	19 (17.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.150
eGFR Low and UP High	Double-blind period	Sparsentan	71	13 (18.3)	0.753 [0.399, 1.420]	0.697 [0.313, 1.556]	-6.0 [-20.7, 8.7]	0.422
		Irbesartan	74	18 (24.3)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	9 (16.4)	1.000 [0.430, 2.328]	1.000 [0.364, 2.746]	0.0 [-15.6, 15.6]	1.000
		Irbesartan	55	9 (16.4)				
eGFR High and UP High	Double-blind period	Sparsentan	37	11 (29.7)	3.568 [1.084, 11.742]	4.654 [1.175, 18.428]	21.4 [1.4, 41.4]	0.035 *
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	6 (15.4)	0.813 [0.301, 2.196]	0.779 [0.235, 2.580]	-3.5 [-23.1, 16.1]	0.766
		Irbesartan	37	7 (18.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.244
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	24 (18.9)	0.841 [0.519, 1.361]	0.803 [0.438, 1.474]	-3.6 [-14.3, 7.1]	0.538
		Irbesartan	129	29 (22.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	12 (24.5)	1.679 [0.723, 3.901]	1.900 [0.676, 5.335]	9.9 [-7.8, 27.6]	0.307
		Irbesartan	48	7 (14.6)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	3 (11.5)	2.885 [0.321, 25.919]	3.130 [0.303, 32.314]	7.5 [-10.9, 25.9]	0.610
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
Full Analysis Set

Hospitalizations								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.329
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	17 (20.7)	0.976 [0.537, 1.773]	0.969 [0.455, 2.065]	-0.5 [-14.3, 13.3]	1.000
		Irbesartan	80	17 (21.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	7 (15.6)	0.635 [0.274, 1.471]	0.568 [0.202, 1.601]	-8.9 [-27.1, 9.2]	0.315
		Irbesartan	49	12 (24.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	12 (24.5)	1.679 [0.723, 3.901]	1.900 [0.676, 5.335]	9.9 [-7.8, 27.6]	0.307
		Irbesartan	48	7 (14.6)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	3 (11.5)	2.885 [0.321, 25.919]	3.130 [0.303, 32.314]	7.5 [-10.9, 25.9]	0.610
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.742
<= 1.75 g/day	Double-blind period	Sparsentan	98	17 (17.3)	1.152 [0.603, 2.203]	1.184 [0.547, 2.564]	2.3 [-9.2, 13.8]	0.699
		Irbesartan	93	14 (15.1)				
> 1.75 g/day	Double-blind period	Sparsentan	104	22 (21.2)	1.003 [0.597, 1.685]	1.003 [0.519, 1.937]	0.1 [-11.9, 12.0]	1.000
		Irbesartan	109	23 (21.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeфф, created on: 20FEB2024

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.928
Yes	Double-blind period	Sparsentan	90	19 (21.1)	1.032 [0.581, 1.832]	1.041 [0.504, 2.147]	0.7 [-12.4, 13.7]	1.000
		Irbesartan	88	18 (20.5)				
No	Double-blind period	Sparsentan	112	20 (17.9)	1.071 [0.605, 1.897]	1.087 [0.545, 2.168]	1.2 [-9.5, 11.9]	0.861
		Irbesartan	114	19 (16.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.001 #
<= 5 years	Double-blind period	Sparsentan	113	18 (15.9)	0.632 [0.376, 1.062]	0.563 [0.296, 1.071]	-9.3 [-20.2, 1.7]	0.082
		Irbesartan	127	32 (25.2)				
> 5 years	Double-blind period	Sparsentan	89	21 (23.6)	3.539 [1.403, 8.932]	4.324 [1.542, 12.120]	16.9 [5.2, 28.6]	0.005 *
		Irbesartan	75	5 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HOS\_FSIM: Patients with hospitalizations by subgroup  
 Full Analysis Set

<u>Hospitalizations</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.266
Yes	Double-blind period	Sparsentan	155	29 (18.7)	0.941 [0.599, 1.479]	0.928 [0.530, 1.623]	-1.2 [-10.5, 8.2]	0.887
		Irbesartan	161	32 (19.9)				
No	Double-blind period	Sparsentan	47	10 (21.3)	1.745 [0.649, 4.688]	1.946 [0.605, 6.254]	9.1 [-8.6, 26.8]	0.395
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024



Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
 Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.909
Male	Double-blind period	Sparsentan	139	9 (6.5)	1.029 [0.421, 2.515]	1.031 [0.397, 2.678]	0.2 [-6.2, 6.6]	1.000
		Irbesartan	143	9 (6.3)				
Female	Double-blind period	Sparsentan	63	4 (6.3)	0.937 [0.245, 3.575]	0.932 [0.222, 3.910]	-0.4 [-10.9, 10.0]	1.000
		Irbesartan	59	4 (6.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
 Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.701
<= 45 years	Double-blind period	Sparsentan	96	9 (9.4)	0.928 [0.394, 2.184]	0.921 [0.357, 2.375]	-0.7 [-10.1, 8.6]	1.000
		Irbesartan	99	10 (10.1)				
> 45 years	Double-blind period	Sparsentan	106	4 (3.8)	1.296 [0.297, 5.647]	1.307 [0.285, 5.990]	0.9 [-5.0, 6.7]	1.000
		Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.557
<= 18 years	Double-blind period	Sparsentan	9	4 (44.4)	2.222 [0.333, 14.845]	3.200 [0.248, 41.208]	24.4 [-38.9, 87.8]	0.580
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	5 (4.9)	0.668 [0.226, 1.975]	0.651 [0.206, 2.058]	-2.4 [-9.8, 5.0]	0.572
		Irbesartan	109	8 (7.3)				
> 40 years	Double-blind period	Sparsentan	91	4 (4.4)	0.967 [0.250, 3.747]	0.966 [0.234, 3.986]	-0.1 [-7.3, 7.0]	1.000
		Irbesartan	88	4 (4.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
 Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.649
North America	Double-blind period	Sparsentan	35	2 (5.7)	2.629 [0.248, 27.835]	2.727 [0.237, 31.357]	3.5 [-7.7, 14.8]	0.575
		Irbesartan	46	1 (2.2)				
Europe	Double-blind period	Sparsentan	98	6 (6.1)	0.782 [0.289, 2.121]	0.768 [0.263, 2.240]	-1.7 [-9.5, 6.1]	0.790
		Irbesartan	115	9 (7.8)				
Asia Pacific	Double-blind period	Sparsentan	69	5 (7.2)	0.990 [0.250, 3.929]	0.990 [0.224, 4.376]	-0.1 [-12.1, 11.9]	1.000
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeфф, created on: 20FEB2024

Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.089
< 27 kg/m**2	Double-blind period	Sparsentan	83	5 (6.0)	0.566 [0.202, 1.590]	0.538 [0.176, 1.645]	-4.6 [-13.8, 4.6]	0.295
		Irbesartan	94	10 (10.6)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	8 (6.7)	2.398 [0.653, 8.807]	2.498 [0.645, 9.672]	3.9 [-2.4, 10.3]	0.222
		Irbesartan	107	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
 Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.154
eGFR Low and UP High	Double-blind period	Sparsentan	71	4 (5.6)	0.417 [0.137, 1.269]	0.382 [0.114, 1.280]	-7.9 [-18.7, 3.0]	0.159
		Irbesartan	74	10 (13.5)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)	2.000 [0.187, 21.420]	2.038 [0.179, 23.151]	1.8 [-6.1, 9.7]	1.000
		Irbesartan	55	1 (1.8)				
eGFR High and UP High	Double-blind period	Sparsentan	37	7 (18.9)	3.405 [0.758, 15.308]	3.967 [0.765, 20.580]	13.4 [-4.0, 30.8]	0.152
		Irbesartan	36	2 (5.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0)	0.950 + [0.019, 46.684]	0.949 + [0.018, 49.075]	0.0 [NE, NE]	NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.156
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	6 (4.7)	0.554 [0.211, 1.453]	0.532 [0.191, 1.485]	-3.8 [-10.7, 3.0]	0.316
		Irbesartan	129	11 (8.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	2.939 [0.624, 13.846]	3.209 [0.614, 16.768]	8.1 [-4.8, 20.9]	0.268
		Irbesartan	48	2 (4.2)				
>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	2.889 + [0.123, 67.752]	3.000 + [0.117, 77.165]	3.8 [-7.5, 15.2]	1.000
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
 Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.281
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	5 (6.1)	0.542 [0.190, 1.547]	0.512 [0.164, 1.601]	-5.2 [-15.0, 4.7]	0.275
		Irbesartan	80	9 (11.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	1 (2.2)	0.544 [0.051, 5.801]	0.534 [0.047, 6.100]	-1.9 [-11.0, 7.3]	1.000
		Irbesartan	49	2 (4.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	2.939 [0.624, 13.846]	3.209 [0.614, 16.768]	8.1 [-4.8, 20.9]	0.268
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	2.889 + [0.123, 67.752]	3.000 + [0.117, 77.165]	3.8 [-7.5, 15.2]	1.000
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024



Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
 Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.184
<= 1.75 g/day	Double-blind period	Sparsentan	98	4 (4.1)	3.796 [0.432, 33.341]	3.915 [0.429, 35.690]	3.0 [-2.5, 8.5]	0.369
		Irbesartan	93	1 (1.1)				
> 1.75 g/day	Double-blind period	Sparsentan	104	9 (8.7)	0.786 [0.346, 1.787]	0.766 [0.308, 1.901]	-2.4 [-11.3, 6.6]	0.649
		Irbesartan	109	12 (11.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.449
Yes	Double-blind period	Sparsentan	90	8 (8.9)	1.304 [0.472, 3.604]	1.333 [0.443, 4.013]	2.1 [-6.9, 11.1]	0.782
		Irbesartan	88	6 (6.8)				
No	Double-blind period	Sparsentan	112	5 (4.5)	0.727 [0.238, 2.223]	0.714 [0.220, 2.321]	-1.7 [-8.4, 5.0]	0.768
		Irbesartan	114	7 (6.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.

N = number of patients in analysis set. n = number of patients with event.

95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).

p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.

RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeff, created on: 20FEB2024

Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
 Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.363
<= 5 years	Double-blind period	Sparsentan	113	7 (6.2)	0.787 [0.310, 1.998]	0.773 [0.284, 2.102]	-1.7 [-9.0, 5.6]	0.802
		Irbesartan	127	10 (7.9)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	1.685 [0.436, 6.511]	1.735 [0.419, 7.188]	2.7 [-5.3, 10.8]	0.510
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2HOK\_FSIM: Patients with hospitalization related to kidney by subgroup  
 Full Analysis Set

<u>Hospitalizations related to kidney</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.846
Yes	Double-blind period	Sparsentan	155	10 (6.5)	1.039 [0.445, 2.426]	1.041 [0.421, 2.576]	0.2 [-5.8, 6.2]	1.000
		Irbesartan	161	10 (6.2)				
No	Double-blind period	Sparsentan	47	3 (6.4)	0.872 [0.186, 4.088]	0.864 [0.165, 4.534]	-0.9 [-13.8, 12.0]	1.000
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeff, created on: 20FEB2024

Table PT2HNS\_FSNM: Number of hospitalizations by subgroup  
Full Analysis Set

Number of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Sex	Sparsentan								0.559
Male	Sparsentan	139	139 (100.0)	304.9	33	0.11	0.988	(0.580, 1.685)	0.965
	Irbesartan	143	143 (100.0)	310.9	34	0.11			
Female	Sparsentan	63	63 (100.0)	138.9	12	0.09	0.838	(0.360, 1.950)	0.682
	Irbesartan	59	59 (100.0)	126.3	15	0.12			
Age	Sparsentan								0.981
<= 45 years	Sparsentan	96	96 (100.0)	210.4	23	0.11	0.934	(0.494, 1.764)	0.833
	Irbesartan	99	99 (100.0)	215.6	26	0.12			
> 45 years	Sparsentan	106	106 (100.0)	233.3	22	0.09	0.904	(0.482, 1.697)	0.755
	Irbesartan	103	103 (100.0)	221.6	23	0.10			
Age at IgAN diagnosis	Sparsentan								0.367
<= 18 years	Sparsentan	9	9 (100.0)	20.0	7	0.35	3.397	(0.418, 27.617)	0.253
	Irbesartan	5	5 (100.0)	10.9	1	0.09			
> 18 to 40 years	Sparsentan	102	102 (100.0)	223.4	17	0.08	0.813	(0.381, 1.734)	0.593
	Irbesartan	109	109 (100.0)	234.0	23	0.10			
> 40 years	Sparsentan	91	91 (100.0)	200.3	21	0.10	0.835	(0.458, 1.522)	0.557
	Irbesartan	88	88 (100.0)	192.3	25	0.13			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.

Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.

A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.

\* = significant treatment effect. # = significant interaction.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024

Table PT2HNS\_FSNM: Number of hospitalizations by subgroup  
Full Analysis Set

Number of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Geographic region	Sparsentan								0.651
	North America								
Europe	Sparsentan	35	35 (100.0)	77.2	6	0.08	0.573	(0.209, 1.573)	0.280
	Irbesartan	46	46 (100.0)	97.8	13	0.13			
Asia Pacific	Sparsentan	98	98 (100.0)	214.6	19	0.09	0.832	(0.434, 1.597)	0.581
	Irbesartan	115	115 (100.0)	251.8	26	0.10			
Baseline BMI < 27 kg/m**2	Sparsentan	69	69 (100.0)	151.9	20	0.13	1.092	(0.456, 2.613)	0.844
	Irbesartan	41	41 (100.0)	87.6	10	0.11			
>= 27 kg/m**2	Sparsentan								0.900
	Sparsentan	83	83 (100.0)	182.3	20	0.11	0.965	(0.530, 1.757)	0.907
Randomization strata eGFR Low and UP High	Irbesartan	94	94 (100.0)	202.2	24	0.12			
	Sparsentan	119	119 (100.0)	261.5	25	0.10	0.896	(0.494, 1.624)	0.717
eGFR Low and UP Low	Irbesartan	107	107 (100.0)	232.7	25	0.11			
	Sparsentan								0.218
eGFR High and UP High	Sparsentan	71	71 (100.0)	154.1	16	0.10	0.764	(0.384, 1.520)	0.444
	Irbesartan	74	74 (100.0)	162.9	22	0.14			
eGFR High and UP Low	Sparsentan	55	55 (100.0)	122.3	10	0.08	0.800	(0.314, 2.041)	0.641
	Irbesartan	55	55 (100.0)	117.4	12	0.10			
Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.	Sparsentan	37	37 (100.0)	81.8	13	0.16	2.486	(0.791, 7.808)	0.119
	Irbesartan	36	36 (100.0)	78.1	5	0.06			
Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.	Sparsentan	39	39 (100.0)	85.6	6	0.07	0.554	(0.182, 1.687)	0.299
	Irbesartan	37	37 (100.0)	78.7	10	0.13			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HNS\_FSNM: Number of hospitalizations by subgroup  
Full Analysis Set

Number of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline eGFR Group 1 < 60 mL/min/1.73 m**2	Sparsentan								0.488
	Sparsentan	127	127 (100.0)	278.6	28	0.10	0.802	(0.472, 1.364)	0.416
	Irbesartan	129	129 (100.0)	279.9	38	0.14			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	107.9	14	0.13	1.551	(0.618, 3.896)	0.350
	Irbesartan	48	48 (100.0)	105.2	9	0.09			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	57.2	3	0.05	1.375	(0.230, 8.230)	0.727
	Irbesartan	25	25 (100.0)	52.1	2	0.04			
Baseline eGFR Group 2 < 45 mL/min/1.73 m**2	Sparsentan								0.399
	Sparsentan	82	82 (100.0)	179.4	21	0.12	0.991	(0.519, 1.891)	0.978
	Irbesartan	80	80 (100.0)	172.7	21	0.12			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	45 (100.0)	99.2	7	0.07	0.536	(0.209, 1.373)	0.194
	Irbesartan	49	49 (100.0)	107.3	17	0.16			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	107.9	14	0.13	1.551	(0.618, 3.896)	0.350
	Irbesartan	48	48 (100.0)	105.2	9	0.09			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	57.2	3	0.05	1.375	(0.230, 8.230)	0.727
	Irbesartan	25	25 (100.0)	52.1	2	0.04			
Baseline urine protein excretion <= 1.75 g/day	Sparsentan								0.989
	Sparsentan	98	98 (100.0)	216.3	18	0.08	0.908	(0.453, 1.822)	0.787
	Irbesartan	93	93 (100.0)	198.9	18	0.09			
	Sparsentan	104	104 (100.0)	227.4	27	0.12	0.928	(0.516, 1.668)	0.803
> 1.75 g/day	Sparsentan	104	104 (100.0)	227.4	27	0.12			
	Irbesartan	109	109 (100.0)	238.3	31	0.13			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HNS\_FSNM: Number of hospitalizations by subgroup  
Full Analysis Set

Number of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan								0.689
Yes	Sparsentan	90	90 (100.0)	197.1	23	0.12	1.010	(0.535, 1.905)	0.977
	Irbesartan	88	88 (100.0)	191.2	22	0.12			
No	Sparsentan	112	112 (100.0)	246.7	22	0.09	0.842	(0.444, 1.598)	0.598
	Irbesartan	114	114 (100.0)	246.0	27	0.11			
Time since renal biopsy	Sparsentan								<0.001 #
<= 5 years	Sparsentan	113	113 (100.0)	247.7	21	0.08	0.531	(0.296, 0.952)	0.034 *
	Irbesartan	127	127 (100.0)	274.4	43	0.16			
> 5 years	Sparsentan	89	89 (100.0)	196.1	24	0.12	3.371	(1.370, 8.294)	0.008 *
	Irbesartan	75	75 (100.0)	162.8	6	0.04			
History of hypertension	Sparsentan								0.329
Yes	Sparsentan	155	155 (100.0)	339.9	33	0.10	0.806	(0.486, 1.336)	0.404
	Irbesartan	161	161 (100.0)	349.5	42	0.12			
No	Sparsentan	47	47 (100.0)	103.9	12	0.12	1.551	(0.560, 4.292)	0.398
	Irbesartan	41	41 (100.0)	87.7	7	0.08			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.

Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.

\* = significant treatment effect. # = significant interaction.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024



Table PT2HNK\_FSNM: Number of hospitalizations related to kidney by subgroup  
 Full Analysis Set

Number of hospitalizations related to kidney							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Sex	Sparsentan								0.505
Male	Sparsentan	139	139 (100.0)	304.9	9	0.03	0.726	(0.278, 1.892)	0.512
	Irbesartan	143	143 (100.0)	310.9	12	0.04			
Female	Sparsentan	63	63 (100.0)	138.9	5	0.04	1.643	(0.397, 6.803)	0.493
	Irbesartan	59	59 (100.0)	126.3	4	0.03			
Age	Sparsentan								0.944
<= 45 years	Sparsentan	96	96 (100.0)	210.4	10	0.05	0.904	(0.366, 2.231)	0.827
	Irbesartan	99	99 (100.0)	215.6	12	0.06			
> 45 years	Sparsentan	106	106 (100.0)	233.3	4	0.02	0.944	(0.198, 4.500)	0.942
	Irbesartan	103	103 (100.0)	221.6	4	0.02			
Age at IgAN diagnosis	Sparsentan								0.516
<= 18 years	Sparsentan	9	9 (100.0)	20.0	5	0.25	2.392	(0.279, 20.479)	0.426
	Irbesartan	5	5 (100.0)	10.9	1	0.09			
> 18 to 40 years	Sparsentan	102	102 (100.0)	223.4	5	0.02	0.555	(0.172, 1.791)	0.325
	Irbesartan	109	109 (100.0)	234.0	10	0.04			
> 40 years	Sparsentan	91	91 (100.0)	200.3	4	0.02	0.835	(0.202, 3.457)	0.803
	Irbesartan	88	88 (100.0)	192.3	5	0.03			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
 NE = not evaluable.

Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.

A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.

\* = significant treatment effect. # = significant interaction.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024

Table PT2HNK\_FSNM: Number of hospitalizations related to kidney by subgroup  
Full Analysis Set

Number of hospitalizations related to kidney							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Geographic region	Sparsentan								0.555
	North America								
Europe	Sparsentan	35	35 (100.0)	77.2	2	0.03	2.942	(0.266, 32.534)	0.379
	Irbesartan	46	46 (100.0)	97.8	1	0.01			
Asia Pacific	Sparsentan	98	98 (100.0)	214.6	7	0.03	0.665	(0.225, 1.963)	0.460
	Irbesartan	115	115 (100.0)	251.8	12	0.05			
Baseline BMI	Sparsentan	69	69 (100.0)	151.9	5	0.03	1.024	(0.242, 4.332)	0.975
	Irbesartan	41	41 (100.0)	87.6	3	0.03			
< 27 kg/m**2	Sparsentan								0.027 #
	Sparsentan	83	83 (100.0)	182.3	5	0.03	0.426	(0.139, 1.305)	0.135
>= 27 kg/m**2	Irbesartan	94	94 (100.0)	202.2	13	0.06			
	Sparsentan	119	119 (100.0)	261.5	9	0.03	2.776	(0.729, 10.580)	0.135
Randomization strata	Irbesartan	107	107 (100.0)	232.7	3	0.01			
	eGFR Low and UP High								NE
eGFR Low and UP High	Sparsentan	71	71 (100.0)	154.1	5	0.03	0.480	(0.157, 1.464)	0.197
	Irbesartan	74	74 (100.0)	162.9	11	0.07			
eGFR Low and UP Low	Sparsentan	55	55 (100.0)	122.3	2	0.02	1.920	(0.174, 21.176)	0.594
	Irbesartan	55	55 (100.0)	117.4	1	0.01			
eGFR High and UP High	Sparsentan	37	37 (100.0)	81.8	7	0.09	1.667	(0.415, 6.685)	0.471
	Irbesartan	36	36 (100.0)	78.1	4	0.05			
eGFR High and UP Low	Sparsentan	39	39 (100.0)	85.6	0	0.00			NE
	Irbesartan	37	37 (100.0)	78.7	0	0.00			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.

Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.

\* = significant treatment effect. # = significant interaction.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024

Table PT2HNK\_FSNM: Number of hospitalizations related to kidney by subgroup  
Full Analysis Set

Number of hospitalizations related to kidney							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline eGFR Group 1 < 60 mL/min/1.73 m**2	Sparsentan								NE
	Sparsentan	127	127 (100.0)	278.6	7	0.03	0.589	(0.222, 1.563)	0.288
	Irbesartan	129	129 (100.0)	279.9	12	0.04			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	107.9	6	0.06	1.464	(0.346, 6.189)	0.604
	Irbesartan	48	48 (100.0)	105.2	4	0.04			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	57.2	1	0.02			NE
	Irbesartan	25	25 (100.0)	52.1	0	0.00			
Baseline eGFR Group 2 < 45 mL/min/1.73 m**2	Sparsentan								NE
	Sparsentan	82	82 (100.0)	179.4	6	0.03	0.615	(0.212, 1.785)	0.372
	Irbesartan	80	80 (100.0)	172.7	10	0.06			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	45 (100.0)	99.2	1	0.01	0.497	(0.045, 5.478)	0.568
	Irbesartan	49	49 (100.0)	107.3	2	0.02			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	107.9	6	0.06	1.464	(0.346, 6.189)	0.604
	Irbesartan	48	48 (100.0)	105.2	4	0.04			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	57.2	1	0.02			NE
	Irbesartan	25	25 (100.0)	52.1	0	0.00			
Baseline urine protein excretion <= 1.75 g/day	Sparsentan								0.135
	Sparsentan	98	98 (100.0)	216.3	4	0.02	3.646	(0.406, 32.723)	0.248
	Irbesartan	93	93 (100.0)	198.9	1	0.01			
	Irbesartan	93	93 (100.0)	198.9	1	0.01			
> 1.75 g/day	Sparsentan	104	104 (100.0)	227.4	10	0.04	0.684	(0.281, 1.668)	0.404
	Irbesartan	109	109 (100.0)	238.3	15	0.06			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HNK\_FSNM: Number of hospitalizations related to kidney by subgroup  
Full Analysis Set

Number of hospitalizations related to kidney							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan								0.430
Yes	Sparsentan	90	90 (100.0)	197.1	9	0.05	1.184	(0.411, 3.414)	0.754
	Irbesartan	88	88 (100.0)	191.2	7	0.04			
No	Sparsentan	112	112 (100.0)	246.7	5	0.02	0.601	(0.180, 2.007)	0.408
	Irbesartan	114	114 (100.0)	246.0	9	0.04			
Time since renal biopsy	Sparsentan								0.165
<= 5 years	Sparsentan	113	113 (100.0)	247.7	7	0.03	0.609	(0.222, 1.673)	0.336
	Irbesartan	127	127 (100.0)	274.4	13	0.05			
> 5 years	Sparsentan	89	89 (100.0)	196.1	7	0.04	1.998	(0.488, 8.185)	0.336
	Irbesartan	75	75 (100.0)	162.8	3	0.02			
History of hypertension	Sparsentan								0.915
Yes	Sparsentan	155	155 (100.0)	339.9	10	0.03	0.832	(0.337, 2.052)	0.689
	Irbesartan	161	161 (100.0)	349.5	13	0.04			
No	Sparsentan	47	47 (100.0)	103.9	4	0.04	1.117	(0.221, 5.650)	0.894
	Irbesartan	41	41 (100.0)	87.7	3	0.03			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.

Number of events = number of considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.

\* = significant treatment effect. # = significant interaction.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024

Table PT2HDS\_FSNM: Duration of hospitalizations by subgroup  
Full Analysis Set

Duration of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Sex	Sparsentan								0.709
Male	Sparsentan	139	139 (100.0)	304.9	198	0.65	0.644	(0.231, 1.794)	0.399
	Irbesartan	143	143 (100.0)	310.9	361	1.16			
Female	Sparsentan	63	63 (100.0)	138.9	80	0.58	1.780	(0.425, 7.462)	0.430
	Irbesartan	59	59 (100.0)	126.3	67	0.53			
Age	Sparsentan								0.850
<= 45 years	Sparsentan	96	96 (100.0)	210.4	183	0.87	0.786	(0.225, 2.747)	0.706
	Irbesartan	99	99 (100.0)	215.6	278	1.29			
> 45 years	Sparsentan	106	106 (100.0)	233.3	95	0.41	0.713	(0.235, 2.159)	0.550
	Irbesartan	103	103 (100.0)	221.6	150	0.68			
Age at IgAN diagnosis	Sparsentan								0.783
<= 18 years	Sparsentan	9	9 (100.0)	20.0	53	2.65	2.540	(0.088, 73.222)	0.587
	Irbesartan	5	5 (100.0)	10.9	11	1.01			
> 18 to 40 years	Sparsentan	102	102 (100.0)	223.4	137	0.61	0.713	(0.160, 3.168)	0.657
	Irbesartan	109	109 (100.0)	234.0	245	1.05			
> 40 years	Sparsentan	91	91 (100.0)	200.3	88	0.44	0.536	(0.187, 1.541)	0.247
	Irbesartan	88	88 (100.0)	192.3	172	0.89			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HDS\_FSNM: Duration of hospitalizations by subgroup  
 Full Analysis Set

Duration of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Geographic region	Sparsentan								0.955
	North America								0.461
Europe	Sparsentan	35	35 (100.0)	77.2	23	0.30	0.553	(0.115, 2.668)	
	Irbesartan	46	46 (100.0)	97.8	59	0.60			0.791
Asia Pacific	Sparsentan	98	98 (100.0)	214.6	178	0.83	0.841	(0.233, 3.032)	
	Irbesartan	115	115 (100.0)	251.8	315	1.25			0.889
Baseline BMI < 27 kg/m**2	Sparsentan	69	69 (100.0)	151.9	77	0.51	0.906	(0.227, 3.620)	
	Irbesartan	41	41 (100.0)	87.6	54	0.62			0.598
>= 27 kg/m**2	Sparsentan	83	83 (100.0)	182.3	135	0.74	1.326	(0.365, 4.813)	0.668
	Irbesartan	94	94 (100.0)	202.2	152	0.75			0.787
Randomization strata eGFR Low and UP High	Sparsentan	119	119 (100.0)	261.5	143	0.55	0.850	(0.262, 2.757)	
	Irbesartan	107	107 (100.0)	232.7	276	1.19			0.216
eGFR Low and UP Low	Sparsentan	71	71 (100.0)	154.1	90	0.58	0.529	(0.163, 1.714)	0.288
	Irbesartan	74	74 (100.0)	162.9	169	1.04			0.812
eGFR High and UP High	Sparsentan	55	55 (100.0)	122.3	60	0.49	1.197	(0.273, 5.244)	
	Irbesartan	55	55 (100.0)	117.4	48	0.41			0.256
eGFR High and UP Low	Sparsentan	37	37 (100.0)	81.8	89	1.09	2.791	(0.475, 16.398)	
	Irbesartan	36	36 (100.0)	78.1	31	0.40			0.132
	Sparsentan	39	39 (100.0)	85.6	39	0.46	0.212	(0.028, 1.594)	
	Irbesartan	37	37 (100.0)	78.7	180	2.29			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
 NE = not evaluable.  
 Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
 A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
 \* = significant treatment effect. # = significant interaction.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2HDS\_FSNM: Duration of hospitalizations by subgroup  
Full Analysis Set

Duration of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline eGFR Group 1 < 60 mL/min/1.73 m**2	Sparsentan								0.419
	Sparsentan	127	127 (100.0)	278.6	157	0.56	0.817	(0.332, 2.011)	0.660
	Irbesartan	129	129 (100.0)	279.9	244	0.87			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	107.9	95	0.88	1.829	(0.380, 8.807)	0.451
	Irbesartan	48	48 (100.0)	105.2	60	0.57			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	57.2	26	0.45	0.766	(0.002, 382.922)	0.933
	Irbesartan	25	25 (100.0)	52.1	124	2.38			
Baseline eGFR Group 2 < 45 mL/min/1.73 m**2	Sparsentan								0.272
	Sparsentan	82	82 (100.0)	179.4	133	0.74	1.316	(0.398, 4.349)	0.652
	Irbesartan	80	80 (100.0)	172.7	125	0.72			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	45 (100.0)	99.2	24	0.24	0.274	(0.067, 1.121)	0.072
	Irbesartan	49	49 (100.0)	107.3	119	1.11			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	107.9	95	0.88	1.829	(0.380, 8.807)	0.451
	Irbesartan	48	48 (100.0)	105.2	60	0.57			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	57.2	26	0.45	0.766	(0.002, 382.922)	0.933
	Irbesartan	25	25 (100.0)	52.1	124	2.38			
Baseline urine protein excretion <= 1.75 g/day	Sparsentan								0.561
	Sparsentan	98	98 (100.0)	216.3	96	0.44	0.535	(0.140, 2.048)	0.361
	Irbesartan	93	93 (100.0)	198.9	212	1.07			
	Sparsentan	104	104 (100.0)	227.4	182	0.80	0.880	(0.306, 2.530)	0.813
> 1.75 g/day	Sparsentan	109	109 (100.0)	238.3	216	0.91			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HDS\_FSNM: Duration of hospitalizations by subgroup  
Full Analysis Set

Duration of hospitalizations							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan								0.285
Yes	Sparsentan	90	90 (100.0)	197.1	166	0.84	1.001	(0.303, 3.305)	0.999
	Irbesartan	88	88 (100.0)	191.2	152	0.79			
No	Sparsentan	112	112 (100.0)	246.7	112	0.45	0.421	(0.128, 1.386)	0.155
	Irbesartan	114	114 (100.0)	246.0	276	1.12			
Time since renal biopsy	Sparsentan								0.048 #
<= 5 years	Sparsentan	113	113 (100.0)	247.7	134	0.54	0.319	(0.107, 0.956)	0.041 *
	Irbesartan	127	127 (100.0)	274.4	356	1.30			
> 5 years	Sparsentan	89	89 (100.0)	196.1	144	0.73	3.024	(0.673, 13.585)	0.149
	Irbesartan	75	75 (100.0)	162.8	72	0.44			
History of hypertension	Sparsentan								0.399
Yes	Sparsentan	155	155 (100.0)	339.9	192	0.56	0.554	(0.208, 1.476)	0.237
	Irbesartan	161	161 (100.0)	349.5	390	1.12			
No	Sparsentan	47	47 (100.0)	103.9	86	0.83	3.962	(0.413, 37.968)	0.233
	Irbesartan	41	41 (100.0)	87.7	38	0.43			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.

Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.

\* = significant treatment effect. # = significant interaction.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

Source Data: aeFF, created on: 20FEB2024



Table PT2HDK\_FSNM: Duration of hospitalizations related to kidney by subgroup  
 Full Analysis Set

Duration of hospitalizations related to kidney							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Sex	Sparsentan								0.036 #
Male	Sparsentan	139	139 (100.0)	304.9	35	0.11	0.213	(0.041, 1.119)	0.068
	Irbesartan	143	143 (100.0)	310.9	96	0.31			
Female	Sparsentan	63	63 (100.0)	138.9	51	0.37	14.622	(0.406, 526.863)	0.142
	Irbesartan	59	59 (100.0)	126.3	27	0.21			
Age	Sparsentan								0.538
<= 45 years	Sparsentan	96	96 (100.0)	210.4	68	0.32	1.043	(0.203, 5.364)	0.960
	Irbesartan	99	99 (100.0)	215.6	100	0.46			
> 45 years	Sparsentan	106	106 (100.0)	233.3	18	0.08	2.393	(0.034, 170.982)	0.689
	Irbesartan	103	103 (100.0)	221.6	23	0.10			
Age at IgAN diagnosis	Sparsentan								0.686
<= 18 years	Sparsentan	9	9 (100.0)	20.0	51	2.55	2.562	(0.048, 136.943)	0.643
	Irbesartan	5	5 (100.0)	10.9	11	1.01			
> 18 to 40 years	Sparsentan	102	102 (100.0)	223.4	17	0.08	0.397	(0.038, 4.120)	0.439
	Irbesartan	109	109 (100.0)	234.0	83	0.35			
> 40 years	Sparsentan	91	91 (100.0)	200.3	18	0.09	1.947	(0.035, 107.159)	0.745
	Irbesartan	88	88 (100.0)	192.3	29	0.15			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
 NE = not evaluable.  
 Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
 A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
 \* = significant treatment effect. # = significant interaction.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aeFF, created on: 20FEB2024

Table PT2HDK\_FSNM: Duration of hospitalizations related to kidney by subgroup  
Full Analysis Set

Duration of hospitalizations related to kidney							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Geographic region	Sparsentan								0.985
	North America								0.785
Europe	Sparsentan	35	35 (100.0)	77.2	6	0.08	2.016	(0.013, 308.597)	
	Irbesartan	46	46 (100.0)	97.8	7	0.07			0.726
Asia Pacific	Sparsentan	98	98 (100.0)	214.6	59	0.27	1.456	(0.178, 11.909)	
	Irbesartan	115	115 (100.0)	251.8	99	0.39			0.991
Baseline BMI	Sparsentan	69	69 (100.0)	151.9	21	0.14	1.018	(0.043, 23.918)	
	< 27 kg/m**2	41	41 (100.0)	87.6	17	0.19			0.106
>= 27 kg/m**2	Sparsentan	83	83 (100.0)	182.3	20	0.11	0.526	(0.050, 5.488)	0.591
	Irbesartan	94	94 (100.0)	202.2	108	0.53			0.126
Randomization strata	Sparsentan	119	119 (100.0)	261.5	66	0.25	5.597	(0.616, 50.872)	
	eGFR Low and UP High	107	107 (100.0)	232.7	15	0.06			NE
eGFR Low and UP Low	Sparsentan	71	71 (100.0)	154.1	45	0.29	0.474	(0.070, 3.207)	0.444
	Irbesartan	74	74 (100.0)	162.9	94	0.58			0.167
eGFR High and UP High	Sparsentan	55	55 (100.0)	122.3	14	0.11	12.909	(0.344, 484.408)	
	Irbesartan	55	55 (100.0)	117.4	1	0.01			0.941
eGFR High and UP Low	Sparsentan	37	37 (100.0)	81.8	27	0.33	0.921	(0.104, 8.120)	
	Irbesartan	36	36 (100.0)	78.1	28	0.36			NE
	Sparsentan	39	39 (100.0)	85.6	0	0.00			
	Irbesartan	37	37 (100.0)	78.7	0	0.00			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeff, created on: 20FEB2024

Table PT2HDK\_FSNM: Duration of hospitalizations related to kidney by subgroup  
Full Analysis Set

Duration of hospitalizations related to kidney							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline eGFR Group 1 < 60 mL/min/1.73 m**2	Sparsentan								NE
	Sparsentan	127	127 (100.0)	278.6	59	0.21	1.343	(0.181, 9.945)	0.773
	Irbesartan	129	129 (100.0)	279.9	95	0.34			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	107.9	22	0.20	0.769	(0.079, 7.457)	0.820
	Irbesartan	48	48 (100.0)	105.2	28	0.27			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	57.2	5	0.09			NE
	Irbesartan	25	25 (100.0)	52.1	0	0.00			
Baseline eGFR Group 2 < 45 mL/min/1.73 m**2	Sparsentan								NE
	Sparsentan	82	82 (100.0)	179.4	57	0.32	1.778	(0.166, 19.052)	0.634
	Irbesartan	80	80 (100.0)	172.7	72	0.42			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	45 (100.0)	99.2	2	0.02	0.070	(0.001, 3.741)	0.190
	Irbesartan	49	49 (100.0)	107.3	23	0.21			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	49 (100.0)	107.9	22	0.20	0.769	(0.079, 7.457)	0.820
	Irbesartan	48	48 (100.0)	105.2	28	0.27			
>= 90 mL/min/1.73 m**2	Sparsentan	26	26 (100.0)	57.2	5	0.09			NE
	Irbesartan	25	25 (100.0)	52.1	0	0.00			
Baseline urine protein excretion <= 1.75 g/day	Sparsentan								0.093
	Sparsentan	98	98 (100.0)	216.3	11	0.05	10.367	(0.682, 157.613)	0.092
	Irbesartan	93	93 (100.0)	198.9	1	0.01			
	Sparsentan	104	104 (100.0)	227.4	75	0.33	0.751	(0.146, 3.868)	0.732
> 1.75 g/day	Sparsentan	104	104 (100.0)	227.4	75	0.33			
	Irbesartan	109	109 (100.0)	238.3	122	0.51			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2HDK\_FSNM: Duration of hospitalizations related to kidney by subgroup  
Full Analysis Set

Duration of hospitalizations related to kidney							Negative binomial model		
Subgroup	Treatment	N	nval (%)	Time at risk (years)	Number of events	Crude rate	Rate ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan								0.818
Yes	Sparsentan	90	90 (100.0)	197.1	59	0.30	2.020	(0.233, 17.501)	0.523
	Irbesartan	88	88 (100.0)	191.2	61	0.32			
No	Sparsentan	112	112 (100.0)	246.7	27	0.11	0.775	(0.062, 9.703)	0.843
	Irbesartan	114	114 (100.0)	246.0	62	0.25			
Time since renal biopsy	Sparsentan								0.226
<= 5 years	Sparsentan	113	113 (100.0)	247.7	23	0.09	0.452	(0.058, 3.531)	0.449
	Irbesartan	127	127 (100.0)	274.4	96	0.35			
> 5 years	Sparsentan	89	89 (100.0)	196.1	63	0.32	3.765	(0.276, 51.393)	0.320
	Irbesartan	75	75 (100.0)	162.8	27	0.17			
History of hypertension	Sparsentan								0.892
Yes	Sparsentan	155	155 (100.0)	339.9	45	0.13	0.680	(0.099, 4.652)	0.694
	Irbesartan	161	161 (100.0)	349.5	102	0.29			
No	Sparsentan	47	47 (100.0)	103.9	41	0.39	2.823	(0.107, 74.260)	0.534
	Irbesartan	41	41 (100.0)	87.7	21	0.24			

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. nval (%) = number of patients included in the analysis (i.e., with post-baseline data).  
NE = not evaluable.  
Number of events = number of days with considered hospitalizations. 95% CI = 95% confidence interval for rate ratio.  
A negative binomial model was applied with factors treatment, randomisation strata, and log of offset. The study duration of double blind period was used as offset. Randomisation stratum will be included only once in the model, if this stratum is investigated subgroup.  
\* = significant treatment effect. # = significant interaction.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aeFF, created on: 20FEB2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Sex														
Male	EQ-5D VAS	Baseline	Sparsentan	139	132 (95.0)	81.11 (11.21)	40.0	74.50	81.00	90.00	100.0			
			Irbesartan	143	132 (92.3)	79.64 (14.68)	19.0	75.00	80.00	90.00	100.0			
		Week 24	Sparsentan	139	114 (82.0)	84.02 (10.82)	50.0	78.00	85.00	91.00	100.0			
			Irbesartan	143	103 (72.0)	80.04 (18.26)	4.0	73.00	83.00	91.00	100.0			
		Week 48	Sparsentan	139	118 (84.9)	83.19 (10.61)	49.0	80.00	84.00	90.00	100.0			
			Irbesartan	143	94 (65.7)	82.32 (13.44)	40.0	75.00	85.00	91.00	100.0			
		Week 70	Sparsentan	139	119 (85.6)	81.27 (10.56)	50.0	75.00	81.00	90.00	100.0			
			Irbesartan	143	97 (67.8)	82.01 (15.41)	20.0	75.00	85.00	93.00	100.0			
		Week 94	Sparsentan	139	109 (78.4)	83.09 (12.02)	34.0	79.00	85.00	91.00	100.0			
			Irbesartan	143	99 (69.2)	82.59 (14.49)	34.0	76.00	86.00	91.00	100.0			
		Week 110	Sparsentan	139	106 (76.3)	82.97 (11.21)	50.0	78.00	83.00	91.00	100.0			
			Irbesartan	143	89 (62.2)	81.40 (13.41)	35.0	76.00	84.00	90.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	139	114 (82.0)	2.54 (11.36)	-31.0	-3.00	2.00	10.00	44.0	0.10 [-0.17, 0.37]
					Irbesartan	143	103 (72.0)	1.11 (17.25)	-90.0	-5.00	0.00	10.00	52.0	
			Week 48	Sparsentan	139	118 (84.9)	1.90 (10.49)	-30.0	-3.00	1.00	7.00	41.0	-0.15 [-0.42, 0.12]	
				Irbesartan	143	94 (65.7)	3.56 (12.20)	-35.0	-4.00	3.00	10.00	43.0		
			Week 70	Sparsentan	139	119 (85.6)	-0.20 (11.69)	-32.0	-9.00	0.00	8.00	40.0	-0.27 [-0.54, -0.00]	
				Irbesartan	143	97 (67.8)	3.13 (12.82)	-60.0	-3.00	1.00	10.00	40.0		
			Week 94	Sparsentan	139	109 (78.4)	1.78 (12.93)	-61.0	-3.00	2.00	10.00	40.0	-0.17 [-0.45, 0.10]	
				Irbesartan	143	99 (69.2)	4.15 (14.58)	-41.0	-3.00	3.00	10.00	50.0		
	Week 110	Sparsentan	139	106 (76.3)	1.10 (10.05)	-31.0	-4.00	1.00	7.00	21.0	-0.18 [-0.46, 0.10]			
		Irbesartan	143	89 (62.2)	3.10 (12.02)	-26.0	-4.00	1.00	10.00	39.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Female	EQ-5D VAS	Baseline	Sparsentan	63	59 (93.7)	79.07 (19.44)	7.0	75.00	82.00	90.00	100.0			
			Irbesartan	59	52 (88.1)	82.29 (12.91)	37.0	74.00	83.00	91.50	100.0			
		Week 24	Sparsentan	63	50 (79.4)	84.60 (10.46)	51.0	80.00	88.50	91.00	100.0			
			Irbesartan	59	40 (67.8)	83.80 (9.93)	48.0	80.00	84.50	90.50	99.0			
		Week 48	Sparsentan	63	50 (79.4)	83.44 (12.18)	49.0	78.00	82.50	92.00	100.0			
			Irbesartan	59	38 (64.4)	82.21 (12.94)	50.0	80.00	82.00	90.00	100.0			
		Week 70	Sparsentan	63	47 (74.6)	81.49 (14.77)	27.0	75.00	82.00	90.00	100.0			
			Irbesartan	59	39 (66.1)	80.82 (16.48)	20.0	76.00	84.00	91.00	100.0			
		Week 94	Sparsentan	63	47 (74.6)	79.74 (15.23)	41.0	70.00	81.00	90.00	100.0			
			Irbesartan	59	37 (62.7)	81.81 (13.91)	40.0	78.00	82.00	90.00	100.0			
		Week 110	Sparsentan	63	46 (73.0)	81.07 (12.00)	48.0	76.00	81.00	90.00	100.0			
			Irbesartan	59	32 (54.2)	82.88 (9.41)	60.0	79.50	81.00	90.00	100.0			
		Change from baseline in EQ-5D VAS	EQ-5D VAS	Week 24	Sparsentan	63	50 (79.4)	4.68 (18.98)	-25.0	-9.00	0.00	12.00	75.0	0.26 [-0.16, 0.68]
					Irbesartan	59	40 (67.8)	0.43 (11.91)	-37.0	-4.50	1.00	7.00	23.0	
				Week 48	Sparsentan	63	50 (79.4)	3.54 (20.84)	-29.0	-9.00	0.00	11.00	87.0	0.19 [-0.23, 0.62]
					Irbesartan	59	38 (64.4)	0.11 (12.53)	-31.0	-8.00	1.50	7.00	23.0	
				Week 70	Sparsentan	63	47 (74.6)	0.74 (17.49)	-51.0	-6.00	0.00	9.00	59.0	0.26 [-0.17, 0.68]
					Irbesartan	59	39 (66.1)	-3.26 (12.83)	-49.0	-10.00	-1.00	7.00	18.0	
				Week 94	Sparsentan	63	47 (74.6)	-0.30 (18.22)	-34.0	-12.00	0.00	10.00	45.0	0.04 [-0.39, 0.47]
					Irbesartan	59	37 (62.7)	-0.95 (14.41)	-45.0	-7.00	0.00	10.00	22.0	
Week 110	Sparsentan			63	46 (73.0)	2.83 (23.17)	-40.0	-10.00	0.00	11.00	88.0	0.11 [-0.34, 0.56]		
	Irbesartan			59	32 (54.2)	0.69 (10.07)	-21.0	-7.00	1.50	5.00	27.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age														
<= 45 years	EQ-5D VAS	Baseline	Sparsentan	96	89 (92.7)	79.93 (14.66)	7.0	75.00	81.00	90.00	100.0			
			Irbesartan	99	92 (92.9)	81.66 (14.94)	19.0	75.00	82.00	90.50	100.0			
		Week 24	Sparsentan	96	75 (78.1)	84.88 (10.79)	59.0	78.00	88.00	91.00	100.0			
			Irbesartan	99	72 (72.7)	81.65 (17.51)	10.0	72.50	85.00	92.50	100.0			
		Week 48	Sparsentan	96	79 (82.3)	83.62 (11.66)	49.0	79.00	86.00	91.00	100.0			
			Irbesartan	99	63 (63.6)	82.43 (13.86)	40.0	75.00	85.00	91.00	100.0			
		Week 70	Sparsentan	96	76 (79.2)	82.32 (11.82)	27.0	77.50	83.50	90.00	100.0			
			Irbesartan	99	64 (64.6)	83.45 (16.03)	20.0	78.50	88.00	94.00	100.0			
		Week 94	Sparsentan	96	71 (74.0)	81.52 (14.32)	34.0	77.00	84.00	91.00	100.0			
			Irbesartan	99	65 (65.7)	81.71 (15.46)	40.0	71.00	84.00	92.00	100.0			
		Week 110	Sparsentan	96	71 (74.0)	83.46 (11.20)	48.0	80.00	84.00	90.00	100.0			
			Irbesartan	99	59 (59.6)	83.73 (12.60)	47.0	79.00	87.00	92.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	96	75 (78.1)	4.85 (15.83)	-31.0	-4.00	2.00	12.00	75.0	0.39 [0.06, 0.72]
					Irbesartan	99	72 (72.7)	-1.42 (16.32)	-90.0	-6.00	-1.00	9.00	23.0	
			Week 48	Sparsentan	96	79 (82.3)	2.78 (14.37)	-30.0	-4.00	1.00	10.00	64.0	0.24 [-0.09, 0.58]	
				Irbesartan	99	63 (63.6)	-0.51 (12.23)	-35.0	-6.00	0.00	6.00	27.0		
			Week 70	Sparsentan	96	76 (79.2)	1.28 (13.02)	-51.0	-6.00	0.00	9.00	41.0	0.09 [-0.24, 0.42]	
				Irbesartan	99	64 (64.6)	0.13 (12.36)	-49.0	-5.50	0.00	10.00	29.0		
			Week 94	Sparsentan	96	71 (74.0)	0.38 (17.36)	-61.0	-9.00	1.00	11.00	45.0	0.05 [-0.29, 0.38]	
				Irbesartan	99	65 (65.7)	-0.37 (15.64)	-45.0	-6.00	0.00	9.00	44.0		
	Week 110	Sparsentan	96	71 (74.0)	2.85 (16.95)	-40.0	-4.00	1.00	10.00	88.0	0.07 [-0.28, 0.41]			
		Irbesartan	99	59 (59.6)	1.85 (11.38)	-26.0	-5.00	0.00	8.00	30.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 45 years	EQ-5D VAS	Baseline	Sparsentan	106	102 (96.2)	80.96 (13.92)	11.0	75.00	82.00	90.00	100.0			
			Irbesartan	103	92 (89.3)	79.11 (13.42)	34.0	74.00	80.00	89.00	100.0			
		Week 24	Sparsentan	106	89 (84.0)	83.62 (10.61)	50.0	78.00	84.00	90.00	100.0			
			Irbesartan	103	71 (68.9)	80.52 (15.34)	4.0	78.00	81.00	90.00	100.0			
		Week 48	Sparsentan	106	89 (84.0)	82.94 (10.56)	49.0	79.00	84.00	90.00	100.0			
			Irbesartan	103	69 (67.0)	82.16 (12.77)	50.0	79.00	83.00	90.00	100.0			
		Week 70	Sparsentan	106	90 (84.9)	80.50 (11.89)	40.0	75.00	81.00	90.00	100.0			
			Irbesartan	103	72 (69.9)	80.08 (15.29)	20.0	73.50	82.00	90.50	100.0			
		Week 94	Sparsentan	106	85 (80.2)	82.55 (12.08)	40.0	79.00	83.00	91.00	100.0			
			Irbesartan	103	71 (68.9)	82.99 (13.22)	34.0	79.00	84.00	90.00	100.0			
		Week 110	Sparsentan	106	81 (76.4)	81.46 (11.65)	50.0	75.00	81.00	90.00	100.0			
			Irbesartan	103	62 (60.2)	79.95 (12.14)	35.0	76.00	81.00	89.00	100.0			
		Change from baseline in EQ-5D VAS		Week 24	Sparsentan	106	89 (84.0)	1.80 (12.37)	-29.0	-4.00	0.00	7.00	44.0	-0.11 [-0.42, 0.20]
					Irbesartan	103	71 (68.9)	3.28 (15.20)	-71.0	-1.00	2.00	10.00	52.0	
				Week 48	Sparsentan	106	89 (84.0)	2.03 (14.35)	-29.0	-3.00	0.00	7.00	87.0	-0.25 [-0.57, 0.06]
					Irbesartan	103	69 (67.0)	5.38 (11.86)	-19.0	-1.00	5.00	10.00	43.0	
				Week 70	Sparsentan	106	90 (84.9)	-0.96 (13.94)	-45.0	-10.00	-1.00	6.00	59.0	-0.24 [-0.55, 0.07]
					Irbesartan	103	72 (69.9)	2.35 (13.72)	-60.0	-4.50	0.50	9.50	40.0	
				Week 94	Sparsentan	106	85 (80.2)	1.80 (12.09)	-37.0	-3.00	1.00	8.00	40.0	-0.30 [-0.62, 0.01]
					Irbesartan	103	71 (68.9)	5.63 (13.17)	-41.0	-3.00	5.00	11.00	50.0	
Week 110	Sparsentan			106	81 (76.4)	0.56 (13.49)	-31.0	-5.00	1.00	8.00	70.0	-0.20 [-0.53, 0.14]		
	Irbesartan			103	62 (60.2)	3.05 (11.77)	-23.0	-4.00	1.50	10.00	39.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024



Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age at IgAN diagnosis														
<= 18 years	EQ-5D VAS	Baseline	Sparsentan	9	9 (100.0)	74.67 (15.03)	51.0	62.00	79.00	82.00	95.0			
			Irbesartan	5	5 (100.0)	76.40 (11.59)	60.0	73.00	74.00	86.00	89.0			
		Week 24	Sparsentan	9	5 (55.6)	86.00 (16.73)	60.0	80.00	90.00	100.00	100.0			
			Irbesartan	5	5 (100.0)	72.60 (25.04)	31.0	69.00	83.00	85.00	95.0			
		Week 48	Sparsentan	9	7 (77.8)	82.00 (11.79)	66.0	71.00	81.00	92.00	100.0			
			Irbesartan	5	3 (60.0)	79.67 (11.93)	66.0	66.00	85.00	88.00	88.0			
		Week 70	Sparsentan	9	7 (77.8)	82.29 (11.61)	66.0	70.00	85.00	90.00	100.0			
			Irbesartan	5	4 (80.0)	74.50 (17.25)	55.0	60.00	76.50	89.00	90.0			
		Week 94	Sparsentan	9	5 (55.6)	69.40 (27.40)	34.0	50.00	73.00	90.00	100.0			
			Irbesartan	5	4 (80.0)	74.25 (24.58)	49.0	53.50	74.00	95.00	100.0			
		Week 110	Sparsentan	9	4 (44.4)	87.75 (9.67)	80.0	80.00	85.50	95.50	100.0			
			Irbesartan	5	2 (40.0)	53.50 (9.19)	47.0	47.00	53.50	60.00	60.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	9	5 (55.6)	9.60 (7.57)	0.0	7.00	10.00	10.00	21.0	0.80 [-0.49, 2.09]
					Irbesartan	5	5 (100.0)	-3.80 (22.42)	-42.0	-6.00	9.00	9.00	11.0	
			Week 48	Sparsentan	9	7 (77.8)	6.14 (8.53)	-9.0	-2.00	10.00	11.00	15.0	0.09 [-1.26, 1.45]	
				Irbesartan	5	3 (60.0)	5.33 (9.02)	-4.0	-4.00	6.00	14.00	14.0		
			Week 70	Sparsentan	9	7 (77.8)	6.43 (8.89)	-10.0	0.00	8.00	15.00	15.0	0.94 [-0.35, 2.23]	
				Irbesartan	5	4 (80.0)	-2.50 (10.66)	-18.0	-9.50	1.50	4.50	5.0		
			Week 94	Sparsentan	9	5 (55.6)	-10.20 (34.35)	-61.0	-30.00	7.00	13.00	20.0	-0.27 [-1.59, 1.05]	
				Irbesartan	5	4 (80.0)	-2.75 (15.13)	-24.0	-13.00	1.00	7.50	11.0		
	Week 110	Sparsentan	9	4 (44.4)	9.00 (8.29)	0.0	3.50	8.00	14.50	20.0	1.89 [-0.12, 3.89]			
		Irbesartan	5	2 (40.0)	-13.00 (18.38)	-26.0	-26.00	-13.00	0.00	0.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 VAS = visual analogue scale.  
 Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 18 to 40 years	EQ-5D VAS	Baseline	Sparsentan	102	95 (93.1)	80.85 (14.44)	7.0	75.00	82.00	90.00	100.0		
			Irbesartan	109	100 (91.7)	81.14 (15.01)	19.0	75.00	82.00	90.00	100.0		
	Week 24	Sparsentan	102	82 (80.4)	84.83 (10.16)	59.0	78.00	87.00	91.00	100.0			
		Irbesartan	109	75 (68.8)	81.83 (17.09)	10.0	74.00	85.00	92.00	100.0			
	Week 48	Sparsentan	102	85 (83.3)	83.93 (10.76)	49.0	80.00	86.00	91.00	100.0			
		Irbesartan	109	69 (63.3)	81.87 (14.37)	40.0	79.00	85.00	92.00	100.0			
	Week 70	Sparsentan	102	81 (79.4)	82.41 (11.41)	27.0	77.00	83.00	90.00	100.0			
		Irbesartan	109	71 (65.1)	83.44 (15.49)	20.0	79.00	87.00	94.00	100.0			
	Week 94	Sparsentan	102	77 (75.5)	83.01 (12.15)	41.0	80.00	85.00	91.00	100.0			
		Irbesartan	109	71 (65.1)	81.52 (14.97)	39.0	71.00	84.00	91.00	100.0			
	Week 110	Sparsentan	102	75 (73.5)	83.07 (11.39)	48.0	79.00	84.00	90.00	100.0			
		Irbesartan	109	65 (59.6)	83.35 (13.26)	35.0	79.00	88.00	91.00	100.0			
	Change from baseline in EQ-5D VAS		Week 24	Sparsentan	102	82 (80.4)	4.37 (16.12)	-31.0	-3.00	1.50	11.00	75.0	0.29 [-0.02, 0.61]
				Irbesartan	109	75 (68.8)	-0.31 (15.83)	-90.0	-6.00	-1.00	10.00	23.0	
			Week 48	Sparsentan	102	85 (83.3)	2.48 (14.53)	-30.0	-4.00	1.00	10.00	64.0	0.17 [-0.15, 0.49]
				Irbesartan	109	69 (63.3)	0.16 (12.35)	-35.0	-5.00	0.00	7.00	27.0	
			Week 70	Sparsentan	102	81 (79.4)	0.98 (13.41)	-51.0	-6.00	0.00	8.00	41.0	-0.00 [-0.32, 0.32]
				Irbesartan	109	71 (65.1)	0.99 (11.97)	-49.0	-6.00	0.00	10.00	29.0	
			Week 94	Sparsentan	102	77 (75.5)	1.88 (15.27)	-34.0	-5.00	2.00	9.00	45.0	0.10 [-0.22, 0.43]
				Irbesartan	109	71 (65.1)	0.28 (15.44)	-45.0	-9.00	0.00	10.00	44.0	
Week 110			Sparsentan	102	75 (73.5)	2.05 (16.59)	-40.0	-5.00	1.00	7.00	88.0	-0.02 [-0.35, 0.31]	
			Irbesartan	109	65 (59.6)	2.35 (11.57)	-23.0	-4.00	0.00	11.00	30.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 40 years	EQ-5D VAS	Baseline	Sparsentan	91	87 (95.6)	80.68 (13.98)	11.0	72.00	81.00	90.00	100.0			
			Irbesartan	88	79 (89.8)	79.68 (13.39)	40.0	73.00	80.00	90.00	100.0			
		Week 24	Sparsentan	91	77 (84.6)	83.40 (10.89)	50.0	78.00	83.00	90.00	100.0			
			Irbesartan	88	63 (71.6)	80.89 (14.90)	4.0	78.00	80.00	90.00	100.0			
		Week 48	Sparsentan	91	76 (83.5)	82.63 (11.44)	49.0	78.50	84.00	90.00	100.0			
			Irbesartan	88	60 (68.2)	82.90 (12.09)	50.0	77.00	82.50	90.00	100.0			
		Week 70	Sparsentan	91	78 (85.7)	80.13 (12.36)	40.0	75.00	81.00	90.00	100.0			
			Irbesartan	88	61 (69.3)	80.08 (15.75)	20.0	74.00	82.00	91.00	100.0			
		Week 94	Sparsentan	91	74 (81.3)	81.97 (12.58)	40.0	77.00	82.50	91.00	100.0			
			Irbesartan	88	61 (69.3)	83.90 (12.66)	34.0	80.00	85.00	90.00	100.0			
		Week 110	Sparsentan	91	73 (80.2)	81.41 (11.61)	50.0	75.00	81.00	90.00	100.0			
			Irbesartan	88	54 (61.4)	80.96 (10.19)	40.0	76.00	81.00	89.00	100.0			
		Change from baseline in EQ-5D VAS	EQ-5D VAS	Week 24	Sparsentan	91	77 (84.6)	1.53 (11.81)	-29.0	-6.00	0.00	7.00	40.0	-0.09 [-0.42, 0.24]
					Irbesartan	88	63 (71.6)	2.75 (15.50)	-71.0	-1.00	1.00	9.00	52.0	
				Week 48	Sparsentan	91	76 (83.5)	1.93 (14.59)	-29.0	-4.50	0.00	5.00	87.0	-0.24 [-0.58, 0.10]
					Irbesartan	88	60 (68.2)	5.20 (12.08)	-19.0	0.00	5.00	10.00	43.0	
				Week 70	Sparsentan	91	78 (85.7)	-1.45 (13.88)	-45.0	-10.00	-1.50	6.00	59.0	-0.24 [-0.57, 0.10]
					Irbesartan	88	61 (69.3)	1.92 (14.55)	-60.0	-4.00	0.00	9.00	40.0	
				Week 94	Sparsentan	91	74 (81.3)	1.16 (11.90)	-37.0	-5.00	0.00	10.00	31.0	-0.39 [-0.73, -0.05]
					Irbesartan	88	61 (69.3)	6.02 (13.19)	-41.0	-1.00	5.00	10.00	50.0	
Week 110	Sparsentan			91	73 (80.2)	0.78 (13.95)	-31.0	-5.00	1.00	9.00	70.0	-0.19 [-0.54, 0.17]		
	Irbesartan			88	54 (61.4)	3.17 (11.16)	-14.0	-4.00	2.00	9.00	39.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Geographic region													
North America	EQ-5D VAS	Baseline	Sparsentan	35	33 (94.3)	79.61 (18.57)	7.0	75.00	85.00	93.00	100.0		
			Irbesartan	46	44 (95.7)	81.18 (15.14)	34.0	75.00	82.50	92.50	100.0		
	Week 24	Sparsentan	35	24 (68.6)	86.88 (9.29)	70.0	81.00	90.00	93.50	100.0			
		Irbesartan	46	38 (82.6)	79.74 (22.03)	4.0	74.00	85.00	94.00	100.0			
	Week 48	Sparsentan	35	26 (74.3)	83.77 (12.62)	49.0	79.00	88.00	92.00	99.0			
		Irbesartan	46	35 (76.1)	83.06 (13.44)	50.0	75.00	82.00	96.00	100.0			
	Week 70	Sparsentan	35	24 (68.6)	83.42 (10.29)	60.0	76.00	85.00	90.50	100.0			
		Irbesartan	46	32 (69.6)	85.53 (14.18)	35.0	80.00	90.00	95.00	100.0			
	Week 94	Sparsentan	35	24 (68.6)	83.25 (14.03)	41.0	73.00	87.00	94.50	100.0			
		Irbesartan	46	30 (65.2)	82.90 (16.24)	34.0	80.00	84.00	95.00	100.0			
	Week 110	Sparsentan	35	21 (60.0)	85.95 (10.78)	65.0	76.00	90.00	95.00	100.0			
		Irbesartan	46	31 (67.4)	83.39 (13.26)	35.0	79.00	87.00	90.00	100.0			
		Change from baseline in EQ-5D VAS	Week 24	Sparsentan	35	24 (68.6)	4.63 (18.47)	-21.0	-2.00	0.50	9.00	75.0	0.32 [-0.20, 0.83]
				Irbesartan	46	38 (82.6)	-1.97 (21.94)	-90.0	-5.00	1.00	9.00	23.0	
	Week 48		Sparsentan	35	26 (74.3)	4.65 (15.79)	-15.0	-4.00	1.50	7.00	64.0	0.10 [-0.41, 0.61]	
			Irbesartan	46	35 (76.1)	3.17 (14.27)	-26.0	-4.00	2.00	9.00	38.0		
	Week 70		Sparsentan	35	24 (68.6)	-0.38 (9.77)	-12.0	-7.50	-2.50	2.50	24.0	-0.33 [-0.86, 0.21]	
			Irbesartan	46	32 (69.6)	3.28 (12.09)	-20.0	-3.00	1.50	8.00	40.0		
Week 94	Sparsentan		35	24 (68.6)	-0.63 (11.97)	-34.0	-6.50	-1.00	2.50	23.0	-0.24 [-0.78, 0.30]		
	Irbesartan		46	30 (65.2)	2.47 (13.52)	-41.0	-4.00	4.00	10.00	27.0			
Week 110	Sparsentan		35	21 (60.0)	3.71 (20.43)	-12.0	-4.00	1.00	4.00	88.0	0.02 [-0.53, 0.58]		
	Irbesartan		46	31 (67.4)	3.35 (12.00)	-13.0	-4.00	2.00	9.00	34.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 VAS = visual analogue scale.  
 Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Europe	EQ-5D VAS	Baseline	Sparsentan	98	91 (92.9)	80.81 (13.55)	11.0	75.00	81.00	90.00	100.0			
			Irbesartan	115	99 (86.1)	81.15 (13.69)	40.0	75.00	81.00	90.00	100.0			
		Week 24	Sparsentan	98	79 (80.6)	83.08 (11.65)	50.0	76.00	84.00	90.00	100.0			
			Irbesartan	115	68 (59.1)	82.32 (14.34)	31.0	78.00	85.00	91.50	100.0			
		Week 48	Sparsentan	98	79 (80.6)	84.33 (11.17)	50.0	79.00	85.00	92.00	100.0			
			Irbesartan	115	62 (53.9)	82.31 (14.44)	40.0	79.00	85.00	92.00	100.0			
		Week 70	Sparsentan	98	78 (79.6)	81.41 (11.11)	50.0	75.00	81.00	90.00	100.0			
			Irbesartan	115	72 (62.6)	79.76 (17.61)	20.0	70.50	83.50	93.00	100.0			
		Week 94	Sparsentan	98	72 (73.5)	81.53 (14.17)	34.0	76.50	83.00	91.00	100.0			
			Irbesartan	115	73 (63.5)	82.18 (14.91)	39.0	72.00	87.00	91.00	100.0			
		Week 110	Sparsentan	98	70 (71.4)	81.69 (10.94)	48.0	78.00	81.00	90.00	100.0			
			Irbesartan	115	61 (53.0)	80.46 (13.44)	40.0	74.00	81.00	90.00	100.0			
		Change from baseline in EQ-5D VAS	EQ-5D VAS	Week 24	Sparsentan	98	79 (80.6)	1.76 (12.73)	-31.0	-5.00	1.00	9.00	40.0	0.03 [-0.29, 0.36]
					Irbesartan	115	68 (59.1)	1.34 (13.05)	-42.0	-5.50	0.00	10.00	52.0	
				Week 48	Sparsentan	98	79 (80.6)	2.75 (14.61)	-30.0	-3.00	1.00	10.00	87.0	0.03 [-0.31, 0.36]
					Irbesartan	115	62 (53.9)	2.39 (11.64)	-35.0	-4.00	1.00	8.00	43.0	
				Week 70	Sparsentan	98	78 (79.6)	0.15 (12.89)	-32.0	-9.00	0.00	8.00	59.0	0.03 [-0.29, 0.35]
					Irbesartan	115	72 (62.6)	-0.28 (15.03)	-60.0	-6.00	0.00	8.00	40.0	
				Week 94	Sparsentan	98	72 (73.5)	1.25 (15.27)	-61.0	-4.00	3.00	10.00	45.0	-0.07 [-0.40, 0.26]
					Irbesartan	115	73 (63.5)	2.33 (15.31)	-38.0	-4.00	1.00	10.00	50.0	
Week 110	Sparsentan			98	70 (71.4)	1.60 (14.89)	-40.0	-7.00	1.00	10.00	70.0	0.02 [-0.32, 0.36]		
	Irbesartan			115	61 (53.0)	1.34 (12.58)	-26.0	-5.00	0.00	10.00	39.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Asia Pacific	EQ-5D VAS	Baseline	Sparsentan	69	67 (97.1)	80.46 (12.87)	40.0	73.00	81.00	90.00	100.0		
			Irbesartan	41	41 (100.0)	77.68 (14.49)	19.0	72.00	80.00	85.00	100.0		
	Week 24	Sparsentan	69	61 (88.4)	84.59 (9.79)	60.0	80.00	87.00	91.00	100.0			
		Irbesartan	41	37 (90.2)	80.22 (13.26)	30.0	79.00	80.00	90.00	100.0			
	Week 48	Sparsentan	69	63 (91.3)	81.71 (10.22)	49.0	79.00	81.00	90.00	100.0			
		Irbesartan	41	35 (85.4)	81.49 (10.97)	54.0	75.00	83.00	90.00	100.0			
	Week 70	Sparsentan	69	64 (92.8)	80.45 (13.28)	27.0	75.00	81.00	90.00	100.0			
		Irbesartan	41	32 (78.0)	82.09 (11.60)	40.0	78.00	83.50	90.00	100.0			
	Week 94	Sparsentan	69	60 (87.0)	82.28 (11.50)	40.0	80.00	83.50	90.00	100.0			
		Irbesartan	41	33 (80.5)	82.33 (11.06)	40.0	79.00	82.00	90.00	100.0			
	Week 110	Sparsentan	69	61 (88.4)	81.98 (12.18)	48.0	79.00	82.00	90.00	100.0			
		Irbesartan	41	29 (70.7)	82.90 (9.03)	60.0	80.00	83.00	90.00	100.0			
	Change from baseline in EQ-5D VAS		Week 24	Sparsentan	69	61 (88.4)	4.49 (13.91)	-25.0	-3.00	5.00	12.00	44.0	0.10 [-0.31, 0.51]
				Irbesartan	41	37 (90.2)	3.11 (13.07)	-37.0	-1.00	0.00	10.00	34.0	
			Week 48	Sparsentan	69	63 (91.3)	1.00 (13.38)	-29.0	-8.00	0.00	10.00	46.0	-0.10 [-0.51, 0.31]
				Irbesartan	41	35 (85.4)	2.29 (11.86)	-31.0	-4.00	5.00	10.00	25.0	
			Week 70	Sparsentan	69	64 (92.8)	0.13 (15.54)	-51.0	-6.00	0.00	9.00	41.0	-0.20 [-0.63, 0.22]
				Irbesartan	41	32 (78.0)	2.88 (8.43)	-15.0	-3.00	0.00	10.00	18.0	
			Week 94	Sparsentan	69	60 (87.0)	1.75 (15.13)	-37.0	-6.00	0.00	8.50	44.0	-0.15 [-0.58, 0.27]
				Irbesartan	41	33 (80.5)	4.00 (14.55)	-45.0	-1.00	5.00	10.00	32.0	
Week 110			Sparsentan	69	61 (88.4)	0.93 (13.60)	-31.0	-4.00	1.00	9.00	36.0	-0.24 [-0.68, 0.20]	
			Irbesartan	41	29 (70.7)	3.86 (8.52)	-10.0	0.00	2.00	8.00	30.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline BMI														
< 27 kg/m**2	EQ-5D VAS	Baseline	Sparsentan	83	77 (92.8)	78.96 (14.98)	11.0	72.00	81.00	89.00	100.0			
			Irbesartan	94	87 (92.6)	84.05 (10.58)	52.0	77.00	83.00	91.00	100.0			
		Week 24	Sparsentan	83	68 (81.9)	82.84 (11.69)	51.0	74.00	82.50	90.00	100.0			
			Irbesartan	94	72 (76.6)	83.15 (13.29)	31.0	79.00	85.00	91.00	100.0			
		Week 48	Sparsentan	83	69 (83.1)	81.17 (11.72)	49.0	77.00	81.00	90.00	100.0			
			Irbesartan	94	63 (67.0)	83.59 (12.41)	40.0	80.00	85.00	91.00	100.0			
		Week 70	Sparsentan	83	68 (81.9)	79.29 (13.34)	27.0	70.00	80.50	90.00	100.0			
			Irbesartan	94	63 (67.0)	82.10 (15.27)	20.0	75.00	85.00	92.00	100.0			
		Week 94	Sparsentan	83	65 (78.3)	79.58 (13.65)	40.0	72.00	82.00	89.00	100.0			
			Irbesartan	94	65 (69.1)	83.15 (13.09)	49.0	76.00	85.00	92.00	100.0			
		Week 110	Sparsentan	83	64 (77.1)	80.84 (12.13)	48.0	75.50	81.00	90.00	100.0			
			Irbesartan	94	57 (60.6)	81.61 (12.20)	47.0	75.00	81.00	90.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	83	68 (81.9)	2.46 (12.39)	-25.0	-3.50	0.50	9.50	40.0	0.23 [-0.10, 0.56]
					Irbesartan	94	72 (76.6)	-0.19 (10.85)	-42.0	-5.00	-0.50	8.50	19.0	
				Week 48	Sparsentan	83	69 (83.1)	1.71 (16.66)	-30.0	-6.00	0.00	9.00	87.0	0.13 [-0.21, 0.47]
					Irbesartan	94	63 (67.0)	-0.13 (11.29)	-35.0	-5.00	0.00	7.00	27.0	
				Week 70	Sparsentan	83	68 (81.9)	-0.22 (15.54)	-51.0	-7.50	0.00	8.00	59.0	0.10 [-0.25, 0.44]
					Irbesartan	94	63 (67.0)	-1.59 (12.49)	-60.0	-5.00	0.00	5.00	29.0	
				Week 94	Sparsentan	83	65 (78.3)	0.15 (14.08)	-37.0	-7.00	0.00	7.00	44.0	0.04 [-0.30, 0.39]
					Irbesartan	94	65 (69.1)	-0.40 (12.54)	-38.0	-5.00	0.00	6.00	38.0	
		Week 110	Sparsentan	83	64 (77.1)	0.75 (15.04)	-31.0	-6.50	0.00	6.00	70.0	0.17 [-0.19, 0.52]		
			Irbesartan	94	57 (60.6)	-1.40 (10.03)	-26.0	-7.00	0.00	2.00	28.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
>= 27 kg/m**2	EQ-5D VAS	Baseline	Sparsentan	119	114 (95.8)	81.51 (13.69)	7.0	75.00	82.00	91.00	100.0			
			Irbesartan	107	96 (89.7)	77.49 (15.82)	19.0	72.50	80.00	88.50	100.0			
		Week 24	Sparsentan	119	96 (80.7)	85.16 (9.86)	50.0	80.00	88.00	91.00	100.0			
			Irbesartan	107	71 (66.4)	79.00 (18.95)	4.0	73.00	81.00	91.00	100.0			
		Week 48	Sparsentan	119	99 (83.2)	84.72 (10.40)	50.0	80.00	86.00	91.00	100.0			
			Irbesartan	107	69 (64.5)	81.10 (13.96)	50.0	74.00	83.00	90.00	100.0			
		Week 70	Sparsentan	119	98 (82.4)	82.74 (10.55)	50.0	79.00	82.00	90.00	100.0			
			Irbesartan	107	73 (68.2)	81.30 (16.10)	20.0	75.00	85.00	93.00	100.0			
		Week 94	Sparsentan	119	91 (76.5)	83.87 (12.48)	34.0	80.00	87.00	92.00	100.0			
			Irbesartan	107	70 (65.4)	81.63 (15.48)	34.0	78.00	84.00	90.00	100.0			
		Week 110	Sparsentan	119	88 (73.9)	83.52 (10.86)	48.0	78.50	84.50	90.00	100.0			
			Irbesartan	107	63 (58.9)	82.40 (12.37)	35.0	79.00	85.00	90.00	100.0			
		Change from baseline in EQ-5D VAS		Week 24	Sparsentan	119	96 (80.7)	3.72 (15.23)	-31.0	-4.50	1.00	10.00	75.0	0.10 [-0.21, 0.40]
					Irbesartan	107	71 (66.4)	2.04 (19.77)	-90.0	-4.00	2.00	13.00	52.0	
				Week 48	Sparsentan	119	99 (83.2)	2.86 (12.50)	-29.0	-3.00	1.00	10.00	64.0	-0.17 [-0.48, 0.14]
					Irbesartan	107	69 (64.5)	5.03 (12.83)	-22.0	-1.00	5.00	10.00	43.0	
				Week 70	Sparsentan	119	98 (82.4)	0.27 (12.02)	-32.0	-8.00	0.00	8.00	40.0	-0.28 [-0.59, 0.02]
					Irbesartan	107	73 (68.2)	3.79 (13.18)	-49.0	-4.00	4.00	11.00	40.0	
				Week 94	Sparsentan	119	91 (76.5)	1.87 (15.15)	-61.0	-4.00	2.00	11.00	45.0	-0.21 [-0.53, 0.10]
					Irbesartan	107	70 (65.4)	5.11 (15.33)	-45.0	-3.00	5.00	12.00	50.0	
Week 110	Sparsentan			119	88 (73.9)	2.26 (15.36)	-40.0	-4.00	1.00	9.00	88.0	-0.25 [-0.58, 0.07]		
	Irbesartan			107	63 (58.9)	5.78 (11.84)	-21.0	-2.00	5.00	11.00	39.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024



Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Randomization strata														
eGFR Low and UP High	EQ-5D VAS	Baseline	Sparsentan	71	66 (93.0)	82.14 (10.50)	51.0	79.00	81.50	90.00	100.0			
			Irbesartan	74	64 (86.5)	78.67 (15.16)	37.0	75.00	80.00	87.50	100.0			
		Week 24	Sparsentan	71	54 (76.1)	83.89 (11.03)	50.0	76.00	85.00	91.00	100.0			
			Irbesartan	74	47 (63.5)	77.17 (19.06)	4.0	70.00	81.00	90.00	100.0			
		Week 48	Sparsentan	71	55 (77.5)	82.80 (10.43)	50.0	78.00	81.00	90.00	100.0			
			Irbesartan	74	41 (55.4)	82.27 (14.52)	50.0	79.00	85.00	90.00	100.0			
		Week 70	Sparsentan	71	58 (81.7)	79.91 (11.62)	50.0	72.00	80.50	89.00	100.0			
			Irbesartan	74	41 (55.4)	79.24 (15.87)	20.0	73.00	82.00	90.00	100.0			
		Week 94	Sparsentan	71	52 (73.2)	83.12 (12.31)	50.0	76.00	84.50	93.00	100.0			
			Irbesartan	74	40 (54.1)	79.50 (17.24)	34.0	75.00	84.00	90.50	100.0			
		Week 110	Sparsentan	71	53 (74.6)	82.21 (13.61)	48.0	75.00	84.00	91.00	100.0			
			Irbesartan	74	36 (48.6)	78.42 (14.19)	40.0	71.00	81.00	89.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	71	54 (76.1)	0.35 (11.30)	-31.0	-5.00	0.00	8.00	35.0	0.19 [-0.20, 0.58]
					Irbesartan	74	47 (63.5)	-2.57 (18.90)	-71.0	-10.00	-1.00	8.00	52.0	
				Week 48	Sparsentan	71	55 (77.5)	-0.51 (9.33)	-29.0	-6.00	-1.00	4.00	28.0	-0.43 [-0.84, -0.02]
					Irbesartan	74	41 (55.4)	4.78 (15.43)	-31.0	-1.00	3.00	10.00	43.0	
				Week 70	Sparsentan	71	58 (81.7)	-3.64 (10.26)	-32.0	-10.00	-2.50	4.00	19.0	-0.31 [-0.71, 0.10]
					Irbesartan	74	41 (55.4)	0.46 (16.94)	-60.0	-6.00	0.00	8.00	40.0	
				Week 94	Sparsentan	71	52 (73.2)	-0.04 (12.91)	-37.0	-4.50	0.50	7.50	45.0	-0.17 [-0.58, 0.24]
					Irbesartan	74	40 (54.1)	2.70 (19.40)	-45.0	-5.00	0.50	13.50	50.0	
		Week 110	Sparsentan	71	53 (74.6)	-0.51 (12.27)	-40.0	-3.00	0.00	5.00	35.0	-0.24 [-0.66, 0.18]		
			Irbesartan	74	36 (48.6)	2.69 (14.82)	-26.0	-4.50	1.50	9.00	39.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR Low and UP Low	EQ-5D VAS	Baseline	Sparsentan	55	52 (94.5)	79.10 (17.39)	11.0	71.00	81.00	90.00	100.0			
			Irbesartan	55	51 (92.7)	81.61 (13.00)	34.0	74.00	80.00	90.00	100.0			
		Week 24	Sparsentan	55	46 (83.6)	82.74 (10.57)	51.0	78.00	82.00	90.00	100.0			
			Irbesartan	55	40 (72.7)	81.20 (16.32)	10.0	80.00	82.00	90.00	100.0			
		Week 48	Sparsentan	55	43 (78.2)	83.26 (10.87)	49.0	80.00	83.00	91.00	100.0			
			Irbesartan	55	38 (69.1)	82.11 (13.07)	50.0	80.00	83.00	91.00	100.0			
		Week 70	Sparsentan	55	44 (80.0)	83.20 (13.03)	40.0	79.50	86.50	91.00	100.0			
			Irbesartan	55	41 (74.5)	80.56 (16.86)	20.0	75.00	83.00	92.00	100.0			
		Week 94	Sparsentan	55	42 (76.4)	81.79 (14.41)	40.0	80.00	84.00	91.00	100.0			
			Irbesartan	55	40 (72.7)	81.90 (13.78)	39.0	76.00	83.00	90.00	100.0			
		Week 110	Sparsentan	55	38 (69.1)	80.87 (10.90)	59.0	75.00	81.50	88.00	100.0			
			Irbesartan	55	34 (61.8)	81.26 (13.23)	35.0	76.00	82.00	90.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	55	46 (83.6)	3.70 (14.17)	-25.0	-4.00	0.00	10.00	44.0	0.20 [-0.22, 0.63]
		Irbesartan			55	40 (72.7)	0.45 (17.72)	-90.0	-2.00	1.00	10.00	23.0		
		Week 48		Sparsentan	55	43 (78.2)	3.53 (17.62)	-22.0	-2.00	1.00	10.00	87.0	0.10 [-0.34, 0.53]	
				Irbesartan	55	38 (69.1)	2.13 (10.45)	-22.0	-5.00	2.50	10.00	23.0		
		Week 70		Sparsentan	55	44 (80.0)	3.50 (16.72)	-45.0	-6.00	1.50	10.50	59.0	0.18 [-0.24, 0.61]	
				Irbesartan	55	41 (74.5)	0.80 (12.56)	-49.0	-5.00	0.00	9.00	21.0		
		Week 94		Sparsentan	55	42 (76.4)	3.60 (14.17)	-34.0	-4.00	1.00	12.00	40.0	0.10 [-0.34, 0.53]	
				Irbesartan	55	40 (72.7)	2.38 (11.16)	-22.0	-3.50	3.50	10.00	24.0		
Week 110	Sparsentan	55		38 (69.1)	0.84 (16.12)	-28.0	-8.00	0.50	6.00	70.0	-0.08 [-0.54, 0.39]			
	Irbesartan	55		34 (61.8)	1.91 (10.46)	-21.0	-4.00	0.50	11.00	20.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP High	EQ-5D VAS	Baseline	Sparsentan	37	35 (94.6)	78.66 (18.53)	7.0	70.00	82.00	91.00	100.0			
			Irbesartan	36	33 (91.7)	81.24 (11.45)	50.0	73.00	80.00	90.00	100.0			
		Week 24	Sparsentan	37	29 (78.4)	83.62 (11.12)	60.0	78.00	86.00	90.00	100.0			
			Irbesartan	36	27 (75.0)	85.44 (10.56)	69.0	72.00	90.00	95.00	99.0			
		Week 48	Sparsentan	37	34 (91.9)	81.94 (11.61)	50.0	78.00	83.00	89.00	100.0			
			Irbesartan	36	25 (69.4)	82.24 (14.51)	40.0	80.00	85.00	90.00	100.0			
		Week 70	Sparsentan	37	31 (83.8)	81.84 (9.82)	59.0	75.00	81.00	90.00	98.0			
			Irbesartan	36	25 (69.4)	81.80 (14.83)	40.0	71.00	85.00	91.00	100.0			
		Week 94	Sparsentan	37	27 (73.0)	80.52 (13.44)	34.0	78.00	84.00	90.00	96.0			
			Irbesartan	36	25 (69.4)	85.00 (11.80)	58.0	79.00	87.00	95.00	100.0			
		Week 110	Sparsentan	37	26 (70.3)	84.88 (8.44)	60.0	80.00	84.50	90.00	100.0			
			Irbesartan	36	22 (61.1)	83.18 (10.65)	60.0	79.00	86.50	90.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	37	29 (78.4)	4.52 (18.92)	-25.0	-2.00	5.00	8.00	75.0	-0.07 [-0.60, 0.45]
		Irbesartan			36	27 (75.0)	5.63 (11.02)	-21.0	-1.00	5.00	11.00	34.0		
		Week 48		Sparsentan	37	34 (91.9)	3.29 (18.43)	-30.0	-9.00	2.00	11.00	64.0	0.02 [-0.49, 0.54]	
				Irbesartan	36	25 (69.4)	2.92 (11.91)	-35.0	0.00	5.00	9.00	25.0		
		Week 70		Sparsentan	37	31 (83.8)	2.06 (12.41)	-20.0	-6.00	1.00	10.00	41.0	0.04 [-0.49, 0.56]	
				Irbesartan	36	25 (69.4)	1.68 (8.74)	-20.0	-3.00	3.00	8.00	18.0		
		Week 94		Sparsentan	37	27 (73.0)	-0.41 (19.17)	-61.0	-8.00	1.00	10.00	44.0	-0.34 [-0.89, 0.21]	
				Irbesartan	36	25 (69.4)	4.92 (10.55)	-11.0	-2.00	2.00	10.00	32.0		
Week 110	Sparsentan	37		26 (70.3)	6.85 (21.11)	-30.0	-5.00	4.50	14.00	88.0	0.23 [-0.34, 0.80]			
	Irbesartan	36		22 (61.1)	2.95 (9.88)	-13.0	-3.00	4.50	9.00	30.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP Low	EQ-5D VAS	Baseline	Sparsentan	39	38 (97.4)	81.18 (10.21)	50.0	75.00	80.50	90.00	100.0			
			Irbesartan	37	36 (97.3)	80.92 (16.54)	19.0	75.00	84.00	90.00	100.0			
		Week 24	Sparsentan	39	35 (89.7)	87.06 (9.81)	64.0	80.00	89.00	91.00	100.0			
			Irbesartan	37	29 (78.4)	83.24 (15.72)	30.0	80.00	84.00	95.00	100.0			
		Week 48	Sparsentan	39	36 (92.3)	85.22 (11.90)	49.0	80.00	87.50	93.00	100.0			
			Irbesartan	37	28 (75.7)	82.61 (10.87)	60.0	75.00	80.00	90.00	100.0			
		Week 70	Sparsentan	39	33 (84.6)	80.85 (12.53)	27.0	77.00	80.00	90.00	100.0			
			Irbesartan	37	29 (78.4)	86.55 (13.95)	40.0	80.00	86.00	100.00	100.0			
		Week 94	Sparsentan	39	35 (89.7)	82.11 (12.80)	49.0	77.00	82.00	92.00	100.0			
			Irbesartan	37	31 (83.8)	84.58 (12.35)	50.0	80.00	86.00	94.00	100.0			
		Week 110	Sparsentan	39	35 (89.7)	82.49 (10.49)	48.0	78.00	81.00	90.00	98.0			
			Irbesartan	37	29 (78.4)	85.55 (9.54)	60.0	80.00	85.00	92.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	39	35 (89.7)	5.83 (13.10)	-20.0	-1.00	5.00	18.00	29.0	0.25 [-0.24, 0.75]
					Irbesartan	37	29 (78.4)	2.83 (9.93)	-10.0	-4.00	0.00	9.00	30.0	
			Week 48	Sparsentan	39	36 (92.3)	4.58 (11.45)	-29.0	-1.00	3.00	11.50	27.0	0.46 [-0.04, 0.96]	
				Irbesartan	37	28 (75.7)	-0.39 (9.73)	-25.0	-5.00	-0.50	5.00	27.0		
			Week 70	Sparsentan	39	33 (84.6)	0.12 (13.85)	-51.0	-7.00	2.00	8.00	25.0	-0.22 [-0.72, 0.28]	
				Irbesartan	37	29 (78.4)	2.86 (11.06)	-30.0	-1.00	0.00	10.00	29.0		
			Week 94	Sparsentan	39	35 (89.7)	1.20 (14.13)	-31.0	-7.00	3.00	11.00	30.0	-0.03 [-0.51, 0.45]	
				Irbesartan	37	31 (83.8)	1.61 (14.89)	-38.0	-5.00	0.00	10.00	38.0		
	Week 110	Sparsentan	39	35 (89.7)	1.83 (12.47)	-30.0	-4.00	3.00	11.00	30.0	-0.05 [-0.55, 0.44]			
		Irbesartan	37	29 (78.4)	2.45 (9.70)	-13.0	-5.00	0.00	9.00	28.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 1														
< 60 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	127	119 (93.7)	80.61 (13.81)	11.0	75.00	81.00	90.00	100.0			
			Irbesartan	129	116 (89.9)	79.86 (13.70)	34.0	75.00	80.00	89.50	100.0			
		Week 24	Sparsentan	127	102 (80.3)	83.27 (10.57)	50.0	78.00	84.00	91.00	100.0			
			Irbesartan	129	89 (69.0)	78.97 (17.68)	4.0	74.00	81.00	90.00	100.0			
		Week 48	Sparsentan	127	99 (78.0)	83.22 (10.65)	49.0	80.00	83.00	91.00	100.0			
			Irbesartan	129	81 (62.8)	81.89 (13.60)	50.0	79.00	83.00	91.00	100.0			
		Week 70	Sparsentan	127	103 (81.1)	81.17 (12.12)	40.0	75.00	82.00	90.00	100.0			
			Irbesartan	129	84 (65.1)	79.49 (16.70)	20.0	74.00	82.50	90.50	100.0			
		Week 94	Sparsentan	127	94 (74.0)	82.18 (12.83)	40.0	77.00	83.00	91.00	100.0			
			Irbesartan	129	82 (63.6)	81.46 (15.20)	34.0	78.00	84.00	90.00	100.0			
		Week 110	Sparsentan	127	93 (73.2)	81.84 (12.27)	48.0	75.00	82.00	90.00	100.0			
			Irbesartan	129	72 (55.8)	79.99 (13.40)	35.0	73.00	81.00	90.00	100.0			
		Change from baseline in EQ-5D VAS		Week 24	Sparsentan	127	102 (80.3)	2.09 (12.72)	-31.0	-5.00	0.00	10.00	44.0	0.16 [-0.13, 0.44]
					Irbesartan	129	89 (69.0)	-0.36 (18.30)	-90.0	-6.00	0.00	10.00	52.0	
				Week 48	Sparsentan	127	99 (78.0)	1.72 (13.82)	-29.0	-4.00	0.00	7.00	87.0	-0.13 [-0.43, 0.16]
					Irbesartan	129	81 (62.8)	3.49 (12.90)	-31.0	-1.00	3.00	10.00	43.0	
				Week 70	Sparsentan	127	103 (81.1)	-0.50 (13.73)	-45.0	-9.00	0.00	7.00	59.0	-0.07 [-0.36, 0.21]
					Irbesartan	129	84 (65.1)	0.55 (14.73)	-60.0	-6.00	0.00	8.50	40.0	
Week 94	Sparsentan			127	94 (74.0)	1.49 (13.42)	-37.0	-4.00	1.00	8.00	45.0	-0.11 [-0.41, 0.18]		
	Irbesartan			129	82 (63.6)	3.15 (15.68)	-45.0	-4.00	3.50	10.00	50.0			
Week 110	Sparsentan			127	93 (73.2)	0.29 (13.79)	-40.0	-5.00	1.00	6.00	70.0	-0.15 [-0.46, 0.16]		
	Irbesartan			129	72 (55.8)	2.31 (12.76)	-26.0	-4.50	1.00	10.00	39.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	49	47 (95.9)	80.83 (16.08)	7.0	75.00	82.00	90.00	100.0		
		Week 24	Irbesartan	48	44 (91.7)	80.59 (13.76)	37.0	71.50	81.00	90.00	100.0		
			Sparsentan	49	40 (81.6)	85.60 (11.14)	60.0	78.00	89.00	93.00	100.0		
		Week 48	Irbesartan	48	34 (70.8)	85.21 (10.00)	65.0	80.00	84.50	95.00	100.0		
			Sparsentan	49	45 (91.8)	83.89 (10.93)	49.0	79.00	85.00	90.00	100.0		
		Week 70	Irbesartan	48	33 (68.8)	82.12 (13.83)	40.0	75.00	85.00	90.00	100.0		
			Sparsentan	49	40 (81.6)	81.73 (13.23)	27.0	75.00	81.00	90.00	100.0		
		Week 94	Irbesartan	48	32 (66.7)	83.22 (14.99)	40.0	72.50	86.00	95.50	100.0		
			Sparsentan	49	41 (83.7)	80.63 (14.95)	34.0	73.00	82.00	92.00	100.0		
		Week 110	Irbesartan	48	35 (72.9)	82.51 (13.87)	50.0	70.00	85.00	95.00	100.0		
			Sparsentan	49	37 (75.5)	82.38 (11.13)	48.0	78.00	82.00	90.00	98.0		
		Change from baseline in EQ-5D VAS	Week 24	Irbesartan	48	34 (70.8)	3.09 (10.97)	-11.0	-4.00	-0.50	9.00	34.0	
				Sparsentan	49	45 (91.8)	3.44 (14.86)	-29.0	-4.00	2.00	11.00	64.0	0.09 [-0.37, 0.54]
	Week 48		Irbesartan	48	33 (68.8)	1.52 (11.56)	-35.0	-5.00	0.00	9.00	25.0		
			Sparsentan	49	40 (81.6)	-0.13 (12.99)	-51.0	-6.00	0.00	8.00	23.0	0.14 [-0.31, 0.59]	
	Week 70		Irbesartan	48	32 (66.7)	1.31 (10.39)	-30.0	-4.00	0.00	10.00	18.0		
			Sparsentan	49	41 (83.7)	-1.27 (16.87)	-61.0	-8.00	0.00	10.00	30.0	-0.12 [-0.59, 0.34]	
	Week 94		Irbesartan	48	35 (72.9)	2.40 (12.46)	-33.0	-4.00	0.00	10.00	31.0		
		Sparsentan	49	37 (75.5)	2.92 (19.21)	-30.0	-5.00	1.00	9.00	88.0	-0.24 [-0.70, 0.21]		
	Week 110	Irbesartan	48	32 (66.7)	1.69 (10.06)	-13.0	-5.00	0.00	8.00	27.0			
Sparsentan		49	37 (75.5)	2.92 (19.21)	-30.0	-5.00	1.00	9.00	88.0	0.08 [-0.39, 0.55]			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	26	25 (96.2)	79.20 (13.04)	40.0	71.00	81.00	89.00	97.0		
			Irbesartan	25	24 (96.0)	82.54 (17.58)	19.0	78.00	86.00	92.50	100.0		
		Week 24	Sparsentan	26	22 (84.6)	85.91 (10.32)	60.0	80.00	88.50	91.00	100.0		
			Irbesartan	25	20 (80.0)	83.55 (18.30)	30.0	74.50	90.50	95.50	100.0		
		Week 48	Sparsentan	26	24 (92.3)	82.25 (13.24)	50.0	77.50	84.50	92.00	100.0		
			Irbesartan	25	18 (72.0)	84.39 (10.84)	66.0	75.00	80.50	92.00	100.0		
		Week 70	Sparsentan	26	23 (88.5)	81.39 (7.95)	66.0	77.00	80.00	89.00	95.0		
			Irbesartan	25	20 (80.0)	88.35 (9.39)	70.0	80.50	90.00	97.00	100.0		
		Week 94	Sparsentan	26	21 (80.8)	84.48 (10.47)	50.0	80.00	85.00	90.00	97.0		
			Irbesartan	25	19 (76.0)	86.05 (10.49)	62.0	80.00	90.00	94.00	100.0		
		Week 110	Sparsentan	26	22 (84.6)	84.77 (7.95)	62.0	81.00	84.50	90.00	100.0		
			Irbesartan	25	17 (68.0)	87.94 (8.07)	70.0	84.00	90.00	91.00	100.0		
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	26	22 (84.6)	6.23 (14.25)	-20.0	-1.00	5.50	16.00	40.0
				Irbesartan	25	20 (80.0)	2.90 (10.39)	-21.0	-4.00	2.50	10.50	21.0	
			Week 48	Sparsentan	26	24 (92.3)	3.17 (15.77)	-30.0	-5.50	3.50	10.50	46.0	0.20 [-0.41, 0.81]
				Irbesartan	25	18 (72.0)	0.33 (11.36)	-25.0	-4.00	0.00	5.00	27.0	
			Week 70	Sparsentan	26	23 (88.5)	2.96 (13.77)	-22.0	-6.00	5.00	9.00	41.0	-0.13 [-0.73, 0.47]
				Irbesartan	25	20 (80.0)	4.45 (9.07)	-12.0	-1.00	4.00	10.00	29.0	
			Week 94	Sparsentan	26	21 (80.8)	4.38 (15.59)	-30.0	-4.00	5.00	11.00	44.0	0.17 [-0.45, 0.79]
				Irbesartan	25	19 (76.0)	1.79 (14.48)	-38.0	-3.00	0.00	9.00	38.0	
		Week 110	Sparsentan	26	22 (84.6)	5.09 (13.09)	-19.0	-4.00	5.50	11.00	36.0	0.04 [-0.59, 0.68]	
			Irbesartan	25	17 (68.0)	4.59 (8.70)	-7.0	-2.00	5.00	9.00	28.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 2														
< 45 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	82	75 (91.5)	79.11 (15.04)	11.0	71.00	81.00	88.00	100.0			
			Irbesartan	80	70 (87.5)	77.90 (14.78)	34.0	73.00	80.00	85.00	100.0			
		Week 24	Sparsentan	82	64 (78.0)	83.00 (10.89)	50.0	79.00	83.00	90.50	100.0			
			Irbesartan	80	52 (65.0)	75.65 (18.49)	4.0	70.00	80.00	90.00	99.0			
		Week 48	Sparsentan	82	63 (76.8)	83.68 (10.95)	49.0	80.00	84.00	91.00	100.0			
			Irbesartan	80	48 (60.0)	80.73 (14.16)	50.0	79.00	83.00	89.50	100.0			
		Week 70	Sparsentan	82	65 (79.3)	81.11 (13.00)	40.0	75.00	81.00	90.00	100.0			
			Irbesartan	80	50 (62.5)	75.88 (17.61)	20.0	70.00	79.50	89.00	100.0			
		Week 94	Sparsentan	82	61 (74.4)	82.08 (13.61)	40.0	76.00	83.00	91.00	100.0			
			Irbesartan	80	50 (62.5)	78.78 (17.14)	34.0	76.00	81.50	90.00	100.0			
		Week 110	Sparsentan	82	57 (69.5)	81.21 (13.67)	48.0	72.00	82.00	91.00	100.0			
			Irbesartan	80	44 (55.0)	77.16 (14.25)	35.0	71.00	80.00	87.50	100.0			
		Change from baseline in EQ-5D VAS		Week 24	Sparsentan	82	64 (78.0)	3.19 (12.42)	-29.0	-4.50	0.50	10.00	44.0	0.24 [-0.13, 0.60]
					Irbesartan	80	52 (65.0)	-0.52 (18.91)	-71.0	-8.00	0.00	10.00	52.0	
				Week 48	Sparsentan	82	63 (76.8)	4.00 (14.80)	-29.0	-2.00	1.00	7.00	87.0	-0.03 [-0.41, 0.34]
					Irbesartan	80	48 (60.0)	4.50 (14.10)	-31.0	0.00	5.00	10.00	43.0	
				Week 70	Sparsentan	82	65 (79.3)	0.72 (14.75)	-45.0	-7.00	0.00	7.00	59.0	0.11 [-0.26, 0.48]
					Irbesartan	80	50 (62.5)	-1.08 (17.13)	-60.0	-10.00	0.00	8.00	40.0	
				Week 94	Sparsentan	82	61 (74.4)	2.56 (13.69)	-37.0	-3.00	1.00	9.00	45.0	0.02 [-0.35, 0.39]
					Irbesartan	80	50 (62.5)	2.24 (17.85)	-45.0	-6.00	4.50	10.00	50.0	
Week 110	Sparsentan			82	57 (69.5)	0.72 (15.57)	-40.0	-6.00	1.00	7.00	70.0	-0.10 [-0.49, 0.29]		
	Irbesartan			80	44 (55.0)	2.18 (13.55)	-26.0	-4.50	1.00	11.00	39.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024



Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
45 to < 60 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	45	44 (97.8)	83.18 (11.11)	50.0	78.50	81.50	91.00	100.0	
		Week 24	Irbesartan	49	46 (93.9)	82.85 (11.40)	50.0	75.00	82.00	90.00	100.0	
			Sparsentan	45	38 (84.4)	83.74 (10.14)	59.0	75.00	86.00	91.00	100.0	
		Week 48	Irbesartan	49	37 (75.5)	83.62 (15.56)	10.0	80.00	85.00	91.00	100.0	
			Sparsentan	45	36 (80.0)	82.42 (10.19)	55.0	76.50	81.00	90.50	100.0	
		Week 70	Irbesartan	49	33 (67.3)	83.58 (12.75)	51.0	80.00	83.00	94.00	100.0	
			Sparsentan	45	38 (84.4)	81.26 (10.59)	60.0	75.00	82.00	90.00	100.0	
		Week 94	Irbesartan	49	34 (69.4)	84.79 (13.86)	40.0	80.00	90.00	95.00	100.0	
			Sparsentan	45	33 (73.3)	82.36 (11.43)	49.0	80.00	85.00	91.00	97.0	
		Week 110	Irbesartan	49	32 (65.3)	85.66 (10.48)	55.0	80.00	88.50	93.00	100.0	
			Sparsentan	45	36 (80.0)	82.83 (9.76)	60.0	75.50	82.00	90.00	100.0	
		Change from baseline in EQ-5D VAS	Week 24	Sparsentan	45	38 (84.4)	0.24 (13.17)	-31.0	-9.00	0.00	8.00	35.0
	Irbesartan			49	37 (75.5)	-0.14 (17.67)	-90.0	-3.00	1.00	9.00	22.0	
	Week 48		Sparsentan	45	36 (80.0)	-2.28 (10.99)	-23.0	-9.50	-1.00	2.00	19.0	-0.39 [-0.87, 0.08]
			Irbesartan	49	33 (67.3)	2.03 (10.98)	-22.0	-5.00	0.00	9.00	37.0	
	Week 70		Sparsentan	45	38 (84.4)	-2.61 (11.69)	-32.0	-12.00	-1.50	7.00	19.0	-0.51 [-0.98, -0.04]
			Irbesartan	49	34 (69.4)	2.94 (10.01)	-15.0	-4.00	1.50	10.00	27.0	
	Week 94		Sparsentan	45	33 (73.3)	-0.48 (12.86)	-31.0	-6.00	1.00	7.00	30.0	-0.41 [-0.90, 0.08]
			Irbesartan	49	32 (65.3)	4.56 (11.66)	-16.0	-2.50	2.50	10.50	32.0	
	Week 110	Sparsentan	45	36 (80.0)	-0.39 (10.54)	-28.0	-4.50	0.00	4.50	22.0	-0.26 [-0.76, 0.23]	
Irbesartan		49	28 (57.1)	2.50 (11.63)	-15.0	-5.50	1.50	8.50	30.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
60 to < 90 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	49	47 (95.9)	80.83 (16.08)	7.0	75.00	82.00	90.00	100.0			
			Irbesartan	48	44 (91.7)	80.59 (13.76)	37.0	71.50	81.00	90.00	100.0			
		Week 24	Sparsentan	49	40 (81.6)	85.60 (11.14)	60.0	78.00	89.00	93.00	100.0			
			Irbesartan	48	34 (70.8)	85.21 (10.00)	65.0	80.00	84.50	95.00	100.0			
		Week 48	Sparsentan	49	45 (91.8)	83.89 (10.93)	49.0	79.00	85.00	90.00	100.0			
			Irbesartan	48	33 (68.8)	82.12 (13.83)	40.0	75.00	85.00	90.00	100.0			
		Week 70	Sparsentan	49	40 (81.6)	81.73 (13.23)	27.0	75.00	81.00	90.00	100.0			
			Irbesartan	48	32 (66.7)	83.22 (14.99)	40.0	72.50	86.00	95.50	100.0			
		Week 94	Sparsentan	49	41 (83.7)	80.63 (14.95)	34.0	73.00	82.00	92.00	100.0			
			Irbesartan	48	35 (72.9)	82.51 (13.87)	50.0	70.00	85.00	95.00	100.0			
		Week 110	Sparsentan	49	37 (75.5)	82.38 (11.13)	48.0	78.00	82.00	90.00	98.0			
			Irbesartan	48	32 (66.7)	82.59 (11.26)	60.0	78.50	83.00	90.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	49	40 (81.6)	4.35 (17.11)	-25.0	-2.50	2.00	10.50	75.0	0.09 [-0.37, 0.54]
					Irbesartan	48	34 (70.8)	3.09 (10.97)	-11.0	-4.00	-0.50	9.00	34.0	
				Week 48	Sparsentan	49	45 (91.8)	3.44 (14.86)	-29.0	-4.00	2.00	11.00	64.0	0.14 [-0.31, 0.59]
					Irbesartan	48	33 (68.8)	1.52 (11.56)	-35.0	-5.00	0.00	9.00	25.0	
				Week 70	Sparsentan	49	40 (81.6)	-0.13 (12.99)	-51.0	-6.00	0.00	8.00	23.0	-0.12 [-0.59, 0.34]
					Irbesartan	48	32 (66.7)	1.31 (10.39)	-30.0	-4.00	0.00	10.00	18.0	
				Week 94	Sparsentan	49	41 (83.7)	-1.27 (16.87)	-61.0	-8.00	0.00	10.00	30.0	-0.24 [-0.70, 0.21]
				Irbesartan	48	35 (72.9)	2.40 (12.46)	-33.0	-4.00	0.00	10.00	31.0		
		Week 110	Sparsentan	49	37 (75.5)	2.92 (19.21)	-30.0	-5.00	1.00	9.00	88.0	0.08 [-0.39, 0.55]		
			Irbesartan	48	32 (66.7)	1.69 (10.06)	-13.0	-5.00	0.00	8.00	27.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
>= 90 mL/min/1.73 m**2	EQ-5D VAS	Baseline	Sparsentan	26	25 (96.2)	79.20 (13.04)	40.0	71.00	81.00	89.00	97.0			
			Irbesartan	25	24 (96.0)	82.54 (17.58)	19.0	78.00	86.00	92.50	100.0			
		Week 24	Sparsentan	26	22 (84.6)	85.91 (10.32)	60.0	80.00	88.50	91.00	100.0			
			Irbesartan	25	20 (80.0)	83.55 (18.30)	30.0	74.50	90.50	95.50	100.0			
		Week 48	Sparsentan	26	24 (92.3)	82.25 (13.24)	50.0	77.50	84.50	92.00	100.0			
			Irbesartan	25	18 (72.0)	84.39 (10.84)	66.0	75.00	80.50	92.00	100.0			
		Week 70	Sparsentan	26	23 (88.5)	81.39 (7.95)	66.0	77.00	80.00	89.00	95.0			
			Irbesartan	25	20 (80.0)	88.35 (9.39)	70.0	80.50	90.00	97.00	100.0			
		Week 94	Sparsentan	26	21 (80.8)	84.48 (10.47)	50.0	80.00	85.00	90.00	97.0			
			Irbesartan	25	19 (76.0)	86.05 (10.49)	62.0	80.00	90.00	94.00	100.0			
		Week 110	Sparsentan	26	22 (84.6)	84.77 (7.95)	62.0	81.00	84.50	90.00	100.0			
			Irbesartan	25	17 (68.0)	87.94 (8.07)	70.0	84.00	90.00	91.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	26	22 (84.6)	6.23 (14.25)	-20.0	-1.00	5.50	16.00	40.0	0.26 [-0.34, 0.87]
					Irbesartan	25	20 (80.0)	2.90 (10.39)	-21.0	-4.00	2.50	10.50	21.0	
				Week 48	Sparsentan	26	24 (92.3)	3.17 (15.77)	-30.0	-5.50	3.50	10.50	46.0	0.20 [-0.41, 0.81]
					Irbesartan	25	18 (72.0)	0.33 (11.36)	-25.0	-4.00	0.00	5.00	27.0	
				Week 70	Sparsentan	26	23 (88.5)	2.96 (13.77)	-22.0	-6.00	5.00	9.00	41.0	-0.13 [-0.73, 0.47]
					Irbesartan	25	20 (80.0)	4.45 (9.07)	-12.0	-1.00	4.00	10.00	29.0	
				Week 94	Sparsentan	26	21 (80.8)	4.38 (15.59)	-30.0	-4.00	5.00	11.00	44.0	0.17 [-0.45, 0.79]
				Irbesartan	25	19 (76.0)	1.79 (14.48)	-38.0	-3.00	0.00	9.00	38.0		
		Week 110	Sparsentan	26	22 (84.6)	5.09 (13.09)	-19.0	-4.00	5.50	11.00	36.0	0.04 [-0.59, 0.68]		
			Irbesartan	25	17 (68.0)	4.59 (8.70)	-7.0	-2.00	5.00	9.00	28.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline urine protein excretion														
<= 1.75 g/day	EQ-5D VAS	Baseline	Sparsentan	98	94 (95.9)	79.36 (16.33)	7.0	73.00	82.00	90.00	100.0			
			Irbesartan	93	83 (89.2)	80.96 (14.54)	19.0	74.00	81.00	90.00	100.0			
		Week 24	Sparsentan	98	84 (85.7)	83.60 (10.74)	51.0	78.00	82.50	90.00	100.0			
			Irbesartan	93	60 (64.5)	83.52 (12.82)	30.0	80.00	84.50	91.00	100.0			
		Week 48	Sparsentan	98	84 (85.7)	83.73 (11.37)	49.0	80.00	85.00	91.00	100.0			
			Irbesartan	93	59 (63.4)	83.44 (10.47)	51.0	79.00	83.00	91.00	100.0			
		Week 70	Sparsentan	98	80 (81.6)	81.61 (12.84)	27.0	75.00	81.50	90.00	100.0			
			Irbesartan	93	62 (66.7)	84.73 (12.72)	40.0	79.00	87.00	94.00	100.0			
		Week 94	Sparsentan	98	80 (81.6)	81.35 (12.79)	40.0	76.00	82.00	90.00	100.0			
			Irbesartan	93	65 (69.9)	85.32 (10.54)	59.0	80.00	88.00	91.00	100.0			
		Week 110	Sparsentan	98	78 (79.6)	80.55 (11.15)	48.0	76.00	81.00	90.00	100.0			
			Irbesartan	93	57 (61.3)	84.02 (10.88)	54.0	79.00	85.00	91.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	98	84 (85.7)	4.61 (15.97)	-25.0	-4.50	1.00	11.50	75.0	0.22 [-0.11, 0.55]
					Irbesartan	93	60 (64.5)	1.62 (8.98)	-20.0	-4.50	0.00	9.00	21.0	
				Week 48	Sparsentan	98	84 (85.7)	4.30 (16.08)	-29.0	-2.00	1.00	10.00	87.0	0.18 [-0.15, 0.51]
					Irbesartan	93	59 (63.4)	1.76 (10.78)	-25.0	-5.00	2.00	9.00	27.0	
				Week 70	Sparsentan	98	80 (81.6)	1.29 (15.19)	-51.0	-6.50	0.50	8.50	59.0	-0.06 [-0.40, 0.27]
					Irbesartan	93	62 (66.7)	2.16 (10.83)	-30.0	-4.00	0.00	10.00	29.0	
				Week 94	Sparsentan	98	80 (81.6)	1.83 (13.70)	-31.0	-7.00	1.00	11.00	40.0	-0.21 [-0.54, 0.12]
					Irbesartan	93	65 (69.9)	4.54 (12.17)	-22.0	-3.00	4.00	10.00	44.0	
		Week 110	Sparsentan	98	78 (79.6)	1.28 (18.13)	-40.0	-7.00	1.00	9.00	88.0	-0.09 [-0.43, 0.26]		
			Irbesartan	93	57 (61.3)	2.61 (11.15)	-21.0	-5.00	0.00	11.00	28.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 VAS = visual analogue scale.  
 Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 1.75 g/day	EQ-5D VAS	Baseline	Sparsentan	104	97 (93.3)	81.57 (11.86)	40.0	75.00	81.00	91.00	100.0		
			Irbesartan	109	101 (92.7)	79.91 (14.01)	34.0	75.00	80.00	89.00	100.0		
	Week 24	Sparsentan	104	80 (76.9)	84.83 (10.65)	50.0	78.00	87.00	91.00	100.0			
		Irbesartan	109	83 (76.1)	79.34 (18.47)	4.0	71.00	83.00	91.00	100.0			
	Week 48	Sparsentan	104	84 (80.8)	82.80 (10.80)	50.0	78.50	82.00	90.00	100.0			
		Irbesartan	109	73 (67.0)	81.36 (15.14)	40.0	74.00	83.00	90.00	100.0			
	Week 70	Sparsentan	104	86 (82.7)	81.07 (10.94)	50.0	75.00	81.00	90.00	100.0			
		Irbesartan	109	74 (67.9)	79.11 (17.45)	20.0	71.00	82.50	91.00	100.0			
	Week 94	Sparsentan	104	76 (73.1)	82.86 (13.48)	34.0	79.00	85.50	92.00	100.0			
		Irbesartan	109	71 (65.1)	79.68 (16.64)	34.0	71.00	83.00	90.00	100.0			
	Week 110	Sparsentan	104	74 (71.2)	84.34 (11.52)	50.0	79.00	86.00	92.00	100.0			
		Irbesartan	109	64 (58.7)	79.81 (13.49)	35.0	73.00	81.00	89.50	100.0			
	Change from baseline in EQ-5D VAS		Week 24	Sparsentan	104	80 (76.9)	1.71 (11.74)	-31.0	-3.50	1.00	8.00	40.0	0.08 [-0.23, 0.39]
				Irbesartan	109	83 (76.1)	0.41 (19.47)	-90.0	-5.00	0.00	10.00	52.0	
			Week 48	Sparsentan	104	84 (80.8)	0.48 (12.11)	-30.0	-6.50	0.00	7.00	46.0	-0.21 [-0.53, 0.10]
				Irbesartan	109	73 (67.0)	3.22 (13.53)	-35.0	-1.00	3.00	9.00	43.0	
			Week 70	Sparsentan	104	86 (82.7)	-1.07 (11.76)	-32.0	-10.00	-0.50	6.00	41.0	-0.12 [-0.44, 0.19]
				Irbesartan	109	74 (67.9)	0.58 (14.77)	-60.0	-6.00	0.00	8.00	40.0	
			Week 94	Sparsentan	104	76 (73.1)	0.45 (15.73)	-61.0	-4.00	1.00	8.50	45.0	-0.04 [-0.37, 0.28]
				Irbesartan	109	71 (65.1)	1.14 (16.54)	-45.0	-4.00	0.00	9.00	50.0	
Week 110			Sparsentan	104	74 (71.2)	1.99 (11.42)	-31.0	-4.00	1.00	9.00	36.0	-0.03 [-0.36, 0.31]	
			Irbesartan	109	64 (58.7)	2.33 (11.97)	-26.0	-3.00	1.50	7.50	39.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline use of antihypertensives														
Yes	EQ-5D VAS	Baseline	Sparsentan	90	82 (91.1)	78.51 (16.81)	7.0	71.00	81.00	90.00	100.0			
			Irbesartan	88	78 (88.6)	78.95 (13.71)	37.0	72.00	80.00	90.00	100.0			
		Week 24	Sparsentan	90	71 (78.9)	84.14 (11.19)	50.0	78.00	89.00	90.00	100.0			
			Irbesartan	88	55 (62.5)	80.76 (13.99)	4.0	74.00	81.00	90.00	99.0			
		Week 48	Sparsentan	90	70 (77.8)	82.93 (12.35)	49.0	78.00	84.50	92.00	100.0			
			Irbesartan	88	51 (58.0)	81.51 (12.57)	50.0	75.00	83.00	90.00	100.0			
		Week 70	Sparsentan	90	71 (78.9)	81.13 (11.34)	50.0	75.00	81.00	90.00	100.0			
			Irbesartan	88	52 (59.1)	77.02 (17.68)	20.0	70.00	80.00	89.00	100.0			
		Week 94	Sparsentan	90	67 (74.4)	82.70 (14.05)	40.0	77.00	86.00	92.00	100.0			
			Irbesartan	88	58 (65.9)	82.12 (12.95)	34.0	76.00	84.00	90.00	100.0			
		Week 110	Sparsentan	90	65 (72.2)	82.62 (12.18)	50.0	76.00	83.00	92.00	100.0			
			Irbesartan	88	51 (58.0)	79.98 (10.43)	54.0	71.00	81.00	89.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	90	71 (78.9)	4.99 (15.45)	-29.0	-2.00	6.00	10.00	75.0	0.19 [-0.17, 0.54]
					Irbesartan	88	55 (62.5)	2.00 (16.86)	-71.0	-6.00	0.00	10.00	52.0	
		Week 48		Sparsentan	90	70 (77.8)	4.53 (16.27)	-29.0	-2.00	2.50	7.00	87.0	-0.04 [-0.40, 0.32]	
				Irbesartan	88	51 (58.0)	5.06 (12.25)	-19.0	0.00	5.00	10.00	43.0		
		Week 70		Sparsentan	90	71 (78.9)	0.70 (13.82)	-32.0	-9.00	0.00	8.00	59.0	0.09 [-0.27, 0.45]	
				Irbesartan	88	52 (59.1)	-0.63 (16.01)	-60.0	-6.00	0.00	8.00	40.0		
Week 94	Sparsentan	90		67 (74.4)	3.73 (13.74)	-34.0	-4.00	3.00	12.00	45.0	-0.14 [-0.49, 0.22]			
	Irbesartan	88		58 (65.9)	5.76 (15.99)	-41.0	-4.00	5.00	12.00	50.0				
Week 110	Sparsentan	90		65 (72.2)	3.72 (18.35)	-31.0	-4.00	1.00	10.00	88.0	-0.00 [-0.37, 0.37]			
	Irbesartan	88		51 (58.0)	3.73 (12.40)	-23.0	-4.00	1.00	11.00	39.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
No	EQ-5D VAS	Baseline	Sparsentan	112	109 (97.3)	81.96 (11.82)	40.0	78.00	82.00	90.00	100.0			
			Irbesartan	114	106 (93.0)	81.44 (14.55)	19.0	75.00	83.00	90.00	100.0			
		Week 24	Sparsentan	112	93 (83.0)	84.24 (10.34)	59.0	78.00	84.00	91.00	100.0			
			Irbesartan	114	88 (77.2)	81.30 (17.84)	10.0	78.50	85.00	92.50	100.0			
		Week 48	Sparsentan	112	98 (87.5)	83.50 (10.11)	49.0	80.00	83.50	90.00	100.0			
			Irbesartan	114	81 (71.1)	82.78 (13.72)	40.0	79.00	83.00	92.00	100.0			
		Week 70	Sparsentan	112	95 (84.8)	81.48 (12.29)	27.0	75.00	82.00	90.00	100.0			
			Irbesartan	114	84 (73.7)	84.55 (13.62)	35.0	79.00	86.50	95.00	100.0			
		Week 94	Sparsentan	112	89 (79.5)	81.62 (12.41)	34.0	80.00	83.00	90.00	100.0			
			Irbesartan	114	78 (68.4)	82.56 (15.29)	39.0	78.00	85.00	95.00	100.0			
		Week 110	Sparsentan	112	87 (77.7)	82.23 (10.94)	48.0	78.00	82.00	90.00	100.0			
			Irbesartan	114	70 (61.4)	83.11 (13.67)	35.0	79.00	85.00	91.00	100.0			
		Change from baseline in EQ-5D VAS	EQ-5D VAS	Week 24	Sparsentan	112	93 (83.0)	1.83 (12.89)	-31.0	-4.00	0.00	8.00	40.0	0.11 [-0.18, 0.40]
					Irbesartan	114	88 (77.2)	0.24 (15.32)	-90.0	-4.00	0.00	9.50	23.0	
				Week 48	Sparsentan	112	98 (87.5)	0.86 (12.61)	-30.0	-6.00	0.00	10.00	46.0	-0.01 [-0.31, 0.28]
					Irbesartan	114	81 (71.1)	1.00 (12.23)	-35.0	-5.00	1.00	8.00	38.0	
				Week 70	Sparsentan	112	95 (84.8)	-0.41 (13.37)	-51.0	-7.00	0.00	8.00	41.0	-0.24 [-0.53, 0.06]
					Irbesartan	114	84 (73.7)	2.50 (10.85)	-20.0	-3.50	1.00	10.00	40.0	
				Week 94	Sparsentan	112	89 (79.5)	-0.79 (15.16)	-61.0	-7.00	0.00	5.00	44.0	-0.09 [-0.40, 0.21]
					Irbesartan	114	78 (68.4)	0.54 (13.26)	-45.0	-6.00	0.00	10.00	38.0	
Week 110	Sparsentan			112	87 (77.7)	0.06 (12.20)	-40.0	-6.00	0.00	6.00	36.0	-0.13 [-0.44, 0.19]		
	Irbesartan			114	70 (61.4)	1.54 (10.89)	-26.0	-5.00	0.00	7.00	34.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Time since renal biopsy														
<= 5 years	EQ-5D VAS	Baseline	Sparsentan	113	107 (94.7)	80.36 (13.96)	11.0	74.00	81.00	90.00	100.0			
			Irbesartan	127	119 (93.7)	80.84 (14.67)	19.0	75.00	80.00	90.00	100.0			
		Week 24	Sparsentan	113	94 (83.2)	84.89 (11.18)	50.0	78.00	89.00	91.00	100.0			
			Irbesartan	127	94 (74.0)	81.62 (17.02)	4.0	79.00	85.00	91.00	100.0			
		Week 48	Sparsentan	113	95 (84.1)	82.75 (12.19)	49.0	78.00	85.00	91.00	100.0			
			Irbesartan	127	89 (70.1)	82.17 (13.77)	40.0	75.00	83.00	90.00	100.0			
		Week 70	Sparsentan	113	94 (83.2)	80.60 (12.80)	27.0	75.00	81.00	90.00	100.0			
			Irbesartan	127	89 (70.1)	83.22 (15.92)	20.0	78.00	86.00	94.00	100.0			
		Week 94	Sparsentan	113	89 (78.8)	82.54 (13.49)	40.0	78.00	85.00	91.00	100.0			
			Irbesartan	127	89 (70.1)	83.28 (13.88)	34.0	79.00	85.00	91.00	100.0			
		Week 110	Sparsentan	113	89 (78.8)	83.71 (10.96)	48.0	79.00	84.00	91.00	100.0			
			Irbesartan	127	80 (63.0)	83.66 (11.17)	40.0	79.00	84.50	90.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	113	94 (83.2)	4.23 (13.03)	-29.0	-3.00	3.50	10.00	40.0	0.24 [-0.05, 0.53]
					Irbesartan	127	94 (74.0)	0.54 (17.30)	-90.0	-5.00	0.00	10.00	52.0	
		Week 48		Sparsentan	113	95 (84.1)	2.26 (15.11)	-30.0	-6.00	1.00	10.00	87.0	0.06 [-0.23, 0.35]	
				Irbesartan	127	89 (70.1)	1.40 (13.55)	-35.0	-5.00	1.00	9.00	43.0		
		Week 70		Sparsentan	113	94 (83.2)	-0.65 (15.32)	-51.0	-10.00	0.00	8.00	59.0	-0.19 [-0.48, 0.10]	
				Irbesartan	127	89 (70.1)	2.19 (14.70)	-60.0	-4.00	2.00	10.00	40.0		
Week 94	Sparsentan	113		89 (78.8)	2.34 (14.22)	-34.0	-5.00	2.00	11.00	45.0	-0.05 [-0.34, 0.25]			
	Irbesartan	127		89 (70.1)	3.03 (14.89)	-41.0	-4.00	1.00	10.00	50.0				
Week 110	Sparsentan	113		89 (78.8)	3.28 (14.13)	-30.0	-4.00	2.00	11.00	70.0	-0.04 [-0.34, 0.27]			
	Irbesartan	127		80 (63.0)	3.75 (11.11)	-19.0	-3.50	2.00	10.00	39.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024



Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 5 years	EQ-5D VAS	Baseline	Sparsentan	89	84 (94.4)	80.63 (14.68)	7.0	77.50	82.00	90.00	100.0		
			Irbesartan	75	65 (86.7)	79.55 (13.43)	34.0	73.00	80.00	89.00	100.0		
		Week 24	Sparsentan	89	70 (78.7)	83.26 (9.98)	59.0	78.00	82.00	90.00	100.0		
			Irbesartan	75	49 (65.3)	80.08 (15.31)	31.0	73.00	81.00	90.00	100.0		
		Week 48	Sparsentan	89	73 (82.0)	83.93 (9.44)	51.0	80.00	83.00	90.00	100.0		
			Irbesartan	75	43 (57.3)	82.53 (12.27)	50.0	79.00	85.00	91.00	100.0		
		Week 70	Sparsentan	89	72 (80.9)	82.29 (10.52)	59.0	75.00	82.50	90.00	100.0		
			Irbesartan	75	47 (62.7)	78.72 (14.91)	35.0	70.00	82.00	90.00	100.0		
		Week 94	Sparsentan	89	67 (75.3)	81.48 (12.66)	34.0	77.00	82.00	90.00	100.0		
			Irbesartan	75	47 (62.7)	80.66 (15.04)	39.0	72.00	84.00	91.00	100.0		
		Week 110	Sparsentan	89	63 (70.8)	80.54 (11.96)	48.0	75.00	81.00	90.00	100.0		
			Irbesartan	75	41 (54.7)	78.15 (14.10)	35.0	71.00	81.00	89.00	100.0		
		Change from baseline in EQ-5D VAS	Week 24	Sparsentan	89	70 (78.7)	1.80 (15.40)	-31.0	-6.00	0.00	8.00	75.0	0.01 [-0.35, 0.38]
				Irbesartan	75	49 (65.3)	1.63 (12.93)	-42.0	-2.00	1.00	9.00	34.0	
			Week 48	Sparsentan	89	73 (82.0)	2.55 (13.33)	-22.0	-4.00	0.00	7.00	64.0	-0.20 [-0.58, 0.17]
				Irbesartan	75	43 (57.3)	4.98 (9.05)	-20.0	-1.00	5.00	10.00	25.0	
			Week 70	Sparsentan	89	72 (80.9)	1.00 (10.80)	-20.0	-6.00	0.00	8.00	40.0	0.14 [-0.23, 0.50]
				Irbesartan	75	47 (62.7)	-0.38 (9.27)	-18.0	-9.00	0.00	5.00	21.0	
	Week 94		Sparsentan	89	67 (75.3)	-0.42 (15.26)	-61.0	-7.00	1.00	8.00	40.0	-0.18 [-0.55, 0.19]	
			Irbesartan	75	47 (62.7)	2.26 (14.35)	-45.0	-7.00	4.00	10.00	32.0		
Week 110	Sparsentan	89	63 (70.8)	-0.71 (16.41)	-40.0	-7.00	0.00	6.00	88.0	-0.04 [-0.44, 0.35]			
	Irbesartan	75	41 (54.7)	-0.05 (12.11)	-26.0	-9.00	0.00	5.00	30.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: History of hypertension														
Yes	EQ-5D VAS	Baseline	Sparsentan	155	145 (93.5)	80.06 (13.64)	11.0	71.00	81.00	90.00	100.0			
			Irbesartan	161	144 (89.4)	79.76 (14.12)	34.0	74.00	80.00	90.00	100.0			
		Week 24	Sparsentan	155	125 (80.6)	84.22 (10.79)	50.0	78.00	86.00	91.00	100.0			
			Irbesartan	161	111 (68.9)	80.56 (16.66)	4.0	74.00	83.00	91.00	100.0			
		Week 48	Sparsentan	155	126 (81.3)	83.26 (11.39)	49.0	79.00	84.00	91.00	100.0			
			Irbesartan	161	102 (63.4)	81.80 (13.68)	40.0	75.00	83.50	90.00	100.0			
		Week 70	Sparsentan	155	128 (82.6)	81.56 (11.55)	40.0	75.00	81.00	90.00	100.0			
			Irbesartan	161	105 (65.2)	80.21 (16.58)	20.0	74.00	84.00	92.00	100.0			
		Week 94	Sparsentan	155	119 (76.8)	81.93 (13.39)	34.0	78.00	83.00	91.00	100.0			
			Irbesartan	161	105 (65.2)	81.75 (14.26)	34.0	76.00	84.00	90.00	100.0			
		Week 110	Sparsentan	155	114 (73.5)	82.32 (11.17)	50.0	76.00	82.00	90.00	100.0			
			Irbesartan	161	92 (57.1)	80.75 (12.67)	35.0	75.00	82.00	90.00	100.0			
			Change from baseline in EQ-5D VAS	Week 24	Sparsentan	155	125 (80.6)	3.39 (12.67)	-29.0	-4.00	2.00	10.00	44.0	0.15 [-0.10, 0.41]
					Irbesartan	161	111 (68.9)	1.12 (16.99)	-90.0	-5.00	0.00	10.00	52.0	
			Week 48	Sparsentan	155	126 (81.3)	2.89 (13.83)	-30.0	-2.00	1.00	10.00	87.0	-0.05 [-0.31, 0.21]	
				Irbesartan	161	102 (63.4)	3.51 (12.63)	-35.0	-1.00	3.00	10.00	43.0		
			Week 70	Sparsentan	155	128 (82.6)	0.94 (13.18)	-45.0	-7.50	0.00	8.50	59.0	-0.04 [-0.30, 0.22]	
				Irbesartan	161	105 (65.2)	1.50 (14.04)	-60.0	-5.00	0.00	10.00	40.0		
			Week 94	Sparsentan	155	119 (76.8)	1.73 (15.23)	-61.0	-4.00	2.00	11.00	45.0	-0.14 [-0.40, 0.12]	
				Irbesartan	161	105 (65.2)	3.85 (14.73)	-41.0	-4.00	3.00	10.00	50.0		
	Week 110	Sparsentan	155	114 (73.5)	1.72 (13.32)	-31.0	-5.00	1.00	9.00	70.0	-0.11 [-0.39, 0.16]			
		Irbesartan	161	92 (57.1)	3.16 (12.15)	-26.0	-4.00	1.00	10.50	39.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSHM: Course of EQ-5D VAS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
No	EQ-5D VAS	Baseline	Sparsentan	47	46 (97.9)	81.80 (16.09)	7.0	80.00	83.50	90.00	100.0			
			Irbesartan	41	40 (97.6)	82.65 (14.53)	19.0	75.00	85.00	90.50	100.0			
		Week 24	Sparsentan	47	39 (83.0)	84.10 (10.45)	59.0	78.00	86.00	90.00	100.0			
			Irbesartan	41	32 (78.0)	82.94 (15.65)	30.0	80.00	85.00	93.00	100.0			
		Week 48	Sparsentan	47	42 (89.4)	83.26 (10.15)	49.0	80.00	84.50	90.00	100.0			
			Irbesartan	41	30 (73.2)	83.93 (11.75)	54.0	79.00	83.00	92.00	100.0			
		Week 70	Sparsentan	47	38 (80.9)	80.55 (12.99)	27.0	74.00	80.00	90.00	96.0			
			Irbesartan	41	31 (75.6)	86.61 (10.92)	55.0	80.00	88.00	96.00	100.0			
		Week 94	Sparsentan	47	37 (78.7)	82.57 (12.32)	40.0	78.00	85.00	91.00	97.0			
			Irbesartan	41	31 (75.6)	84.48 (14.42)	40.0	80.00	86.00	95.00	100.0			
		Week 110	Sparsentan	47	38 (80.9)	82.61 (12.42)	48.0	80.00	82.50	90.00	99.0			
			Irbesartan	41	29 (70.7)	85.10 (11.34)	55.0	80.00	88.00	92.00	100.0			
		Change from baseline in EQ-5D VAS		Week 24	Sparsentan	47	39 (83.0)	2.56 (18.11)	-31.0	-6.00	0.00	10.00	75.0	0.15 [-0.32, 0.62]
					Irbesartan	41	32 (78.0)	0.22 (11.51)	-37.0	-4.00	0.00	9.00	22.0	
				Week 48	Sparsentan	47	42 (89.4)	0.88 (15.78)	-29.0	-8.00	-1.00	6.00	64.0	0.11 [-0.36, 0.58]
					Irbesartan	41	30 (73.2)	-0.63 (10.96)	-31.0	-5.00	0.00	7.00	19.0	
				Week 70	Sparsentan	47	38 (80.9)	-2.87 (14.47)	-51.0	-9.00	-0.50	4.00	41.0	-0.28 [-0.76, 0.19]
					Irbesartan	41	31 (75.6)	0.65 (9.38)	-18.0	-6.00	0.00	10.00	21.0	
				Week 94	Sparsentan	47	37 (78.7)	-0.70 (12.82)	-28.0	-8.00	-1.00	5.00	44.0	0.01 [-0.46, 0.49]
					Irbesartan	41	31 (75.6)	-0.90 (14.03)	-45.0	-6.00	1.00	10.00	20.0	
Week 110	Sparsentan			47	38 (80.9)	1.34 (20.01)	-40.0	-6.00	0.00	7.00	88.0	0.07 [-0.42, 0.55]		
	Irbesartan			41	29 (70.7)	0.24 (9.22)	-21.0	-5.00	0.00	5.00	21.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

VAS = visual analogue scale.

Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.559
Male	Week 24	Sparsentan	139	114 (82.0)	3.13 (1.04)	(1.08, 5.17)	2.84 (1.51)	(-0.13, 5.80)	0.061
		Irbesartan	143	103 (72.0)	0.29 (1.09)	(-1.86, 2.44)			
	Week 48	Sparsentan	139	118 (84.9)	2.41 (1.02)	(0.40, 4.42)	-0.28 (1.52)	(-3.26, 2.71)	0.854
		Irbesartan	143	94 (65.7)	2.69 (1.12)	(0.49, 4.89)			
	Week 70	Sparsentan	139	119 (85.6)	0.42 (1.02)	(-1.59, 2.43)	-2.15 (1.52)	(-5.13, 0.83)	0.158
		Irbesartan	143	97 (67.8)	2.57 (1.12)	(0.37, 4.77)			
	Week 94	Sparsentan	139	109 (78.4)	2.26 (1.06)	(0.19, 4.34)	-1.01 (1.54)	(-4.03, 2.01)	0.512
		Irbesartan	143	99 (69.2)	3.27 (1.12)	(1.08, 5.46)			
	Week 110	Sparsentan	139	106 (76.3)	1.72 (1.08)	(-0.40, 3.84)	-0.62 (1.60)	(-3.76, 2.51)	0.695
		Irbesartan	143	89 (62.2)	2.34 (1.17)	(0.04, 4.64)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
 Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Female	Week 24	Sparsentan	63	50 (79.4)	3.74 (1.71)	(0.37, 7.11)	1.56 (2.58)	(-3.51, 6.63)	0.546
		Irbesartan	59	40 (67.8)	2.18 (1.92)	(-1.59, 5.96)			
	Week 48	Sparsentan	63	50 (79.4)	2.27 (1.71)	(-1.09, 5.63)	1.02 (2.59)	(-4.08, 6.12)	0.694
		Irbesartan	59	38 (64.4)	1.25 (1.95)	(-2.58, 5.08)			
	Week 70	Sparsentan	63	47 (74.6)	0.45 (1.75)	(-2.99, 3.90)	1.75 (2.62)	(-3.40, 6.90)	0.504
		Irbesartan	59	39 (66.1)	-1.30 (1.94)	(-5.11, 2.52)			
	Week 94	Sparsentan	63	47 (74.6)	-0.88 (1.76)	(-4.34, 2.58)	-0.79 (2.66)	(-6.02, 4.43)	0.765
		Irbesartan	59	37 (62.7)	-0.09 (1.99)	(-4.00, 3.82)			
	Week 110	Sparsentan	63	46 (73.0)	1.04 (1.79)	(-2.48, 4.56)	0.34 (2.79)	(-5.14, 5.82)	0.902
		Irbesartan	59	32 (54.2)	0.70 (2.13)	(-3.49, 4.88)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
 Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.172
<= 45 years	Week 24	Sparsentan	96	75 (78.1)	3.58 (1.41)	(0.80, 6.36)	4.15 (2.03)	(0.17, 8.13)	0.041 *
		Irbesartan	99	72 (72.7)	-0.57 (1.45)	(-3.41, 2.27)			
	Week 48	Sparsentan	96	79 (82.3)	2.38 (1.38)	(-0.33, 5.09)	2.35 (2.04)	(-1.67, 6.36)	0.251
		Irbesartan	99	63 (63.6)	0.04 (1.51)	(-2.92, 2.99)			
	Week 70	Sparsentan	96	76 (79.2)	1.03 (1.40)	(-1.72, 3.79)	0.30 (2.06)	(-3.75, 4.35)	0.886
		Irbesartan	99	64 (64.6)	0.74 (1.51)	(-2.23, 3.71)			
	Week 94	Sparsentan	96	71 (74.0)	0.01 (1.44)	(-2.82, 2.83)	0.38 (2.09)	(-3.72, 4.47)	0.857
		Irbesartan	99	65 (65.7)	-0.37 (1.51)	(-3.34, 2.60)			
	Week 110	Sparsentan	96	71 (74.0)	1.70 (1.46)	(-1.16, 4.55)	0.50 (2.15)	(-3.72, 4.72)	0.817
		Irbesartan	99	59 (59.6)	1.20 (1.58)	(-1.91, 4.30)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 45 years	Week 24	Sparsentan	106	89 (84.0)	3.17 (1.15)	(0.91, 5.43)	1.24 (1.73)	(-2.16, 4.64)	0.474
		Irbesartan	103	71 (68.9)	1.93 (1.29)	(-0.60, 4.45)			
	Week 48	Sparsentan	106	89 (84.0)	2.49 (1.15)	(0.24, 4.74)	-1.59 (1.73)	(-4.98, 1.81)	
		Irbesartan	103	69 (67.0)	4.08 (1.29)	(1.54, 6.61)			
	Week 70	Sparsentan	106	90 (84.9)	0.03 (1.15)	(-2.22, 2.28)	-1.68 (1.72)	(-5.06, 1.69)	
		Irbesartan	103	72 (69.9)	1.72 (1.27)	(-0.78, 4.22)			
	Week 94	Sparsentan	106	85 (80.2)	2.53 (1.17)	(0.23, 4.84)	-1.87 (1.74)	(-5.29, 1.56)	
		Irbesartan	103	71 (68.9)	4.40 (1.29)	(1.87, 6.93)			
	Week 110	Sparsentan	106	81 (76.4)	1.57 (1.20)	(-0.80, 3.93)	-0.49 (1.83)	(-4.08, 3.10)	
		Irbesartan	103	62 (60.2)	2.05 (1.37)	(-0.64, 4.75)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.066
<= 18 years	Week 24	Sparsentan	9	5 (55.6)	8.76 (7.92)	(-8.27, 25.80)	13.61 (12.32)	(-13.61, 40.83)	0.294
		Irbesartan	5	5 (100.0)	-4.85 (8.91)	(-24.81, 15.12)			
	Week 48	Sparsentan	9	7 (77.8)	7.82 (7.16)	(-8.03, 23.68)	9.95 (12.36)	(-17.22, 37.13)	0.438
		Irbesartan	5	3 (60.0)	-2.13 (9.65)	(-23.06, 18.80)			
	Week 70	Sparsentan	9	7 (77.8)	7.75 (7.29)	(-8.35, 23.85)	8.48 (12.35)	(-18.78, 35.74)	0.507
		Irbesartan	5	4 (80.0)	-0.73 (9.48)	(-21.52, 20.06)			
	Week 94	Sparsentan	9	5 (55.6)	-7.08 (7.93)	(-24.16, 10.01)	-5.35 (13.04)	(-33.77, 23.07)	0.689
		Irbesartan	5	4 (80.0)	-1.73 (9.81)	(-23.16, 19.71)			
	Week 110	Sparsentan	9	4 (44.4)	7.32 (8.70)	(-11.15, 25.79)	11.92 (14.91)	(-19.68, 43.51)	0.436
		Irbesartan	5	2 (40.0)	-4.59 (11.78)	(-29.37, 20.19)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024



Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 18 to 40 years	Week 24	Sparsentan	102	82 (80.4)	3.71 (1.30)	(1.16, 6.26)	3.60 (1.87)	(-0.08, 7.28)	0.055
		Irbesartan	109	75 (68.8)	0.11 (1.35)	(-2.54, 2.77)			
	Week 48	Sparsentan	102	85 (83.3)	2.56 (1.28)	(0.05, 5.07)	2.18 (1.88)	(-1.51, 5.88)	0.247
		Irbesartan	109	69 (63.3)	0.38 (1.38)	(-2.33, 3.09)			
	Week 70	Sparsentan	102	81 (79.4)	1.11 (1.30)	(-1.44, 3.67)	-0.21 (1.90)	(-3.93, 3.52)	0.914
		Irbesartan	109	71 (65.1)	1.32 (1.38)	(-1.39, 4.03)			
	Week 94	Sparsentan	102	77 (75.5)	1.68 (1.33)	(-0.93, 4.28)	1.76 (1.92)	(-2.01, 5.53)	0.360
		Irbesartan	109	71 (65.1)	-0.09 (1.39)	(-2.81, 2.64)			
	Week 110	Sparsentan	102	75 (73.5)	1.73 (1.36)	(-0.93, 4.39)	0.53 (1.98)	(-3.37, 4.42)	0.789
		Irbesartan	109	65 (59.6)	1.20 (1.44)	(-1.64, 4.04)			
> 40 years	Week 24	Sparsentan	91	77 (84.6)	2.76 (1.24)	(0.32, 5.21)	1.14 (1.86)	(-2.51, 4.78)	0.540
		Irbesartan	88	63 (71.6)	1.63 (1.37)	(-1.07, 4.32)			
	Week 48	Sparsentan	91	76 (83.5)	2.14 (1.25)	(-0.30, 4.59)	-2.08 (1.87)	(-5.76, 1.60)	0.267
		Irbesartan	88	60 (68.2)	4.23 (1.40)	(1.48, 6.97)			
	Week 70	Sparsentan	91	78 (85.7)	-0.53 (1.24)	(-2.96, 1.90)	-1.79 (1.86)	(-5.46, 1.87)	0.337
		Irbesartan	88	61 (69.3)	1.26 (1.39)	(-1.47, 4.00)			
	Week 94	Sparsentan	91	74 (81.3)	1.80 (1.26)	(-0.68, 4.28)	-3.13 (1.89)	(-6.84, 0.58)	0.098
		Irbesartan	88	61 (69.3)	4.93 (1.40)	(2.19, 7.67)			
	Week 110	Sparsentan	91	73 (80.2)	1.42 (1.28)	(-1.09, 3.92)	-1.09 (1.96)	(-4.94, 2.76)	0.579
		Irbesartan	88	54 (61.4)	2.50 (1.48)	(-0.41, 5.42)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.854
North America	Week 24	Sparsentan	35	24 (68.6)	3.98 (2.61)	(-1.17, 9.13)	6.38 (3.36)	(-0.24, 13.00)	0.059
		Irbesartan	46	38 (82.6)	-2.40 (2.10)	(-6.54, 1.75)			
	Week 48	Sparsentan	35	26 (74.3)	3.58 (2.52)	(-1.40, 8.56)	0.40 (3.32)	(-6.15, 6.95)	0.904
		Irbesartan	46	35 (76.1)	3.18 (2.15)	(-1.06, 7.42)			
	Week 70	Sparsentan	35	24 (68.6)	1.47 (2.60)	(-3.66, 6.59)	-1.46 (3.42)	(-8.22, 5.29)	0.669
		Irbesartan	46	32 (69.6)	2.93 (2.23)	(-1.47, 7.33)			
	Week 94	Sparsentan	35	24 (68.6)	1.33 (2.62)	(-3.83, 6.50)	-0.36 (3.50)	(-7.26, 6.54)	0.918
		Irbesartan	46	30 (65.2)	1.69 (2.31)	(-2.87, 6.26)			
	Week 110	Sparsentan	35	21 (60.0)	4.17 (2.78)	(-1.32, 9.66)	1.59 (3.63)	(-5.58, 8.75)	0.663
		Irbesartan	46	31 (67.4)	2.58 (2.33)	(-2.02, 7.18)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Europe	Week 24	Sparsentan	98	79 (80.6)	2.23 (1.32)	(-0.36, 4.81)	0.87 (1.93)	(-2.93, 4.67)	0.653	
		Irbesartan	115	68 (59.1)	1.36 (1.42)	(-1.43, 4.14)				
	Week 48	Sparsentan	98	79 (80.6)	3.54 (1.31)	(0.96, 6.12)	1.84 (1.96)	(-2.01, 5.69)	0.349	
		Irbesartan	115	62 (53.9)	1.70 (1.46)	(-1.17, 4.56)				
	Week 70	Sparsentan	98	78 (79.6)	0.63 (1.33)	(-1.98, 3.24)	1.11 (1.92)	(-2.65, 4.88)	0.561	
		Irbesartan	115	72 (62.6)	-0.49 (1.38)	(-3.19, 2.22)				
	Week 94	Sparsentan	98	72 (73.5)	1.10 (1.37)	(-1.59, 3.80)	-0.65 (1.94)	(-4.46, 3.16)	0.737	
		Irbesartan	115	73 (63.5)	1.75 (1.37)	(-0.94, 4.45)				
	Week 110	Sparsentan	98	70 (71.4)	1.36 (1.40)	(-1.39, 4.10)	1.30 (2.04)	(-2.71, 5.31)	0.524	
		Irbesartan	115	61 (53.0)	0.05 (1.49)	(-2.86, 2.97)				
Asia Pacific	Week 24	Sparsentan	69	61 (88.4)	4.97 (1.35)	(2.32, 7.62)	3.29 (2.20)	(-1.04, 7.62)	0.136	
		Irbesartan	41	37 (90.2)	1.68 (1.74)	(-1.74, 5.09)				
	Week 48	Sparsentan	69	63 (91.3)	1.03 (1.33)	(-1.58, 3.64)	-0.09 (2.21)	(-4.45, 4.26)	0.966	
		Irbesartan	41	35 (85.4)	1.12 (1.76)	(-2.35, 4.59)				
	Week 70	Sparsentan	69	64 (92.8)	0.42 (1.32)	(-2.18, 3.02)	-2.14 (2.27)	(-6.60, 2.32)	0.346	
		Irbesartan	41	32 (78.0)	2.56 (1.83)	(-1.05, 6.17)				
	Week 94	Sparsentan	69	60 (87.0)	2.39 (1.35)	(-0.27, 5.04)	-0.06 (2.28)	(-4.56, 4.43)	0.978	
		Irbesartan	41	33 (80.5)	2.45 (1.84)	(-1.17, 6.06)				
	Week 110	Sparsentan	69	61 (88.4)	1.79 (1.35)	(-0.87, 4.45)	-1.45 (2.36)	(-6.10, 3.20)	0.539	
		Irbesartan	41	29 (70.7)	3.25 (1.93)	(-0.56, 7.05)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.346
< 27 kg/m**2	Week 24	Sparsentan	83	68 (81.9)	1.90 (1.37)	(-0.79, 4.58)	1.17 (1.92)	(-2.60, 4.93)	0.543
		Irbesartan	94	72 (76.6)	0.73 (1.34)	(-1.89, 3.36)			
	Week 48	Sparsentan	83	69 (83.1)	0.21 (1.36)	(-2.46, 2.88)	-0.60 (1.96)	(-4.45, 3.25)	0.759
		Irbesartan	94	63 (67.0)	0.81 (1.40)	(-1.95, 3.56)			
	Week 70	Sparsentan	83	68 (81.9)	-1.50 (1.37)	(-4.19, 1.20)	-0.98 (1.97)	(-4.86, 2.89)	0.618
		Irbesartan	94	63 (67.0)	-0.51 (1.41)	(-3.29, 2.26)			
	Week 94	Sparsentan	83	65 (78.3)	-1.06 (1.40)	(-3.80, 1.69)	-1.48 (1.98)	(-5.38, 2.42)	0.456
		Irbesartan	94	65 (69.1)	0.43 (1.40)	(-2.33, 3.18)			
	Week 110	Sparsentan	83	64 (77.1)	0.01 (1.41)	(-2.77, 2.78)	0.89 (2.05)	(-3.15, 4.93)	0.665
		Irbesartan	94	57 (60.6)	-0.88 (1.49)	(-3.81, 2.04)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
>= 27 kg/m**2	Week 24	Sparsentan	119	96 (80.7)	4.69 (1.19)	(2.35, 7.03)	4.73 (1.83)	(1.13, 8.33)	0.010	*
		Irbesartan	107	71 (66.4)	-0.03 (1.39)	(-2.75, 2.69)				
	Week 48	Sparsentan	119	99 (83.2)	4.37 (1.17)	(2.06, 6.67)	1.67 (1.82)	(-1.91, 5.24)	0.360	
		Irbesartan	107	69 (64.5)	2.70 (1.38)	(-0.02, 5.42)				
	Week 70	Sparsentan	119	98 (82.4)	2.26 (1.18)	(-0.06, 4.59)	0.06 (1.81)	(-3.50, 3.62)	0.972	
		Irbesartan	107	73 (68.2)	2.20 (1.37)	(-0.48, 4.88)				
	Week 94	Sparsentan	119	91 (76.5)	3.47 (1.22)	(1.08, 5.86)	0.58 (1.85)	(-3.06, 4.22)	0.753	
		Irbesartan	107	70 (65.4)	2.89 (1.39)	(0.15, 5.62)				
	Week 110	Sparsentan	119	88 (73.9)	3.07 (1.24)	(0.63, 5.52)	-0.53 (1.92)	(-4.31, 3.25)	0.782	
		Irbesartan	107	63 (58.9)	3.61 (1.46)	(0.74, 6.48)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan						Interaction:	0.880
eGFR Low and UP High	Week 24	Sparsentan	71	54 (76.1)	1.67 (1.71)	(-1.69, 5.03)	5.41 (2.51)	(0.48, 10.34)	0.032 *
		Irbesartan	74	47 (63.5)	-3.74 (1.83)	(-7.33, -0.14)			
	Week 48	Sparsentan	71	55 (77.5)	1.12 (1.68)	(-2.19, 4.43)	-1.73 (2.55)	(-6.75, 3.29)	0.498
		Irbesartan	74	41 (55.4)	2.85 (1.90)	(-0.90, 6.59)			
	Week 70	Sparsentan	71	58 (81.7)	-1.93 (1.66)	(-5.20, 1.34)	-1.26 (2.56)	(-6.29, 3.77)	0.622
		Irbesartan	74	41 (55.4)	-0.66 (1.93)	(-4.46, 3.13)			
	Week 94	Sparsentan	71	52 (73.2)	0.77 (1.72)	(-2.62, 4.15)	0.04 (2.63)	(-5.13, 5.21)	0.987
		Irbesartan	74	40 (54.1)	0.72 (1.97)	(-3.15, 4.60)			
	Week 110	Sparsentan	71	53 (74.6)	0.12 (1.73)	(-3.29, 3.52)	-0.31 (2.72)	(-5.66, 5.04)	0.909
		Irbesartan	74	36 (48.6)	0.43 (2.08)	(-3.66, 4.52)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis			
						Change from Baseline		Treatment Difference	
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI
eGFR Low and UP Low	Week 24	Sparsentan	55	46 (83.6)	3.36 (1.68)	(0.05, 6.67)	2.16 (2.47)	(-2.70, 7.01)	0.383
		Irbesartan	55	40 (72.7)	1.20 (1.80)	(-2.35, 4.75)			
	Week 48	Sparsentan	55	43 (78.2)	2.91 (1.72)	(-0.47, 6.30)	0.09 (2.51)	(-4.85, 5.02)	0.972
		Irbesartan	55	38 (69.1)	2.82 (1.82)	(-0.76, 6.41)			
	Week 70	Sparsentan	55	44 (80.0)	3.03 (1.72)	(-0.35, 6.40)	1.72 (2.47)	(-3.14, 6.57)	0.487
		Irbesartan	55	41 (74.5)	1.31 (1.77)	(-2.18, 4.80)			
	Week 94	Sparsentan	55	42 (76.4)	2.83 (1.75)	(-0.62, 6.27)	0.40 (2.51)	(-4.54, 5.34)	0.875
		Irbesartan	55	40 (72.7)	2.43 (1.80)	(-1.11, 5.97)			
	Week 110	Sparsentan	55	38 (69.1)	1.77 (1.84)	(-1.84, 5.39)	0.26 (2.67)	(-5.00, 5.51)	0.923
		Irbesartan	55	34 (61.8)	1.52 (1.94)	(-2.30, 5.33)			
eGFR High and UP High	Week 24	Sparsentan	37	29 (78.4)	4.19 (2.02)	(0.21, 8.16)	-1.77 (2.92)	(-7.53, 4.00)	0.546
		Irbesartan	36	27 (75.0)	5.95 (2.12)	(1.78, 10.13)			
	Week 48	Sparsentan	37	34 (91.9)	2.83 (1.89)	(-0.90, 6.55)	0.11 (2.88)	(-5.57, 5.79)	0.970
		Irbesartan	36	25 (69.4)	2.72 (2.18)	(-1.57, 7.00)			
	Week 70	Sparsentan	37	31 (83.8)	2.67 (1.96)	(-1.20, 6.54)	0.54 (2.94)	(-5.25, 6.33)	0.854
		Irbesartan	36	25 (69.4)	2.13 (2.18)	(-2.17, 6.43)			
	Week 94	Sparsentan	37	27 (73.0)	1.29 (2.08)	(-2.81, 5.40)	-3.47 (3.01)	(-9.41, 2.47)	0.251
		Irbesartan	36	25 (69.4)	4.76 (2.18)	(0.46, 9.05)			
	Week 110	Sparsentan	37	26 (70.3)	4.79 (2.14)	(0.56, 9.01)	1.42 (3.15)	(-4.79, 7.63)	0.653
		Irbesartan	36	22 (61.1)	3.37 (2.31)	(-1.18, 7.92)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
eGFR High and UP Low	Week 24	Sparsentan	39	35 (89.7)	5.65 (1.77)	(2.17, 9.13)	3.33 (2.61)	(-1.82, 8.48)	0.204
		Irbesartan	37	29 (78.4)	2.32 (1.93)	(-1.48, 6.11)			
	Week 48	Sparsentan	39	36 (92.3)	3.99 (1.75)	(0.54, 7.43)	3.78 (2.61)	(-1.36, 8.93)	
		Irbesartan	37	28 (75.7)	0.20 (1.94)	(-3.61, 4.02)			
	Week 70	Sparsentan	39	33 (84.6)	-0.48 (1.80)	(-4.03, 3.07)	-3.99 (2.64)	(-9.19, 1.21)	
		Irbesartan	37	29 (78.4)	3.51 (1.93)	(-0.29, 7.30)			
	Week 94	Sparsentan	39	35 (89.7)	0.88 (1.77)	(-2.61, 4.37)	-1.21 (2.59)	(-6.31, 3.88)	
		Irbesartan	37	31 (83.8)	2.09 (1.88)	(-1.62, 5.81)			
	Week 110	Sparsentan	39	35 (89.7)	1.37 (1.77)	(-2.13, 4.86)	-1.31 (2.63)	(-6.48, 3.87)	
		Irbesartan	37	29 (78.4)	2.67 (1.93)	(-1.14, 6.48)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
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Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.772
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	102 (80.3)	2.60 (1.18)	(0.28, 4.92)	3.68 (1.73)	(0.27, 7.08)	0.034 *
		Irbesartan	129	89 (69.0)	-1.08 (1.27)	(-3.56, 1.41)			
	Week 48	Sparsentan	127	99 (78.0)	2.41 (1.19)	(0.08, 4.75)	-0.10 (1.77)	(-3.56, 3.37)	0.956
		Irbesartan	129	81 (62.8)	2.51 (1.30)	(-0.04, 5.07)			
	Week 70	Sparsentan	127	103 (81.1)	0.33 (1.18)	(-1.99, 2.64)	0.38 (1.75)	(-3.06, 3.83)	0.827
		Irbesartan	129	84 (65.1)	-0.06 (1.29)	(-2.60, 2.48)			
	Week 94	Sparsentan	127	94 (74.0)	1.74 (1.22)	(-0.66, 4.13)	-0.42 (1.80)	(-3.95, 3.10)	0.814
		Irbesartan	129	82 (63.6)	2.16 (1.31)	(-0.42, 4.74)			
	Week 110	Sparsentan	127	93 (73.2)	1.27 (1.24)	(-1.16, 3.70)	0.45 (1.87)	(-3.22, 4.13)	0.809
		Irbesartan	129	72 (55.8)	0.82 (1.40)	(-1.92, 3.56)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
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Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis			
						Change from Baseline		Treatment Difference	
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	40 (81.6)	4.94 (1.78)	(1.43, 8.44)	1.17 (2.64)	(-4.02, 6.37)	0.657
		Irbesartan	48	34 (70.8)	3.76 (1.94)	(-0.05, 7.58)			
	Week 48	Sparsentan	49	45 (91.8)	3.29 (1.70)	(-0.06, 6.64)	2.37 (2.59)	(-2.73, 7.47)	0.361
		Irbesartan	48	33 (68.8)	0.92 (1.95)	(-2.91, 4.75)			
	Week 70	Sparsentan	49	40 (81.6)	1.01 (1.77)	(-2.48, 4.50)	-0.63 (2.65)	(-5.85, 4.60)	0.814
		Irbesartan	48	32 (66.7)	1.64 (1.97)	(-2.24, 5.51)			
	Week 94	Sparsentan	49	41 (83.7)	0.06 (1.77)	(-3.43, 3.55)	-1.19 (2.61)	(-6.33, 3.95)	0.650
		Irbesartan	48	35 (72.9)	1.25 (1.91)	(-2.52, 5.01)			
	Week 110	Sparsentan	49	37 (75.5)	1.65 (1.86)	(-2.00, 5.31)	-0.06 (2.72)	(-5.42, 5.30)	0.982
		Irbesartan	48	32 (66.7)	1.71 (1.99)	(-2.20, 5.62)			
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	22 (84.6)	5.09 (2.15)	(0.84, 9.33)	2.40 (3.11)	(-3.74, 8.54)	0.442
		Irbesartan	25	20 (80.0)	2.69 (2.25)	(-1.76, 7.13)			
	Week 48	Sparsentan	26	24 (92.3)	2.03 (2.07)	(-2.07, 6.13)	-0.12 (3.13)	(-6.30, 6.07)	0.970
		Irbesartan	25	18 (72.0)	2.14 (2.34)	(-2.47, 6.76)			
	Week 70	Sparsentan	26	23 (88.5)	1.22 (2.11)	(-2.96, 5.39)	-4.56 (3.11)	(-10.70, 1.59)	0.145
		Irbesartan	25	20 (80.0)	5.78 (2.27)	(1.29, 10.26)			
	Week 94	Sparsentan	26	21 (80.8)	3.61 (2.19)	(-0.72, 7.93)	0.23 (3.19)	(-6.08, 6.54)	0.942
		Irbesartan	25	19 (76.0)	3.37 (2.31)	(-1.20, 7.95)			
	Week 110	Sparsentan	26	22 (84.6)	4.13 (2.16)	(-0.15, 8.40)	-0.95 (3.26)	(-7.39, 5.49)	0.771
		Irbesartan	25	17 (68.0)	5.08 (2.43)	(0.28, 9.88)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.097
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	64 (78.0)	3.91 (1.58)	(0.80, 7.01)	5.59 (2.36)	(0.95, 10.23)	0.018 *
		Irbesartan	80	52 (65.0)	-1.69 (1.75)	(-5.13, 1.76)			
	Week 48	Sparsentan	82	63 (76.8)	4.44 (1.58)	(1.32, 7.55)	1.34 (2.38)	(-3.35, 6.03)	0.575
		Irbesartan	80	48 (60.0)	3.10 (1.78)	(-0.40, 6.60)			
	Week 70	Sparsentan	82	65 (79.3)	1.93 (1.57)	(-1.16, 5.02)	3.57 (2.36)	(-1.08, 8.21)	0.132
		Irbesartan	80	50 (62.5)	-1.64 (1.76)	(-5.10, 1.82)			
	Week 94	Sparsentan	82	61 (74.4)	3.25 (1.61)	(0.08, 6.41)	2.07 (2.40)	(-2.65, 6.78)	0.389
		Irbesartan	80	50 (62.5)	1.18 (1.78)	(-2.31, 4.67)			
	Week 110	Sparsentan	82	57 (69.5)	2.34 (1.67)	(-0.94, 5.62)	2.05 (2.52)	(-2.92, 7.01)	0.418
		Irbesartan	80	44 (55.0)	0.29 (1.89)	(-3.42, 4.00)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	38 (84.4)	0.41 (1.72)	(-2.97, 3.80)	0.60 (2.45)	(-4.22, 5.43)	0.805	
		Irbesartan	49	37 (75.5)	-0.19 (1.74)	(-3.62, 3.24)				
	Week 48	Sparsentan	45	36 (80.0)	-1.08 (1.75)	(-4.52, 2.36)	-2.87 (2.53)	(-7.86, 2.12)	0.259	
		Irbesartan	49	33 (67.3)	1.79 (1.82)	(-1.80, 5.38)				
	Week 70	Sparsentan	45	38 (84.4)	-2.40 (1.72)	(-5.79, 1.00)	-4.71 (2.52)	(-9.67, 0.24)	0.062	
		Irbesartan	49	34 (69.4)	2.32 (1.83)	(-1.28, 5.91)				
	Week 94	Sparsentan	45	33 (73.3)	-0.87 (1.81)	(-4.44, 2.71)	-4.54 (2.62)	(-9.69, 0.61)	0.084	
		Irbesartan	49	32 (65.3)	3.67 (1.88)	(-0.02, 7.37)				
	Week 110	Sparsentan	45	36 (80.0)	-0.46 (1.77)	(-3.95, 3.03)	-2.03 (2.68)	(-7.30, 3.25)	0.450	
		Irbesartan	49	28 (57.1)	1.57 (2.00)	(-2.37, 5.51)				
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	40 (81.6)	4.94 (1.78)	(1.43, 8.44)	1.17 (2.64)	(-4.02, 6.37)	0.657	
		Irbesartan	48	34 (70.8)	3.76 (1.94)	(-0.05, 7.58)				
	Week 48	Sparsentan	49	45 (91.8)	3.29 (1.70)	(-0.06, 6.64)	2.37 (2.59)	(-2.73, 7.47)	0.361	
		Irbesartan	48	33 (68.8)	0.92 (1.95)	(-2.91, 4.75)				
	Week 70	Sparsentan	49	40 (81.6)	1.01 (1.77)	(-2.48, 4.50)	-0.63 (2.65)	(-5.85, 4.60)	0.814	
		Irbesartan	48	32 (66.7)	1.64 (1.97)	(-2.24, 5.51)				
	Week 94	Sparsentan	49	41 (83.7)	0.06 (1.77)	(-3.43, 3.55)	-1.19 (2.61)	(-6.33, 3.95)	0.650	
		Irbesartan	48	35 (72.9)	1.25 (1.91)	(-2.52, 5.01)				
	Week 110	Sparsentan	49	37 (75.5)	1.65 (1.86)	(-2.00, 5.31)	-0.06 (2.72)	(-5.42, 5.30)	0.982	
		Irbesartan	48	32 (66.7)	1.71 (1.99)	(-2.20, 5.62)				

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
 Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	22 (84.6)	5.09 (2.15)	(0.84, 9.33)	2.40 (3.11)	(-3.74, 8.54)	0.442
		Irbesartan	25	20 (80.0)	2.69 (2.25)	(-1.76, 7.13)			
	Week 48	Sparsentan	26	24 (92.3)	2.03 (2.07)	(-2.07, 6.13)	-0.12 (3.13)	(-6.30, 6.07)	0.970
		Irbesartan	25	18 (72.0)	2.14 (2.34)	(-2.47, 6.76)			
	Week 70	Sparsentan	26	23 (88.5)	1.22 (2.11)	(-2.96, 5.39)	-4.56 (3.11)	(-10.70, 1.59)	0.145
		Irbesartan	25	20 (80.0)	5.78 (2.27)	(1.29, 10.26)			
	Week 94	Sparsentan	26	21 (80.8)	3.61 (2.19)	(-0.72, 7.93)	0.23 (3.19)	(-6.08, 6.54)	0.942
		Irbesartan	25	19 (76.0)	3.37 (2.31)	(-1.20, 7.95)			
	Week 110	Sparsentan	26	22 (84.6)	4.13 (2.16)	(-0.15, 8.40)	-0.95 (3.26)	(-7.39, 5.49)	0.771
		Irbesartan	25	17 (68.0)	5.08 (2.43)	(0.28, 9.88)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
 Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.075
<= 1.75 g/day	Week 24	Sparsentan	98	84 (85.7)	3.64 (1.13)	(1.42, 5.85)	1.32 (1.74)	(-2.11, 4.74)	0.450
		Irbesartan	93	60 (64.5)	2.32 (1.33)	(-0.29, 4.93)			
	Week 48	Sparsentan	98	84 (85.7)	3.16 (1.12)	(0.95, 5.37)	0.44 (1.73)	(-2.97, 3.84)	0.801
		Irbesartan	93	59 (63.4)	2.72 (1.32)	(0.13, 5.31)			
	Week 70	Sparsentan	98	80 (81.6)	1.00 (1.15)	(-1.25, 3.26)	-2.42 (1.74)	(-5.83, 1.00)	0.165
		Irbesartan	93	62 (66.7)	3.42 (1.30)	(0.86, 5.98)			
	Week 94	Sparsentan	98	80 (81.6)	1.58 (1.15)	(-0.68, 3.85)	-3.05 (1.73)	(-6.45, 0.35)	0.078
		Irbesartan	93	65 (69.9)	4.63 (1.29)	(2.10, 7.17)			
	Week 110	Sparsentan	98	78 (79.6)	0.77 (1.17)	(-1.53, 3.07)	-2.49 (1.80)	(-6.02, 1.03)	0.165
		Irbesartan	93	57 (61.3)	3.26 (1.36)	(0.59, 5.94)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 1.75 g/day	Week 24	Sparsentan	104	80 (76.9)	3.06 (1.39)	(0.33, 5.80)	3.57 (1.96)	(-0.29, 7.43)	0.070
		Irbesartan	109	83 (76.1)	-0.51 (1.38)	(-3.21, 2.20)			
	Week 48	Sparsentan	104	84 (80.8)	1.64 (1.36)	(-1.04, 4.31)	-0.08 (1.99)	(-3.98, 3.82)	0.969
		Irbesartan	109	73 (67.0)	1.71 (1.44)	(-1.11, 4.54)			
	Week 70	Sparsentan	104	86 (82.7)	-0.06 (1.35)	(-2.72, 2.59)	0.37 (1.98)	(-3.52, 4.27)	0.851
		Irbesartan	109	74 (67.9)	-0.44 (1.44)	(-3.27, 2.40)			
	Week 94	Sparsentan	104	76 (73.1)	1.25 (1.42)	(-1.55, 4.04)	1.18 (2.05)	(-2.85, 5.20)	0.567
		Irbesartan	109	71 (65.1)	0.07 (1.47)	(-2.82, 2.96)			
	Week 110	Sparsentan	104	74 (71.2)	2.63 (1.45)	(-0.23, 5.48)	2.18 (2.13)	(-2.00, 6.36)	0.306
		Irbesartan	109	64 (58.7)	0.45 (1.55)	(-2.60, 3.49)			

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
 Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.414
Yes	Week 24	Sparsentan	90	71 (78.9)	5.25 (1.39)	(2.53, 7.98)	3.00 (2.10)	(-1.12, 7.12)	0.153
		Irbesartan	88	55 (62.5)	2.25 (1.57)	(-0.84, 5.33)			
	Week 48	Sparsentan	90	70 (77.8)	4.54 (1.39)	(1.81, 7.27)	0.17 (2.13)	(-4.01, 4.36)	0.935
		Irbesartan	88	51 (58.0)	4.37 (1.61)	(1.20, 7.53)			
	Week 70	Sparsentan	90	71 (78.9)	2.03 (1.39)	(-0.69, 4.75)	2.76 (2.12)	(-1.41, 6.93)	0.195
		Irbesartan	88	52 (59.1)	-0.72 (1.60)	(-3.87, 2.43)			
	Week 94	Sparsentan	90	67 (74.4)	4.11 (1.42)	(1.32, 6.90)	-0.55 (2.09)	(-4.66, 3.56)	0.794
		Irbesartan	88	58 (65.9)	4.66 (1.54)	(1.64, 7.68)			
	Week 110	Sparsentan	90	65 (72.2)	4.02 (1.45)	(1.17, 6.86)	1.25 (2.18)	(-3.03, 5.54)	0.566
		Irbesartan	88	51 (58.0)	2.76 (1.63)	(-0.43, 5.96)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
 Source Data: aqs, created on: 05MAR2024



Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
No	Week 24	Sparsentan	112	93 (83.0)	2.13 (1.16)	(-0.16, 4.41)	2.48 (1.67)	(-0.81, 5.76)	0.139
		Irbesartan	114	88 (77.2)	-0.35 (1.20)	(-2.71, 2.01)			
	Week 48	Sparsentan	112	98 (87.5)	0.93 (1.14)	(-1.30, 3.17)	0.38 (1.67)	(-2.90, 3.67)	0.819
		Irbesartan	114	81 (71.1)	0.55 (1.23)	(-1.86, 2.96)			
	Week 70	Sparsentan	112	95 (84.8)	-0.58 (1.16)	(-2.85, 1.69)	-2.84 (1.68)	(-6.15, 0.47)	0.092
		Irbesartan	114	84 (73.7)	2.26 (1.23)	(-0.14, 4.67)			
	Week 94	Sparsentan	112	89 (79.5)	-0.59 (1.19)	(-2.92, 1.74)	-0.75 (1.74)	(-4.16, 2.66)	0.666
		Irbesartan	114	78 (68.4)	0.16 (1.27)	(-2.33, 2.65)			
	Week 110	Sparsentan	112	87 (77.7)	-0.06 (1.21)	(-2.43, 2.32)	-0.82 (1.80)	(-4.36, 2.71)	0.647
		Irbesartan	114	70 (61.4)	0.77 (1.34)	(-1.85, 3.39)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.781
<= 5 years	Week 24	Sparsentan	113	94 (83.2)	4.08 (1.22)	(1.68, 6.48)	3.54 (1.73)	(0.15, 6.93)	0.041 *
		Irbesartan	127	94 (74.0)	0.54 (1.22)	(-1.86, 2.94)			
	Week 48	Sparsentan	113	95 (84.1)	1.99 (1.21)	(-0.38, 4.37)	0.30 (1.73)	(-3.10, 3.70)	0.862
		Irbesartan	127	89 (70.1)	1.69 (1.24)	(-0.74, 4.13)			
	Week 70	Sparsentan	113	94 (83.2)	-0.37 (1.22)	(-2.76, 2.02)	-2.49 (1.74)	(-5.91, 0.93)	0.154
		Irbesartan	127	89 (70.1)	2.11 (1.25)	(-0.33, 4.56)			
	Week 94	Sparsentan	113	89 (78.8)	1.91 (1.25)	(-0.53, 4.36)	-0.71 (1.76)	(-4.17, 2.75)	0.687
		Irbesartan	127	89 (70.1)	2.63 (1.25)	(0.17, 5.08)			
	Week 110	Sparsentan	113	89 (78.8)	3.13 (1.26)	(0.66, 5.59)	0.15 (1.82)	(-3.42, 3.72)	0.934
		Irbesartan	127	80 (63.0)	2.97 (1.32)	(0.39, 5.56)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 5 years	Week 24	Sparsentan	89	70 (78.7)	2.69 (1.32)	(0.10, 5.28)	1.83 (2.07)	(-2.24, 5.91)	0.377
		Irbesartan	75	49 (65.3)	0.86 (1.59)	(-2.27, 3.99)			
	Week 48	Sparsentan	89	73 (82.0)	3.26 (1.30)	(0.71, 5.81)	0.41 (2.09)	(-3.71, 4.53)	0.844
		Irbesartan	75	43 (57.3)	2.85 (1.64)	(-0.38, 6.07)			
	Week 70	Sparsentan	89	72 (80.9)	1.86 (1.31)	(-0.72, 4.44)	2.29 (2.08)	(-1.80, 6.39)	0.271
		Irbesartan	75	47 (62.7)	-0.43 (1.61)	(-3.60, 2.73)			
	Week 94	Sparsentan	89	67 (75.3)	1.03 (1.35)	(-1.62, 3.68)	-0.05 (2.12)	(-4.21, 4.11)	0.982
		Irbesartan	75	47 (62.7)	1.08 (1.63)	(-2.12, 4.27)			
	Week 110	Sparsentan	89	63 (70.8)	-0.16 (1.39)	(-2.89, 2.56)	0.76 (2.21)	(-3.60, 5.11)	0.733
		Irbesartan	75	41 (54.7)	-0.92 (1.72)	(-4.30, 2.46)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.554
Yes	Week 24	Sparsentan	155	125 (80.6)	4.05 (1.04)	(2.01, 6.10)	3.15 (1.52)	(0.17, 6.12)	0.038 *
		Irbesartan	161	111 (68.9)	0.91 (1.10)	(-1.26, 3.07)			
	Week 48	Sparsentan	155	126 (81.3)	3.15 (1.03)	(1.13, 5.18)	0.22 (1.53)	(-2.79, 3.23)	0.885
		Irbesartan	161	102 (63.4)	2.93 (1.13)	(0.71, 5.16)			
	Week 70	Sparsentan	155	128 (82.6)	1.47 (1.03)	(-0.55, 3.49)	0.30 (1.53)	(-2.70, 3.30)	0.845
		Irbesartan	161	105 (65.2)	1.17 (1.13)	(-1.04, 3.38)			
	Week 94	Sparsentan	155	119 (76.8)	1.98 (1.06)	(-0.10, 4.06)	-0.93 (1.55)	(-3.98, 2.11)	0.548
		Irbesartan	161	105 (65.2)	2.91 (1.13)	(0.69, 5.13)			
	Week 110	Sparsentan	155	114 (73.5)	2.25 (1.09)	(0.12, 4.38)	0.35 (1.62)	(-2.83, 3.54)	0.828
		Irbesartan	161	92 (57.1)	1.90 (1.20)	(-0.47, 4.26)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Table PT2VSC\_FSCM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Change from baseline in EQ-5D VAS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
No	Week 24	Sparsentan	47	39 (83.0)	0.95 (1.80)	(-2.59, 4.49)	1.25 (2.68)	(-4.03, 6.54)	0.640
		Irbesartan	41	32 (78.0)	-0.30 (1.98)	(-4.21, 3.61)			
	Week 48	Sparsentan	47	42 (89.4)	0.07 (1.75)	(-3.38, 3.52)	0.86 (2.67)	(-4.40, 6.12)	0.746
		Irbesartan	41	30 (73.2)	-0.79 (2.00)	(-4.74, 3.15)			
	Week 70	Sparsentan	47	38 (80.9)	-2.61 (1.80)	(-6.17, 0.95)	-4.13 (2.71)	(-9.46, 1.21)	0.129
		Irbesartan	41	31 (75.6)	1.52 (2.00)	(-2.43, 5.47)			
	Week 94	Sparsentan	47	37 (78.7)	-0.36 (1.84)	(-3.98, 3.26)	0.01 (2.75)	(-5.42, 5.44)	0.997
		Irbesartan	41	31 (75.6)	-0.37 (2.03)	(-4.38, 3.64)			
	Week 110	Sparsentan	47	38 (80.9)	-0.33 (1.84)	(-3.96, 3.30)	-1.17 (2.81)	(-6.70, 4.36)	0.677
		Irbesartan	41	29 (70.7)	0.84 (2.10)	(-3.30, 4.98)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. VAS = visual analogue scale.  
Source Data: aqs, created on: 05MAR2024

Figure PF2VSC\_FSGM: Change from baseline in EQ-5D VAS by subgroup  
Full Analysis Set

Not done: No significant subgroup found.

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
eGFR Low =  $30 - < 60 \text{ ml/min/1.73m}^2$ , eGFR High  $\geq 60 \text{ ml/min/1.73m}^2$ , UP Low =  $\leq 1.75 \text{ g/day}$ , UP High =  $> 1.75 \text{ g/day}$ .  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
VAS = visual analogue scale.  
Reference table: PT2VSC\_FSCM

Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.938
Male	Sparsentan	139	28 (20.1)	NE		1.011	(0.589, 1.735)	0.968
	Irbesartan	143	31 (21.7)	NE				
Female	Sparsentan	63	17 (27.0)	NE		1.159	(0.517, 2.600)	0.720
	Irbesartan	59	12 (20.3)	117.1	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.116
<= 45 years	Sparsentan	96	26 (27.1)	NE		1.354	(0.742, 2.471)	0.324
	Irbesartan	99	20 (20.2)	117.1	(NE, NE)			
> 45 years	Sparsentan	106	19 (17.9)	NE		0.738	(0.396, 1.374)	0.338
	Irbesartan	103	23 (22.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024



Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	4 (44.4) No events in 1 group	110.3	(25.0, NE)	NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	23 (22.5)	NE		1.173	(0.645, 2.133)	0.601
	Irbesartan	109	24 (22.0)	117.1	(117.1, NE)			
> 40 years	Sparsentan	91	18 (19.8)	NE		0.795	(0.402, 1.570)	0.509
	Irbesartan	88	19 (21.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.298
North America	Sparsentan	35	9 (25.7)	NE		0.749	(0.302, 1.861)	0.534
	Irbesartan	46	12 (26.1)	NE				
Europe	Sparsentan	98	22 (22.4)	NE		0.994	(0.542, 1.821)	0.984
	Irbesartan	115	23 (20.0)	117.1	(117.1, NE)			
Asia Pacific	Sparsentan	69	14 (20.3)	NE		2.014	(0.713, 5.692)	0.187
	Irbesartan	41	8 (19.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.764
< 27 kg/m**2	Sparsentan	83	19 (22.9)	NE		0.953	(0.450, 2.019)	0.901
	Irbesartan	94	13 (13.8)	NE				
≥ 27 kg/m**2	Sparsentan	119	26 (21.8)	NE		1.000	(0.575, 1.741)	0.999
	Irbesartan	107	29 (27.1)	117.1	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan							Interaction test: 0.078
eGFR Low and UP High	Sparsentan	71	7 (9.9)	NE		0.731	(0.253, 2.107)	0.561
	Irbesartan	74	13 (17.6)	NE				
eGFR Low and UP Low	Sparsentan	55	13 (23.6)	NE		0.708	(0.323, 1.551)	0.388
	Irbesartan	55	14 (25.5)	NE				
eGFR High and UP High	Sparsentan	37	12 (32.4)	NE		0.929	(0.360, 2.392)	0.878
	Irbesartan	36	8 (22.2)	NE				
eGFR High and UP Low	Sparsentan	39	13 (33.3)	NE		3.011	(1.039, 8.729)	0.042 *
	Irbesartan	37	8 (21.6)	117.1	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.181
< 60 mL/min/1.73 m**2	Sparsentan	127	22 (17.3)	NE		0.810	(0.453, 1.448)	0.478
	Irbesartan	129	27 (20.9)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	15 (30.6)	NE		1.151	(0.507, 2.611)	0.737
	Irbesartan	48	12 (25.0)	117.1	(NE, NE)			
≥ 90 mL/min/1.73 m**2	Sparsentan	26	8 (30.8)	NE		2.543	(0.628, 10.294)	0.191
	Irbesartan	25	4 (16.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.182
< 45 mL/min/1.73 m**2	Sparsentan	82	13 (15.9)	NE		0.626	(0.302, 1.298)	0.208
	Irbesartan	80	19 (23.8)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	9 (20.0)	NE		1.522	(0.530, 4.371)	0.435
	Irbesartan	49	8 (16.3)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	15 (30.6)	NE		1.151	(0.507, 2.611)	0.737
	Irbesartan	48	12 (25.0)	117.1	(NE, NE)			
≥ 90 mL/min/1.73 m**2	Sparsentan	26	8 (30.8)	NE		2.543	(0.628, 10.294)	0.191
	Irbesartan	25	4 (16.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.689
<= 1.75 g/day	Sparsentan	98	25 (25.5)	NE		1.006	(0.550, 1.840)	0.986
	Irbesartan	93	22 (23.7)	117.1	(NE, NE)			
> 1.75 g/day	Sparsentan	104	20 (19.2)	NE		1.277	(0.655, 2.490)	0.473
	Irbesartan	109	21 (19.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.048 #
Yes	Sparsentan	90	19 (21.1)	NE		0.636	(0.326, 1.243)	0.186
	Irbesartan	88	20 (22.7)	NE				
No	Sparsentan	112	26 (23.2)	NE		1.539	(0.838, 2.827)	0.165
	Irbesartan	114	23 (20.2)	117.1	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.526
<= 5 years	Sparsentan	113	28 (24.8)	NE		1.158	(0.674, 1.992)	0.595
	Irbesartan	127	29 (22.8)	NE				
> 5 years	Sparsentan	89	17 (19.1)	NE		1.024	(0.484, 2.168)	0.950
	Irbesartan	75	14 (18.7)	117.1	(117.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

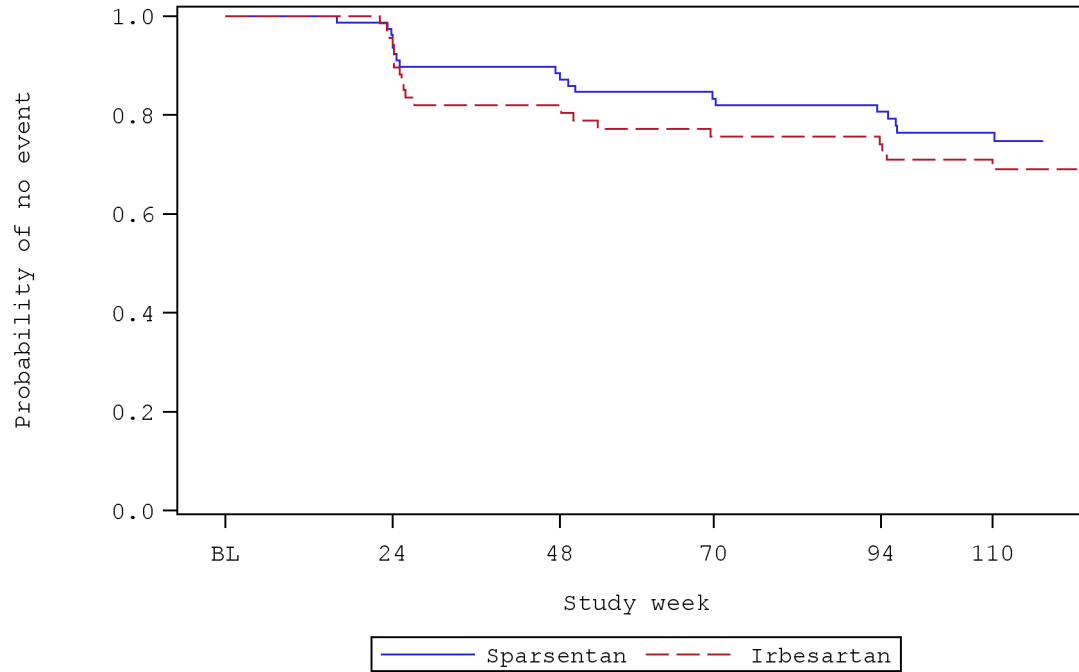
Table PT2VSIT\_FSTM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to increase in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension						Interaction test: 0.176		
Yes	Sparsentan	155	36 (23.2)	NE		0.945	(0.595, 1.501)	0.810
	Irbesartan	161	40 (24.8)	117.1	(117.1, NE)			
No	Sparsentan	47	9 (19.1)	NE		2.503	(0.648, 9.675)	0.183
	Irbesartan	41	3 (7.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Figure PF2VSIT\_FSKM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

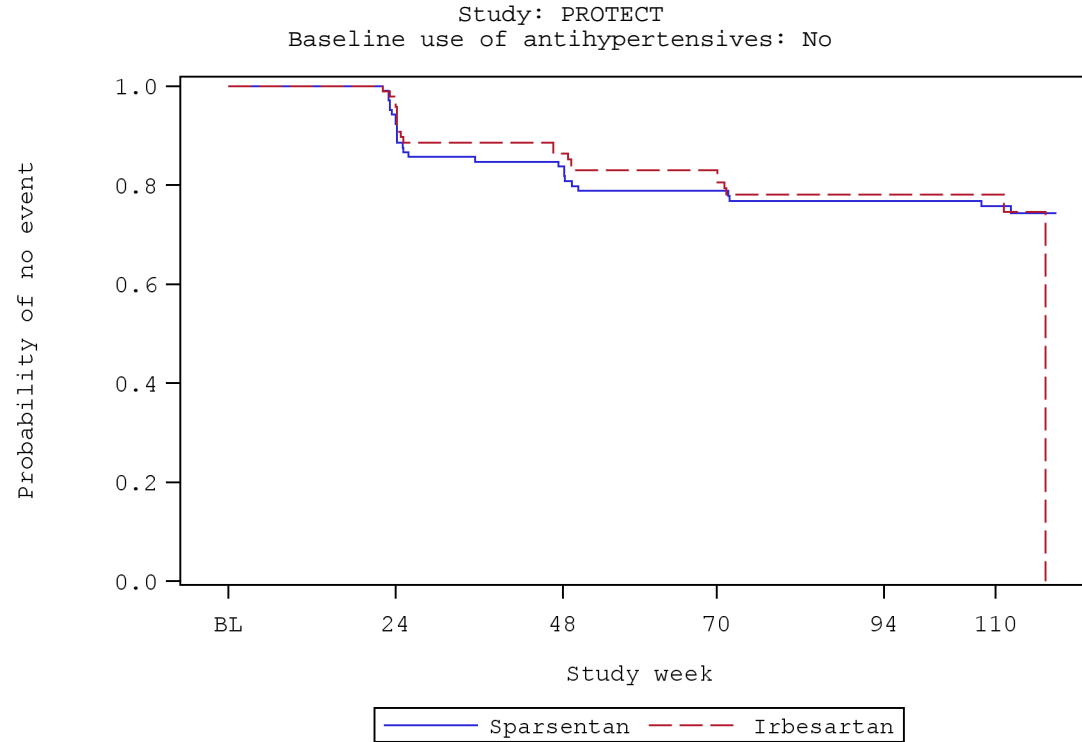
Study: PROTECT  
 Baseline use of antihypertensives: Yes



Sparsentan	90	75	69	64	57	49
Irbesartan	88	65	53	48	47	36

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2VSIT\_FSTM

Figure PF2VSIT\_FSKM: Time to increase in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	112	99	86	80	74	64
Irbesartan	114	94	78	68	62	54

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2VSIT\_FSTM

Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.941
Male	Sparsentan	139	29 (20.9)	NE		1.268	(0.707, 2.275)	0.426
	Irbesartan	143	19 (13.3)	118.1	(NE, NE)			
Female	Sparsentan	63	20 (31.7)	NE		1.323	(0.653, 2.679)	0.437
	Irbesartan	59	13 (22.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.510
<= 45 years	Sparsentan	96	24 (25.0)	NE		1.186	(0.651, 2.162)	0.578
	Irbesartan	99	20 (20.2)	NE				
> 45 years	Sparsentan	106	25 (23.6)	NE		1.621	(0.810, 3.244)	0.172
	Irbesartan	103	12 (11.7)	118.1	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	0.648
<= 18 years	Sparsentan	9	2 (22.2)	NE		0.473	(0.025, 8.929)	0.618
	Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Sparsentan	102	23 (22.5)	NE		1.170	(0.639, 2.139)	0.611
	Irbesartan	109	21 (19.3)	118.1	(NE, NE)			
> 40 years	Sparsentan	91	24 (26.4)	NE		1.714	(0.810, 3.630)	0.159
	Irbesartan	88	10 (11.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.298
North America	Sparsentan	35	5 (14.3)	NE		0.796	(0.258, 2.453)	0.691
	Irbesartan	46	8 (17.4)	NE				
Europe	Sparsentan	98	25 (25.5)	NE		1.256	(0.693, 2.278)	0.453
	Irbesartan	115	20 (17.4)	118.1	(NE, NE)			
Asia Pacific	Sparsentan	69	19 (27.5)	NE		2.253	(0.752, 6.752)	0.147
	Irbesartan	41	4 (9.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024



Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan							Interaction test: 0.087
< 27 kg/m**2	Sparsentan	83	25 (30.1)	NE		2.031	(1.069, 3.860)	0.031 *
	Irbesartan	94	17 (18.1)	118.1	(NE, NE)			
≥ 27 kg/m**2	Sparsentan	119	24 (20.2)	NE		0.906	(0.468, 1.752)	0.769
	Irbesartan	107	15 (14.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan							Interaction test: 0.159
eGFR Low and UP High	Sparsentan	71	17 (23.9)	NE		0.764	(0.386, 1.515)	0.441
	Irbesartan	74	17 (23.0)	118.1	(NE, NE)			
eGFR Low and UP Low	Sparsentan	55	12 (21.8)	NE		1.637	(0.644, 4.163)	0.301
	Irbesartan	55	7 (12.7)	NE				
eGFR High and UP High	Sparsentan	37	9 (24.3)	NE		2.496	(0.670, 9.301)	0.173
	Irbesartan	36	3 (8.3)	NE				
eGFR High and UP Low	Sparsentan	39	11 (28.2)	NE		2.569	(0.875, 7.544)	0.086
	Irbesartan	37	5 (13.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.272
< 60 mL/min/1.73 m**2	Sparsentan	127	29 (22.8)	NE		1.037	(0.597, 1.802)	0.898
	Irbesartan	129	23 (17.8)	118.1	(NE, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	11 (22.4)	NE		1.757	(0.636, 4.852)	0.277
	Irbesartan	48	6 (12.5)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	9 (34.6)	NE		3.688	(0.907, 15.000)	0.068
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: 0.009 #
< 45 mL/min/1.73 m**2	Sparsentan	82	15 (18.3)	NE		0.549	(0.272, 1.106)	0.093
	Irbesartan	80	18 (22.5)	118.1	(NE, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	14 (31.1)	NE		3.421	(1.172, 9.981)	0.024 *
	Irbesartan	49	5 (10.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	11 (22.4)	NE		1.757	(0.636, 4.852)	0.277
	Irbesartan	48	6 (12.5)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	9 (34.6)	NE		3.688	(0.907, 15.000)	0.068
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.020 #
<= 1.75 g/day	Sparsentan	98	26 (26.5)	NE		2.995	(1.340, 6.691)	0.007 *
	Irbesartan	93	8 (8.6)	NE				
> 1.75 g/day	Sparsentan	104	23 (22.1)	NE		0.882	(0.494, 1.577)	0.673
	Irbesartan	109	24 (22.0)	118.1	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.793
Yes	Sparsentan	90	19 (21.1)	NE		1.236	(0.593, 2.576)	0.572
	Irbesartan	88	12 (13.6)	118.1	(NE, NE)			
No	Sparsentan	112	30 (26.8)	NE		1.399	(0.792, 2.472)	0.248
	Irbesartan	114	20 (17.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.557
<= 5 years	Sparsentan	113	30 (26.5)	NE		1.510	(0.860, 2.651)	0.151
	Irbesartan	127	21 (16.5)	NE				
> 5 years	Sparsentan	89	19 (21.3)	NE		1.169	(0.551, 2.477)	0.684
	Irbesartan	75	11 (14.7)	118.1	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

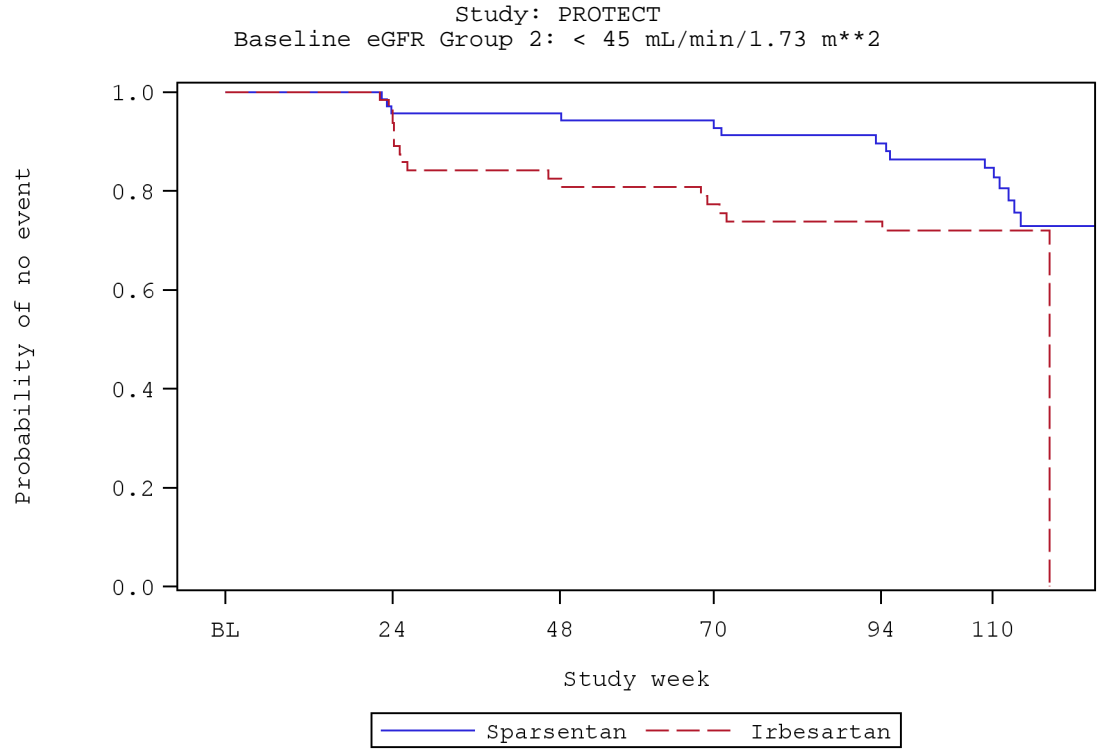
Table PT2VSDT\_FSTM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set

Time to decrease in EQ-5D VAS by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension		Sparsentan				Interaction test: 0.532		
Yes	Sparsentan	155	37 (23.9)	NE		1.461	(0.861, 2.480)	0.160
	Irbesartan	161	22 (13.7)	118.1	(NE, NE)			
No	Sparsentan	47	12 (25.5)	NE		1.154	(0.475, 2.799)	0.752
	Irbesartan	41	10 (24.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



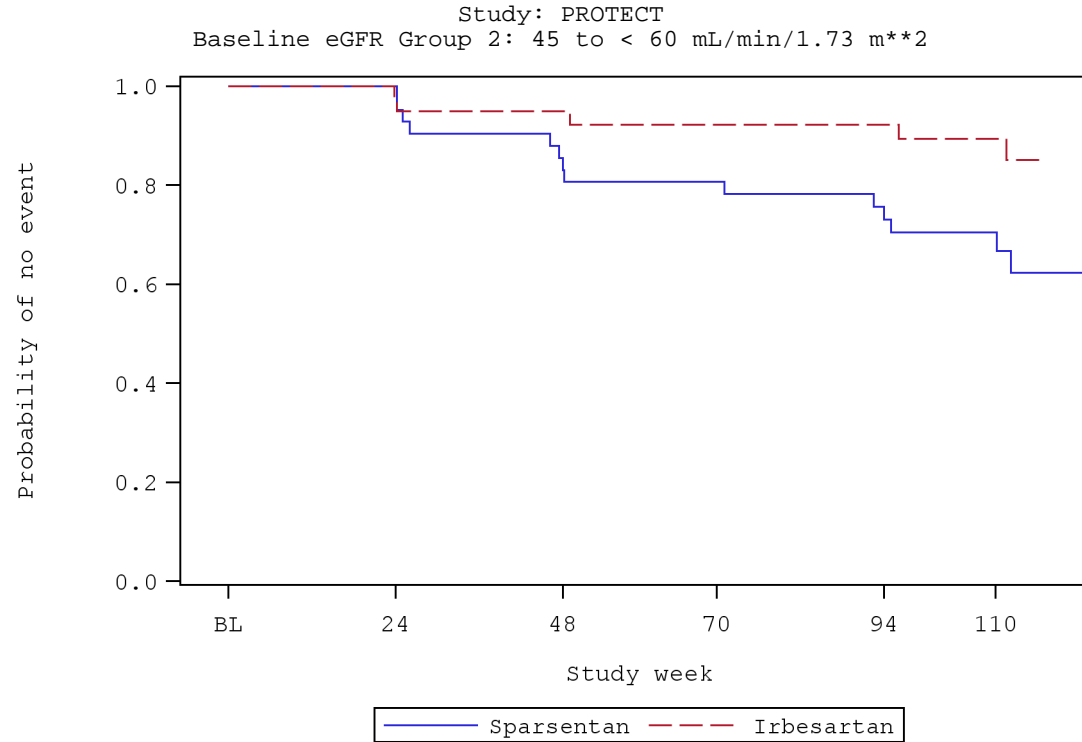
Figure PF2VSDT\_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	82	67	66	64	56	46
Irbesartan	80	62	49	44	42	34

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2VSDT\_FSTM

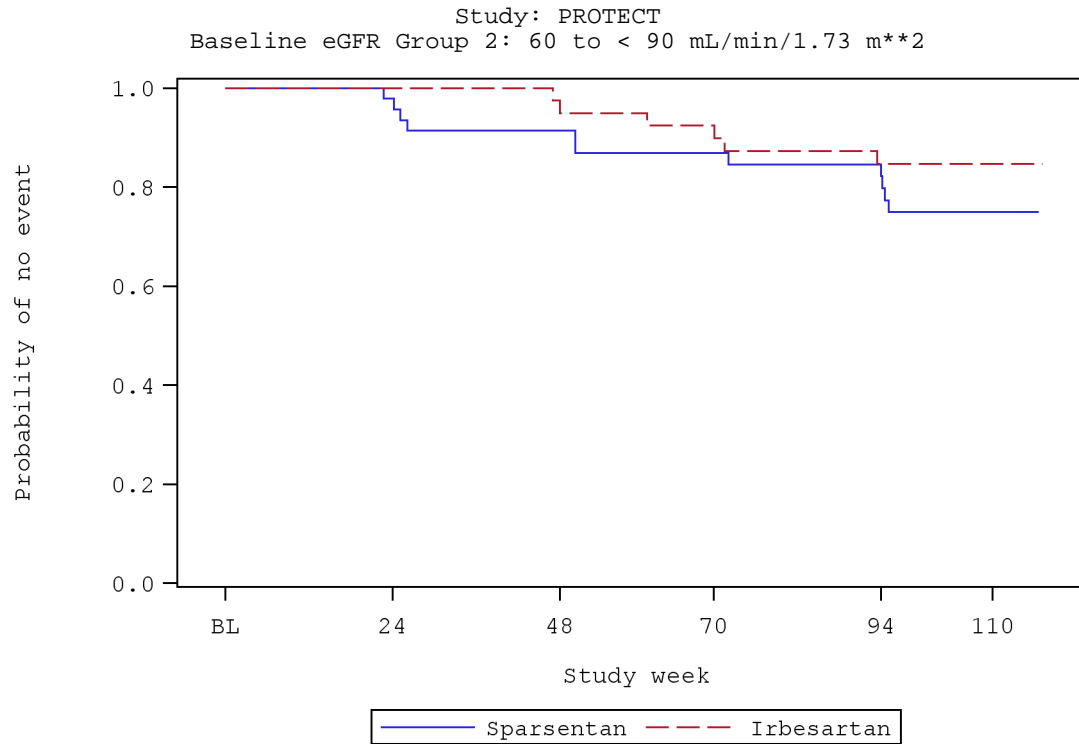
Figure PF2VSDT\_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	45	42	35	33	29	22
Irbesartan	49	39	35	34	32	28

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2VSDT\_FSTM

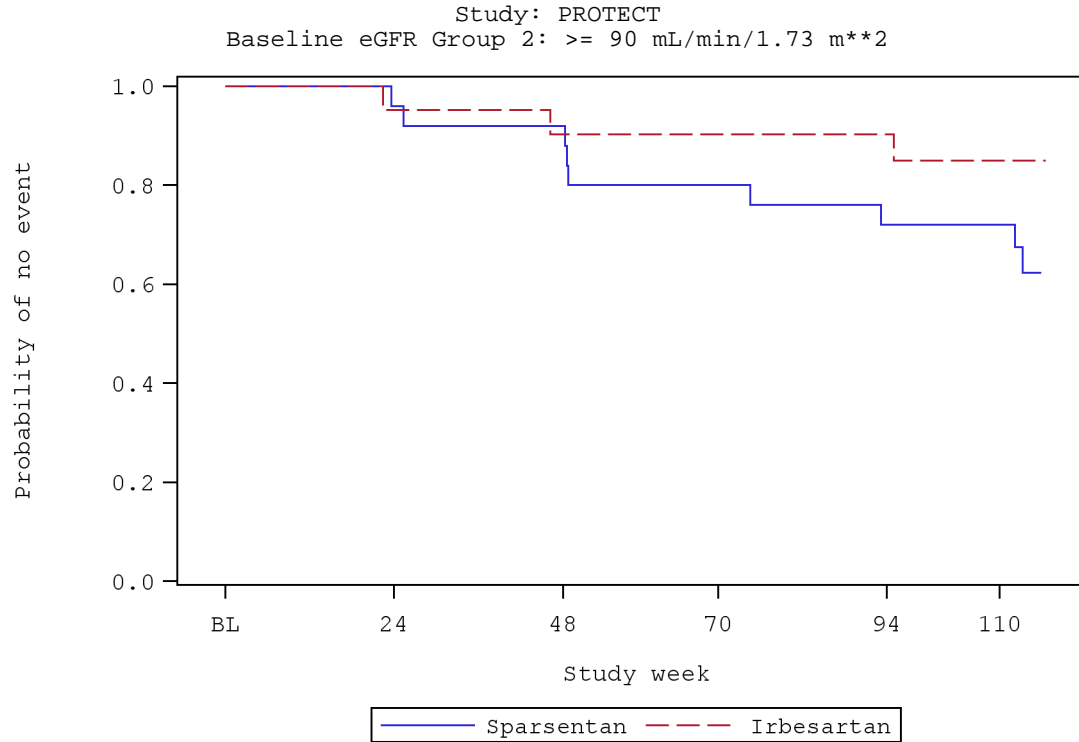
Figure PF2VSDT\_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	49	46	42	39	35	29
Irbesartan	48	41	39	36	33	28

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2VSDT\_FSTM

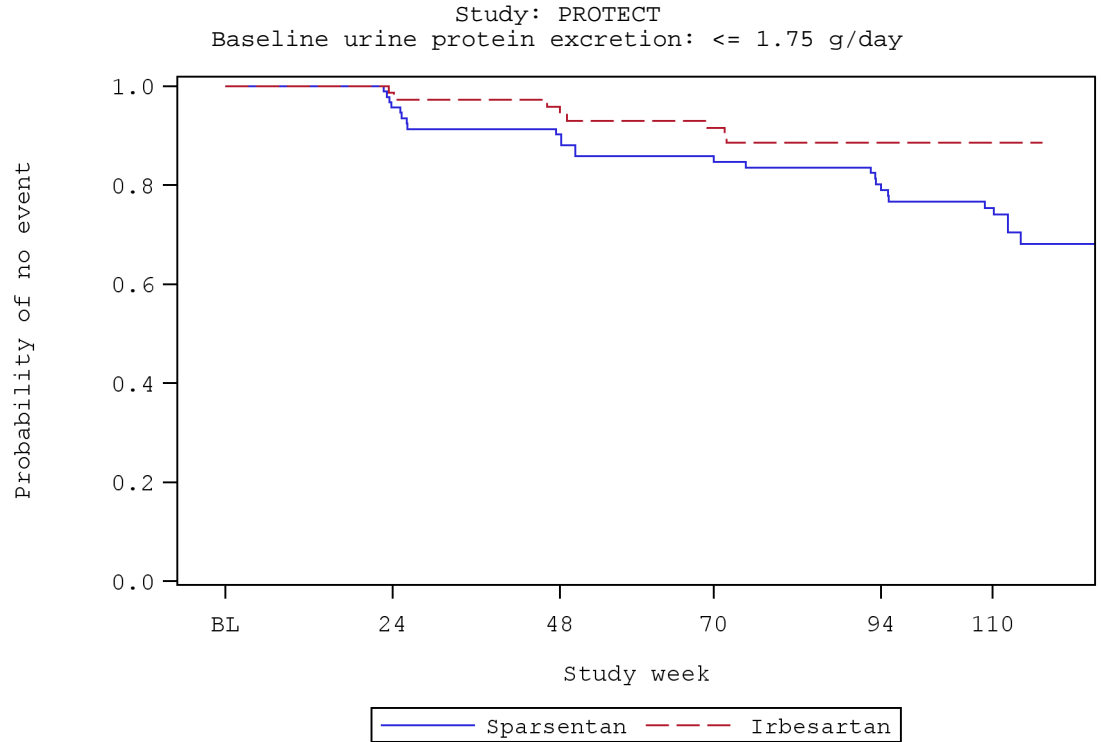
Figure PF2VSDT\_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	26	24	23	20	18	17
Irbesartan	25	20	18	18	17	14

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2VSDT\_FSTM

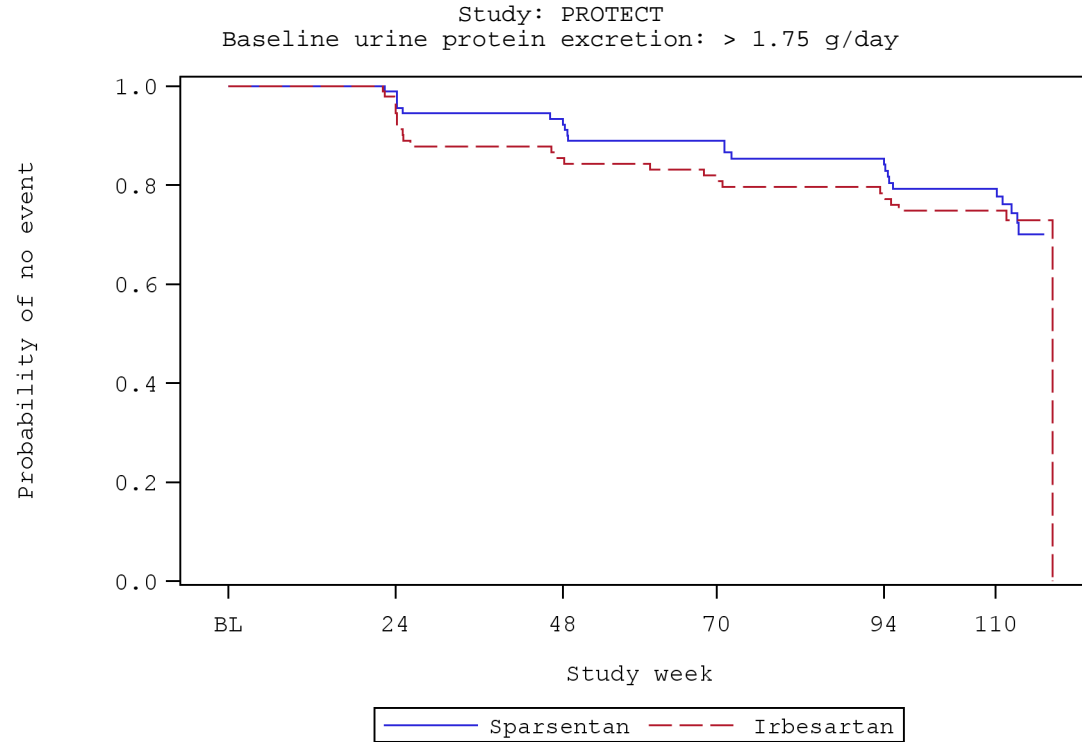
Figure PF2VSDT\_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	98	89	82	78	69	57
Irbesartan	93	73	68	63	58	47

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2VSDT\_FSTM

Figure PF2VSDT\_FSKM: Time to decrease in EQ-5D VAS by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	104	90	84	78	69	57
Irbesartan	109	89	73	69	66	57

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2VSDT\_FSTM

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Sex													
Male	KDQOL: burden of kidney disease	Baseline	Sparsentan	139	129 (92.8)	72.09 (22.63)	6.3	62.50	75.00	87.50	100.0		
			Irbesartan	143	131 (91.6)	79.29 (22.24)	6.3	68.75	87.50	100.00	100.0		
		Week 24	Sparsentan	139	109 (78.4)	76.38 (22.25)	0.0	68.75	81.25	93.75	100.0		
			Irbesartan	143	93 (65.0)	76.88 (23.92)	0.0	68.75	81.25	100.00	100.0		
		Week 48	Sparsentan	139	112 (80.6)	77.23 (22.74)	0.0	68.75	81.25	100.00	100.0		
			Irbesartan	143	87 (60.8)	81.11 (21.35)	0.0	75.00	87.50	100.00	100.0		
		Week 70	Sparsentan	139	114 (82.0)	78.89 (21.51)	6.3	68.75	81.25	93.75	100.0		
			Irbesartan	143	92 (64.3)	79.48 (22.72)	12.5	68.75	87.50	100.00	100.0		
		Week 94	Sparsentan	139	105 (75.5)	78.51 (21.48)	0.0	75.00	81.25	93.75	100.0		
			Irbesartan	143	96 (67.1)	75.91 (23.46)	12.5	68.75	78.13	96.88	100.0		
		Week 110	Sparsentan	139	103 (74.1)	77.97 (21.82)	25.0	62.50	81.25	100.00	100.0		
			Irbesartan	143	89 (62.2)	75.42 (23.72)	12.5	62.50	75.00	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	139	109 (78.4)	6.02 (20.94)	-50.0	-6.25	6.25	12.50	62.5	0.43 [0.15, 0.71]
				Irbesartan	143	93 (65.0)	-2.35 (17.68)	-50.0	-12.50	0.00	6.25	50.0	
	Week 48		Sparsentan	139	112 (80.6)	6.03 (20.01)	-68.8	0.00	6.25	18.75	50.0	0.24 [-0.04, 0.52]	
			Irbesartan	143	87 (60.8)	1.51 (17.83)	-62.5	-6.25	0.00	12.50	56.3		
	Week 70		Sparsentan	139	114 (82.0)	7.13 (18.65)	-50.0	0.00	6.25	18.75	56.3	0.30 [0.02, 0.57]	
			Irbesartan	143	92 (64.3)	1.77 (17.44)	-43.8	-6.25	0.00	9.38	56.3		
Week 94	Sparsentan	139	105 (75.5)	7.50 (23.93)	-87.5	0.00	6.25	25.00	68.8	0.43 [0.15, 0.71]			
	Irbesartan	143	96 (67.1)	-1.89 (19.22)	-75.0	-12.50	0.00	6.25	56.3				
Week 110	Sparsentan	139	103 (74.1)	6.67 (21.78)	-50.0	-6.25	6.25	18.75	62.5	0.44 [0.16, 0.73]			
	Irbesartan	143	89 (62.2)	-2.53 (19.62)	-62.5	-12.50	0.00	6.25	56.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Female	KDQOL: burden of kidney disease	Baseline	Sparsentan	63	58 (92.1)	65.52 (29.99)	0.0	50.00	71.88	93.75	100.0	
			Irbesartan	59	51 (86.4)	70.83 (23.31)	18.8	56.25	75.00	87.50	100.0	
		Week 24	Sparsentan	63	48 (76.2)	77.60 (22.47)	18.8	68.75	81.25	100.00	100.0	
			Irbesartan	59	38 (64.4)	83.06 (19.27)	25.0	75.00	87.50	100.00	100.0	
		Week 48	Sparsentan	63	50 (79.4)	77.88 (23.26)	12.5	62.50	84.38	100.00	100.0	
			Irbesartan	59	37 (62.7)	77.53 (21.27)	6.3	68.75	81.25	100.00	100.0	
		Week 70	Sparsentan	63	46 (73.0)	77.45 (22.92)	25.0	62.50	81.25	100.00	100.0	
			Irbesartan	59	38 (64.4)	79.77 (18.75)	25.0	75.00	81.25	100.00	100.0	
		Week 94	Sparsentan	63	46 (73.0)	71.06 (25.19)	0.0	56.25	75.00	93.75	100.0	
			Irbesartan	59	34 (57.6)	78.86 (18.72)	37.5	68.75	75.00	100.00	100.0	
		Week 110	Sparsentan	63	46 (73.0)	74.46 (20.83)	18.8	62.50	75.00	93.75	100.0	
			Irbesartan	59	32 (54.2)	78.13 (17.53)	25.0	68.75	78.13	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	63	48 (76.2)	11.72 (25.09)	-81.3	0.00	6.25	25.00	81.3	0.17 [-0.25, 0.60]
			Irbesartan	59	38 (64.4)	8.06 (14.88)	-18.8	0.00	6.25	18.75	56.3	
		Week 48	Sparsentan	63	50 (79.4)	11.50 (21.22)	-25.0	0.00	6.25	25.00	62.5	0.38 [-0.05, 0.81]
			Irbesartan	59	37 (62.7)	3.89 (18.48)	-50.0	-6.25	0.00	12.50	50.0	
		Week 70	Sparsentan	63	46 (73.0)	11.28 (28.12)	-43.8	0.00	0.00	25.00	87.5	0.26 [-0.17, 0.69]
			Irbesartan	59	38 (64.4)	4.93 (19.45)	-25.0	-6.25	0.00	12.50	50.0	
		Week 94	Sparsentan	63	46 (73.0)	6.66 (26.33)	-50.0	-12.50	0.00	25.00	75.0	0.09 [-0.35, 0.54]
			Irbesartan	59	34 (57.6)	4.23 (24.80)	-50.0	-6.25	0.00	18.75	50.0	
Week 110	Sparsentan	63	46 (73.0)	9.10 (29.86)	-56.3	-6.25	6.25	18.75	87.5	0.23 [-0.22, 0.69]		
	Irbesartan	59	32 (54.2)	2.73 (22.89)	-43.8	-12.50	0.00	21.88	43.8			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age												
<= 45 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	96	87 (90.6)	67.10 (26.85)	0.0	50.00	75.00	87.50	100.0	
			Irbesartan	99	92 (92.9)	71.20 (25.00)	6.3	53.13	75.00	93.75	100.0	
		Week 24	Sparsentan	96	71 (74.0)	76.23 (22.70)	18.8	62.50	81.25	100.00	100.0	
			Irbesartan	99	67 (67.7)	76.31 (25.50)	0.0	62.50	81.25	100.00	100.0	
		Week 48	Sparsentan	96	75 (78.1)	75.83 (23.46)	6.3	62.50	81.25	93.75	100.0	
			Irbesartan	99	62 (62.6)	77.82 (26.07)	0.0	62.50	87.50	100.00	100.0	
		Week 70	Sparsentan	96	72 (75.0)	76.91 (19.84)	25.0	68.75	75.00	93.75	100.0	
			Irbesartan	99	62 (62.6)	78.02 (23.70)	12.5	68.75	84.38	100.00	100.0	
		Week 94	Sparsentan	96	68 (70.8)	73.62 (23.92)	0.0	62.50	78.13	93.75	100.0	
			Irbesartan	99	64 (64.6)	75.49 (25.35)	12.5	62.50	81.25	100.00	100.0	
		Week 110	Sparsentan	96	69 (71.9)	75.91 (21.36)	25.0	62.50	75.00	93.75	100.0	
			Irbesartan	99	59 (59.6)	73.31 (24.94)	25.0	56.25	75.00	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	96	71 (74.0)	10.83 (27.16)	-81.3	0.00	6.25	25.00	81.3	0.36 [0.02, 0.70]
			Irbesartan	99	67 (67.7)	2.15 (20.40)	-50.0	-6.25	0.00	12.50	56.3	
		Week 48	Sparsentan	96	75 (78.1)	9.33 (20.20)	-62.5	0.00	6.25	25.00	50.0	0.32 [-0.02, 0.65]
			Irbesartan	99	62 (62.6)	2.92 (20.36)	-62.5	-6.25	0.00	12.50	56.3	
		Week 70	Sparsentan	96	72 (75.0)	9.72 (25.41)	-43.8	-6.25	6.25	25.00	87.5	0.24 [-0.10, 0.58]
			Irbesartan	99	62 (62.6)	4.23 (18.83)	-43.8	-6.25	3.13	12.50	43.8	
Week 94	Sparsentan	96	68 (70.8)	6.80 (27.03)	-62.5	-9.38	6.25	25.00	75.0	0.16 [-0.18, 0.50]		
	Irbesartan	99	64 (64.6)	3.03 (20.32)	-37.5	-12.50	6.25	18.75	56.3			
Week 110	Sparsentan	96	69 (71.9)	8.42 (29.19)	-56.3	-12.50	0.00	18.75	87.5	0.35 [-0.00, 0.70]		
	Irbesartan	99	59 (59.6)	-0.53 (20.64)	-56.3	-12.50	0.00	6.25	50.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 45 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	106	100 (94.3)	72.63 (23.60)	6.3	56.25	75.00	90.63	100.0		
			Irbesartan	103	90 (87.4)	82.78 (18.70)	18.8	75.00	87.50	100.00	100.0		
		Week 24	Sparsentan	106	86 (81.1)	77.18 (22.00)	0.0	68.75	81.25	93.75	100.0		
			Irbesartan	103	64 (62.1)	81.15 (19.40)	18.8	68.75	81.25	100.00	100.0		
		Week 48	Sparsentan	106	87 (82.1)	78.81 (22.32)	0.0	75.00	81.25	100.00	100.0		
			Irbesartan	103	62 (60.2)	82.26 (15.02)	37.5	75.00	81.25	93.75	100.0		
		Week 70	Sparsentan	106	88 (83.0)	79.76 (23.42)	6.3	75.00	87.50	100.00	100.0		
			Irbesartan	103	68 (66.0)	80.97 (19.48)	12.5	75.00	84.38	100.00	100.0		
		Week 94	Sparsentan	106	83 (78.3)	78.39 (21.83)	0.0	75.00	81.25	100.00	100.0		
			Irbesartan	103	66 (64.1)	77.84 (18.96)	25.0	68.75	75.00	93.75	100.0		
		Week 110	Sparsentan	106	80 (75.5)	77.73 (21.74)	18.8	62.50	81.25	93.75	100.0		
			Irbesartan	103	62 (60.2)	78.83 (19.08)	12.5	75.00	81.25	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	106	86 (81.1)	5.23 (17.20)	-43.8	-6.25	0.00	12.50	43.8	0.39 [0.06, 0.71]
				Irbesartan	103	64 (62.1)	-0.88 (13.86)	-31.3	-9.38	0.00	6.25	37.5	
	Week 48		Sparsentan	106	87 (82.1)	6.32 (20.74)	-68.8	0.00	6.25	18.75	62.5	0.26 [-0.07, 0.58]	
			Irbesartan	103	62 (60.2)	1.51 (15.38)	-50.0	-6.25	0.00	6.25	43.8		
	Week 70		Sparsentan	106	88 (83.0)	7.17 (18.36)	-50.0	0.00	6.25	18.75	50.0	0.33 [0.01, 0.65]	
			Irbesartan	103	68 (66.0)	1.29 (17.30)	-37.5	-6.25	0.00	6.25	56.3		
	Week 94		Sparsentan	106	83 (78.3)	7.61 (22.58)	-87.5	0.00	6.25	25.00	68.8	0.51 [0.18, 0.83]	
			Irbesartan	103	66 (64.1)	-3.50 (21.09)	-75.0	-12.50	0.00	6.25	50.0		
Week 110	Sparsentan	106	80 (75.5)	6.56 (19.69)	-43.8	-6.25	6.25	18.75	56.3	0.41 [0.08, 0.75]			
	Irbesartan	103	62 (60.2)	-1.71 (20.66)	-62.5	-12.50	0.00	6.25	56.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: Age at IgAN diagnosis												
<= 18 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	9	9 (100.0)	58.33 (30.94)	6.3	37.50	62.50	68.75	100.0	
			Irbesartan	5	5 (100.0)	72.50 (31.12)	18.8	75.00	81.25	93.75	93.8	
		Week 24	Sparsentan	9	5 (55.6)	91.25 (13.69)	68.8	87.50	100.00	100.00	100.0	
			Irbesartan	5	5 (100.0)	77.50 (28.16)	31.3	75.00	81.25	100.00	100.0	
		Week 48	Sparsentan	9	7 (77.8)	64.29 (33.80)	6.3	31.25	75.00	93.75	100.0	
			Irbesartan	5	3 (60.0)	87.50 (10.83)	81.3	81.25	81.25	100.00	100.0	
		Week 70	Sparsentan	9	7 (77.8)	68.75 (24.47)	43.8	43.75	68.75	100.00	100.0	
			Irbesartan	5	4 (80.0)	76.56 (35.49)	25.0	53.13	90.63	100.00	100.0	
		Week 94	Sparsentan	9	5 (55.6)	71.25 (21.01)	37.5	68.75	75.00	81.25	93.8	
			Irbesartan	5	4 (80.0)	68.75 (34.99)	25.0	40.63	75.00	96.88	100.0	
		Week 110	Sparsentan	9	4 (44.4)	81.25 (16.14)	62.5	68.75	81.25	93.75	100.0	
			Irbesartan	5	2 (40.0)	50.00 (26.52)	31.3	31.25	50.00	68.75	68.8	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	9	5 (55.6)	27.50 (19.06)	0.0	18.75	31.25	37.50	50.0	1.61 [0.18, 3.04]
			Irbesartan	5	5 (100.0)	5.00 (5.23)	0.0	0.00	6.25	6.25	12.5	
		Week 48	Sparsentan	9	7 (77.8)	3.57 (32.65)	-62.5	-6.25	12.50	25.00	37.5	-0.02 [-1.37, 1.33]
			Irbesartan	5	3 (60.0)	4.17 (3.61)	0.0	0.00	6.25	6.25	6.3	
		Week 70	Sparsentan	9	7 (77.8)	8.04 (13.84)	0.0	0.00	0.00	12.50	37.5	0.29 [-0.94, 1.53]
			Irbesartan	5	4 (80.0)	4.69 (3.13)	0.0	3.13	6.25	6.25	6.3	
Week 94		Sparsentan	9	5 (55.6)	7.50 (26.66)	-31.3	-6.25	12.50	31.25	31.3	0.47 [-0.86, 1.81]	
		Irbesartan	5	4 (80.0)	-3.13 (14.88)	-25.0	-12.50	3.13	6.25	6.3		
Week 110		Sparsentan	9	4 (44.4)	9.38 (21.35)	-12.5	-6.25	6.25	25.00	37.5	0.46 [-1.26, 2.17]	
		Irbesartan	5	2 (40.0)	0.00 (17.68)	-12.5	-12.50	0.00	12.50	12.5		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 18 to 40 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	102	94 (92.2)	69.95 (25.14)	0.0	62.50	75.00	87.50	100.0	
		Week 24	Irbesartan	109	99 (90.8)	73.23 (24.39)	6.3	56.25	75.00	93.75	100.0	
			Sparsentan	102	79 (77.5)	76.34 (21.76)	18.8	62.50	81.25	93.75	100.0	
		Week 48	Irbesartan	109	71 (65.1)	76.76 (24.94)	0.0	62.50	81.25	100.00	100.0	
			Sparsentan	102	82 (80.4)	76.37 (22.82)	0.0	62.50	81.25	93.75	100.0	
		Week 70	Irbesartan	109	67 (61.5)	77.24 (25.51)	0.0	62.50	87.50	100.00	100.0	
			Sparsentan	102	78 (76.5)	79.57 (19.18)	25.0	68.75	81.25	93.75	100.0	
		Week 94	Irbesartan	109	68 (62.4)	78.77 (23.06)	12.5	68.75	87.50	100.00	100.0	
			Sparsentan	102	73 (71.6)	74.91 (23.33)	0.0	68.75	81.25	93.75	100.0	
		Week 110	Irbesartan	109	69 (63.3)	76.81 (24.63)	12.5	62.50	81.25	100.00	100.0	
			Sparsentan	102	72 (70.6)	75.52 (21.41)	25.0	62.50	75.00	96.88	100.0	
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	102	79 (77.5)	8.54 (25.70)	-81.3	0.00	6.25	18.75	81.3
	Irbesartan			109	71 (65.1)	0.97 (19.47)	-50.0	-6.25	0.00	6.25	56.3	
	Week 48		Sparsentan	102	82 (80.4)	7.24 (21.64)	-68.8	0.00	3.13	18.75	50.0	0.25 [-0.07, 0.57]
			Irbesartan	109	67 (61.5)	2.15 (18.86)	-62.5	-6.25	0.00	12.50	56.3	
	Week 70		Sparsentan	102	78 (76.5)	10.10 (25.29)	-43.8	-6.25	6.25	25.00	87.5	0.28 [-0.05, 0.61]
			Irbesartan	109	68 (62.4)	3.77 (18.80)	-43.8	-6.25	0.00	12.50	50.0	
	Week 94	Sparsentan	102	73 (71.6)	6.76 (25.51)	-62.5	-6.25	6.25	18.75	75.0	0.17 [-0.16, 0.50]	
Irbesartan		109	69 (63.3)	2.72 (20.85)	-50.0	-6.25	0.00	12.50	56.3			
Week 110	Sparsentan	102	72 (70.6)	8.33 (28.69)	-56.3	-12.50	3.13	18.75	87.5	0.36 [0.02, 0.69]		
	Irbesartan	109	65 (59.6)	-0.58 (20.18)	-56.3	-12.50	0.00	6.25	50.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	91	84 (92.3)	71.43 (24.72)	6.3	50.00	75.00	90.63	100.0	
			Irbesartan	88	78 (88.6)	81.89 (19.26)	18.8	75.00	87.50	100.00	100.0	
		Week 24	Sparsentan	91	73 (80.2)	76.20 (23.12)	0.0	68.75	81.25	93.75	100.0	
			Irbesartan	88	55 (62.5)	81.25 (19.28)	18.8	68.75	81.25	100.00	100.0	
		Week 48	Sparsentan	91	73 (80.2)	79.88 (21.46)	6.3	75.00	81.25	100.00	100.0	
			Irbesartan	88	54 (61.4)	83.10 (14.70)	50.0	75.00	81.25	100.00	100.0	
		Week 70	Sparsentan	91	75 (82.4)	78.25 (24.19)	6.3	75.00	81.25	100.00	100.0	
			Irbesartan	88	58 (65.9)	80.71 (18.92)	12.5	75.00	78.13	100.00	100.0	
		Week 94	Sparsentan	91	73 (80.2)	77.91 (22.63)	0.0	75.00	81.25	93.75	100.0	
			Irbesartan	88	57 (64.8)	77.08 (18.35)	25.0	68.75	75.00	93.75	100.0	
		Week 110	Sparsentan	91	73 (80.2)	78.00 (22.00)	18.8	62.50	81.25	93.75	100.0	
			Irbesartan	88	54 (61.4)	78.13 (19.07)	12.5	75.00	81.25	93.75	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	91	73 (80.2)	5.57 (17.76)	-43.8	-6.25	0.00	12.50	43.8	0.34 [-0.02, 0.69]
			Irbesartan	88	55 (62.5)	-0.11 (15.57)	-31.3	-12.50	0.00	6.25	43.8	
		Week 48	Sparsentan	91	73 (80.2)	8.65 (17.88)	-43.8	0.00	6.25	18.75	62.5	0.36 [0.01, 0.72]
			Irbesartan	88	54 (61.4)	2.20 (17.52)	-50.0	-6.25	0.00	12.50	50.0	
		Week 70	Sparsentan	91	75 (82.4)	6.50 (18.22)	-50.0	0.00	6.25	18.75	50.0	0.29 [-0.06, 0.63]
			Irbesartan	88	58 (65.9)	1.29 (17.78)	-37.5	-6.25	0.00	6.25	56.3	
Week 94	Sparsentan	91	73 (80.2)	7.71 (23.88)	-87.5	0.00	6.25	25.00	68.8	0.50 [0.15, 0.86]		
	Irbesartan	88	57 (64.8)	-3.73 (21.02)	-75.0	-18.75	0.00	6.25	50.0			
Week 110	Sparsentan	91	73 (80.2)	6.42 (19.98)	-43.8	-6.25	6.25	18.75	56.3	0.40 [0.05, 0.76]		
	Irbesartan	88	54 (61.4)	-1.85 (21.43)	-62.5	-18.75	0.00	6.25	56.3			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Geographic region													
North America	KDQOL: burden of kidney disease	Baseline	Sparsentan	35	31 (88.6)	72.38 (25.61)	6.3	62.50	75.00	93.75	100.0		
			Irbesartan	46	43 (93.5)	82.99 (17.11)	43.8	68.75	81.25	100.00	100.0		
		Week 24	Sparsentan	35	23 (65.7)	80.71 (18.84)	25.0	68.75	81.25	100.00	100.0		
			Irbesartan	46	35 (76.1)	83.21 (21.37)	6.3	75.00	87.50	100.00	100.0		
		Week 48	Sparsentan	35	25 (71.4)	78.00 (24.94)	12.5	68.75	81.25	100.00	100.0		
			Irbesartan	46	32 (69.6)	79.49 (25.11)	0.0	71.88	84.38	100.00	100.0		
		Week 70	Sparsentan	35	22 (62.9)	79.55 (19.31)	31.3	75.00	81.25	93.75	100.0		
			Irbesartan	46	30 (65.2)	82.29 (23.56)	12.5	75.00	87.50	100.00	100.0		
		Week 94	Sparsentan	35	23 (65.7)	73.91 (27.67)	0.0	62.50	81.25	100.00	100.0		
			Irbesartan	46	30 (65.2)	80.63 (23.92)	12.5	68.75	87.50	100.00	100.0		
		Week 110	Sparsentan	35	21 (60.0)	76.19 (19.63)	37.5	56.25	75.00	93.75	100.0		
			Irbesartan	46	31 (67.4)	82.46 (19.86)	25.0	75.00	87.50	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	35	23 (65.7)	6.52 (24.75)	-37.5	-12.50	0.00	12.50	81.3	0.38 [-0.15, 0.91]
				Irbesartan	46	35 (76.1)	-1.07 (15.79)	-37.5	-12.50	0.00	6.25	31.3	
	Week 48		Sparsentan	35	25 (71.4)	8.75 (17.02)	-12.5	0.00	0.00	18.75	56.3	0.65 [0.11, 1.19]	
			Irbesartan	46	32 (69.6)	-3.13 (19.18)	-62.5	-6.25	0.00	6.25	37.5		
	Week 70		Sparsentan	35	22 (62.9)	4.83 (17.67)	-25.0	-6.25	0.00	12.50	43.8	0.38 [-0.17, 0.94]	
			Irbesartan	46	30 (65.2)	-1.46 (15.46)	-31.3	-12.50	0.00	6.25	31.3		
Week 94	Sparsentan	35	23 (65.7)	0.00 (24.06)	-62.5	-12.50	0.00	12.50	68.8	0.21 [-0.33, 0.76]			
	Irbesartan	46	30 (65.2)	-4.58 (19.07)	-75.0	-12.50	0.00	6.25	25.0				
Week 110	Sparsentan	35	21 (60.0)	1.19 (20.02)	-43.8	-6.25	0.00	6.25	56.3	0.20 [-0.36, 0.75]			
	Irbesartan	46	31 (67.4)	-2.22 (15.01)	-31.3	-12.50	0.00	6.25	31.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Europe	KDQOL: burden of kidney disease	Baseline	Sparsentan	98	89 (90.8)	76.26 (21.90)	6.3	68.75	81.25	93.75	100.0		
			Irbesartan	115	98 (85.2)	75.64 (24.14)	18.8	56.25	87.50	93.75	100.0		
		Week 24	Sparsentan	98	73 (74.5)	80.57 (20.56)	12.5	75.00	87.50	100.00	100.0		
			Irbesartan	115	62 (53.9)	77.72 (21.04)	25.0	62.50	81.25	93.75	100.0		
		Week 48	Sparsentan	98	76 (77.6)	79.44 (22.33)	0.0	75.00	81.25	100.00	100.0		
			Irbesartan	115	59 (51.3)	80.72 (18.87)	37.5	68.75	81.25	100.00	100.0		
		Week 70	Sparsentan	98	75 (76.5)	82.50 (18.95)	12.5	75.00	87.50	100.00	100.0		
			Irbesartan	115	69 (60.0)	79.44 (20.59)	25.0	68.75	87.50	100.00	100.0		
		Week 94	Sparsentan	98	68 (69.4)	79.69 (20.04)	6.3	68.75	81.25	93.75	100.0		
			Irbesartan	115	70 (60.9)	74.46 (23.88)	12.5	62.50	75.00	93.75	100.0		
		Week 110	Sparsentan	98	67 (68.4)	80.32 (19.86)	25.0	68.75	81.25	100.00	100.0		
			Irbesartan	115	61 (53.0)	75.72 (22.68)	25.0	62.50	75.00	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	98	73 (74.5)	4.45 (23.28)	-81.3	-6.25	0.00	12.50	75.0	0.19 [-0.15, 0.53]
				Irbesartan	115	62 (53.9)	0.40 (18.24)	-50.0	-12.50	0.00	6.25	56.3	
	Week 48		Sparsentan	98	76 (77.6)	3.29 (20.45)	-68.8	-3.13	6.25	12.50	50.0	-0.03 [-0.37, 0.31]	
			Irbesartan	115	59 (51.3)	3.92 (16.61)	-50.0	-6.25	0.00	18.75	50.0		
	Week 70		Sparsentan	98	75 (76.5)	5.42 (19.83)	-50.0	0.00	0.00	18.75	87.5	0.04 [-0.29, 0.36]	
			Irbesartan	115	69 (60.0)	4.71 (18.11)	-37.5	-6.25	0.00	12.50	56.3		
	Week 94		Sparsentan	98	68 (69.4)	3.86 (21.49)	-50.0	-6.25	0.00	12.50	75.0	0.18 [-0.15, 0.52]	
			Irbesartan	115	70 (60.9)	-0.09 (21.40)	-50.0	-12.50	0.00	12.50	50.0		
Week 110	Sparsentan	98	67 (68.4)	3.64 (22.83)	-56.3	-6.25	0.00	12.50	62.5	0.13 [-0.22, 0.47]			
	Irbesartan	115	61 (53.0)	0.92 (20.34)	-56.3	-12.50	0.00	12.50	56.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Asia Pacific	KDQOL: burden of kidney disease	Baseline	Sparsentan	69	67 (97.1)	60.73 (26.76)	0.0	43.75	62.50	87.50	100.0	
			Irbesartan	41	41 (100.0)	73.63 (24.02)	6.3	62.50	75.00	93.75	100.0	
		Week 24	Sparsentan	69	61 (88.4)	70.70 (24.27)	0.0	56.25	75.00	87.50	100.0	
			Irbesartan	41	34 (82.9)	75.74 (26.86)	0.0	68.75	81.25	100.00	100.0	
		Week 48	Sparsentan	69	61 (88.4)	74.69 (22.66)	6.3	62.50	75.00	93.75	100.0	
			Irbesartan	41	33 (80.5)	79.36 (22.07)	6.3	75.00	81.25	100.00	100.0	
		Week 70	Sparsentan	69	63 (91.3)	73.31 (24.98)	6.3	68.75	75.00	93.75	100.0	
			Irbesartan	41	31 (75.6)	77.22 (22.10)	12.5	75.00	75.00	93.75	100.0	
		Week 94	Sparsentan	69	60 (87.0)	73.23 (23.67)	0.0	62.50	75.00	90.63	100.0	
			Irbesartan	41	30 (73.2)	77.92 (15.81)	25.0	75.00	75.00	81.25	100.0	
		Week 110	Sparsentan	69	61 (88.4)	73.36 (23.52)	18.8	56.25	75.00	93.75	100.0	
			Irbesartan	41	29 (70.7)	70.26 (22.58)	12.5	62.50	75.00	81.25	100.0	
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	69	61 (88.4)	12.19 (19.78)	-25.0	0.00	6.25	31.25	62.5
	Irbesartan			41	34 (82.9)	2.94 (18.09)	-37.5	0.00	0.00	6.25	50.0	
	Week 48		Sparsentan	69	61 (88.4)	12.81 (20.87)	-50.0	0.00	12.50	25.00	62.5	0.42 [-0.01, 0.85]
			Irbesartan	41	33 (80.5)	4.36 (18.65)	-37.5	0.00	0.00	6.25	56.3	
	Week 70		Sparsentan	69	63 (91.3)	13.00 (24.60)	-37.5	0.00	12.50	31.25	87.5	0.46 [0.03, 0.90]
			Irbesartan	41	31 (75.6)	2.22 (19.93)	-43.8	-6.25	0.00	12.50	37.5	
	Week 94		Sparsentan	69	60 (87.0)	13.85 (26.84)	-87.5	0.00	18.75	31.25	62.5	0.41 [-0.03, 0.85]
		Irbesartan	41	30 (73.2)	3.54 (21.32)	-31.3	0.00	0.00	18.75	56.3		
Week 110	Sparsentan	69	61 (88.4)	13.73 (26.49)	-43.8	0.00	12.50	31.25	87.5	0.69 [0.23, 1.14]		
	Irbesartan	41	29 (70.7)	-4.31 (25.78)	-62.5	-25.00	0.00	6.25	50.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline BMI														
< 27 kg/m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	83	75 (90.4)	64.08 (25.83)	6.3	43.75	68.75	87.50	100.0			
			Irbesartan	94	86 (91.5)	72.75 (22.36)	18.8	56.25	75.00	93.75	100.0			
		Week 24	Sparsentan	83	66 (79.5)	72.82 (23.88)	0.0	62.50	75.00	87.50	100.0			
			Irbesartan	94	66 (70.2)	76.14 (23.69)	6.3	68.75	81.25	93.75	100.0			
		Week 48	Sparsentan	83	66 (79.5)	72.82 (24.67)	6.3	50.00	75.00	93.75	100.0			
			Irbesartan	94	60 (63.8)	74.17 (23.03)	0.0	65.63	81.25	87.50	100.0			
		Week 70	Sparsentan	83	64 (77.1)	74.32 (24.55)	6.3	59.38	81.25	93.75	100.0			
			Irbesartan	94	59 (62.8)	73.52 (22.78)	12.5	62.50	75.00	93.75	100.0			
		Week 94	Sparsentan	83	63 (75.9)	74.01 (21.69)	0.0	62.50	75.00	87.50	100.0			
			Irbesartan	94	62 (66.0)	76.21 (19.10)	12.5	68.75	75.00	93.75	100.0			
		Week 110	Sparsentan	83	63 (75.9)	74.70 (22.63)	18.8	56.25	81.25	93.75	100.0			
			Irbesartan	94	57 (60.6)	75.00 (21.49)	25.0	68.75	75.00	100.00	100.0			
			KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	83	66 (79.5)	9.00 (22.66)	-43.8	0.00	6.25	18.75	81.3	0.25 [-0.09, 0.59]
					Irbesartan	94	66 (70.2)	3.79 (18.83)	-37.5	-12.50	0.00	12.50	56.3	
		Week 48		Sparsentan	83	66 (79.5)	9.94 (21.20)	-50.0	0.00	6.25	25.00	62.5	0.38 [0.03, 0.73]	
				Irbesartan	94	60 (63.8)	1.98 (20.54)	-62.5	-6.25	0.00	15.63	56.3		
		Week 70		Sparsentan	83	64 (77.1)	9.47 (23.78)	-37.5	0.00	6.25	25.00	87.5	0.38 [0.02, 0.74]	
				Irbesartan	94	59 (62.8)	1.27 (19.10)	-37.5	-6.25	0.00	12.50	43.8		
Week 94	Sparsentan	83		63 (75.9)	10.52 (23.73)	-37.5	0.00	6.25	25.00	68.8	0.40 [0.05, 0.76]			
	Irbesartan	94		62 (66.0)	1.81 (19.19)	-37.5	-6.25	0.00	12.50	56.3				
Week 110	Sparsentan	83		63 (75.9)	9.52 (26.58)	-43.8	-6.25	6.25	25.00	87.5	0.39 [0.03, 0.75]			
	Irbesartan	94		57 (60.6)	0.33 (19.35)	-56.3	-12.50	0.00	6.25	50.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 27 kg/m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	119	112 (94.1)	74.05 (24.14)	0.0	62.50	78.13	93.75	100.0		
			Irbesartan	107	95 (88.8)	81.32 (21.85)	6.3	68.75	87.50	100.00	100.0		
		Week 24	Sparsentan	119	91 (76.5)	79.60 (20.66)	12.5	68.75	87.50	100.00	100.0		
			Irbesartan	107	65 (60.7)	81.25 (21.68)	0.0	68.75	87.50	100.00	100.0		
		Week 48	Sparsentan	119	96 (80.7)	80.60 (21.02)	0.0	75.00	84.38	100.00	100.0		
			Irbesartan	107	64 (59.8)	85.55 (18.06)	31.3	75.00	93.75	100.00	100.0		
		Week 70	Sparsentan	119	96 (80.7)	81.25 (19.51)	12.5	75.00	81.25	100.00	100.0		
			Irbesartan	107	71 (66.4)	84.60 (19.24)	12.5	75.00	87.50	100.00	100.0		
		Week 94	Sparsentan	119	88 (73.9)	77.84 (23.63)	0.0	71.88	81.25	100.00	100.0		
			Irbesartan	107	67 (62.6)	78.08 (23.85)	25.0	68.75	81.25	100.00	100.0		
		Week 110	Sparsentan	119	86 (72.3)	78.49 (20.64)	25.0	62.50	81.25	100.00	100.0		
			Irbesartan	107	63 (58.9)	77.98 (22.16)	12.5	62.50	81.25	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	119	91 (76.5)	6.87 (22.23)	-81.3	-6.25	6.25	18.75	75.0	0.47 [0.15, 0.80]
			Irbesartan	107	65 (60.7)	-2.50 (15.58)	-50.0	-6.25	0.00	6.25	43.8		
	Week 48		Sparsentan	119	96 (80.7)	6.18 (19.94)	-68.8	-6.25	3.13	18.75	56.3	0.21 [-0.11, 0.52]	
			Irbesartan	107	64 (59.8)	2.44 (15.37)	-37.5	-6.25	0.00	6.25	50.0		
	Week 70		Sparsentan	119	96 (80.7)	7.55 (20.44)	-50.0	0.00	6.25	18.75	87.5	0.19 [-0.11, 0.50]	
			Irbesartan	107	71 (66.4)	3.87 (17.14)	-43.8	0.00	0.00	12.50	56.3		
	Week 94		Sparsentan	119	88 (73.9)	4.90 (25.08)	-87.5	-6.25	6.25	18.75	75.0	0.29 [-0.03, 0.61]	
			Irbesartan	107	67 (62.6)	-2.15 (22.47)	-75.0	-18.75	0.00	12.50	50.0		
Week 110	Sparsentan	119	86 (72.3)	5.89 (22.86)	-56.3	-6.25	0.00	18.75	75.0	0.38 [0.05, 0.71]			
	Irbesartan	107	63 (58.9)	-2.58 (21.81)	-62.5	-18.75	0.00	6.25	56.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Randomization strata													
eGFR Low and UP High	KDQOL: burden of kidney disease	Baseline	Sparsentan	71	63 (88.7)	70.73 (24.25)	6.3	50.00	75.00	87.50	100.0		
			Irbesartan	74	63 (85.1)	72.82 (24.13)	18.8	56.25	75.00	93.75	100.0		
		Week 24	Sparsentan	71	54 (76.1)	76.97 (21.92)	12.5	62.50	81.25	93.75	100.0		
			Irbesartan	74	40 (54.1)	73.28 (24.72)	6.3	62.50	75.00	93.75	100.0		
		Week 48	Sparsentan	71	51 (71.8)	75.74 (21.71)	18.8	62.50	75.00	93.75	100.0		
			Irbesartan	74	37 (50.0)	77.20 (25.31)	0.0	75.00	81.25	93.75	100.0		
		Week 70	Sparsentan	71	57 (80.3)	75.33 (23.28)	12.5	62.50	75.00	93.75	100.0		
			Irbesartan	74	37 (50.0)	74.32 (21.84)	12.5	68.75	75.00	87.50	100.0		
		Week 94	Sparsentan	71	51 (71.8)	71.32 (24.94)	0.0	56.25	75.00	93.75	100.0		
			Irbesartan	74	37 (50.0)	69.26 (24.27)	12.5	62.50	75.00	81.25	100.0		
		Week 110	Sparsentan	71	53 (74.6)	75.24 (22.86)	25.0	50.00	81.25	93.75	100.0		
			Irbesartan	74	36 (48.6)	73.44 (21.46)	25.0	59.38	75.00	90.63	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	71	54 (76.1)	7.87 (22.75)	-50.0	-6.25	6.25	25.00	62.5	0.53 [0.11, 0.94]
			Irbesartan	74	40 (54.1)	-2.50 (14.42)	-37.5	-12.50	0.00	3.13	37.5		
	Week 48		Sparsentan	71	51 (71.8)	6.74 (19.24)	-50.0	-6.25	0.00	25.00	43.8	0.32 [-0.11, 0.74]	
			Irbesartan	74	37 (50.0)	0.84 (17.38)	-62.5	-6.25	0.00	6.25	31.3		
	Week 70		Sparsentan	71	57 (80.3)	5.37 (20.40)	-50.0	0.00	6.25	18.75	56.3	0.25 [-0.17, 0.66]	
			Irbesartan	74	37 (50.0)	0.51 (18.18)	-37.5	-6.25	0.00	6.25	56.3		
Week 94	Sparsentan	71	51 (71.8)	1.35 (26.41)	-87.5	-6.25	0.00	18.75	56.3	0.17 [-0.25, 0.60]			
	Irbesartan	74	37 (50.0)	-2.70 (18.90)	-50.0	-12.50	0.00	6.25	31.3				
Week 110	Sparsentan	71	53 (74.6)	4.72 (22.83)	-56.3	-6.25	6.25	18.75	50.0	0.23 [-0.20, 0.65]			
	Irbesartan	74	36 (48.6)	-0.35 (21.07)	-43.8	-12.50	0.00	9.38	56.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR Low and UP Low	KDQOL: burden of kidney disease	Baseline	Sparsentan	55	52 (94.5)	67.55 (25.64)	0.0	50.00	71.88	87.50	100.0		
			Irbesartan	55	50 (90.9)	78.88 (22.26)	25.0	62.50	84.38	100.00	100.0		
		Week 24	Sparsentan	55	43 (78.2)	71.95 (25.96)	0.0	56.25	75.00	93.75	100.0		
			Irbesartan	55	37 (67.3)	82.77 (17.46)	37.5	68.75	87.50	100.00	100.0		
		Week 48	Sparsentan	55	43 (78.2)	79.80 (21.08)	6.3	75.00	81.25	100.00	100.0		
			Irbesartan	55	35 (63.6)	80.54 (19.10)	31.3	75.00	81.25	100.00	100.0		
		Week 70	Sparsentan	55	42 (76.4)	76.79 (23.03)	6.3	75.00	81.25	93.75	100.0		
			Irbesartan	55	39 (70.9)	79.65 (20.57)	12.5	75.00	81.25	100.00	100.0		
		Week 94	Sparsentan	55	41 (74.5)	79.88 (19.84)	18.8	68.75	81.25	100.00	100.0		
			Irbesartan	55	39 (70.9)	76.92 (20.34)	25.0	68.75	75.00	93.75	100.0		
		Week 110	Sparsentan	55	37 (67.3)	75.17 (22.36)	18.8	62.50	75.00	93.75	100.0		
			Irbesartan	55	34 (61.8)	76.84 (20.67)	25.0	68.75	78.13	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	55	43 (78.2)	5.96 (23.89)	-81.3	0.00	0.00	18.75	62.5	0.11 [-0.33, 0.55]
				Irbesartan	55	37 (67.3)	3.72 (17.33)	-31.3	-6.25	0.00	12.50	50.0	
	Week 48		Sparsentan	55	43 (78.2)	10.61 (18.58)	-25.0	0.00	6.25	25.00	62.5	0.38 [-0.07, 0.83]	
			Irbesartan	55	35 (63.6)	3.57 (18.95)	-50.0	-6.25	0.00	18.75	56.3		
	Week 70		Sparsentan	55	42 (76.4)	8.78 (22.30)	-43.8	-6.25	6.25	25.00	62.5	0.27 [-0.17, 0.71]	
			Irbesartan	55	39 (70.9)	3.21 (18.80)	-43.8	-6.25	0.00	12.50	50.0		
	Week 94		Sparsentan	55	41 (74.5)	13.87 (23.78)	-50.0	0.00	12.50	25.00	68.8	0.59 [0.15, 1.04]	
			Irbesartan	55	39 (70.9)	-0.48 (24.52)	-37.5	-12.50	0.00	12.50	56.3		
Week 110	Sparsentan	55	37 (67.3)	7.77 (28.66)	-50.0	-12.50	6.25	31.25	62.5	0.19 [-0.28, 0.65]			
	Irbesartan	55	34 (61.8)	3.13 (20.43)	-31.3	-12.50	0.00	12.50	50.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR High and UP High	KDQOL: burden of kidney disease	Baseline	Sparsentan	37	35 (94.6)	67.68 (27.47)	6.3	50.00	75.00	87.50	100.0		
			Irbesartan	36	33 (91.7)	76.89 (21.39)	31.3	62.50	81.25	93.75	100.0		
		Week 24	Sparsentan	37	26 (70.3)	75.48 (18.79)	25.0	68.75	78.13	87.50	100.0		
			Irbesartan	36	26 (72.2)	78.85 (23.05)	18.8	68.75	84.38	100.00	100.0		
		Week 48	Sparsentan	37	33 (89.2)	72.92 (24.74)	6.3	62.50	75.00	93.75	100.0		
			Irbesartan	36	24 (66.7)	79.95 (21.17)	31.3	71.88	81.25	100.00	100.0		
		Week 70	Sparsentan	37	31 (83.8)	79.64 (22.88)	25.0	75.00	87.50	100.00	100.0		
			Irbesartan	36	25 (69.4)	76.75 (27.33)	12.5	62.50	87.50	100.00	100.0		
		Week 94	Sparsentan	37	27 (73.0)	73.61 (21.74)	25.0	68.75	75.00	87.50	100.0		
			Irbesartan	36	24 (66.7)	80.47 (21.44)	25.0	71.88	87.50	100.00	100.0		
		Week 110	Sparsentan	37	26 (70.3)	76.44 (21.74)	25.0	68.75	75.00	100.00	100.0		
			Irbesartan	36	22 (61.1)	77.27 (26.55)	25.0	62.50	90.63	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	37	26 (70.3)	9.13 (23.00)	-31.3	-6.25	6.25	18.75	75.0	0.43 [-0.12, 0.98]
				Irbesartan	36	26 (72.2)	-0.24 (21.03)	-50.0	-6.25	0.00	6.25	50.0	
	Week 48		Sparsentan	37	33 (89.2)	6.25 (24.05)	-62.5	0.00	6.25	18.75	56.3	0.02 [-0.50, 0.55]	
			Irbesartan	36	24 (66.7)	5.73 (19.93)	-37.5	-6.25	0.00	18.75	50.0		
	Week 70		Sparsentan	37	31 (83.8)	12.70 (25.39)	-25.0	0.00	6.25	18.75	87.5	0.46 [-0.08, 0.99]	
			Irbesartan	36	25 (69.4)	2.25 (19.08)	-37.5	-6.25	0.00	6.25	43.8		
	Week 94		Sparsentan	37	27 (73.0)	7.64 (23.34)	-31.3	0.00	0.00	18.75	75.0	0.14 [-0.41, 0.69]	
			Irbesartan	36	24 (66.7)	4.95 (13.91)	-25.0	-3.13	6.25	15.63	25.0		
Week 110	Sparsentan	37	26 (70.3)	11.30 (23.52)	-25.0	0.00	6.25	18.75	87.5	0.61 [0.03, 1.19]			
	Irbesartan	36	22 (61.1)	-1.14 (15.98)	-31.3	-12.50	0.00	6.25	37.5				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR High and UP Low	KDQOL: burden of kidney disease	Baseline	Sparsentan	39	37 (94.9)	74.66 (24.47)	0.0	62.50	81.25	93.75	100.0		
			Irbesartan	37	36 (97.3)	81.42 (22.08)	6.3	75.00	90.63	96.88	100.0		
		Week 24	Sparsentan	39	34 (87.2)	83.46 (19.15)	25.0	75.00	87.50	100.00	100.0		
			Irbesartan	37	28 (75.7)	80.80 (25.40)	0.0	75.00	90.63	100.00	100.0		
		Week 48	Sparsentan	39	35 (89.7)	81.25 (24.58)	0.0	75.00	87.50	100.00	100.0		
			Irbesartan	37	28 (75.7)	83.26 (18.72)	25.0	75.00	87.50	100.00	100.0		
		Week 70	Sparsentan	39	30 (76.9)	85.63 (14.32)	50.0	81.25	87.50	100.00	100.0		
			Irbesartan	37	29 (78.4)	88.58 (13.68)	56.3	75.00	93.75	100.00	100.0		
		Week 94	Sparsentan	39	32 (82.1)	81.64 (22.89)	0.0	75.00	87.50	100.00	100.0		
			Irbesartan	37	30 (81.1)	82.50 (21.30)	25.0	68.75	93.75	100.00	100.0		
		Week 110	Sparsentan	39	33 (84.6)	81.82 (18.04)	37.5	75.00	87.50	93.75	100.0		
			Irbesartan	37	29 (78.4)	77.80 (22.20)	12.5	68.75	75.00	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	39	34 (87.2)	8.82 (19.95)	-18.8	0.00	6.25	12.50	81.3	0.35 [-0.15, 0.86]
				Irbesartan	37	28 (75.7)	2.01 (18.32)	-31.3	-12.50	0.00	9.38	56.3	
	Week 48		Sparsentan	39	35 (89.7)	6.96 (21.37)	-68.8	0.00	6.25	18.75	50.0	0.40 [-0.10, 0.90]	
			Irbesartan	37	28 (75.7)	-0.67 (15.99)	-43.8	-6.25	0.00	6.25	31.3		
	Week 70		Sparsentan	39	30 (76.9)	8.75 (19.74)	-12.5	-6.25	0.00	18.75	50.0	0.20 [-0.31, 0.71]	
			Irbesartan	37	29 (78.4)	5.17 (16.37)	-25.0	0.00	0.00	18.75	31.3		
	Week 94		Sparsentan	39	32 (82.1)	7.81 (22.39)	-43.8	-6.25	3.13	21.88	62.5	0.40 [-0.10, 0.90]	
			Irbesartan	37	30 (81.1)	-1.25 (22.94)	-75.0	-12.50	0.00	12.50	37.5		
Week 110	Sparsentan	39	33 (84.6)	8.33 (23.32)	-18.8	-6.25	0.00	18.75	75.0	0.67 [0.16, 1.18]			
	Irbesartan	37	29 (78.4)	-7.11 (22.70)	-62.5	-25.00	0.00	6.25	31.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 1														
< 60 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	127	116 (91.3)	70.10 (25.09)	0.0	50.00	75.00	87.50	100.0			
			Irbesartan	129	114 (88.4)	77.41 (22.84)	18.8	62.50	81.25	100.00	100.0			
		Week 24	Sparsentan	127	98 (77.2)	74.74 (23.69)	0.0	62.50	81.25	93.75	100.0			
			Irbesartan	129	80 (62.0)	77.89 (21.97)	6.3	68.75	81.25	100.00	100.0			
		Week 48	Sparsentan	127	95 (74.8)	77.76 (21.43)	6.3	68.75	81.25	100.00	100.0			
			Irbesartan	129	74 (57.4)	79.31 (22.80)	0.0	75.00	81.25	100.00	100.0			
		Week 70	Sparsentan	127	100 (78.7)	76.13 (23.05)	6.3	68.75	81.25	93.75	100.0			
			Irbesartan	129	78 (60.5)	78.37 (21.53)	12.5	68.75	78.13	100.00	100.0			
		Week 94	Sparsentan	127	92 (72.4)	75.07 (23.27)	0.0	65.63	78.13	93.75	100.0			
			Irbesartan	129	77 (59.7)	74.19 (22.81)	12.5	62.50	75.00	93.75	100.0			
		Week 110	Sparsentan	127	92 (72.4)	76.36 (22.41)	18.8	62.50	81.25	93.75	100.0			
			Irbesartan	129	72 (55.8)	76.56 (20.39)	25.0	65.63	81.25	96.88	100.0			
			KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	127	98 (77.2)	6.31 (23.12)	-81.3	-6.25	0.00	18.75	62.5	0.34 [0.04, 0.63]
					Irbesartan	129	80 (62.0)	-0.47 (15.65)	-37.5	-12.50	0.00	6.25	50.0	
				Week 48	Sparsentan	127	95 (74.8)	7.70 (18.56)	-50.0	0.00	6.25	18.75	62.5	0.35 [0.04, 0.66]
					Irbesartan	129	74 (57.4)	1.18 (18.61)	-62.5	-6.25	0.00	12.50	56.3	
				Week 70	Sparsentan	127	100 (78.7)	6.06 (20.97)	-50.0	-6.25	6.25	18.75	62.5	0.20 [-0.10, 0.50]
					Irbesartan	129	78 (60.5)	2.08 (18.26)	-43.8	-6.25	0.00	12.50	56.3	
		Week 94	Sparsentan	127	92 (72.4)	6.18 (26.31)	-87.5	-6.25	6.25	25.00	68.8	0.34 [0.04, 0.65]		
			Irbesartan	129	77 (59.7)	-2.27 (22.92)	-75.0	-12.50	0.00	6.25	56.3			
		Week 110	Sparsentan	127	92 (72.4)	6.05 (24.94)	-56.3	-6.25	6.25	21.88	62.5	0.22 [-0.09, 0.53]		
			Irbesartan	129	72 (55.8)	0.95 (19.98)	-43.8	-12.50	0.00	9.38	56.3			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	49	47 (95.9)	72.61 (23.91)	0.0	62.50	75.00	87.50	100.0	
			Irbesartan	48	44 (91.7)	78.27 (21.33)	18.8	65.63	87.50	93.75	100.0	
		Week 24	Sparsentan	49	38 (77.6)	79.28 (19.45)	25.0	68.75	81.25	93.75	100.0	
			Irbesartan	48	31 (64.6)	85.28 (16.89)	37.5	75.00	93.75	100.00	100.0	
		Week 48	Sparsentan	49	44 (89.8)	78.13 (23.67)	6.3	75.00	81.25	96.88	100.0	
			Irbesartan	48	32 (66.7)	79.88 (20.00)	25.0	68.75	81.25	100.00	100.0	
		Week 70	Sparsentan	49	38 (77.6)	81.41 (19.63)	25.0	75.00	84.38	100.00	100.0	
			Irbesartan	48	32 (66.7)	80.47 (22.21)	25.0	71.88	87.50	100.00	100.0	
		Week 94	Sparsentan	49	39 (79.6)	77.56 (23.55)	0.0	68.75	81.25	100.00	100.0	
			Irbesartan	48	34 (70.8)	77.02 (20.34)	25.0	68.75	75.00	93.75	100.0	
		Week 110	Sparsentan	49	36 (73.5)	74.83 (20.02)	37.5	56.25	78.13	93.75	100.0	
			Irbesartan	48	32 (66.7)	75.59 (22.47)	25.0	65.63	75.00	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	49	38 (77.6)	7.89 (21.34)	-31.3	-6.25	6.25	18.75	81.3	0.19 [-0.29, 0.66]
			Irbesartan	48	31 (64.6)	4.23 (17.48)	-25.0	-6.25	0.00	12.50	50.0	
		Week 48	Sparsentan	49	44 (89.8)	6.68 (20.77)	-62.5	0.00	3.13	21.88	50.0	0.21 [-0.25, 0.66]
			Irbesartan	48	32 (66.7)	2.54 (18.64)	-43.8	-6.25	0.00	9.38	50.0	
		Week 70	Sparsentan	49	38 (77.6)	7.73 (17.88)	-25.0	0.00	0.00	18.75	50.0	0.31 [-0.16, 0.78]
			Irbesartan	48	32 (66.7)	2.15 (18.28)	-25.0	-9.38	0.00	9.38	43.8	
		Week 94	Sparsentan	49	39 (79.6)	6.73 (19.15)	-31.3	0.00	6.25	18.75	62.5	0.47 [0.01, 0.94]
			Irbesartan	48	34 (70.8)	-1.84 (16.68)	-37.5	-12.50	0.00	6.25	25.0	
Week 110	Sparsentan	49	36 (73.5)	6.25 (21.02)	-25.0	-9.38	0.00	15.63	75.0	0.56 [0.07, 1.04]		
	Irbesartan	48	32 (66.7)	-5.47 (20.99)	-56.3	-25.00	0.00	6.25	37.5			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	26	24 (92.3)	64.84 (28.66)	6.3	50.00	75.00	87.50	100.0		
			Irbesartan	25	24 (96.0)	72.14 (25.47)	6.3	53.13	75.00	93.75	100.0		
		Week 24	Sparsentan	26	21 (80.8)	81.55 (19.61)	31.3	75.00	87.50	100.00	100.0		
			Irbesartan	25	20 (80.0)	71.56 (31.05)	0.0	50.00	81.25	100.00	100.0		
		Week 48	Sparsentan	26	23 (88.5)	74.73 (27.34)	0.0	56.25	87.50	100.00	100.0		
			Irbesartan	25	18 (72.0)	83.33 (17.55)	43.8	75.00	84.38	100.00	100.0		
		Week 70	Sparsentan	26	22 (84.6)	84.09 (18.97)	25.0	81.25	87.50	100.00	100.0		
			Irbesartan	25	20 (80.0)	82.81 (21.35)	12.5	71.88	87.50	100.00	100.0		
		Week 94	Sparsentan	26	20 (76.9)	79.06 (20.00)	25.0	75.00	81.25	100.00	100.0		
			Irbesartan	25	19 (76.0)	86.18 (21.91)	25.0	75.00	100.00	100.00	100.0		
		Week 110	Sparsentan	26	21 (80.8)	82.74 (19.85)	25.0	75.00	87.50	100.00	100.0		
			Irbesartan	25	17 (68.0)	75.37 (29.60)	12.5	56.25	93.75	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	26	21 (80.8)	14.29 (20.27)	-6.3	0.00	6.25	18.75	75.0	0.66 [0.03, 1.29]
			Irbesartan	25	20 (80.0)	-0.31 (23.86)	-50.0	-9.38	0.00	9.38	56.3		
	Week 48		Sparsentan	26	23 (88.5)	9.78 (27.36)	-68.8	0.00	12.50	25.00	56.3	0.17 [-0.45, 0.79]	
			Irbesartan	25	18 (72.0)	5.90 (14.13)	-18.8	-6.25	6.25	12.50	31.3		
	Week 70		Sparsentan	26	22 (84.6)	19.60 (28.24)	-12.5	0.00	12.50	31.25	87.5	0.58 [-0.04, 1.20]	
			Irbesartan	25	20 (80.0)	5.94 (17.26)	-37.5	0.00	6.25	18.75	31.3		
	Week 94		Sparsentan	26	20 (76.9)	13.13 (26.20)	-18.8	-3.13	3.13	28.13	75.0	0.12 [-0.51, 0.75]	
			Irbesartan	25	19 (76.0)	10.53 (16.01)	-25.0	0.00	6.25	25.00	37.5		
Week 110	Sparsentan	26	21 (80.8)	15.48 (27.43)	-18.8	0.00	6.25	18.75	87.5	0.69 [0.03, 1.34]			
	Irbesartan	25	17 (68.0)	-1.84 (22.18)	-62.5	-12.50	6.25	12.50	31.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 2														
< 45 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	82	73 (89.0)	67.29 (25.03)	6.3	50.00	75.00	87.50	100.0			
		Week 24	Irbesartan	80	69 (86.3)	73.82 (24.05)	18.8	62.50	81.25	93.75	100.0			
			Sparsentan	82	64 (78.0)	75.10 (24.00)	12.5	62.50	81.25	100.00	100.0			
		Week 48	Irbesartan	80	48 (60.0)	76.69 (24.39)	6.3	62.50	81.25	100.00	100.0			
			Sparsentan	82	61 (74.4)	76.13 (23.32)	6.3	62.50	81.25	93.75	100.0			
		Week 70	Irbesartan	80	43 (53.8)	78.63 (24.44)	0.0	75.00	81.25	100.00	100.0			
			Sparsentan	82	65 (79.3)	75.87 (24.13)	6.3	68.75	81.25	93.75	100.0			
		Week 94	Irbesartan	80	45 (56.3)	76.25 (21.76)	12.5	68.75	75.00	93.75	100.0			
			Sparsentan	82	60 (73.2)	75.31 (22.85)	0.0	68.75	81.25	93.75	100.0			
		Week 110	Irbesartan	80	47 (58.8)	70.88 (23.47)	12.5	62.50	75.00	93.75	100.0			
			Sparsentan	82	56 (68.3)	76.45 (23.57)	18.8	59.38	81.25	96.88	100.0			
			Irbesartan	80	44 (55.0)	75.00 (19.90)	25.0	62.50	78.13	90.63	100.0			
			KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	82	64 (78.0)	8.01 (22.73)	-81.3	0.00	6.25	18.75	62.5	0.37 [-0.01, 0.74]
				Week 48	Irbesartan	80	48 (60.0)	0.78 (14.79)	-37.5	-6.25	0.00	12.50	37.5	
					Sparsentan	82	61 (74.4)	8.81 (20.71)	-50.0	0.00	6.25	25.00	62.5	0.29 [-0.10, 0.68]
				Week 70	Irbesartan	80	43 (53.8)	3.20 (17.38)	-62.5	-6.25	0.00	18.75	43.8	
					Sparsentan	82	65 (79.3)	7.98 (19.78)	-43.8	0.00	6.25	18.75	56.3	0.23 [-0.15, 0.61]
				Week 94	Irbesartan	80	45 (56.3)	3.47 (20.01)	-37.5	-6.25	0.00	12.50	56.3	
	Sparsentan	82			60 (73.2)	7.19 (25.70)	-87.5	-6.25	6.25	25.00	68.8	0.39 [0.00, 0.77]		
	Week 110	Irbesartan		80	47 (58.8)	-2.39 (23.59)	-50.0	-18.75	0.00	12.50	50.0			
		Sparsentan		82	56 (68.3)	7.81 (24.23)	-56.3	0.00	6.25	25.00	62.5	0.26 [-0.13, 0.66]		
	Irbesartan	80		44 (55.0)	1.85 (20.69)	-43.8	-12.50	0.00	9.38	56.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
45 to < 60 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	45	43 (95.6)	74.85 (24.76)	0.0	62.50	81.25	93.75	100.0		
			Irbesartan	49	45 (91.8)	82.92 (19.87)	25.0	68.75	93.75	100.00	100.0		
		Week 24	Sparsentan	45	34 (75.6)	74.08 (23.44)	0.0	62.50	75.00	93.75	100.0		
			Irbesartan	49	32 (65.3)	79.69 (17.96)	31.3	68.75	81.25	96.88	100.0		
		Week 48	Sparsentan	45	34 (75.6)	80.70 (17.50)	43.8	68.75	81.25	100.00	100.0		
			Irbesartan	49	31 (63.3)	80.24 (20.67)	31.3	75.00	81.25	100.00	100.0		
		Week 70	Sparsentan	45	35 (77.8)	76.61 (21.24)	12.5	62.50	75.00	93.75	100.0		
			Irbesartan	49	33 (67.3)	81.25 (21.19)	12.5	68.75	87.50	100.00	100.0		
		Week 94	Sparsentan	45	32 (71.1)	74.61 (24.38)	18.8	59.38	75.00	96.88	100.0		
			Irbesartan	49	30 (61.2)	79.38 (21.09)	25.0	75.00	81.25	100.00	100.0		
		Week 110	Sparsentan	45	36 (80.0)	76.22 (20.80)	25.0	65.63	78.13	93.75	100.0		
			Irbesartan	49	28 (57.1)	79.02 (21.26)	25.0	71.88	81.25	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	45	34 (75.6)	3.13 (23.85)	-50.0	-12.50	0.00	18.75	50.0	0.26 [-0.22, 0.75]
				Irbesartan	49	32 (65.3)	-2.34 (16.93)	-37.5	-12.50	0.00	0.00	50.0	
	Week 48		Sparsentan	45	34 (75.6)	5.70 (13.96)	-25.0	0.00	3.13	12.50	37.5	0.43 [-0.07, 0.92]	
			Irbesartan	49	31 (63.3)	-1.61 (20.15)	-50.0	-6.25	0.00	6.25	56.3		
	Week 70		Sparsentan	45	35 (77.8)	2.50 (22.90)	-50.0	-12.50	0.00	18.75	62.5	0.12 [-0.36, 0.59]	
			Irbesartan	49	33 (67.3)	0.19 (15.66)	-43.8	-6.25	0.00	6.25	37.5		
	Week 94		Sparsentan	45	32 (71.1)	4.30 (27.74)	-62.5	-15.63	6.25	25.00	56.3	0.25 [-0.25, 0.75]	
			Irbesartan	49	30 (61.2)	-2.08 (22.22)	-75.0	-12.50	0.00	6.25	56.3		
Week 110	Sparsentan	45	36 (80.0)	3.30 (26.11)	-50.0	-12.50	0.00	15.63	62.5	0.16 [-0.33, 0.66]			
	Irbesartan	49	28 (57.1)	-0.45 (19.09)	-31.3	-9.38	0.00	9.38	50.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	49	47 (95.9)	72.61 (23.91)	0.0	62.50	75.00	87.50	100.0	
			Irbesartan	48	44 (91.7)	78.27 (21.33)	18.8	65.63	87.50	93.75	100.0	
		Week 24	Sparsentan	49	38 (77.6)	79.28 (19.45)	25.0	68.75	81.25	93.75	100.0	
			Irbesartan	48	31 (64.6)	85.28 (16.89)	37.5	75.00	93.75	100.00	100.0	
		Week 48	Sparsentan	49	44 (89.8)	78.13 (23.67)	6.3	75.00	81.25	96.88	100.0	
			Irbesartan	48	32 (66.7)	79.88 (20.00)	25.0	68.75	81.25	100.00	100.0	
		Week 70	Sparsentan	49	38 (77.6)	81.41 (19.63)	25.0	75.00	84.38	100.00	100.0	
			Irbesartan	48	32 (66.7)	80.47 (22.21)	25.0	71.88	87.50	100.00	100.0	
		Week 94	Sparsentan	49	39 (79.6)	77.56 (23.55)	0.0	68.75	81.25	100.00	100.0	
			Irbesartan	48	34 (70.8)	77.02 (20.34)	25.0	68.75	75.00	93.75	100.0	
		Week 110	Sparsentan	49	36 (73.5)	74.83 (20.02)	37.5	56.25	78.13	93.75	100.0	
			Irbesartan	48	32 (66.7)	75.59 (22.47)	25.0	65.63	75.00	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	49	38 (77.6)	7.89 (21.34)	-31.3	-6.25	6.25	18.75	81.3	0.19 [-0.29, 0.66]
			Irbesartan	48	31 (64.6)	4.23 (17.48)	-25.0	-6.25	0.00	12.50	50.0	
		Week 48	Sparsentan	49	44 (89.8)	6.68 (20.77)	-62.5	0.00	3.13	21.88	50.0	0.21 [-0.25, 0.66]
			Irbesartan	48	32 (66.7)	2.54 (18.64)	-43.8	-6.25	0.00	9.38	50.0	
		Week 70	Sparsentan	49	38 (77.6)	7.73 (17.88)	-25.0	0.00	0.00	18.75	50.0	0.31 [-0.16, 0.78]
			Irbesartan	48	32 (66.7)	2.15 (18.28)	-25.0	-9.38	0.00	9.38	43.8	
		Week 94	Sparsentan	49	39 (79.6)	6.73 (19.15)	-31.3	0.00	6.25	18.75	62.5	0.47 [0.01, 0.94]
			Irbesartan	48	34 (70.8)	-1.84 (16.68)	-37.5	-12.50	0.00	6.25	25.0	
Week 110	Sparsentan	49	36 (73.5)	6.25 (21.02)	-25.0	-9.38	0.00	15.63	75.0	0.56 [0.07, 1.04]		
	Irbesartan	48	32 (66.7)	-5.47 (20.99)	-56.3	-25.00	0.00	6.25	37.5			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL: burden of kidney disease	Baseline	Sparsentan	26	24 (92.3)	64.84 (28.66)	6.3	50.00	75.00	87.50	100.0		
			Irbesartan	25	24 (96.0)	72.14 (25.47)	6.3	53.13	75.00	93.75	100.0		
		Week 24	Sparsentan	26	21 (80.8)	81.55 (19.61)	31.3	75.00	87.50	100.00	100.0		
			Irbesartan	25	20 (80.0)	71.56 (31.05)	0.0	50.00	81.25	100.00	100.0		
		Week 48	Sparsentan	26	23 (88.5)	74.73 (27.34)	0.0	56.25	87.50	100.00	100.0		
			Irbesartan	25	18 (72.0)	83.33 (17.55)	43.8	75.00	84.38	100.00	100.0		
		Week 70	Sparsentan	26	22 (84.6)	84.09 (18.97)	25.0	81.25	87.50	100.00	100.0		
			Irbesartan	25	20 (80.0)	82.81 (21.35)	12.5	71.88	87.50	100.00	100.0		
		Week 94	Sparsentan	26	20 (76.9)	79.06 (20.00)	25.0	75.00	81.25	100.00	100.0		
			Irbesartan	25	19 (76.0)	86.18 (21.91)	25.0	75.00	100.00	100.00	100.0		
		Week 110	Sparsentan	26	21 (80.8)	82.74 (19.85)	25.0	75.00	87.50	100.00	100.0		
			Irbesartan	25	17 (68.0)	75.37 (29.60)	12.5	56.25	93.75	100.00	100.0		
			KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	26	21 (80.8)	14.29 (20.27)	-6.3	0.00	6.25	18.75	75.0
		Irbesartan		25	20 (80.0)	-0.31 (23.86)	-50.0	-9.38	0.00	9.38	56.3		
	Week 48	Sparsentan		26	23 (88.5)	9.78 (27.36)	-68.8	0.00	12.50	25.00	56.3	0.17 [-0.45, 0.79]	
		Irbesartan		25	18 (72.0)	5.90 (14.13)	-18.8	-6.25	6.25	12.50	31.3		
	Week 70	Sparsentan		26	22 (84.6)	19.60 (28.24)	-12.5	0.00	12.50	31.25	87.5	0.58 [-0.04, 1.20]	
		Irbesartan		25	20 (80.0)	5.94 (17.26)	-37.5	0.00	6.25	18.75	31.3		
	Week 94	Sparsentan		26	20 (76.9)	13.13 (26.20)	-18.8	-3.13	3.13	28.13	75.0	0.12 [-0.51, 0.75]	
		Irbesartan	25	19 (76.0)	10.53 (16.01)	-25.0	0.00	6.25	25.00	37.5			
Week 110	Sparsentan	26	21 (80.8)	15.48 (27.43)	-18.8	0.00	6.25	18.75	87.5	0.69 [0.03, 1.34]			
	Irbesartan	25	17 (68.0)	-1.84 (22.18)	-62.5	-12.50	6.25	12.50	31.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline urine protein excretion														
<= 1.75 g/day	KDQOL: burden of kidney disease	Baseline	Sparsentan	98	93 (94.9)	69.76 (25.04)	0.0	56.25	75.00	87.50	100.0			
			Irbesartan	93	83 (89.2)	78.61 (21.76)	6.3	68.75	87.50	93.75	100.0			
		Week 24	Sparsentan	98	80 (81.6)	77.42 (23.20)	12.5	68.75	81.25	100.00	100.0			
			Irbesartan	93	55 (59.1)	81.82 (22.12)	0.0	68.75	87.50	100.00	100.0			
		Week 48	Sparsentan	98	82 (83.7)	78.89 (22.84)	0.0	75.00	81.25	100.00	100.0			
			Irbesartan	93	57 (61.3)	81.69 (18.86)	25.0	75.00	81.25	100.00	100.0			
		Week 70	Sparsentan	98	75 (76.5)	78.33 (21.73)	6.3	68.75	81.25	100.00	100.0			
			Irbesartan	93	61 (65.6)	83.61 (18.86)	12.5	75.00	87.50	100.00	100.0			
		Week 94	Sparsentan	98	76 (77.6)	76.89 (23.94)	0.0	68.75	81.25	100.00	100.0			
			Irbesartan	93	63 (67.7)	78.08 (22.53)	12.5	68.75	81.25	100.00	100.0			
		Week 110	Sparsentan	98	75 (76.5)	77.00 (21.09)	18.8	62.50	81.25	93.75	100.0			
			Irbesartan	93	57 (61.3)	76.97 (22.94)	12.5	68.75	81.25	100.00	100.0			
			KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	98	80 (81.6)	8.98 (23.32)	-81.3	0.00	6.25	21.88	81.3	0.34 [-0.01, 0.68]
					Irbesartan	93	55 (59.1)	1.93 (16.62)	-31.3	-6.25	0.00	12.50	50.0	
		Week 48	Sparsentan	98	82 (83.7)	8.00 (20.21)	-68.8	-6.25	6.25	25.00	62.5	0.32 [-0.02, 0.66]		
			Irbesartan	93	57 (61.3)	1.86 (17.72)	-50.0	-6.25	0.00	12.50	56.3			
		Week 70	Sparsentan	98	75 (76.5)	7.42 (21.84)	-43.8	-6.25	0.00	25.00	62.5	0.17 [-0.17, 0.51]		
			Irbesartan	93	61 (65.6)	4.00 (16.73)	-43.8	-6.25	0.00	12.50	43.8			
		Week 94	Sparsentan	98	76 (77.6)	8.22 (25.27)	-62.5	-6.25	6.25	25.00	68.8	0.36 [0.03, 0.70]		
			Irbesartan	93	63 (67.7)	-0.10 (19.43)	-37.5	-12.50	0.00	12.50	56.3			
		Week 110	Sparsentan	98	75 (76.5)	8.25 (25.73)	-50.0	-6.25	0.00	25.00	75.0	0.34 [-0.00, 0.69]		
			Irbesartan	93	57 (61.3)	0.22 (19.90)	-62.5	-12.50	0.00	12.50	50.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 1.75 g/day	KDQOL: burden of kidney disease	Baseline	Sparsentan	104	94 (90.4)	70.35 (25.58)	6.3	56.25	75.00	87.50	100.0		
			Irbesartan	109	99 (90.8)	75.51 (23.65)	18.8	56.25	81.25	100.00	100.0		
		Week 24	Sparsentan	104	77 (74.0)	76.06 (21.35)	0.0	68.75	81.25	93.75	100.0		
			Irbesartan	109	76 (69.7)	76.40 (23.11)	6.3	65.63	81.25	93.75	100.0		
		Week 48	Sparsentan	104	80 (76.9)	75.94 (22.87)	6.3	62.50	81.25	96.88	100.0		
			Irbesartan	109	67 (61.5)	78.64 (23.24)	0.0	68.75	81.25	100.00	100.0		
		Week 70	Sparsentan	104	85 (81.7)	78.60 (22.11)	12.5	68.75	81.25	93.75	100.0		
			Irbesartan	109	69 (63.3)	76.00 (23.25)	12.5	68.75	75.00	93.75	100.0		
		Week 94	Sparsentan	104	75 (72.1)	75.58 (21.82)	0.0	68.75	81.25	93.75	100.0		
			Irbesartan	109	67 (61.5)	75.37 (22.13)	12.5	62.50	75.00	93.75	100.0		
		Week 110	Sparsentan	104	74 (71.2)	76.77 (22.08)	25.0	62.50	81.25	93.75	100.0		
			Irbesartan	109	64 (58.7)	75.39 (21.70)	25.0	62.50	75.00	100.00	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	104	77 (74.0)	6.49 (21.40)	-50.0	-6.25	6.25	12.50	75.0	0.34 [0.02, 0.66]
				Irbesartan	109	76 (69.7)	-0.25 (18.18)	-50.0	-9.38	0.00	6.25	56.3	
	Week 48		Sparsentan	104	80 (76.9)	7.42 (20.89)	-62.5	0.00	6.25	18.75	56.3	0.25 [-0.08, 0.57]	
			Irbesartan	109	67 (61.5)	2.52 (18.34)	-62.5	-6.25	0.00	18.75	50.0		
	Week 70		Sparsentan	104	85 (81.7)	9.12 (21.83)	-50.0	0.00	6.25	18.75	87.5	0.37 [0.05, 0.69]	
			Irbesartan	109	69 (63.3)	1.54 (19.16)	-37.5	-6.25	0.00	6.25	56.3		
	Week 94		Sparsentan	104	75 (72.1)	6.25 (24.03)	-87.5	0.00	6.25	18.75	75.0	0.29 [-0.04, 0.62]	
			Irbesartan	109	67 (61.5)	-0.47 (22.33)	-75.0	-12.50	0.00	12.50	50.0		
Week 110	Sparsentan	104	74 (71.2)	6.59 (23.29)	-56.3	0.00	6.25	18.75	87.5	0.40 [0.06, 0.74]			
	Irbesartan	109	64 (58.7)	-2.34 (21.23)	-56.3	-18.75	0.00	6.25	56.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline use of antihypertensives														
Yes	KDQOL: burden of kidney disease	Baseline	Sparsentan	90	79 (87.8)	70.17 (25.28)	6.3	56.25	75.00	87.50	100.0			
			Irbesartan	88	76 (86.4)	77.80 (21.73)	18.8	62.50	84.38	100.00	100.0			
		Week 24	Sparsentan	90	66 (73.3)	77.18 (22.80)	0.0	68.75	81.25	93.75	100.0			
			Irbesartan	88	49 (55.7)	78.57 (23.18)	6.3	68.75	81.25	100.00	100.0			
		Week 48	Sparsentan	90	65 (72.2)	78.17 (21.37)	6.3	68.75	81.25	100.00	100.0			
			Irbesartan	88	47 (53.4)	76.99 (20.32)	25.0	68.75	81.25	93.75	100.0			
		Week 70	Sparsentan	90	68 (75.6)	77.67 (23.69)	6.3	75.00	81.25	93.75	100.0			
			Irbesartan	88	49 (55.7)	75.77 (24.43)	12.5	62.50	75.00	100.00	100.0			
		Week 94	Sparsentan	90	63 (70.0)	77.68 (22.16)	6.3	68.75	81.25	93.75	100.0			
			Irbesartan	88	54 (61.4)	72.45 (24.27)	12.5	62.50	75.00	93.75	100.0			
		Week 110	Sparsentan	90	63 (70.0)	76.29 (22.73)	25.0	62.50	81.25	93.75	100.0			
			Irbesartan	88	51 (58.0)	73.16 (21.55)	25.0	68.75	75.00	87.50	100.0			
			KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	90	66 (73.3)	8.43 (20.58)	-31.3	-6.25	0.00	18.75	75.0	0.49 [0.11, 0.86]
					Irbesartan	88	49 (55.7)	-0.77 (16.12)	-37.5	-6.25	0.00	6.25	37.5	
		Week 48		Sparsentan	90	65 (72.2)	9.90 (19.51)	-50.0	0.00	6.25	25.00	56.3	0.49 [0.11, 0.87]	
				Irbesartan	88	47 (53.4)	0.53 (18.56)	-50.0	-6.25	0.00	6.25	43.8		
		Week 70		Sparsentan	90	68 (75.6)	7.17 (21.93)	-50.0	-6.25	3.13	18.75	87.5	0.34 [-0.03, 0.71]	
				Irbesartan	88	49 (55.7)	0.00 (19.39)	-43.8	-12.50	0.00	12.50	37.5		
Week 94	Sparsentan	90		63 (70.0)	9.62 (24.31)	-62.5	0.00	6.25	25.00	75.0	0.54 [0.17, 0.91]			
	Irbesartan	88		54 (61.4)	-3.24 (23.25)	-75.0	-18.75	0.00	12.50	50.0				
Week 110	Sparsentan	90		63 (70.0)	7.04 (22.10)	-56.3	0.00	6.25	18.75	62.5	0.49 [0.12, 0.87]			
	Irbesartan	88		51 (58.0)	-3.06 (18.51)	-43.8	-12.50	0.00	6.25	43.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL: burden of kidney disease	Baseline	Sparsentan	112	108 (96.4)	69.97 (25.34)	0.0	53.13	75.00	87.50	100.0	
			Irbesartan	114	106 (93.0)	76.30 (23.62)	6.3	62.50	81.25	93.75	100.0	
		Week 24	Sparsentan	112	91 (81.3)	76.44 (21.97)	18.8	62.50	81.25	100.00	100.0	
			Irbesartan	114	82 (71.9)	78.73 (22.67)	0.0	68.75	81.25	100.00	100.0	
		Week 48	Sparsentan	112	97 (86.6)	76.93 (23.86)	0.0	62.50	81.25	100.00	100.0	
			Irbesartan	114	77 (67.5)	81.90 (21.81)	0.0	75.00	87.50	100.00	100.0	
		Week 70	Sparsentan	112	92 (82.1)	79.08 (20.52)	25.0	68.75	81.25	100.00	100.0	
			Irbesartan	114	81 (71.1)	81.87 (19.43)	25.0	75.00	87.50	100.00	100.0	
		Week 94	Sparsentan	112	88 (78.6)	75.21 (23.39)	0.0	68.75	78.13	93.75	100.0	
			Irbesartan	114	76 (66.7)	79.69 (20.39)	25.0	68.75	81.25	100.00	100.0	
		Week 110	Sparsentan	112	86 (76.8)	77.33 (20.70)	18.8	62.50	81.25	100.00	100.0	
			Irbesartan	114	70 (61.4)	78.30 (22.59)	12.5	62.50	81.25	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	112	91 (81.3)	7.28 (23.67)	-81.3	0.00	6.25	18.75	81.3	0.27 [-0.03, 0.57]
			Irbesartan	114	82 (71.9)	1.52 (18.34)	-50.0	-6.25	0.00	6.25	56.3	
		Week 48	Sparsentan	112	97 (86.6)	6.25 (21.08)	-68.8	0.00	0.00	18.75	62.5	0.15 [-0.15, 0.45]
			Irbesartan	114	77 (67.5)	3.25 (17.67)	-62.5	-6.25	0.00	12.50	56.3	
		Week 70	Sparsentan	112	92 (82.1)	9.17 (21.76)	-43.8	0.00	6.25	18.75	87.5	0.25 [-0.05, 0.55]
			Irbesartan	114	81 (71.1)	4.32 (17.08)	-25.0	-6.25	0.00	6.25	56.3	
		Week 94	Sparsentan	112	88 (78.6)	5.54 (24.81)	-87.5	-6.25	6.25	18.75	62.5	0.17 [-0.14, 0.47]
			Irbesartan	114	76 (66.7)	1.81 (18.92)	-50.0	-6.25	0.00	12.50	56.3	
Week 110	Sparsentan	112	86 (76.8)	7.70 (26.21)	-50.0	-6.25	6.25	18.75	87.5	0.30 [-0.01, 0.62]		
	Irbesartan	114	70 (61.4)	0.27 (21.97)	-62.5	-12.50	0.00	12.50	56.3			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Time since renal biopsy														
<= 5 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	113	104 (92.0)	67.91 (26.43)	0.0	50.00	75.00	87.50	100.0			
			Irbesartan	127	118 (92.9)	75.48 (22.96)	6.3	56.25	81.25	93.75	100.0			
		Week 24	Sparsentan	113	88 (77.9)	76.70 (23.13)	0.0	68.75	81.25	100.00	100.0			
			Irbesartan	127	86 (67.7)	77.83 (22.84)	0.0	62.50	81.25	100.00	100.0			
		Week 48	Sparsentan	113	92 (81.4)	76.02 (23.24)	6.3	62.50	81.25	100.00	100.0			
			Irbesartan	127	85 (66.9)	79.56 (22.21)	0.0	75.00	81.25	100.00	100.0			
		Week 70	Sparsentan	113	90 (79.6)	76.46 (23.46)	6.3	68.75	81.25	93.75	100.0			
			Irbesartan	127	86 (67.7)	79.36 (21.48)	12.5	68.75	84.38	100.00	100.0			
		Week 94	Sparsentan	113	86 (76.1)	76.16 (22.82)	0.0	68.75	81.25	93.75	100.0			
			Irbesartan	127	86 (67.7)	77.11 (22.10)	12.5	68.75	78.13	100.00	100.0			
		Week 110	Sparsentan	113	88 (77.9)	76.14 (22.58)	18.8	59.38	81.25	93.75	100.0			
			Irbesartan	127	80 (63.0)	76.64 (22.82)	12.5	62.50	75.00	100.00	100.0			
			KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	113	88 (77.9)	11.15 (21.76)	-50.0	0.00	6.25	25.00	81.3	0.45 [0.15, 0.75]
					Irbesartan	127	86 (67.7)	2.03 (18.48)	-50.0	-6.25	0.00	12.50	56.3	
		Week 48		Sparsentan	113	92 (81.4)	9.71 (18.59)	-43.8	0.00	6.25	25.00	56.3	0.31 [0.01, 0.61]	
				Irbesartan	127	85 (66.9)	3.75 (20.00)	-62.5	-6.25	0.00	12.50	56.3		
		Week 70		Sparsentan	113	90 (79.6)	8.68 (21.87)	-50.0	-6.25	6.25	18.75	87.5	0.28 [-0.02, 0.57]	
				Irbesartan	127	86 (67.7)	3.05 (18.58)	-43.8	-6.25	0.00	12.50	43.8		
Week 94	Sparsentan	113		86 (76.1)	10.10 (25.00)	-62.5	0.00	6.25	25.00	68.8	0.39 [0.09, 0.69]			
	Irbesartan	127		86 (67.7)	1.02 (21.58)	-75.0	-12.50	0.00	12.50	56.3				
Week 110	Sparsentan	113		88 (77.9)	8.38 (25.67)	-56.3	-6.25	6.25	18.75	87.5	0.36 [0.06, 0.67]			
	Irbesartan	127		80 (63.0)	-0.16 (21.19)	-62.5	-9.38	0.00	9.38	50.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 5 years	KDQOL: burden of kidney disease	Baseline	Sparsentan	89	83 (93.3)	72.74 (23.56)	0.0	62.50	75.00	87.50	100.0		
			Irbesartan	75	64 (85.3)	79.59 (22.44)	18.8	68.75	87.50	93.75	100.0		
		Week 24	Sparsentan	89	69 (77.5)	76.81 (21.25)	18.8	68.75	81.25	93.75	100.0		
			Irbesartan	75	45 (60.0)	80.28 (22.81)	6.3	68.75	81.25	100.00	100.0		
		Week 48	Sparsentan	89	70 (78.7)	79.29 (22.31)	0.0	75.00	84.38	100.00	100.0		
			Irbesartan	75	39 (52.0)	81.09 (19.42)	25.0	75.00	81.25	93.75	100.0		
		Week 70	Sparsentan	89	70 (78.7)	81.07 (19.47)	25.0	68.75	87.50	100.00	100.0		
			Irbesartan	75	44 (58.7)	79.97 (21.97)	12.5	75.00	84.38	100.00	100.0		
		Week 94	Sparsentan	89	65 (73.0)	76.35 (23.06)	0.0	68.75	81.25	93.75	100.0		
			Irbesartan	75	44 (58.7)	75.85 (22.86)	12.5	65.63	75.00	96.88	100.0		
		Week 110	Sparsentan	89	61 (68.5)	77.97 (20.01)	31.3	62.50	81.25	93.75	100.0		
			Irbesartan	75	41 (54.7)	75.15 (21.22)	25.0	62.50	81.25	93.75	100.0		
		KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	89	69 (77.5)	3.44 (22.52)	-81.3	-6.25	0.00	12.50	75.0	0.27 [-0.11, 0.65]
			Irbesartan	75	45 (60.0)	-1.94 (15.36)	-37.5	-6.25	0.00	6.25	37.5		
	Week 48		Sparsentan	89	70 (78.7)	5.09 (22.60)	-68.8	0.00	0.00	18.75	62.5	0.32 [-0.08, 0.71]	
			Irbesartan	75	39 (52.0)	-1.12 (12.07)	-37.5	-6.25	0.00	6.25	18.8		
	Week 70		Sparsentan	89	70 (78.7)	7.86 (21.82)	-43.8	0.00	0.00	18.75	87.5	0.29 [-0.09, 0.67]	
			Irbesartan	75	44 (58.7)	1.99 (17.09)	-31.3	-6.25	0.00	6.25	56.3		
	Week 94		Sparsentan	89	65 (73.0)	3.46 (23.72)	-87.5	-6.25	0.00	18.75	75.0	0.29 [-0.10, 0.67]	
			Irbesartan	75	44 (58.7)	-2.84 (19.46)	-50.0	-15.63	0.00	9.38	50.0		
Week 110	Sparsentan	89	61 (68.5)	6.05 (22.79)	-50.0	-6.25	0.00	18.75	62.5	0.42 [0.02, 0.82]			
	Irbesartan	75	41 (54.7)	-3.05 (19.42)	-43.8	-18.75	0.00	6.25	56.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Subgroup: History of hypertension												
Yes	KDQOL: burden of kidney disease	Baseline	Sparsentan	155	141 (91.0)	69.95 (25.90)	0.0	56.25	75.00	87.50	100.0	
			Irbesartan	161	142 (88.2)	78.30 (21.89)	18.8	62.50	84.38	100.00	100.0	
		Week 24	Sparsentan	155	119 (76.8)	76.94 (21.81)	0.0	62.50	81.25	93.75	100.0	
			Irbesartan	161	100 (62.1)	80.38 (21.52)	6.3	68.75	87.50	100.00	100.0	
		Week 48	Sparsentan	155	121 (78.1)	77.12 (23.27)	0.0	62.50	81.25	100.00	100.0	
			Irbesartan	161	95 (59.0)	80.53 (21.14)	0.0	68.75	87.50	100.00	100.0	
		Week 70	Sparsentan	155	123 (79.4)	78.35 (22.08)	12.5	68.75	81.25	93.75	100.0	
			Irbesartan	161	100 (62.1)	79.31 (22.45)	12.5	68.75	84.38	100.00	100.0	
		Week 94	Sparsentan	155	114 (73.5)	76.04 (24.18)	0.0	68.75	81.25	93.75	100.0	
			Irbesartan	161	101 (62.7)	75.06 (23.63)	12.5	62.50	75.00	100.00	100.0	
		Week 110	Sparsentan	155	112 (72.3)	76.95 (21.85)	18.8	62.50	81.25	93.75	100.0	
			Irbesartan	161	92 (57.1)	77.51 (21.31)	25.0	68.75	81.25	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	155	119 (76.8)	8.40 (22.08)	-43.8	-6.25	6.25	18.75	81.3	0.35 [0.08, 0.62]
			Irbesartan	161	100 (62.1)	1.38 (17.53)	-50.0	-6.25	0.00	12.50	56.3	
		Week 48	Sparsentan	155	121 (78.1)	7.95 (20.33)	-68.8	0.00	6.25	18.75	62.5	0.29 [0.02, 0.56]
			Irbesartan	161	95 (59.0)	2.37 (18.19)	-62.5	-6.25	0.00	12.50	50.0	
		Week 70	Sparsentan	155	123 (79.4)	8.64 (20.83)	-50.0	0.00	6.25	18.75	87.5	0.29 [0.03, 0.56]
			Irbesartan	161	100 (62.1)	2.81 (18.54)	-43.8	-6.25	0.00	12.50	56.3	
Week 94	Sparsentan	155	114 (73.5)	7.46 (24.31)	-87.5	-6.25	6.25	25.00	75.0	0.38 [0.11, 0.65]		
	Irbesartan	161	101 (62.7)	-1.36 (21.26)	-75.0	-12.50	0.00	12.50	50.0			
Week 110	Sparsentan	155	112 (72.3)	7.42 (23.93)	-56.3	-6.25	6.25	18.75	75.0	0.33 [0.05, 0.61]		
	Irbesartan	161	92 (57.1)	0.34 (18.17)	-43.8	-12.50	0.00	6.25	56.3			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSHM: KDQOL: Course of burden of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL: burden of kidney disease	Baseline	Sparsentan	47	46 (97.9)	70.38 (23.40)	12.5	50.00	75.00	87.50	100.0	
			Irbesartan	41	40 (97.6)	72.03 (25.48)	6.3	56.25	75.00	93.75	100.0	
		Week 24	Sparsentan	47	38 (80.9)	76.15 (23.87)	12.5	68.75	78.13	100.00	100.0	
			Irbesartan	41	31 (75.6)	73.19 (26.03)	0.0	56.25	75.00	93.75	100.0	
		Week 48	Sparsentan	47	41 (87.2)	78.35 (21.75)	6.3	68.75	87.50	93.75	100.0	
			Irbesartan	41	29 (70.7)	78.45 (22.13)	6.3	75.00	81.25	93.75	100.0	
		Week 70	Sparsentan	47	37 (78.7)	78.89 (21.42)	6.3	68.75	81.25	100.00	100.0	
			Irbesartan	41	30 (73.2)	80.42 (18.62)	31.3	75.00	84.38	93.75	100.0	
		Week 94	Sparsentan	47	37 (78.7)	76.86 (18.39)	37.5	68.75	75.00	93.75	100.0	
			Irbesartan	41	29 (70.7)	82.33 (15.85)	37.5	75.00	81.25	93.75	100.0	
		Week 110	Sparsentan	47	37 (78.7)	76.69 (20.76)	37.5	62.50	75.00	100.00	100.0	
			Irbesartan	41	29 (70.7)	71.77 (24.75)	12.5	62.50	75.00	100.00	100.0	
	KDQOL: Change from baseline in burden of kidney disease	Week 24	Sparsentan	47	38 (80.9)	5.76 (23.40)	-81.3	0.00	6.25	12.50	56.3	0.35 [-0.13, 0.83]
			Irbesartan	41	31 (75.6)	-1.61 (17.53)	-31.3	-12.50	-6.25	0.00	50.0	
		Week 48	Sparsentan	47	41 (87.2)	7.01 (21.16)	-50.0	0.00	6.25	18.75	50.0	0.27 [-0.21, 0.75]
			Irbesartan	41	29 (70.7)	1.72 (17.59)	-43.8	-6.25	0.00	12.50	56.3	
		Week 70	Sparsentan	47	37 (78.7)	7.26 (24.98)	-43.8	-6.25	6.25	18.75	87.5	0.23 [-0.25, 0.71]
			Irbesartan	41	30 (73.2)	2.29 (16.53)	-25.0	-6.25	0.00	6.25	43.8	
		Week 94	Sparsentan	47	37 (78.7)	6.59 (25.81)	-50.0	-6.25	6.25	18.75	62.5	0.14 [-0.35, 0.62]
			Irbesartan	41	29 (70.7)	3.45 (19.45)	-37.5	-6.25	0.00	12.50	56.3	
Week 110	Sparsentan	47	37 (78.7)	7.43 (26.43)	-50.0	-6.25	6.25	18.75	87.5	0.50 [0.01, 0.99]		
	Irbesartan	41	29 (70.7)	-5.82 (26.67)	-62.5	-25.00	0.00	6.25	50.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease	Repeated measures analysis									
	Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Sex	Overall	Sparsentan								Interaction: 0.210
Male	Week 24	Sparsentan	139	109 (78.4)	4.59 (1.70)	(1.25, 7.94)	5.19 (2.52)	(0.25, 10.12)	0.040	*
		Irbesartan	143	93 (65.0)	-0.59 (1.84)	(-4.20, 3.02)				
	Week 48	Sparsentan	139	112 (80.6)	4.56 (1.67)	(1.28, 7.85)	1.01 (2.51)	(-3.92, 5.94)	0.688	
		Irbesartan	143	87 (60.8)	3.55 (1.86)	(-0.10, 7.21)				
	Week 70	Sparsentan	139	114 (82.0)	6.22 (1.67)	(2.94, 9.50)	4.18 (2.49)	(-0.71, 9.07)	0.093	
		Irbesartan	143	92 (64.3)	2.04 (1.84)	(-1.57, 5.64)				
	Week 94	Sparsentan	139	105 (75.5)	5.39 (1.72)	(2.01, 8.77)	6.32 (2.51)	(1.40, 11.25)	0.012	*
		Irbesartan	143	96 (67.1)	-0.93 (1.82)	(-4.50, 2.63)				
	Week 110	Sparsentan	139	103 (74.1)	5.19 (1.75)	(1.75, 8.64)	6.56 (2.58)	(1.50, 11.62)	0.011	*
		Irbesartan	143	89 (62.2)	-1.37 (1.88)	(-5.06, 2.33)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Female	Week 24	Sparsentan	63	48 (76.2)	9.06 (2.58)	(3.98, 14.14)	-2.25 (3.91)	(-9.94, 5.44)	0.565
		Irbesartan	59	38 (64.4)	11.31 (2.92)	(5.57, 17.05)			
	Week 48	Sparsentan	63	50 (79.4)	8.77 (2.53)	(3.80, 13.74)	2.16 (3.87)	(-5.46, 9.78)	0.578
		Irbesartan	59	37 (62.7)	6.61 (2.92)	(0.86, 12.36)			
	Week 70	Sparsentan	63	46 (73.0)	8.21 (2.61)	(3.07, 13.35)	0.54 (3.92)	(-7.19, 8.26)	0.891
		Irbesartan	59	38 (64.4)	7.67 (2.91)	(1.94, 13.40)			
	Week 94	Sparsentan	63	46 (73.0)	3.04 (2.63)	(-2.13, 8.22)	-2.78 (4.04)	(-10.72, 5.16)	0.491
		Irbesartan	59	34 (57.6)	5.83 (3.04)	(-0.15, 11.81)			
	Week 110	Sparsentan	63	46 (73.0)	5.81 (2.65)	(0.59, 11.03)	0.06 (4.14)	(-8.09, 8.21)	0.989
		Irbesartan	59	32 (54.2)	5.75 (3.16)	(-0.46, 11.97)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.988
<= 45 years	Week 24	Sparsentan	96	71 (74.0)	7.94 (2.32)	(3.38, 12.50)	4.23 (3.34)	(-2.33, 10.80)	0.206
		Irbesartan	99	67 (67.7)	3.71 (2.39)	(-0.99, 8.40)			
	Week 48	Sparsentan	96	75 (78.1)	7.04 (2.27)	(2.58, 11.49)	1.75 (3.31)	(-4.76, 8.27)	0.597
		Irbesartan	99	62 (62.6)	5.28 (2.41)	(0.55, 10.02)			
	Week 70	Sparsentan	96	72 (75.0)	7.57 (2.30)	(3.05, 12.08)	2.98 (3.35)	(-3.60, 9.57)	0.374
		Irbesartan	99	62 (62.6)	4.58 (2.43)	(-0.20, 9.36)			
	Week 94	Sparsentan	96	68 (70.8)	4.25 (2.34)	(-0.35, 8.85)	0.81 (3.38)	(-5.83, 7.46)	0.810
		Irbesartan	99	64 (64.6)	3.44 (2.43)	(-1.35, 8.22)			
	Week 110	Sparsentan	96	69 (71.9)	6.20 (2.37)	(1.55, 10.85)	4.59 (3.47)	(-2.22, 11.41)	0.186
		Irbesartan	99	59 (59.6)	1.61 (2.53)	(-3.36, 6.58)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 45 years	Week 24	Sparsentan	106	86 (81.1)	3.99 (1.78)	(0.50, 7.47)	2.02 (2.74)	(-3.36, 7.41)	0.460
		Irbesartan	103	64 (62.1)	1.96 (2.07)	(-2.10, 6.03)			
	Week 48	Sparsentan	106	87 (82.1)	4.78 (1.76)	(1.33, 8.24)	1.22 (2.73)	(-4.15, 6.58)	0.656
		Irbesartan	103	62 (60.2)	3.56 (2.08)	(-0.52, 7.64)			
	Week 70	Sparsentan	106	88 (83.0)	6.06 (1.76)	(2.61, 9.51)	3.51 (2.67)	(-1.73, 8.76)	0.189
		Irbesartan	103	68 (66.0)	2.55 (2.00)	(-1.37, 6.47)			
	Week 94	Sparsentan	106	83 (78.3)	5.14 (1.81)	(1.59, 8.69)	6.70 (2.73)	(1.33, 12.06)	0.014 *
		Irbesartan	103	66 (64.1)	-1.56 (2.03)	(-5.54, 2.43)			
	Week 110	Sparsentan	106	80 (75.5)	4.65 (1.85)	(1.03, 8.28)	4.83 (2.81)	(-0.68, 10.33)	0.086
		Irbesartan	103	62 (60.2)	-0.17 (2.10)	(-4.29, 3.95)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.959
<= 18 years	Week 24	Sparsentan	9	5 (55.6)	23.28 (8.23)	(6.37, 40.18)	17.44 (11.82)	(-7.05, 41.92)	0.154
		Irbesartan	5	5 (100.0)	5.84 (8.18)	(-11.20, 22.88)			
	Week 48	Sparsentan	9	7 (77.8)	2.31 (6.88)	(-11.94, 16.56)	-6.89 (12.51)	(-32.54, 18.76)	0.586
		Irbesartan	5	3 (60.0)	9.20 (10.33)	(-11.88, 30.27)			
	Week 70	Sparsentan	9	7 (77.8)	6.48 (6.89)	(-7.84, 20.79)	1.94 (11.84)	(-22.61, 26.50)	0.871
		Irbesartan	5	4 (80.0)	4.53 (9.34)	(-14.75, 23.82)			
	Week 94	Sparsentan	9	5 (55.6)	7.06 (8.21)	(-9.79, 23.90)	10.48 (12.72)	(-15.85, 36.82)	0.418
		Irbesartan	5	4 (80.0)	-3.43 (9.36)	(-22.88, 16.02)			
	Week 110	Sparsentan	9	4 (44.4)	10.59 (9.26)	(-8.40, 29.58)	11.61 (15.90)	(-20.98, 44.20)	0.471
		Irbesartan	5	2 (40.0)	-1.02 (12.86)	(-27.34, 25.29)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
> 18 to 40 years	Week 24	Sparsentan	102	79 (77.5)	6.87 (2.17)	(2.60, 11.14)	4.65 (3.15)	(-1.55, 10.84)	0.141	
		Irbesartan	109	71 (65.1)	2.22 (2.28)	(-2.25, 6.70)				
	Week 48	Sparsentan	102	82 (80.4)	5.57 (2.13)	(1.38, 9.75)	2.07 (3.13)	(-4.08, 8.23)	0.508	
		Irbesartan	109	67 (61.5)	3.49 (2.29)	(-1.00, 7.99)				
	Week 70	Sparsentan	102	78 (76.5)	8.37 (2.17)	(4.11, 12.63)	4.40 (3.17)	(-1.83, 10.62)	0.166	
		Irbesartan	109	68 (62.4)	3.97 (2.30)	(-0.55, 8.49)				
	Week 94	Sparsentan	102	73 (71.6)	4.06 (2.22)	(-0.31, 8.43)	1.24 (3.21)	(-5.08, 7.55)	0.701	
		Irbesartan	109	69 (63.3)	2.82 (2.31)	(-1.71, 7.36)				
	Week 110	Sparsentan	102	72 (70.6)	5.63 (2.27)	(1.16, 10.10)	4.87 (3.30)	(-1.62, 11.36)	0.141	
		Irbesartan	109	65 (59.6)	0.76 (2.38)	(-3.92, 5.44)				
> 40 years	Week 24	Sparsentan	91	73 (80.2)	4.09 (1.94)	(0.29, 7.90)	1.14 (2.97)	(-4.69, 6.97)	0.702	
		Irbesartan	88	55 (62.5)	2.96 (2.23)	(-1.42, 7.34)				
	Week 48	Sparsentan	91	73 (80.2)	6.71 (1.92)	(2.93, 10.49)	1.77 (2.96)	(-4.04, 7.58)	0.549	
		Irbesartan	88	54 (61.4)	4.94 (2.23)	(0.55, 9.32)				
	Week 70	Sparsentan	91	75 (82.4)	5.30 (1.91)	(1.55, 9.04)	2.50 (2.90)	(-3.19, 8.19)	0.388	
		Irbesartan	88	58 (65.9)	2.80 (2.17)	(-1.46, 7.05)				
	Week 94	Sparsentan	91	73 (80.2)	5.18 (1.93)	(1.39, 8.98)	6.60 (2.93)	(0.85, 12.36)	0.025 *	
		Irbesartan	88	57 (64.8)	-1.42 (2.19)	(-5.71, 2.88)				
	Week 110	Sparsentan	91	73 (80.2)	4.88 (1.94)	(1.07, 8.68)	4.65 (2.98)	(-1.20, 10.50)	0.119	
		Irbesartan	88	54 (61.4)	0.23 (2.25)	(-4.20, 4.65)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.612
North America	Week 24	Sparsentan	35	23 (65.7)	5.23 (3.72)	(-2.11, 12.56)	5.35 (4.83)	(-4.17, 14.87)	0.269
		Irbesartan	46	35 (76.1)	-0.12 (3.02)	(-6.08, 5.83)			
	Week 48	Sparsentan	35	25 (71.4)	5.43 (3.62)	(-1.70, 12.56)	7.13 (4.83)	(-2.38, 16.65)	0.141
		Irbesartan	46	32 (69.6)	-1.70 (3.12)	(-7.85, 4.45)			
	Week 70	Sparsentan	35	22 (62.9)	3.70 (3.79)	(-3.77, 11.17)	5.78 (5.00)	(-4.08, 15.64)	0.249
		Irbesartan	46	30 (65.2)	-2.07 (3.22)	(-8.41, 4.26)			
	Week 94	Sparsentan	35	23 (65.7)	-1.36 (3.74)	(-8.74, 6.02)	2.44 (5.00)	(-7.43, 12.30)	0.627
		Irbesartan	46	30 (65.2)	-3.79 (3.25)	(-10.20, 2.61)			
	Week 110	Sparsentan	35	21 (60.0)	-0.14 (3.90)	(-7.82, 7.54)	0.48 (5.09)	(-9.55, 10.52)	0.925
		Irbesartan	46	31 (67.4)	-0.63 (3.23)	(-6.99, 5.74)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Europe	Week 24	Sparsentan	98	73 (74.5)	4.81 (1.99)	(0.89, 8.72)	3.70 (2.94)	(-2.08, 9.49)	0.209
		Irbesartan	115	62 (53.9)	1.10 (2.17)	(-3.16, 5.36)			
	Week 48	Sparsentan	98	76 (77.6)	2.90 (1.95)	(-0.94, 6.73)	-0.50 (2.92)	(-6.24, 5.24)	0.864
		Irbesartan	115	59 (51.3)	3.39 (2.17)	(-0.86, 7.65)			
	Week 70	Sparsentan	98	75 (76.5)	5.54 (1.98)	(1.65, 9.43)	2.00 (2.86)	(-3.62, 7.62)	0.485
		Irbesartan	115	69 (60.0)	3.54 (2.06)	(-0.51, 7.58)			
	Week 94	Sparsentan	98	68 (69.4)	3.08 (2.06)	(-0.96, 7.12)	4.47 (2.90)	(-1.24, 10.17)	0.125
		Irbesartan	115	70 (60.9)	-1.38 (2.05)	(-5.40, 2.64)			
	Week 110	Sparsentan	98	67 (68.4)	3.97 (2.09)	(-0.14, 8.09)	4.02 (3.02)	(-1.92, 9.95)	0.184
		Irbesartan	115	61 (53.0)	-0.05 (2.17)	(-4.31, 4.22)			
Asia Pacific	Week 24	Sparsentan	69	61 (88.4)	9.45 (2.47)	(4.60, 14.31)	2.98 (4.15)	(-5.18, 11.14)	0.473
		Irbesartan	41	34 (82.9)	6.48 (3.29)	(-0.00, 12.95)			
	Week 48	Sparsentan	69	61 (88.4)	11.37 (2.45)	(6.56, 16.19)	2.36 (4.17)	(-5.84, 10.55)	0.572
		Irbesartan	41	33 (80.5)	9.02 (3.33)	(2.46, 15.57)			
	Week 70	Sparsentan	69	63 (91.3)	11.38 (2.42)	(6.61, 16.15)	5.02 (4.25)	(-3.35, 13.38)	0.239
		Irbesartan	41	31 (75.6)	6.36 (3.45)	(-0.43, 13.14)			
	Week 94	Sparsentan	69	60 (87.0)	10.76 (2.47)	(5.91, 15.61)	3.01 (4.33)	(-5.50, 11.52)	0.487
		Irbesartan	41	30 (73.2)	7.75 (3.51)	(0.85, 14.65)			
	Week 110	Sparsentan	69	61 (88.4)	11.03 (2.47)	(6.17, 15.89)	11.36 (4.39)	(2.73, 20.00)	0.010 *
		Irbesartan	41	29 (70.7)	-0.33 (3.58)	(-7.38, 6.71)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.887
< 27 kg/m**2	Week 24	Sparsentan	83	66 (79.5)	7.23 (2.27)	(2.76, 11.69)	1.47 (3.23)	(-4.87, 7.81)	0.648
		Irbesartan	94	66 (70.2)	5.75 (2.27)	(1.29, 10.22)			
	Week 48	Sparsentan	83	66 (79.5)	6.50 (2.26)	(2.06, 10.94)	2.58 (3.26)	(-3.83, 8.99)	0.430
		Irbesartan	94	60 (63.8)	3.92 (2.33)	(-0.66, 8.50)			
	Week 70	Sparsentan	83	64 (77.1)	7.80 (2.29)	(3.30, 12.31)	5.83 (3.31)	(-0.66, 12.33)	0.078
		Irbesartan	94	59 (62.8)	1.97 (2.36)	(-2.67, 6.61)			
	Week 94	Sparsentan	83	63 (75.9)	6.85 (2.32)	(2.30, 11.40)	3.08 (3.31)	(-3.42, 9.58)	0.352
		Irbesartan	94	62 (66.0)	3.77 (2.34)	(-0.82, 8.36)			
	Week 110	Sparsentan	83	63 (75.9)	8.00 (2.33)	(3.42, 12.58)	5.48 (3.38)	(-1.17, 12.13)	0.106
		Irbesartan	94	57 (60.6)	2.52 (2.44)	(-2.27, 7.31)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 27 kg/m**2	Week 24	Sparsentan	119	91 (76.5)	4.72 (1.83)	(1.13, 8.30)	3.96 (2.85)	(-1.64, 9.56)	0.165
		Irbesartan	107	65 (60.7)	0.75 (2.17)	(-3.51, 5.02)			
	Week 48	Sparsentan	119	96 (80.7)	5.00 (1.78)	(1.51, 8.49)	-0.20 (2.79)	(-5.69, 5.29)	0.943
		Irbesartan	107	64 (59.8)	5.20 (2.15)	(0.98, 9.41)			
	Week 70	Sparsentan	119	96 (80.7)	5.76 (1.79)	(2.25, 9.27)	0.40 (2.75)	(-5.00, 5.79)	0.885
		Irbesartan	107	71 (66.4)	5.36 (2.08)	(1.28, 9.44)			
	Week 94	Sparsentan	119	88 (73.9)	2.75 (1.85)	(-0.87, 6.38)	3.71 (2.83)	(-1.84, 9.26)	0.190
		Irbesartan	107	67 (62.6)	-0.95 (2.13)	(-5.13, 3.22)			
	Week 110	Sparsentan	119	86 (72.3)	3.17 (1.89)	(-0.53, 6.88)	3.88 (2.91)	(-1.83, 9.59)	0.183
		Irbesartan	107	63 (58.9)	-0.70 (2.20)	(-5.02, 3.61)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan						Interaction:	0.748
eGFR Low and UP High	Week 24	Sparsentan	71	54 (76.1)	6.96 (2.51)	(2.02, 11.89)	8.20 (3.83)	(0.67, 15.73)	0.033 *
		Irbesartan	74	40 (54.1)	-1.24 (2.89)	(-6.92, 4.44)			
	Week 48	Sparsentan	71	51 (71.8)	5.71 (2.54)	(0.72, 10.70)	3.20 (3.88)	(-4.42, 10.83)	0.409
		Irbesartan	74	37 (50.0)	2.51 (2.93)	(-3.25, 8.26)			
	Week 70	Sparsentan	71	57 (80.3)	4.38 (2.46)	(-0.46, 9.21)	4.28 (3.84)	(-3.28, 11.84)	0.266
		Irbesartan	74	37 (50.0)	0.10 (2.95)	(-5.71, 5.91)			
	Week 94	Sparsentan	71	51 (71.8)	0.28 (2.55)	(-4.73, 5.30)	3.77 (3.92)	(-3.94, 11.48)	0.337
		Irbesartan	74	37 (50.0)	-3.48 (2.98)	(-9.34, 2.37)			
	Week 110	Sparsentan	71	53 (74.6)	3.02 (2.54)	(-1.98, 8.02)	3.54 (3.97)	(-4.27, 11.35)	0.373
		Irbesartan	74	36 (48.6)	-0.52 (3.05)	(-6.52, 5.48)			

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A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
					Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
eGFR Low and UP Low	Week 24	Sparsentan	55	43 (78.2)	3.63 (2.67)	(-1.63, 8.88)	-3.97 (3.96)	(-11.78, 3.83)	0.317
		Irbesartan	55	37 (67.3)	7.60 (2.90)	(1.90, 13.30)			
	Week 48	Sparsentan	55	43 (78.2)	8.56 (2.64)	(3.35, 13.76)	1.92 (3.93)	(-5.83, 9.67)	0.626
		Irbesartan	55	35 (63.6)	6.64 (2.89)	(0.94, 12.33)			
	Week 70	Sparsentan	55	42 (76.4)	7.06 (2.68)	(1.77, 12.34)	0.91 (3.89)	(-6.76, 8.58)	0.815
		Irbesartan	55	39 (70.9)	6.15 (2.80)	(0.64, 11.66)			
	Week 94	Sparsentan	55	41 (74.5)	10.07 (2.72)	(4.70, 15.43)	8.36 (3.93)	(0.62, 16.11)	0.034 *
		Irbesartan	55	39 (70.9)	1.70 (2.81)	(-3.83, 7.24)			
	Week 110	Sparsentan	55	37 (67.3)	6.02 (2.85)	(0.40, 11.64)	0.92 (4.13)	(-7.22, 9.05)	0.825
		Irbesartan	55	34 (61.8)	5.10 (2.97)	(-0.75, 10.95)			
eGFR High and UP High	Week 24	Sparsentan	37	26 (70.3)	6.72 (3.56)	(-0.29, 13.74)	3.31 (5.13)	(-6.80, 13.43)	0.519
		Irbesartan	36	26 (72.2)	3.41 (3.66)	(-3.81, 10.63)			
	Week 48	Sparsentan	37	33 (89.2)	4.20 (3.23)	(-2.18, 10.57)	-4.22 (4.95)	(-13.99, 5.54)	0.395
		Irbesartan	36	24 (66.7)	8.42 (3.73)	(1.06, 15.78)			
	Week 70	Sparsentan	37	31 (83.8)	10.91 (3.32)	(4.35, 17.47)	6.64 (4.99)	(-3.21, 16.49)	0.185
		Irbesartan	36	25 (69.4)	4.27 (3.70)	(-3.04, 11.58)			
	Week 94	Sparsentan	37	27 (73.0)	5.34 (3.52)	(-1.60, 12.27)	-1.18 (5.17)	(-11.37, 9.01)	0.819
		Irbesartan	36	24 (66.7)	6.52 (3.76)	(-0.90, 13.94)			
	Week 110	Sparsentan	37	26 (70.3)	8.33 (3.62)	(1.18, 15.47)	6.83 (5.37)	(-3.77, 17.42)	0.205
		Irbesartan	36	22 (61.1)	1.50 (3.93)	(-6.25, 9.25)			

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 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis					
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference			
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value	
eGFR High and UP Low	Week 24	Sparsentan	39	34 (87.2)	7.12 (2.97)	(1.27, 12.98)	4.33 (4.41)	(-4.35, 13.01)	0.326	
		Irbesartan	37	28 (75.7)	2.79 (3.25)	(-3.61, 9.19)				
	Week 48	Sparsentan	39	35 (89.7)	4.94 (2.93)	(-0.84, 10.73)	3.32 (4.39)	(-5.32, 11.96)		
		Irbesartan	37	28 (75.7)	1.63 (3.24)	(-4.76, 8.01)				
	Week 70	Sparsentan	39	30 (76.9)	7.42 (3.13)	(1.27, 13.58)	1.47 (4.49)	(-7.37, 10.32)		
		Irbesartan	37	29 (78.4)	5.95 (3.21)	(-0.37, 12.27)				
	Week 94	Sparsentan	39	32 (82.1)	4.74 (3.05)	(-1.27, 10.75)	3.05 (4.41)	(-5.64, 11.73)		
		Irbesartan	37	30 (81.1)	1.69 (3.16)	(-4.54, 7.92)				
	Week 110	Sparsentan	39	33 (84.6)	5.89 (3.02)	(-0.06, 11.83)	9.57 (4.44)	(0.83, 18.31)		0.032 *
		Irbesartan	37	29 (78.4)	-3.68 (3.22)	(-10.03, 2.66)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.607
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	98 (77.2)	4.56 (1.82)	(0.97, 8.14)	2.68 (2.73)	(-2.69, 8.04)	0.327
		Irbesartan	129	80 (62.0)	1.88 (2.02)	(-2.08, 5.84)			
	Week 48	Sparsentan	127	95 (74.8)	6.19 (1.83)	(2.60, 9.78)	2.74 (2.75)	(-2.66, 8.13)	0.319
		Irbesartan	129	74 (57.4)	3.45 (2.04)	(-0.56, 7.46)			
	Week 70	Sparsentan	127	100 (78.7)	4.81 (1.81)	(1.26, 8.37)	2.35 (2.72)	(-2.99, 7.69)	0.387
		Irbesartan	129	78 (60.5)	2.46 (2.02)	(-1.50, 6.42)			
	Week 94	Sparsentan	127	92 (72.4)	3.82 (1.86)	(0.16, 7.48)	5.38 (2.77)	(-0.05, 10.82)	0.052
		Irbesartan	129	77 (59.7)	-1.56 (2.04)	(-5.56, 2.44)			
	Week 110	Sparsentan	127	92 (72.4)	4.78 (1.88)	(1.08, 8.48)	2.89 (2.84)	(-2.69, 8.46)	0.310
		Irbesartan	129	72 (55.8)	1.89 (2.11)	(-2.26, 6.04)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	38 (77.6)	6.50 (2.73)	(1.12, 11.89)	-0.66 (4.10)	(-8.74, 7.42)	0.872
		Irbesartan	48	31 (64.6)	7.17 (3.03)	(1.20, 13.13)			
	Week 48	Sparsentan	49	44 (89.8)	4.91 (2.57)	(-0.16, 9.97)	0.78 (3.94)	(-6.98, 8.54)	0.843
		Irbesartan	48	32 (66.7)	4.12 (2.96)	(-1.70, 9.95)			
	Week 70	Sparsentan	49	38 (77.6)	6.86 (2.71)	(1.53, 12.20)	3.22 (4.03)	(-4.72, 11.16)	0.426
		Irbesartan	48	32 (66.7)	3.64 (2.97)	(-2.20, 9.48)			
	Week 94	Sparsentan	49	39 (79.6)	4.42 (2.71)	(-0.92, 9.76)	4.67 (4.00)	(-3.22, 12.55)	0.245
		Irbesartan	48	34 (70.8)	-0.25 (2.92)	(-6.00, 5.49)			
	Week 110	Sparsentan	49	36 (73.5)	2.49 (2.83)	(-3.09, 8.06)	5.20 (4.16)	(-2.99, 13.39)	0.212
		Irbesartan	48	32 (66.7)	-2.71 (3.00)	(-8.63, 3.20)			
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	21 (80.8)	12.19 (4.20)	(3.89, 20.49)	11.72 (6.02)	(-0.17, 23.60)	0.053
		Irbesartan	25	20 (80.0)	0.47 (4.30)	(-8.03, 8.97)			
	Week 48	Sparsentan	26	23 (88.5)	7.16 (4.04)	(-0.83, 15.14)	-2.04 (6.08)	(-14.04, 9.96)	0.738
		Irbesartan	25	18 (72.0)	9.19 (4.51)	(0.30, 18.09)			
	Week 70	Sparsentan	26	22 (84.6)	16.42 (4.12)	(8.28, 24.57)	7.58 (6.01)	(-4.29, 19.45)	0.209
		Irbesartan	25	20 (80.0)	8.84 (4.33)	(0.29, 17.40)			
	Week 94	Sparsentan	26	20 (76.9)	10.03 (4.30)	(1.54, 18.52)	-3.09 (6.19)	(-15.31, 9.13)	0.618
		Irbesartan	25	19 (76.0)	13.12 (4.43)	(4.38, 21.86)			
	Week 110	Sparsentan	26	21 (80.8)	13.73 (4.22)	(5.40, 22.06)	12.18 (6.31)	(-0.28, 24.65)	0.055
		Irbesartan	25	17 (68.0)	1.54 (4.67)	(-7.68, 10.77)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.595
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	64 (78.0)	6.72 (2.30)	(2.20, 11.24)	3.85 (3.51)	(-3.05, 10.76)	0.273
		Irbesartan	80	48 (60.0)	2.87 (2.64)	(-2.33, 8.06)			
	Week 48	Sparsentan	82	61 (74.4)	6.78 (2.32)	(2.22, 11.33)	1.94 (3.56)	(-5.06, 8.93)	0.586
		Irbesartan	80	43 (53.8)	4.84 (2.69)	(-0.45, 10.12)			
	Week 70	Sparsentan	82	65 (79.3)	6.73 (2.29)	(2.24, 11.23)	3.75 (3.51)	(-3.15, 10.64)	0.286
		Irbesartan	80	45 (56.3)	2.98 (2.65)	(-2.23, 8.20)			
	Week 94	Sparsentan	82	60 (73.2)	5.59 (2.35)	(0.97, 10.21)	8.99 (3.54)	(2.03, 15.94)	0.011 *
		Irbesartan	80	47 (58.8)	-3.40 (2.64)	(-8.58, 1.79)			
	Week 110	Sparsentan	82	56 (68.3)	7.58 (2.43)	(2.79, 12.36)	5.58 (3.67)	(-1.63, 12.79)	0.129
		Irbesartan	80	44 (55.0)	2.00 (2.74)	(-3.38, 7.38)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	34 (75.6)	0.47 (3.02)	(-5.47, 6.42)	0.07 (4.36)	(-8.52, 8.67)	0.987	
		Irbesartan	49	32 (65.3)	0.40 (3.12)	(-5.75, 6.55)				
	Week 48	Sparsentan	45	34 (75.6)	5.41 (2.98)	(-0.46, 11.27)	3.87 (4.34)	(-4.68, 12.42)	0.373	
		Irbesartan	49	31 (63.3)	1.54 (3.14)	(-4.64, 7.72)				
	Week 70	Sparsentan	45	35 (77.8)	1.42 (2.97)	(-4.44, 7.28)	-0.41 (4.32)	(-8.92, 8.10)	0.925	
		Irbesartan	49	33 (67.3)	1.83 (3.11)	(-4.30, 7.96)				
	Week 94	Sparsentan	45	32 (71.1)	0.68 (3.07)	(-5.37, 6.72)	-0.44 (4.46)	(-9.23, 8.34)	0.921	
		Irbesartan	49	30 (61.2)	1.12 (3.21)	(-5.20, 7.44)				
	Week 110	Sparsentan	45	36 (80.0)	0.22 (2.98)	(-5.66, 6.09)	-1.20 (4.49)	(-10.06, 7.65)	0.789	
		Irbesartan	49	28 (57.1)	1.42 (3.33)	(-5.15, 7.99)				
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	38 (77.6)	6.50 (2.73)	(1.12, 11.89)	-0.66 (4.10)	(-8.74, 7.42)	0.872	
		Irbesartan	48	31 (64.6)	7.17 (3.03)	(1.20, 13.13)				
	Week 48	Sparsentan	49	44 (89.8)	4.91 (2.57)	(-0.16, 9.97)	0.78 (3.94)	(-6.98, 8.54)	0.843	
		Irbesartan	48	32 (66.7)	4.12 (2.96)	(-1.70, 9.95)				
	Week 70	Sparsentan	49	38 (77.6)	6.86 (2.71)	(1.53, 12.20)	3.22 (4.03)	(-4.72, 11.16)	0.426	
		Irbesartan	48	32 (66.7)	3.64 (2.97)	(-2.20, 9.48)				
	Week 94	Sparsentan	49	39 (79.6)	4.42 (2.71)	(-0.92, 9.76)	4.67 (4.00)	(-3.22, 12.55)	0.245	
		Irbesartan	48	34 (70.8)	-0.25 (2.92)	(-6.00, 5.49)				
	Week 110	Sparsentan	49	36 (73.5)	2.49 (2.83)	(-3.09, 8.06)	5.20 (4.16)	(-2.99, 13.39)	0.212	
		Irbesartan	48	32 (66.7)	-2.71 (3.00)	(-8.63, 3.20)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	21 (80.8)	12.19 (4.20)	(3.89, 20.49)	11.72 (6.02)	(-0.17, 23.60)	0.053
		Irbesartan	25	20 (80.0)	0.47 (4.30)	(-8.03, 8.97)			
	Week 48	Sparsentan	26	23 (88.5)	7.16 (4.04)	(-0.83, 15.14)	-2.04 (6.08)	(-14.04, 9.96)	
		Irbesartan	25	18 (72.0)	9.19 (4.51)	(0.30, 18.09)			
	Week 70	Sparsentan	26	22 (84.6)	16.42 (4.12)	(8.28, 24.57)	7.58 (6.01)	(-4.29, 19.45)	
		Irbesartan	25	20 (80.0)	8.84 (4.33)	(0.29, 17.40)			
	Week 94	Sparsentan	26	20 (76.9)	10.03 (4.30)	(1.54, 18.52)	-3.09 (6.19)	(-15.31, 9.13)	
		Irbesartan	25	19 (76.0)	13.12 (4.43)	(4.38, 21.86)			
	Week 110	Sparsentan	26	21 (80.8)	13.73 (4.22)	(5.40, 22.06)	12.18 (6.31)	(-0.28, 24.65)	
		Irbesartan	25	17 (68.0)	1.54 (4.67)	(-7.68, 10.77)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.583
<= 1.75 g/day	Week 24	Sparsentan	98	80 (81.6)	6.83 (2.01)	(2.88, 10.78)	2.16 (3.15)	(-4.03, 8.35)	0.493
		Irbesartan	93	55 (59.1)	4.67 (2.41)	(-0.06, 9.40)			
	Week 48	Sparsentan	98	82 (83.7)	6.41 (1.98)	(2.52, 10.29)	2.38 (3.08)	(-3.67, 8.42)	0.441
		Irbesartan	93	57 (61.3)	4.03 (2.34)	(-0.58, 8.64)			
	Week 70	Sparsentan	98	75 (76.5)	6.27 (2.05)	(2.24, 10.29)	0.21 (3.09)	(-5.86, 6.27)	0.947
		Irbesartan	93	61 (65.6)	6.06 (2.30)	(1.55, 10.57)			
	Week 94	Sparsentan	98	76 (77.6)	5.29 (2.05)	(1.27, 9.32)	3.40 (3.07)	(-2.63, 9.43)	0.269
		Irbesartan	93	63 (67.7)	1.89 (2.27)	(-2.57, 6.36)			
	Week 110	Sparsentan	98	75 (76.5)	6.09 (2.08)	(2.01, 10.17)	4.51 (3.16)	(-1.70, 10.72)	0.154
		Irbesartan	93	57 (61.3)	1.58 (2.37)	(-3.08, 6.24)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024



Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 1.75 g/day	Week 24	Sparsentan	104	77 (74.0)	5.29 (2.04)	(1.27, 9.30)	3.87 (2.92)	(-1.87, 9.60)	0.186
		Irbesartan	109	76 (69.7)	1.42 (2.07)	(-2.65, 5.49)			
	Week 48	Sparsentan	104	80 (76.9)	5.47 (2.00)	(1.54, 9.39)	0.96 (2.94)	(-4.82, 6.74)	0.745
		Irbesartan	109	67 (61.5)	4.51 (2.15)	(0.29, 8.73)			
	Week 70	Sparsentan	104	85 (81.7)	7.45 (1.97)	(3.59, 11.32)	6.12 (2.92)	(0.39, 11.85)	0.036 *
		Irbesartan	109	69 (63.3)	1.33 (2.14)	(-2.88, 5.54)			
	Week 94	Sparsentan	104	75 (72.1)	4.30 (2.06)	(0.24, 8.35)	4.27 (3.01)	(-1.64, 10.18)	0.156
		Irbesartan	109	67 (61.5)	0.02 (2.18)	(-4.26, 4.31)			
	Week 110	Sparsentan	104	74 (71.2)	4.95 (2.10)	(0.82, 9.07)	5.49 (3.08)	(-0.56, 11.54)	0.075
		Irbesartan	109	64 (58.7)	-0.55 (2.24)	(-4.96, 3.86)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.128
Yes	Week 24	Sparsentan	90	66 (73.3)	7.21 (2.21)	(2.88, 11.55)	5.48 (3.39)	(-1.18, 12.14)	0.107
		Irbesartan	88	49 (55.7)	1.74 (2.56)	(-3.29, 6.76)			
	Week 48	Sparsentan	90	65 (72.2)	7.53 (2.20)	(3.21, 11.85)	5.06 (3.38)	(-1.58, 11.71)	0.135
		Irbesartan	88	47 (53.4)	2.47 (2.56)	(-2.56, 7.50)			
	Week 70	Sparsentan	90	68 (75.6)	6.56 (2.18)	(2.29, 10.84)	5.85 (3.35)	(-0.73, 12.43)	0.081
		Irbesartan	88	49 (55.7)	0.71 (2.53)	(-4.26, 5.69)			
	Week 94	Sparsentan	90	63 (70.0)	7.14 (2.24)	(2.75, 11.54)	8.87 (3.32)	(2.34, 15.40)	0.008 *
		Irbesartan	88	54 (61.4)	-1.73 (2.44)	(-6.53, 3.08)			
	Week 110	Sparsentan	90	63 (70.0)	5.26 (2.27)	(0.80, 9.72)	6.60 (3.40)	(-0.08, 13.27)	0.053
		Irbesartan	88	51 (58.0)	-1.34 (2.51)	(-6.27, 3.60)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
No	Week 24	Sparsentan	112	91 (81.3)	5.21 (1.88)	(1.52, 8.90)	1.72 (2.74)	(-3.66, 7.10)	0.530
		Irbesartan	114	82 (71.9)	3.49 (1.98)	(-0.40, 7.38)			
	Week 48	Sparsentan	112	97 (86.6)	4.79 (1.82)	(1.21, 8.38)	-0.63 (2.72)	(-5.97, 4.72)	0.818
		Irbesartan	114	77 (67.5)	5.42 (2.01)	(1.47, 9.37)			
	Week 70	Sparsentan	112	92 (82.1)	7.18 (1.87)	(3.51, 10.85)	1.85 (2.73)	(-3.52, 7.21)	0.500
		Irbesartan	114	81 (71.1)	5.34 (1.99)	(1.44, 9.24)			
	Week 94	Sparsentan	112	88 (78.6)	3.12 (1.91)	(-0.62, 6.86)	0.43 (2.81)	(-5.08, 5.94)	0.878
		Irbesartan	114	76 (66.7)	2.69 (2.05)	(-1.34, 6.71)			
	Week 110	Sparsentan	112	86 (76.8)	5.68 (1.94)	(1.87, 9.49)	3.87 (2.90)	(-1.81, 9.56)	0.182
		Irbesartan	114	70 (61.4)	1.81 (2.14)	(-2.39, 6.01)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.941
<= 5 years	Week 24	Sparsentan	113	88 (77.9)	8.97 (1.95)	(5.15, 12.79)	5.27 (2.77)	(-0.17, 10.70)	0.058
		Irbesartan	127	86 (67.7)	3.71 (1.96)	(-0.14, 7.55)			
	Week 48	Sparsentan	113	92 (81.4)	6.71 (1.90)	(2.98, 10.44)	1.03 (2.73)	(-4.32, 6.38)	0.706
		Irbesartan	127	85 (66.9)	5.68 (1.94)	(1.86, 9.50)			
	Week 70	Sparsentan	113	90 (79.6)	6.89 (1.92)	(3.13, 10.65)	3.00 (2.74)	(-2.38, 8.38)	0.274
		Irbesartan	127	86 (67.7)	3.89 (1.95)	(0.06, 7.72)			
	Week 94	Sparsentan	113	86 (76.1)	6.87 (1.95)	(3.03, 10.70)	4.20 (2.77)	(-1.25, 9.64)	0.130
		Irbesartan	127	86 (67.7)	2.67 (1.95)	(-1.17, 6.51)			
	Week 110	Sparsentan	113	88 (77.9)	6.50 (1.96)	(2.65, 10.34)	4.43 (2.83)	(-1.13, 9.98)	0.118
		Irbesartan	127	80 (63.0)	2.07 (2.03)	(-1.92, 6.06)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 5 years	Week 24	Sparsentan	89	69 (77.5)	2.12 (2.11)	(-2.03, 6.27)	0.52 (3.41)	(-6.18, 7.21)	0.879
		Irbesartan	75	45 (60.0)	1.60 (2.66)	(-3.62, 6.82)			
	Week 48	Sparsentan	89	70 (78.7)	4.72 (2.09)	(0.62, 8.83)	2.83 (3.47)	(-3.98, 9.65)	
		Irbesartan	75	39 (52.0)	1.89 (2.76)	(-3.53, 7.30)			
	Week 70	Sparsentan	89	70 (78.7)	6.74 (2.10)	(2.60, 10.87)	3.52 (3.39)	(-3.13, 10.18)	
		Irbesartan	75	44 (58.7)	3.21 (2.64)	(-1.98, 8.41)			
	Week 94	Sparsentan	89	65 (73.0)	1.85 (2.17)	(-2.41, 6.11)	4.10 (3.44)	(-2.67, 10.86)	
		Irbesartan	75	44 (58.7)	-2.25 (2.66)	(-7.48, 2.98)			
	Week 110	Sparsentan	89	61 (68.5)	4.00 (2.24)	(-0.41, 8.41)	6.23 (3.56)	(-0.77, 13.23)	
		Irbesartan	75	41 (54.7)	-2.24 (2.75)	(-7.64, 3.17)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.440
Yes	Week 24	Sparsentan	155	119 (76.8)	6.40 (1.62)	(3.23, 9.57)	2.45 (2.40)	(-2.26, 7.16)	0.308
		Irbesartan	161	100 (62.1)	3.95 (1.76)	(0.49, 7.41)			
	Week 48	Sparsentan	155	121 (78.1)	5.80 (1.59)	(2.67, 8.93)	1.29 (2.39)	(-3.41, 5.98)	0.591
		Irbesartan	161	95 (59.0)	4.51 (1.78)	(1.03, 8.00)			
	Week 70	Sparsentan	155	123 (79.4)	6.87 (1.59)	(3.75, 9.99)	3.06 (2.37)	(-1.60, 7.71)	0.198
		Irbesartan	161	100 (62.1)	3.81 (1.75)	(0.37, 7.25)			
	Week 94	Sparsentan	155	114 (73.5)	4.82 (1.64)	(1.61, 8.03)	5.01 (2.40)	(0.30, 9.73)	0.037 *
		Irbesartan	161	101 (62.7)	-0.19 (1.75)	(-3.63, 3.24)			
	Week 110	Sparsentan	155	112 (72.3)	5.53 (1.67)	(2.26, 8.80)	3.37 (2.48)	(-1.50, 8.24)	0.175
		Irbesartan	161	92 (57.1)	2.16 (1.83)	(-1.43, 5.75)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KBUC\_FSCM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in burden of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
No	Week 24	Sparsentan	47	38 (80.9)	4.63 (3.11)	(-1.51, 10.77)	5.13 (4.66)	(-4.05, 14.31)	0.272
		Irbesartan	41	31 (75.6)	-0.50 (3.44)	(-7.29, 6.29)			
	Week 48	Sparsentan	47	41 (87.2)	6.02 (3.02)	(0.06, 11.99)	2.33 (4.63)	(-6.79, 11.46)	0.615
		Irbesartan	41	29 (70.7)	3.69 (3.48)	(-3.17, 10.55)			
	Week 70	Sparsentan	47	37 (78.7)	6.97 (3.12)	(0.81, 13.13)	4.30 (4.69)	(-4.96, 13.56)	0.361
		Irbesartan	41	30 (73.2)	2.67 (3.48)	(-4.18, 9.53)			
	Week 94	Sparsentan	47	37 (78.7)	4.45 (3.16)	(-1.78, 10.68)	-0.01 (4.79)	(-9.46, 9.43)	0.998
		Irbesartan	41	29 (70.7)	4.46 (3.56)	(-2.56, 11.49)			
	Week 110	Sparsentan	47	37 (78.7)	5.09 (3.19)	(-1.21, 11.38)	9.60 (4.86)	(0.02, 19.18)	0.049 *
		Irbesartan	41	29 (70.7)	-4.51 (3.62)	(-11.64, 2.62)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Figure PF2KBUC\_FSGM: KDQOL: Change from baseline in burden of kidney disease by subgroup  
Full Analysis Set

Not done: No significant subgroup found.

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Reference table: PT2KBUC\_FSCM



Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.086
Male	Sparsentan	139	53 (38.1)	NE		1.711	(1.081, 2.706)	0.022 *
	Irbesartan	143	29 (20.3)	NE				
Female	Sparsentan	63	29 (46.0)	87.7	(48.1, NE)	0.860	(0.465, 1.593)	0.633
	Irbesartan	59	20 (33.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan							Interaction test: 0.921
<= 45 years	Sparsentan	96	40 (41.7)	114.6	(69.0, NE)	1.336	(0.812, 2.196)	0.254
	Irbesartan	99	28 (28.3)	NE				
> 45 years	Sparsentan	106	42 (39.6)	NE		1.312	(0.767, 2.244)	0.322
	Irbesartan	103	21 (20.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	5 (55.6) No events in 1 group	37.1	(22.7, NE)	NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	41 (40.2)	114.6	(72.0, NE)	1.369	(0.843, 2.224)	0.204
	Irbesartan	109	29 (26.6)	NE				
> 40 years	Sparsentan	91	36 (39.6)	NE		1.201	(0.679, 2.125)	0.529
	Irbesartan	88	20 (22.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.189
North America	Sparsentan	35	14 (40.0)	114.6	(49.1, NE)	0.679	(0.286, 1.611)	0.379
	Irbesartan	46	14 (30.4)	NE				
Europe	Sparsentan	98	27 (27.6)	NE		1.759	(0.981, 3.153)	0.058
	Irbesartan	115	22 (19.1)	NE				
Asia Pacific	Sparsentan	69	41 (59.4)	60.3	(48.1, 111.0)	1.588	(0.837, 3.010)	0.157
	Irbesartan	41	13 (31.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.691
< 27 kg/m**2	Sparsentan	83	39 (47.0)	93.0	(49.3, NE)	1.151	(0.676, 1.958)	0.604
	Irbesartan	94	26 (27.7)	NE				
≥ 27 kg/m**2	Sparsentan	119	43 (36.1)	NE		1.385	(0.829, 2.314)	0.214
	Irbesartan	107	23 (21.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.665
eGFR Low and UP High	Sparsentan	71	26 (36.6)	NE		1.757	(0.944, 3.271)	0.075
	Irbesartan	74	17 (23.0)	NE				
eGFR Low and UP Low	Sparsentan	55	24 (43.6)	114.1	(49.7, NE)	1.192	(0.597, 2.379)	0.619
	Irbesartan	55	14 (25.5)	NE				
eGFR High and UP High	Sparsentan	37	17 (45.9)	109.0	(50.6, NE)	1.078	(0.454, 2.559)	0.865
	Irbesartan	36	9 (25.0)	NE				
eGFR High and UP Low	Sparsentan	39	15 (38.5)	NE		1.366	(0.593, 3.146)	0.464
	Irbesartan	37	9 (24.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.676
< 60 mL/min/1.73 m**2	Sparsentan	127	48 (37.8)	NE		1.462	(0.920, 2.323)	0.108
	Irbesartan	129	30 (23.3)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	20 (40.8)	NE		1.429	(0.624, 3.271)	0.398
	Irbesartan	48	10 (20.8)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	14 (53.8)	91.8	(47.0, NE)	1.049	(0.426, 2.585)	0.917
	Irbesartan	25	9 (36.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.840
< 45 mL/min/1.73 m**2	Sparsentan	82	33 (40.2)	114.1	(52.0, NE)	1.430	(0.815, 2.508)	0.213
	Irbesartan	80	20 (25.0)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	15 (33.3)	NE		1.380	(0.584, 3.260)	0.463
	Irbesartan	49	10 (20.4)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	20 (40.8)	NE		1.429	(0.624, 3.271)	0.398
	Irbesartan	48	10 (20.8)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	14 (53.8)	91.8	(47.0, NE)	1.049	(0.426, 2.585)	0.917
	Irbesartan	25	9 (36.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.284
<= 1.75 g/day	Sparsentan	98	42 (42.9)	114.6	(52.0, NE)	1.807	(1.051, 3.109)	0.032 *
	Irbesartan	93	20 (21.5)	NE				
> 1.75 g/day	Sparsentan	104	40 (38.5)	NE		1.133	(0.693, 1.851)	0.619
	Irbesartan	109	29 (26.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.199
Yes	Sparsentan	90	35 (38.9)	NE		1.911	(1.045, 3.495)	0.035 *
	Irbesartan	88	16 (18.2)	NE				
No	Sparsentan	112	47 (42.0)	114.6	(87.7, NE)	1.119	(0.712, 1.761)	0.625
	Irbesartan	114	33 (28.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.154
<= 5 years	Sparsentan	113	50 (44.2)	114.1	(70.1, NE)	1.175	(0.763, 1.810)	0.464
	Irbesartan	127	38 (29.9)	NE				
> 5 years	Sparsentan	89	32 (36.0)	NE		2.192	(1.071, 4.487)	0.032 *
	Irbesartan	75	11 (14.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUIT\_FSTM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.551
Yes	Sparsentan	155	60 (38.7)	NE		1.259	(0.829, 1.911)	0.280
	Irbesartan	161	38 (23.6)	NE				
No	Sparsentan	47	22 (46.8)	114.6	(49.1, NE)	1.625	(0.778, 3.398)	0.197
	Irbesartan	41	11 (26.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Figure PF2KBUIT\_FSKM: KDQOL: Time to increase in burden of kidney disease by at least 15 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_   
An increase reflects an improvement of the status of the patient.  
Reference table: PT2KBUIT\_FSTM

Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.922
Male	Sparsentan	139	35 (25.2)	NE		0.685	(0.438, 1.072)	0.098
	Irbesartan	143	47 (32.9)	NE				
Female	Sparsentan	63	14 (22.2)	NE		0.717	(0.344, 1.494)	0.375
	Irbesartan	59	18 (30.5)	116.0	(95.6, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan							Interaction test: 0.991
<= 45 years	Sparsentan	96	22 (22.9)	NE		0.674	(0.383, 1.186)	0.171
	Irbesartan	99	29 (29.3)	NE				
> 45 years	Sparsentan	106	27 (25.5)	NE		0.701	(0.419, 1.174)	0.177
	Irbesartan	103	36 (35.0)	116.0	(108.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: 0.741
<= 18 years	Sparsentan	9	1 (11.1)	NE		35.739	(0.050, NE)	NE
	Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Sparsentan	102	24 (23.5)	NE		0.736	(0.425, 1.274)	0.274
	Irbesartan	109	31 (28.4)	NE				
> 40 years	Sparsentan	91	24 (26.4)	NE		0.595	(0.348, 1.015)	0.057
	Irbesartan	88	33 (37.5)	114.1	(94.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.980
North America	Sparsentan	35	9 (25.7)	NE		0.477	(0.200, 1.137)	0.095
	Irbesartan	46	18 (39.1)	NE				
Europe	Sparsentan	98	24 (24.5)	NE		0.636	(0.375, 1.077)	0.092
	Irbesartan	115	34 (29.6)	116.0	(113.1, NE)			
Asia Pacific	Sparsentan	69	16 (23.2)	NE		0.705	(0.326, 1.528)	0.376
	Irbesartan	41	13 (31.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.530
< 27 kg/m**2	Sparsentan	83	21 (25.3)	NE		0.747	(0.423, 1.319)	0.314
	Irbesartan	94	32 (34.0)	116.0	(108.1, NE)			
≥ 27 kg/m**2	Sparsentan	119	28 (23.5)	NE		0.642	(0.383, 1.077)	0.093
	Irbesartan	107	33 (30.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.985
eGFR Low and UP High	Sparsentan	71	19 (26.8)	NE		0.672	(0.369, 1.225)	0.194
	Irbesartan	74	26 (35.1)	114.1	(94.6, NE)			
eGFR Low and UP Low	Sparsentan	55	11 (20.0)	NE		0.729	(0.334, 1.591)	0.428
	Irbesartan	55	16 (29.1)	NE				
eGFR High and UP High	Sparsentan	37	8 (21.6)	NE		0.655	(0.245, 1.754)	0.400
	Irbesartan	36	9 (25.0)	NE				
eGFR High and UP Low	Sparsentan	39	11 (28.2)	NE		0.580	(0.259, 1.298)	0.185
	Irbesartan	37	14 (37.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							
							Interaction test:	0.947
< 60 mL/min/1.73 m**2	Sparsentan	127	31 (24.4)	NE		0.701	(0.438, 1.123)	0.140
	Irbesartan	129	43 (33.3)	116.0	(110.1, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	13 (26.5)	NE		0.591	(0.274, 1.275)	0.180
	Irbesartan	48	15 (31.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	5 (19.2)	NE		0.542	(0.168, 1.753)	0.307
	Irbesartan	25	7 (28.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: 0.379
< 45 mL/min/1.73 m**2	Sparsentan	82	15 (18.3)	NE		0.493	(0.259, 0.938)	0.031 *
	Irbesartan	80	28 (35.0)	116.0	(95.3, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	16 (35.6)	NE		1.100	(0.534, 2.266)	0.796
	Irbesartan	49	15 (30.6)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	13 (26.5)	NE		0.591	(0.274, 1.275)	0.180
	Irbesartan	48	15 (31.3)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	5 (19.2)	NE		0.542	(0.168, 1.753)	0.307
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 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.883
<= 1.75 g/day	Sparsentan	98	23 (23.5)	NE		0.701	(0.397, 1.238)	0.221
	Irbesartan	93	26 (28.0)	NE				
> 1.75 g/day	Sparsentan	104	26 (25.0)	NE		0.655	(0.394, 1.088)	0.102
	Irbesartan	109	39 (35.8)	114.3	(96.1, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.406
Yes	Sparsentan	90	20 (22.2)	NE		0.547	(0.307, 0.974)	0.041 *
	Irbesartan	88	30 (34.1)	116.0	(95.1, NE)			
No	Sparsentan	112	29 (25.9)	NE		0.736	(0.444, 1.219)	0.234
	Irbesartan	114	35 (30.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.840
<= 5 years	Sparsentan	113	27 (23.9)	NE		0.628	(0.384, 1.025)	0.063
	Irbesartan	127	42 (33.1)	116.0	(113.1, NE)			
> 5 years	Sparsentan	89	22 (24.7)	NE		0.732	(0.398, 1.345)	0.315
	Irbesartan	75	23 (30.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KBUDT\_FSTM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to decrease in burden of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.824
Yes	Sparsentan	155	37 (23.9)	NE		0.674	(0.437, 1.038)	0.073
	Irbesartan	161	49 (30.4)	NE				
No	Sparsentan	47	12 (25.5)	NE		0.573	(0.253, 1.299)	0.182
	Irbesartan	41	16 (39.0)	114.1	(95.3, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Figure PF2KBUDT\_FSKM: KDQOL: Time to decrease in burden of kidney disease by at least 15 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_ A decrease reflects a worsening of the status of the patient.  
Reference table: PT2KBUDT\_FSTM

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Sex														
Male	KDQOL: effect of kidney disease	Baseline	Sparsentan	139	129 (92.8)	90.41 (9.58)	50.0	84.38	93.75	96.88	100.0			
			Irbesartan	143	131 (91.6)	89.72 (12.96)	37.5	87.50	93.75	100.00	100.0			
		Week 24	Sparsentan	139	109 (78.4)	89.28 (14.44)	9.4	84.38	93.75	100.00	100.0			
			Irbesartan	143	93 (65.0)	88.88 (14.59)	28.1	84.38	96.88	100.00	100.0			
		Week 48	Sparsentan	139	112 (80.6)	90.79 (12.62)	0.0	87.50	93.75	100.00	100.0			
			Irbesartan	143	87 (60.8)	92.64 (10.03)	50.0	90.63	96.88	100.00	100.0			
		Week 70	Sparsentan	139	114 (82.0)	90.27 (11.92)	43.8	87.50	93.75	100.00	100.0			
			Irbesartan	143	92 (64.3)	90.83 (14.38)	15.6	87.50	96.88	100.00	100.0			
		Week 94	Sparsentan	139	105 (75.5)	90.71 (10.10)	56.3	84.38	93.75	100.00	100.0			
			Irbesartan	143	96 (67.1)	91.08 (12.00)	43.8	87.50	93.75	100.00	100.0			
		Week 110	Sparsentan	139	103 (74.1)	90.81 (10.22)	43.8	84.38	93.75	100.00	100.0			
			Irbesartan	143	89 (62.2)	90.59 (11.70)	46.9	84.38	96.88	100.00	100.0			
			KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	139	109 (78.4)	-0.52 (14.00)	-75.0	-3.13	0.00	6.25	25.0	0.06 [-0.21, 0.34]
				Irbesartan	143	93 (65.0)	-1.31 (11.10)	-53.1	-3.13	0.00	3.13	31.3		
		Week 48		Sparsentan	139	112 (80.6)	0.53 (13.17)	-100.0	-3.13	0.00	6.25	28.1	-0.05 [-0.33, 0.23]	
				Irbesartan	143	87 (60.8)	1.08 (8.88)	-21.9	-3.13	0.00	6.25	25.0		
		Week 70		Sparsentan	139	114 (82.0)	-0.16 (10.83)	-43.8	-3.13	0.00	6.25	21.9	-0.05 [-0.33, 0.22]	
				Irbesartan	143	92 (64.3)	0.48 (14.50)	-71.9	-3.13	0.00	6.25	40.6		
Week 94	Sparsentan	139	105 (75.5)	0.63 (9.50)	-40.6	-3.13	0.00	3.13	21.9	-0.09 [-0.36, 0.19]				
	Irbesartan	143	96 (67.1)	1.43 (9.14)	-21.9	-3.13	0.00	6.25	40.6					
Week 110	Sparsentan	139	103 (74.1)	0.64 (9.38)	-28.1	-3.13	0.00	6.25	21.9	-0.01 [-0.29, 0.27]				
	Irbesartan	143	89 (62.2)	0.74 (10.90)	-28.1	-3.13	0.00	6.25	43.8					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Female	KDQOL: effect of kidney disease	Baseline	Sparsentan	63	58 (92.1)	84.00 (18.07)	28.1	71.88	90.63	96.88	100.0	
		Week 24	Irbesartan	59	51 (86.4)	87.01 (12.88)	46.9	78.13	90.63	100.00	100.0	
			Sparsentan	63	48 (76.2)	91.67 (9.15)	59.4	87.50	93.75	100.00	100.0	
		Week 48	Irbesartan	59	38 (64.4)	89.31 (18.65)	0.0	87.50	93.75	100.00	100.0	
			Sparsentan	63	50 (79.4)	91.44 (9.55)	62.5	87.50	93.75	100.00	100.0	
		Week 70	Irbesartan	59	37 (62.7)	89.44 (12.74)	40.6	84.38	90.63	100.00	100.0	
			Sparsentan	63	46 (73.0)	89.20 (12.77)	50.0	84.38	92.19	100.00	100.0	
		Week 94	Irbesartan	59	38 (64.4)	91.28 (10.43)	56.3	87.50	93.75	100.00	100.0	
			Sparsentan	63	46 (73.0)	89.95 (12.13)	46.9	81.25	93.75	100.00	100.0	
		Week 110	Irbesartan	59	34 (57.6)	92.00 (9.55)	59.4	87.50	93.75	100.00	100.0	
			Sparsentan	63	46 (73.0)	90.15 (10.35)	56.3	87.50	92.19	96.88	100.0	
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	63	48 (76.2)	6.97 (14.20)	-15.6	-3.13	3.13	14.06	43.8
	Irbesartan			59	38 (64.4)	0.00 (18.65)	-100.0	0.00	0.00	6.25	25.0	
	Week 48		Sparsentan	63	50 (79.4)	6.38 (15.77)	-37.5	-3.13	3.13	9.38	56.3	0.35 [-0.08, 0.78]
			Irbesartan	59	37 (62.7)	1.52 (10.54)	-25.0	0.00	0.00	3.13	25.0	
	Week 70		Sparsentan	63	46 (73.0)	5.03 (15.32)	-31.3	-3.13	0.00	9.38	56.3	0.30 [-0.13, 0.73]
			Irbesartan	59	38 (64.4)	1.15 (9.46)	-31.3	-3.13	0.00	6.25	21.9	
	Week 94	Sparsentan	63	46 (73.0)	5.98 (14.26)	-18.8	0.00	3.13	9.38	56.3	0.23 [-0.22, 0.67]	
Irbesartan		59	34 (57.6)	3.22 (8.86)	-15.6	0.00	0.00	9.38	25.0			
Week 110	Sparsentan	63	46 (73.0)	4.35 (14.68)	-21.9	-6.25	3.13	6.25	46.9	0.19 [-0.26, 0.64]		
	Irbesartan	59	32 (54.2)	1.95 (9.04)	-25.0	-1.56	1.56	6.25	25.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Age													
<= 45 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	96	87 (90.6)	86.82 (14.88)	28.1	84.38	90.63	96.88	100.0		
			Irbesartan	99	92 (92.9)	86.58 (14.49)	37.5	79.69	90.63	96.88	100.0		
		Week 24	Sparsentan	96	71 (74.0)	88.25 (15.29)	9.4	81.25	93.75	100.00	100.0		
			Irbesartan	99	67 (67.7)	87.17 (18.45)	0.0	81.25	93.75	96.88	100.0		
		Week 48	Sparsentan	96	75 (78.1)	88.88 (15.03)	0.0	87.50	93.75	96.88	100.0		
			Irbesartan	99	62 (62.6)	90.42 (12.75)	40.6	84.38	96.88	100.00	100.0		
		Week 70	Sparsentan	96	72 (75.0)	88.89 (12.14)	43.8	84.38	93.75	96.88	100.0		
			Irbesartan	99	62 (62.6)	91.13 (11.87)	40.6	87.50	93.75	100.00	100.0		
		Week 94	Sparsentan	96	68 (70.8)	89.52 (11.45)	46.9	81.25	93.75	100.00	100.0		
			Irbesartan	99	64 (64.6)	89.65 (13.86)	43.8	87.50	93.75	100.00	100.0		
		Week 110	Sparsentan	96	69 (71.9)	90.58 (10.58)	43.8	87.50	93.75	100.00	100.0		
			Irbesartan	99	59 (59.6)	88.72 (12.35)	46.9	84.38	90.63	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	96	71 (74.0)	1.54 (17.04)	-75.0	-6.25	3.13	9.38	43.8	0.16 [-0.17, 0.50]
			Irbesartan	99	67 (67.7)	-1.12 (15.75)	-100.0	-3.13	0.00	6.25	31.3		
	Week 48		Sparsentan	96	75 (78.1)	2.21 (18.53)	-100.0	-3.13	3.13	6.25	56.3	0.07 [-0.26, 0.41]	
			Irbesartan	99	62 (62.6)	1.11 (9.64)	-21.9	-3.13	0.00	3.13	25.0		
	Week 70		Sparsentan	96	72 (75.0)	2.26 (13.94)	-31.3	-3.13	0.00	6.25	56.3	0.08 [-0.26, 0.42]	
			Irbesartan	99	62 (62.6)	1.31 (10.43)	-31.3	-3.13	0.00	6.25	21.9		
Week 94	Sparsentan	96	68 (70.8)	2.94 (14.10)	-40.6	-3.13	3.13	9.38	56.3	0.14 [-0.20, 0.48]			
	Irbesartan	99	64 (64.6)	1.27 (9.36)	-21.9	-4.69	0.00	6.25	25.0				
Week 110	Sparsentan	96	69 (71.9)	3.17 (13.53)	-28.1	-3.13	3.13	9.38	46.9	0.22 [-0.13, 0.57]			
	Irbesartan	99	59 (59.6)	0.58 (9.70)	-28.1	-6.25	0.00	6.25	25.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 45 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	106	100 (94.3)	89.81 (11.26)	50.0	84.38	93.75	100.00	100.0		
			Irbesartan	103	90 (87.4)	91.39 (10.72)	46.9	87.50	93.75	100.00	100.0		
		Week 24	Sparsentan	106	86 (81.1)	91.46 (10.79)	50.0	87.50	93.75	100.00	100.0		
			Irbesartan	103	64 (62.1)	90.92 (12.30)	34.4	85.94	96.88	100.00	100.0		
		Week 48	Sparsentan	106	87 (82.1)	92.82 (7.48)	75.0	87.50	93.75	100.00	100.0		
			Irbesartan	103	62 (60.2)	92.94 (8.73)	68.8	87.50	96.88	100.00	100.0		
		Week 70	Sparsentan	106	88 (83.0)	90.84 (12.13)	46.9	87.50	93.75	100.00	100.0		
			Irbesartan	103	68 (66.0)	90.81 (14.58)	15.6	87.50	96.88	100.00	100.0		
		Week 94	Sparsentan	106	83 (78.3)	91.27 (10.09)	56.3	87.50	93.75	100.00	100.0		
			Irbesartan	103	66 (64.1)	92.95 (8.10)	68.8	87.50	96.88	100.00	100.0		
		Week 110	Sparsentan	106	80 (75.5)	90.63 (9.99)	56.3	87.50	93.75	98.44	100.0		
			Irbesartan	103	62 (60.2)	92.39 (9.70)	62.5	87.50	96.88	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	106	86 (81.1)	1.96 (11.97)	-37.5	-3.13	0.00	6.25	31.3	0.23 [-0.09, 0.56]
				Irbesartan	103	64 (62.1)	-0.73 (11.17)	-53.1	-3.13	0.00	6.25	18.8	
	Week 48		Sparsentan	106	87 (82.1)	2.44 (9.16)	-21.9	-3.13	0.00	6.25	34.4	0.12 [-0.20, 0.45]	
			Irbesartan	103	62 (60.2)	1.31 (9.15)	-25.0	0.00	0.00	6.25	25.0		
	Week 70		Sparsentan	106	88 (83.0)	0.57 (11.14)	-43.8	-3.13	0.00	6.25	40.6	0.04 [-0.28, 0.35]	
			Irbesartan	103	68 (66.0)	0.09 (15.35)	-71.9	0.00	0.00	6.25	40.6		
	Week 94		Sparsentan	106	83 (78.3)	1.69 (8.61)	-18.8	-3.13	0.00	3.13	28.1	-0.09 [-0.42, 0.23]	
			Irbesartan	103	66 (64.1)	2.51 (8.79)	-18.8	0.00	0.00	6.25	40.6		
Week 110	Sparsentan	106	80 (75.5)	0.59 (9.01)	-21.9	-3.13	0.00	4.69	34.4	-0.09 [-0.42, 0.24]			
	Irbesartan	103	62 (60.2)	1.51 (11.11)	-25.0	-3.13	0.00	6.25	43.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age at IgAN diagnosis														
<= 18 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	9	9 (100.0)	76.74 (22.54)	28.1	71.88	84.38	90.63	96.9			
			Irbesartan	5	5 (100.0)	86.88 (23.01)	46.9	87.50	100.00	100.00	100.0			
		Week 24	Sparsentan	9	5 (55.6)	75.63 (37.91)	9.4	81.25	87.50	100.00	100.0			
			Irbesartan	5	5 (100.0)	89.38 (15.08)	62.5	93.75	96.88	96.88	96.9			
		Week 48	Sparsentan	9	7 (77.8)	88.39 (7.81)	75.0	81.25	90.63	93.75	96.9			
			Irbesartan	5	3 (60.0)	97.92 (1.80)	96.9	96.88	96.88	100.00	100.0			
		Week 70	Sparsentan	9	7 (77.8)	87.95 (9.95)	75.0	78.13	84.38	96.88	100.0			
			Irbesartan	5	4 (80.0)	79.69 (27.30)	40.6	60.94	89.06	98.44	100.0			
		Week 94	Sparsentan	9	5 (55.6)	92.50 (7.19)	84.4	87.50	90.63	100.00	100.0			
			Irbesartan	5	4 (80.0)	82.81 (24.27)	46.9	68.75	92.19	96.88	100.0			
		Week 110	Sparsentan	9	4 (44.4)	92.97 (8.22)	81.3	87.50	95.31	98.44	100.0			
			Irbesartan	5	2 (40.0)	70.31 (33.15)	46.9	46.88	70.31	93.75	93.8			
			KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	9	5 (55.6)	-8.75 (38.11)	-75.0	-6.25	6.25	15.63	15.6	-0.41 [-1.66, 0.84]
		Irbesartan			5	5 (100.0)	2.50 (8.39)	-3.1	-3.13	-3.13	6.25	15.6		
		Week 48		Sparsentan	9	7 (77.8)	9.38 (17.86)	-6.3	0.00	3.13	15.63	46.9	0.46 [-0.91, 1.83]	
				Irbesartan	5	3 (60.0)	2.08 (6.51)	-3.1	-3.13	0.00	9.38	9.4		
		Week 70		Sparsentan	9	7 (77.8)	8.93 (22.49)	-12.5	-6.25	6.25	12.50	56.3	0.84 [-0.44, 2.12]	
				Irbesartan	5	4 (80.0)	-7.03 (8.22)	-18.8	-12.50	-4.69	-1.56	0.0		
Week 94	Sparsentan	9		5 (55.6)	6.88 (7.46)	-3.1	3.13	6.25	12.50	15.6	1.68 [0.15, 3.21]			
	Irbesartan	5		4 (80.0)	-3.91 (4.69)	-9.4	-7.81	-3.13	0.00	0.0				
Week 110	Sparsentan	9		4 (44.4)	5.47 (12.07)	-6.3	-3.13	3.13	14.06	21.9	0.80 [-0.95, 2.56]			
	Irbesartan	5		2 (40.0)	-3.13 (4.42)	-6.3	-6.25	-3.13	0.00	0.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 18 to 40 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	102	94 (92.2)	89.10 (12.71)	34.4	87.50	93.75	96.88	100.0		
			Irbesartan	109	99 (90.8)	87.31 (13.87)	37.5	84.38	90.63	96.88	100.0		
		Week 24	Sparsentan	102	79 (77.5)	90.31 (11.60)	56.3	87.50	93.75	100.00	100.0		
			Irbesartan	109	71 (65.1)	87.81 (18.38)	0.0	84.38	93.75	100.00	100.0		
		Week 48	Sparsentan	102	82 (80.4)	89.94 (14.56)	0.0	87.50	93.75	100.00	100.0		
			Irbesartan	109	67 (61.5)	90.44 (12.49)	40.6	84.38	96.88	100.00	100.0		
		Week 70	Sparsentan	102	78 (76.5)	89.78 (12.69)	43.8	87.50	93.75	100.00	100.0		
			Irbesartan	109	68 (62.4)	91.08 (11.98)	37.5	87.50	93.75	100.00	100.0		
		Week 94	Sparsentan	102	73 (71.6)	90.20 (11.34)	46.9	84.38	93.75	100.00	100.0		
			Irbesartan	109	69 (63.3)	90.35 (12.74)	43.8	87.50	93.75	100.00	100.0		
		Week 110	Sparsentan	102	72 (70.6)	90.89 (10.66)	43.8	87.50	93.75	100.00	100.0		
			Irbesartan	109	65 (59.6)	90.14 (11.05)	59.4	84.38	93.75	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	102	79 (77.5)	2.14 (13.75)	-37.5	-6.25	0.00	9.38	43.8	0.25 [-0.07, 0.58]
				Irbesartan	109	71 (65.1)	-1.58 (15.49)	-100.0	-3.13	0.00	3.13	31.3	
	Week 48		Sparsentan	102	82 (80.4)	1.30 (17.20)	-100.0	-3.13	1.56	6.25	56.3	0.03 [-0.30, 0.35]	
			Irbesartan	109	67 (61.5)	0.93 (9.28)	-21.9	-3.13	0.00	3.13	25.0		
	Week 70		Sparsentan	102	78 (76.5)	1.24 (12.85)	-43.8	-3.13	0.00	6.25	34.4	0.01 [-0.31, 0.34]	
			Irbesartan	109	68 (62.4)	1.10 (13.01)	-62.5	-3.13	0.00	6.25	37.5		
	Week 94		Sparsentan	102	73 (71.6)	2.61 (13.67)	-40.6	-3.13	3.13	9.38	56.3	0.06 [-0.27, 0.39]	
			Irbesartan	109	69 (63.3)	1.86 (9.92)	-21.9	-3.13	0.00	6.25	40.6		
Week 110	Sparsentan	102	72 (70.6)	2.99 (13.09)	-28.1	-3.13	3.13	7.81	46.9	0.15 [-0.19, 0.48]			
	Irbesartan	109	65 (59.6)	1.25 (10.29)	-28.1	-3.13	0.00	6.25	40.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	91	84 (92.3)	88.91 (11.83)	50.0	84.38	93.75	96.88	100.0	
			Irbesartan	88	78 (88.6)	91.19 (10.66)	46.9	84.38	93.75	100.00	100.0	
		Week 24	Sparsentan	91	73 (80.2)	90.67 (11.38)	50.0	87.50	93.75	100.00	100.0	
			Irbesartan	88	55 (62.5)	90.51 (11.91)	34.4	84.38	93.75	100.00	100.0	
		Week 48	Sparsentan	91	73 (80.2)	92.42 (7.76)	75.0	87.50	93.75	100.00	100.0	
			Irbesartan	88	54 (61.4)	92.88 (8.87)	68.8	84.38	96.88	100.00	100.0	
		Week 70	Sparsentan	91	75 (82.4)	90.33 (11.85)	46.9	87.50	93.75	100.00	100.0	
			Irbesartan	88	58 (65.9)	91.59 (13.54)	15.6	87.50	96.88	100.00	100.0	
		Week 94	Sparsentan	91	73 (80.2)	90.63 (10.38)	56.3	87.50	93.75	96.88	100.0	
			Irbesartan	88	57 (64.8)	93.09 (7.78)	68.8	87.50	93.75	100.00	100.0	
		Week 110	Sparsentan	91	73 (80.2)	90.20 (10.00)	56.3	87.50	93.75	96.88	100.0	
			Irbesartan	88	54 (61.4)	91.90 (9.85)	62.5	84.38	96.88	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	91	73 (80.2)	2.10 (12.58)	-37.5	-3.13	0.00	6.25	31.3	0.21 [-0.14, 0.56]
			Irbesartan	88	55 (62.5)	-0.40 (11.44)	-53.1	-3.13	0.00	6.25	25.0	
		Week 48	Sparsentan	91	73 (80.2)	2.83 (9.35)	-21.9	0.00	0.00	6.25	34.4	0.14 [-0.21, 0.49]
			Irbesartan	88	54 (61.4)	1.50 (9.71)	-25.0	0.00	0.00	6.25	25.0	
		Week 70	Sparsentan	91	75 (82.4)	0.71 (10.72)	-28.1	-3.13	0.00	6.25	40.6	0.00 [-0.34, 0.34]
			Irbesartan	88	58 (65.9)	0.70 (13.68)	-71.9	0.00	0.00	6.25	40.6	
		Week 94	Sparsentan	91	73 (80.2)	1.58 (8.84)	-18.8	-3.13	0.00	3.13	28.1	-0.09 [-0.44, 0.26]
			Irbesartan	88	57 (64.8)	2.36 (8.13)	-18.8	0.00	0.00	6.25	37.5	
Week 110	Sparsentan	91	73 (80.2)	0.39 (9.29)	-21.9	-3.13	0.00	3.13	34.4	-0.06 [-0.41, 0.29]		
	Irbesartan	88	54 (61.4)	0.98 (10.81)	-25.0	-3.13	0.00	6.25	43.8			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Geographic region													
North America	KDQOL: effect of kidney disease	Baseline	Sparsentan	35	31 (88.6)	88.41 (15.21)	28.1	84.38	93.75	96.88	100.0		
			Irbesartan	46	43 (93.5)	91.28 (10.84)	46.9	87.50	96.88	100.00	100.0		
		Week 24	Sparsentan	35	23 (65.7)	87.23 (20.14)	9.4	87.50	93.75	96.88	100.0		
			Irbesartan	46	35 (76.1)	89.91 (17.71)	0.0	87.50	96.88	100.00	100.0		
		Week 48	Sparsentan	35	25 (71.4)	91.88 (8.70)	75.0	90.63	93.75	100.00	100.0		
			Irbesartan	46	32 (69.6)	92.09 (12.90)	50.0	89.06	98.44	100.00	100.0		
		Week 70	Sparsentan	35	22 (62.9)	90.77 (9.42)	68.8	87.50	92.19	96.88	100.0		
			Irbesartan	46	30 (65.2)	93.75 (10.98)	56.3	93.75	100.00	100.00	100.0		
		Week 94	Sparsentan	35	23 (65.7)	89.81 (13.23)	46.9	81.25	96.88	100.00	100.0		
			Irbesartan	46	30 (65.2)	92.92 (9.91)	59.4	87.50	96.88	100.00	100.0		
		Week 110	Sparsentan	35	21 (60.0)	91.37 (10.50)	62.5	87.50	93.75	100.00	100.0		
			Irbesartan	46	31 (67.4)	93.04 (9.87)	59.4	90.63	96.88	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	35	23 (65.7)	-1.90 (22.17)	-75.0	-6.25	0.00	6.25	43.8	0.03 [-0.50, 0.56]
				Irbesartan	46	35 (76.1)	-2.50 (18.53)	-100.0	-3.13	0.00	6.25	15.6	
	Week 48		Sparsentan	35	25 (71.4)	4.88 (11.45)	-15.6	0.00	3.13	6.25	46.9	0.33 [-0.20, 0.86]	
			Irbesartan	46	32 (69.6)	1.56 (8.84)	-21.9	0.00	0.00	4.69	25.0		
	Week 70		Sparsentan	35	22 (62.9)	2.98 (14.61)	-18.8	-3.13	0.00	6.25	56.3	0.29 [-0.27, 0.84]	
			Irbesartan	46	30 (65.2)	-0.21 (7.70)	-31.3	0.00	0.00	3.13	12.5		
Week 94	Sparsentan	35	23 (65.7)	0.82 (9.94)	-21.9	-6.25	0.00	6.25	18.8	-0.08 [-0.62, 0.47]			
	Irbesartan	46	30 (65.2)	1.46 (6.90)	-15.6	0.00	0.00	3.13	18.8				
Week 110	Sparsentan	35	21 (60.0)	1.34 (7.43)	-18.8	0.00	3.13	6.25	15.6	-0.07 [-0.62, 0.48]			
	Irbesartan	46	31 (67.4)	1.81 (6.39)	-12.5	0.00	0.00	6.25	12.5				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Europe	KDQOL: effect of kidney disease	Baseline	Sparsentan	98	89 (90.8)	89.89 (11.68)	43.8	84.38	93.75	96.88	100.0		
			Irbesartan	115	98 (85.2)	87.15 (13.87)	43.8	78.13	90.63	96.88	100.0		
		Week 24	Sparsentan	98	73 (74.5)	92.12 (9.29)	56.3	87.50	93.75	100.00	100.0		
			Irbesartan	115	62 (53.9)	88.36 (14.60)	34.4	84.38	93.75	96.88	100.0		
		Week 48	Sparsentan	98	76 (77.6)	93.01 (8.11)	56.3	90.63	93.75	100.00	100.0		
			Irbesartan	115	59 (51.3)	91.58 (9.56)	65.6	84.38	96.88	100.00	100.0		
		Week 70	Sparsentan	98	75 (76.5)	91.17 (10.48)	43.8	87.50	93.75	100.00	100.0		
			Irbesartan	115	69 (60.0)	88.68 (15.89)	15.6	84.38	93.75	100.00	100.0		
		Week 94	Sparsentan	98	68 (69.4)	91.45 (9.73)	56.3	85.94	93.75	100.00	100.0		
			Irbesartan	115	70 (60.9)	89.46 (12.98)	43.8	87.50	93.75	96.88	100.0		
		Week 110	Sparsentan	98	67 (68.4)	92.26 (9.27)	43.8	90.63	93.75	100.00	100.0		
			Irbesartan	115	61 (53.0)	88.93 (12.56)	46.9	84.38	90.63	100.00	100.0		
			KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	98	73 (74.5)	2.65 (11.86)	-25.0	-3.13	0.00	6.25	31.3
		Irbesartan			115	62 (53.9)	0.05 (13.13)	-53.1	-3.13	0.00	6.25	31.3	
	Week 48	Sparsentan		98	76 (77.6)	3.13 (12.47)	-40.6	-3.13	1.56	6.25	56.3	0.09 [-0.25, 0.43]	
		Irbesartan		115	59 (51.3)	2.12 (9.82)	-21.9	-3.13	0.00	6.25	25.0		
	Week 70	Sparsentan		98	75 (76.5)	0.79 (9.98)	-28.1	-3.13	0.00	6.25	40.6	-0.04 [-0.37, 0.28]	
		Irbesartan		115	69 (60.0)	1.40 (16.78)	-71.9	-3.13	0.00	6.25	40.6		
	Week 94	Sparsentan		98	68 (69.4)	2.25 (11.23)	-15.6	-3.13	0.00	4.69	56.3	-0.06 [-0.39, 0.28]	
		Irbesartan	115	70 (60.9)	2.86 (10.69)	-21.9	-3.13	0.00	6.25	40.6			
Week 110	Sparsentan	98	67 (68.4)	2.01 (11.48)	-28.1	-3.13	0.00	6.25	46.9	-0.01 [-0.36, 0.33]			
	Irbesartan	115	61 (53.0)	2.15 (12.55)	-25.0	-3.13	0.00	6.25	43.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Asia Pacific	KDQOL: effect of kidney disease	Baseline	Sparsentan	69	67 (97.1)	86.47 (13.82)	34.4	81.25	90.63	96.88	100.0	
			Irbesartan	41	41 (100.0)	90.85 (12.35)	37.5	87.50	93.75	100.00	100.0	
		Week 24	Sparsentan	69	61 (88.4)	88.52 (13.46)	50.0	84.38	93.75	100.00	100.0	
			Irbesartan	41	34 (82.9)	89.25 (16.25)	28.1	84.38	95.31	100.00	100.0	
		Week 48	Sparsentan	69	61 (88.4)	88.11 (15.58)	0.0	87.50	90.63	96.88	100.0	
			Irbesartan	41	33 (80.5)	91.48 (11.59)	40.6	90.63	93.75	100.00	100.0	
		Week 70	Sparsentan	69	63 (91.3)	88.24 (14.55)	46.9	81.25	93.75	100.00	100.0	
			Irbesartan	41	31 (75.6)	93.35 (6.83)	75.0	87.50	96.88	100.00	100.0	
		Week 94	Sparsentan	69	60 (87.0)	89.64 (10.85)	56.3	84.38	93.75	96.88	100.0	
			Irbesartan	41	30 (73.2)	94.06 (7.67)	68.8	90.63	96.88	100.00	100.0	
		Week 110	Sparsentan	69	61 (88.4)	88.52 (10.93)	56.3	81.25	90.63	96.88	100.0	
			Irbesartan	41	29 (70.7)	91.49 (8.91)	68.8	84.38	93.75	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	69	61 (88.4)	2.10 (13.65)	-37.5	-3.13	3.13	9.38	34.4	0.27 [-0.15, 0.69]
			Irbesartan	41	34 (82.9)	-1.10 (7.61)	-15.6	-6.25	0.00	3.13	12.5	
		Week 48	Sparsentan	69	61 (88.4)	0.31 (17.01)	-100.0	-3.13	0.00	6.25	34.4	0.07 [-0.35, 0.50]
			Irbesartan	41	33 (80.5)	-0.76 (8.98)	-25.0	-3.13	0.00	3.13	25.0	
		Week 70	Sparsentan	69	63 (91.3)	1.39 (14.35)	-43.8	-3.13	0.00	9.38	34.4	0.12 [-0.31, 0.55]
			Irbesartan	41	31 (75.6)	-0.10 (7.10)	-21.9	-3.13	0.00	3.13	15.6	
		Week 94	Sparsentan	69	60 (87.0)	2.81 (12.19)	-40.6	-1.56	3.13	9.38	34.4	0.26 [-0.18, 0.69]
			Irbesartan	41	30 (73.2)	0.10 (6.33)	-18.8	-3.13	0.00	3.13	18.8	
Week 110	Sparsentan	69	61 (88.4)	1.69 (12.44)	-21.9	-6.25	0.00	6.25	37.5	0.33 [-0.11, 0.78]		
	Irbesartan	41	29 (70.7)	-2.05 (8.41)	-28.1	-6.25	0.00	0.00	15.6			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline BMI														
< 27 kg/m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	83	75 (90.4)	85.83 (13.40)	28.1	78.13	90.63	96.88	100.0			
			Irbesartan	94	86 (91.5)	88.55 (11.30)	46.9	84.38	90.63	100.00	100.0			
		Week 24	Sparsentan	83	66 (79.5)	88.83 (15.61)	9.4	87.50	93.75	100.00	100.0			
			Irbesartan	94	66 (70.2)	87.55 (16.99)	0.0	84.38	93.75	96.88	100.0			
		Week 48	Sparsentan	83	66 (79.5)	89.11 (10.63)	50.0	84.38	92.19	96.88	100.0			
			Irbesartan	94	60 (63.8)	89.74 (12.26)	40.6	84.38	93.75	96.88	100.0			
		Week 70	Sparsentan	83	64 (77.1)	88.62 (13.99)	43.8	84.38	93.75	100.00	100.0			
			Irbesartan	94	59 (62.8)	89.51 (13.28)	37.5	84.38	93.75	100.00	100.0			
		Week 94	Sparsentan	83	63 (75.9)	90.08 (10.52)	46.9	84.38	93.75	100.00	100.0			
			Irbesartan	94	62 (66.0)	90.47 (11.74)	43.8	87.50	93.75	100.00	100.0			
		Week 110	Sparsentan	83	63 (75.9)	88.89 (11.77)	43.8	81.25	93.75	96.88	100.0			
			Irbesartan	94	57 (60.6)	89.91 (11.63)	46.9	84.38	93.75	100.00	100.0			
			KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	83	66 (79.5)	2.23 (16.92)	-75.0	-3.13	3.13	9.38	43.8	0.16 [-0.18, 0.51]
		Irbesartan			94	66 (70.2)	-0.43 (15.53)	-100.0	-3.13	0.00	6.25	31.3		
		Week 48		Sparsentan	83	66 (79.5)	3.17 (12.39)	-40.6	-3.13	1.56	6.25	46.9	0.16 [-0.19, 0.51]	
				Irbesartan	94	60 (63.8)	1.41 (9.00)	-21.9	-1.56	0.00	6.25	25.0		
		Week 70		Sparsentan	83	64 (77.1)	2.34 (14.12)	-43.8	-3.13	3.13	9.38	56.3	0.15 [-0.21, 0.50]	
				Irbesartan	94	59 (62.8)	0.42 (12.08)	-62.5	-3.13	0.00	6.25	18.8		
Week 94	Sparsentan	83		63 (75.9)	3.47 (9.48)	-15.6	-3.13	3.13	9.38	34.4	0.22 [-0.13, 0.57]			
	Irbesartan	94		62 (66.0)	1.51 (8.37)	-18.8	-3.13	0.00	6.25	18.8				
Week 110	Sparsentan	83		63 (75.9)	1.54 (11.20)	-28.1	-3.13	0.00	6.25	31.3	0.05 [-0.30, 0.41]			
	Irbesartan	94		57 (60.6)	0.99 (8.96)	-25.0	-3.13	0.00	6.25	18.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 27 kg/m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	119	112 (94.1)	90.15 (12.69)	34.4	87.50	93.75	100.00	100.0		
			Irbesartan	107	95 (88.8)	89.51 (14.29)	37.5	87.50	96.88	100.00	100.0		
		Week 24	Sparsentan	119	91 (76.5)	90.87 (10.88)	56.3	84.38	93.75	100.00	100.0		
			Irbesartan	107	65 (60.7)	90.48 (14.47)	28.1	87.50	96.88	100.00	100.0		
		Week 48	Sparsentan	119	96 (80.7)	92.29 (12.32)	0.0	90.63	93.75	100.00	100.0		
			Irbesartan	107	64 (59.8)	93.51 (9.31)	59.4	90.63	98.44	100.00	100.0		
		Week 70	Sparsentan	119	96 (80.7)	90.85 (10.71)	46.9	87.50	93.75	96.88	100.0		
			Irbesartan	107	71 (66.4)	92.17 (13.30)	15.6	87.50	96.88	100.00	100.0		
		Week 94	Sparsentan	119	88 (73.9)	90.77 (10.92)	56.3	84.38	93.75	100.00	100.0		
			Irbesartan	107	67 (62.6)	92.72 (9.87)	53.1	87.50	96.88	100.00	100.0		
		Week 110	Sparsentan	119	86 (72.3)	91.86 (8.80)	56.3	87.50	93.75	100.00	100.0		
			Irbesartan	107	63 (58.9)	91.52 (10.61)	59.4	87.50	96.88	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	119	91 (76.5)	1.44 (12.42)	-37.5	-3.13	0.00	6.25	31.3	0.24 [-0.08, 0.56]
			Irbesartan	107	65 (60.7)	-1.44 (11.55)	-53.1	-3.13	0.00	3.13	25.0		
	Week 48		Sparsentan	119	96 (80.7)	1.76 (15.42)	-100.0	-3.13	1.56	6.25	56.3	0.05 [-0.26, 0.37]	
			Irbesartan	107	64 (59.8)	1.03 (9.76)	-25.0	-3.13	0.00	3.13	25.0		
	Week 70		Sparsentan	119	96 (80.7)	0.65 (11.25)	-31.3	-3.13	0.00	6.25	40.6	-0.02 [-0.33, 0.29]	
			Irbesartan	107	71 (66.4)	0.88 (14.14)	-71.9	-3.13	0.00	3.13	40.6		
	Week 94		Sparsentan	119	88 (73.9)	1.38 (12.56)	-40.6	-3.13	0.00	6.25	56.3	-0.11 [-0.43, 0.21]	
			Irbesartan	107	67 (62.6)	2.61 (9.31)	-15.6	0.00	0.00	3.13	40.6		
Week 110	Sparsentan	119	86 (72.3)	1.96 (11.54)	-21.9	-3.13	0.00	6.25	46.9	0.07 [-0.25, 0.40]			
	Irbesartan	107	63 (58.9)	1.14 (11.72)	-28.1	-3.13	0.00	6.25	43.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Randomization strata													
eGFR Low and UP High	KDQOL: effect of kidney disease	Baseline	Sparsentan	71	63 (88.7)	88.94 (11.54)	50.0	84.38	93.75	96.88	100.0		
			Irbesartan	74	63 (85.1)	85.66 (15.38)	43.8	81.25	90.63	96.88	100.0		
		Week 24	Sparsentan	71	54 (76.1)	89.24 (12.17)	56.3	84.38	93.75	96.88	100.0		
			Irbesartan	74	40 (54.1)	85.39 (15.23)	40.6	79.69	87.50	96.88	100.0		
		Week 48	Sparsentan	71	51 (71.8)	90.75 (10.04)	50.0	87.50	93.75	100.00	100.0		
			Irbesartan	74	37 (50.0)	88.60 (14.27)	40.6	81.25	93.75	100.00	100.0		
		Week 70	Sparsentan	71	57 (80.3)	88.43 (14.25)	43.8	84.38	93.75	96.88	100.0		
			Irbesartan	74	37 (50.0)	88.43 (15.32)	37.5	87.50	93.75	96.88	100.0		
		Week 94	Sparsentan	71	51 (71.8)	89.71 (11.22)	56.3	84.38	93.75	100.00	100.0		
			Irbesartan	74	37 (50.0)	87.75 (13.73)	46.9	84.38	90.63	96.88	100.0		
		Week 110	Sparsentan	71	53 (74.6)	89.62 (11.58)	43.8	81.25	93.75	100.00	100.0		
			Irbesartan	74	36 (48.6)	87.93 (14.45)	46.9	82.81	95.31	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	71	54 (76.1)	0.58 (10.27)	-37.5	-3.13	3.13	6.25	18.8	0.30 [-0.11, 0.71]
			Irbesartan	74	40 (54.1)	-2.42 (9.69)	-25.0	-7.81	0.00	3.13	15.6		
	Week 48		Sparsentan	71	51 (71.8)	2.02 (7.68)	-15.6	-3.13	3.13	6.25	28.1	0.16 [-0.26, 0.59]	
			Irbesartan	74	37 (50.0)	0.68 (9.06)	-21.9	-6.25	0.00	6.25	21.9		
	Week 70		Sparsentan	71	57 (80.3)	-0.66 (11.08)	-43.8	-3.13	0.00	6.25	21.9	-0.10 [-0.51, 0.32]	
			Irbesartan	74	37 (50.0)	0.59 (15.60)	-62.5	-3.13	0.00	6.25	40.6		
Week 94	Sparsentan	71	51 (71.8)	0.86 (8.03)	-21.9	-3.13	0.00	3.13	18.8	-0.13 [-0.56, 0.29]			
	Irbesartan	74	37 (50.0)	2.03 (9.61)	-21.9	0.00	0.00	6.25	37.5				
Week 110	Sparsentan	71	53 (74.6)	0.59 (8.52)	-28.1	-3.13	0.00	6.25	18.8	-0.05 [-0.47, 0.38]			
	Irbesartan	74	36 (48.6)	1.04 (11.43)	-18.8	-6.25	0.00	6.25	43.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR Low and UP Low	KDQOL: effect of kidney disease	Baseline	Sparsentan	55	52 (94.5)	86.36 (13.54)	43.8	78.13	90.63	95.31	100.0		
			Irbesartan	55	50 (90.9)	91.56 (8.84)	71.9	87.50	93.75	100.00	100.0		
		Week 24	Sparsentan	55	43 (78.2)	88.52 (13.11)	50.0	81.25	93.75	100.00	100.0		
			Irbesartan	55	37 (67.3)	94.26 (10.06)	50.0	93.75	96.88	100.00	100.0		
		Week 48	Sparsentan	55	43 (78.2)	89.53 (16.03)	0.0	87.50	93.75	100.00	100.0		
			Irbesartan	55	35 (63.6)	92.95 (8.10)	75.0	87.50	96.88	100.00	100.0		
		Week 70	Sparsentan	55	42 (76.4)	89.06 (13.37)	50.0	84.38	93.75	100.00	100.0		
			Irbesartan	55	39 (70.9)	91.11 (10.84)	56.3	87.50	96.88	100.00	100.0		
		Week 94	Sparsentan	55	41 (74.5)	90.17 (10.59)	59.4	84.38	93.75	100.00	100.0		
			Irbesartan	55	39 (70.9)	93.11 (8.30)	71.9	90.63	96.88	100.00	100.0		
		Week 110	Sparsentan	55	37 (67.3)	90.12 (10.66)	56.3	84.38	93.75	96.88	100.0		
			Irbesartan	55	34 (61.8)	91.45 (9.01)	71.9	87.50	93.75	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	55	43 (78.2)	3.27 (14.83)	-37.5	-3.13	3.13	12.50	31.3	0.12 [-0.32, 0.56]
				Irbesartan	55	37 (67.3)	1.77 (9.39)	-37.5	0.00	0.00	6.25	18.8	
	Week 48		Sparsentan	55	43 (78.2)	1.89 (20.47)	-100.0	0.00	0.00	6.25	56.3	0.03 [-0.42, 0.47]	
			Irbesartan	55	35 (63.6)	1.43 (9.14)	-25.0	0.00	0.00	6.25	25.0		
	Week 70		Sparsentan	55	42 (76.4)	2.23 (12.69)	-28.1	-3.13	3.13	9.38	40.6	0.19 [-0.25, 0.62]	
			Irbesartan	55	39 (70.9)	0.16 (9.07)	-31.3	-3.13	0.00	6.25	18.8		
	Week 94		Sparsentan	55	41 (74.5)	4.12 (13.59)	-15.6	-3.13	0.00	9.38	56.3	0.17 [-0.27, 0.61]	
			Irbesartan	55	39 (70.9)	2.24 (7.59)	-15.6	0.00	0.00	6.25	18.8		
Week 110	Sparsentan	55	37 (67.3)	2.62 (13.93)	-18.8	-3.13	0.00	6.25	46.9	0.04 [-0.42, 0.51]			
	Irbesartan	55	34 (61.8)	2.11 (8.59)	-25.0	0.00	0.00	6.25	18.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024



Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR High and UP High	KDQOL: effect of kidney disease	Baseline	Sparsentan	37	35 (94.6)	89.73 (14.15)	28.1	84.38	93.75	100.00	100.0		
			Irbesartan	36	33 (91.7)	88.45 (11.52)	56.3	84.38	90.63	96.88	100.0		
		Week 24	Sparsentan	37	26 (70.3)	91.23 (17.68)	9.4	90.63	95.31	100.00	100.0		
			Irbesartan	36	26 (72.2)	88.46 (13.51)	34.4	84.38	92.19	96.88	100.0		
		Week 48	Sparsentan	37	33 (89.2)	91.86 (10.51)	56.3	90.63	93.75	100.00	100.0		
			Irbesartan	36	24 (66.7)	92.97 (7.39)	75.0	89.06	95.31	100.00	100.0		
		Week 70	Sparsentan	37	31 (83.8)	91.53 (7.95)	68.8	87.50	93.75	100.00	100.0		
			Irbesartan	36	25 (69.4)	88.50 (17.92)	15.6	84.38	96.88	100.00	100.0		
		Week 94	Sparsentan	37	27 (73.0)	92.36 (7.22)	78.1	84.38	93.75	100.00	100.0		
			Irbesartan	36	24 (66.7)	91.54 (12.65)	43.8	87.50	95.31	100.00	100.0		
		Week 110	Sparsentan	37	26 (70.3)	92.79 (7.44)	71.9	90.63	93.75	100.00	100.0		
			Irbesartan	36	22 (61.1)	89.06 (11.36)	62.5	81.25	93.75	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	37	26 (70.3)	0.60 (19.08)	-75.0	-6.25	1.56	9.38	34.4	0.06 [-0.48, 0.61]
				Irbesartan	36	26 (72.2)	-0.48 (15.48)	-53.1	-3.13	0.00	3.13	31.3	
	Week 48		Sparsentan	37	33 (89.2)	2.27 (16.22)	-40.6	-3.13	0.00	6.25	46.9	-0.13 [-0.65, 0.40]	
			Irbesartan	36	24 (66.7)	4.04 (10.45)	-15.6	-1.56	1.56	9.38	25.0		
	Week 70		Sparsentan	37	31 (83.8)	2.52 (14.91)	-31.3	-3.13	0.00	6.25	56.3	0.11 [-0.41, 0.64]	
			Irbesartan	36	25 (69.4)	0.63 (18.97)	-71.9	-3.13	3.13	6.25	37.5		
	Week 94		Sparsentan	37	27 (73.0)	2.31 (10.17)	-18.8	-3.13	0.00	9.38	34.4	-0.11 [-0.66, 0.44]	
			Irbesartan	36	24 (66.7)	3.52 (11.99)	-18.8	-3.13	3.13	7.81	40.6		
Week 110	Sparsentan	37	26 (70.3)	2.52 (10.42)	-21.9	-3.13	3.13	6.25	31.3	0.22 [-0.35, 0.79]			
	Irbesartan	36	22 (61.1)	-0.28 (14.78)	-28.1	-6.25	0.00	6.25	40.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
eGFR High and UP Low	KDQOL: effect of kidney disease	Baseline	Sparsentan	39	37 (94.9)	89.19 (14.20)	34.4	87.50	93.75	100.00	100.0		
			Irbesartan	37	36 (97.3)	91.58 (13.48)	37.5	90.63	96.88	100.00	100.0		
		Week 24	Sparsentan	39	34 (87.2)	92.19 (10.21)	59.4	90.63	93.75	100.00	100.0		
			Irbesartan	37	28 (75.7)	87.72 (22.44)	0.0	85.94	96.88	100.00	100.0		
		Week 48	Sparsentan	39	35 (89.7)	92.32 (8.75)	65.6	90.63	93.75	100.00	100.0		
			Irbesartan	37	28 (75.7)	93.08 (11.29)	50.0	90.63	96.88	100.00	100.0		
		Week 70	Sparsentan	39	30 (76.9)	92.50 (9.09)	65.6	90.63	93.75	100.00	100.0		
			Irbesartan	37	29 (78.4)	96.12 (6.12)	78.1	93.75	100.00	100.00	100.0		
		Week 94	Sparsentan	39	32 (82.1)	90.53 (12.69)	46.9	85.94	93.75	98.44	100.0		
			Irbesartan	37	30 (81.1)	93.23 (10.05)	53.1	90.63	96.88	100.00	100.0		
		Week 110	Sparsentan	39	33 (84.6)	91.00 (9.46)	62.5	87.50	93.75	96.88	100.0		
			Irbesartan	37	29 (78.4)	94.07 (7.58)	68.8	90.63	96.88	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	39	34 (87.2)	2.67 (15.90)	-37.5	-3.13	0.00	9.38	43.8	0.30 [-0.20, 0.81]
				Irbesartan	37	28 (75.7)	-2.79 (20.15)	-100.0	-3.13	0.00	4.69	12.5	
	Week 48		Sparsentan	39	35 (89.7)	3.39 (10.29)	-21.9	-3.13	3.13	6.25	31.3	0.43 [-0.07, 0.93]	
			Irbesartan	37	28 (75.7)	-0.78 (8.95)	-21.9	-1.56	0.00	3.13	15.6		
	Week 70		Sparsentan	39	30 (76.9)	2.60 (12.04)	-31.3	0.00	0.00	3.13	34.4	0.11 [-0.40, 0.62]	
			Irbesartan	37	29 (78.4)	1.51 (8.25)	-18.8	0.00	0.00	3.13	21.9		
	Week 94		Sparsentan	39	32 (82.1)	2.05 (13.81)	-40.6	-3.13	1.56	7.81	34.4	0.18 [-0.32, 0.68]	
			Irbesartan	37	30 (81.1)	0.00 (7.43)	-12.5	-6.25	0.00	3.13	18.8		
Week 110	Sparsentan	39	33 (84.6)	2.18 (13.13)	-21.9	-3.13	0.00	6.25	37.5	0.12 [-0.38, 0.62]			
	Irbesartan	37	29 (78.4)	0.86 (6.93)	-15.6	0.00	0.00	3.13	15.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 1														
< 60 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	127	116 (91.3)	87.90 (12.51)	43.8	84.38	93.75	96.88	100.0			
			Irbesartan	129	114 (88.4)	89.23 (12.28)	43.8	84.38	93.75	100.00	100.0			
		Week 24	Sparsentan	127	98 (77.2)	88.49 (12.84)	50.0	81.25	93.75	100.00	100.0			
			Irbesartan	129	80 (62.0)	90.04 (13.44)	40.6	84.38	96.88	100.00	100.0			
		Week 48	Sparsentan	127	95 (74.8)	90.43 (13.08)	0.0	87.50	93.75	100.00	100.0			
			Irbesartan	129	74 (57.4)	91.51 (11.09)	40.6	84.38	96.88	100.00	100.0			
		Week 70	Sparsentan	127	100 (78.7)	88.41 (13.90)	43.8	84.38	93.75	100.00	100.0			
			Irbesartan	129	78 (60.5)	90.38 (13.07)	37.5	87.50	95.31	100.00	100.0			
		Week 94	Sparsentan	127	92 (72.4)	89.71 (11.41)	56.3	84.38	93.75	100.00	100.0			
			Irbesartan	129	77 (59.7)	91.56 (10.89)	46.9	87.50	93.75	100.00	100.0			
		Week 110	Sparsentan	127	92 (72.4)	89.91 (11.12)	43.8	84.38	93.75	100.00	100.0			
			Irbesartan	129	72 (55.8)	90.76 (11.32)	46.9	87.50	93.75	100.00	100.0			
			KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	127	98 (77.2)	1.28 (13.09)	-37.5	-3.13	1.56	6.25	31.3	0.12 [-0.18, 0.41]
					Irbesartan	129	80 (62.0)	-0.08 (9.55)	-37.5	-3.13	0.00	6.25	18.8	
		Week 48	Sparsentan	127	95 (74.8)	2.04 (14.78)	-100.0	0.00	3.13	6.25	56.3	0.08 [-0.23, 0.38]		
			Irbesartan	129	74 (57.4)	1.06 (8.96)	-25.0	-3.13	0.00	6.25	25.0			
		Week 70	Sparsentan	127	100 (78.7)	0.13 (12.21)	-43.8	-6.25	0.00	6.25	40.6	-0.01 [-0.31, 0.29]		
			Irbesartan	129	78 (60.5)	0.24 (12.28)	-62.5	-3.13	0.00	6.25	40.6			
		Week 94	Sparsentan	127	92 (72.4)	2.07 (11.77)	-40.6	-3.13	0.00	6.25	56.3	-0.01 [-0.31, 0.30]		
			Irbesartan	129	77 (59.7)	2.15 (8.08)	-21.9	0.00	0.00	6.25	37.5			
		Week 110	Sparsentan	127	92 (72.4)	1.29 (11.15)	-28.1	-3.13	0.00	6.25	46.9	-0.02 [-0.33, 0.29]		
			Irbesartan	129	72 (55.8)	1.48 (10.16)	-28.1	-3.13	0.00	6.25	43.8			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	49	47 (95.9)	90.69 (12.66)	34.4	87.50	93.75	100.00	100.0		
			Irbesartan	48	44 (91.7)	87.93 (13.76)	46.9	81.25	92.19	100.00	100.0		
		Week 24	Sparsentan	49	38 (77.6)	90.46 (15.85)	9.4	87.50	93.75	100.00	100.0		
			Irbesartan	48	31 (64.6)	90.02 (12.69)	34.4	84.38	93.75	96.88	100.0		
		Week 48	Sparsentan	49	44 (89.8)	91.69 (9.31)	62.5	90.63	93.75	98.44	100.0		
			Irbesartan	48	32 (66.7)	90.43 (12.17)	50.0	84.38	95.31	100.00	100.0		
		Week 70	Sparsentan	49	38 (77.6)	92.19 (8.40)	68.8	90.63	93.75	100.00	100.0		
			Irbesartan	48	32 (66.7)	90.23 (16.61)	15.6	87.50	96.88	100.00	100.0		
		Week 94	Sparsentan	49	39 (79.6)	90.22 (10.47)	46.9	84.38	93.75	96.88	100.0		
			Irbesartan	48	34 (70.8)	90.17 (12.26)	43.8	87.50	93.75	100.00	100.0		
		Week 110	Sparsentan	49	36 (73.5)	90.45 (9.82)	62.5	87.50	93.75	96.88	100.0		
			Irbesartan	48	32 (66.7)	88.96 (11.69)	59.4	82.81	93.75	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	49	38 (77.6)	-0.25 (18.39)	-75.0	-6.25	0.00	6.25	43.8	-0.05 [-0.52, 0.43]
			Irbesartan	48	31 (64.6)	0.50 (13.74)	-53.1	-3.13	0.00	6.25	31.3		
	Week 48		Sparsentan	49	44 (89.8)	1.28 (10.83)	-37.5	-3.13	0.00	6.25	31.3	-0.05 [-0.51, 0.40]	
			Irbesartan	48	32 (66.7)	1.86 (11.72)	-21.9	-3.13	0.00	9.38	25.0		
	Week 70		Sparsentan	49	38 (77.6)	0.99 (9.61)	-31.3	-3.13	0.00	3.13	34.4	-0.01 [-0.48, 0.46]	
			Irbesartan	48	32 (66.7)	1.07 (17.31)	-71.9	-3.13	0.00	9.38	37.5		
	Week 94		Sparsentan	49	39 (79.6)	0.72 (9.19)	-18.8	-3.13	0.00	3.13	34.4	-0.15 [-0.61, 0.31]	
			Irbesartan	48	34 (70.8)	2.30 (11.47)	-15.6	-6.25	0.00	9.38	40.6		
Week 110	Sparsentan	49	36 (73.5)	0.95 (11.92)	-21.9	-3.13	0.00	4.69	37.5	0.06 [-0.41, 0.54]			
	Irbesartan	48	32 (66.7)	0.20 (12.42)	-25.0	-6.25	0.00	6.25	40.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	26	24 (92.3)	86.46 (16.50)	28.1	84.38	90.63	98.44	100.0		
			Irbesartan	25	24 (96.0)	89.58 (14.96)	37.5	85.94	96.88	100.00	100.0		
		Week 24	Sparsentan	26	21 (80.8)	96.28 (3.91)	87.5	93.75	96.88	100.00	100.0		
			Irbesartan	25	20 (80.0)	83.28 (25.73)	0.0	78.13	93.75	96.88	100.0		
		Week 48	Sparsentan	26	23 (88.5)	91.98 (10.14)	56.3	87.50	93.75	100.00	100.0		
			Irbesartan	25	18 (72.0)	94.62 (7.64)	75.0	90.63	96.88	100.00	100.0		
		Week 70	Sparsentan	26	22 (84.6)	93.18 (7.00)	75.0	90.63	93.75	100.00	100.0		
			Irbesartan	25	20 (80.0)	94.38 (6.69)	81.3	89.06	96.88	100.00	100.0		
		Week 94	Sparsentan	26	20 (76.9)	94.53 (6.72)	78.1	93.75	96.88	100.00	100.0		
			Irbesartan	25	19 (76.0)	92.43 (12.21)	53.1	90.63	96.88	100.00	100.0		
	Week 110	Sparsentan	26	21 (80.8)	93.90 (5.46)	81.3	90.63	93.75	96.88	100.0			
		Irbesartan	25	17 (68.0)	93.01 (9.60)	68.8	90.63	100.00	100.00	100.0			
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	26	21 (80.8)	7.74 (11.08)	-12.5	3.13	3.13	12.50	34.4	0.78 [0.15, 1.42]
			Irbesartan	25	20 (80.0)	-6.56 (23.65)	-100.0	-6.25	0.00	3.13	12.5		
	Week 48		Sparsentan	26	23 (88.5)	5.57 (17.50)	-40.6	-3.13	3.13	15.63	46.9	0.35 [-0.27, 0.98]	
			Irbesartan	25	18 (72.0)	0.69 (6.12)	-12.5	0.00	0.00	3.13	15.6		
	Week 70		Sparsentan	26	22 (84.6)	7.39 (16.27)	-15.6	-3.13	3.13	9.38	56.3	0.43 [-0.19, 1.04]	
			Irbesartan	25	20 (80.0)	1.72 (8.92)	-18.8	-1.56	3.13	4.69	21.9		
	Week 94		Sparsentan	26	20 (76.9)	6.09 (13.08)	-15.6	0.00	3.13	12.50	34.4	0.54 [-0.10, 1.18]	
			Irbesartan	25	19 (76.0)	0.16 (8.23)	-18.8	0.00	0.00	3.13	18.8		
Week 110	Sparsentan		26	21 (80.8)	5.36 (11.19)	-12.5	0.00	6.25	9.38	31.3	0.46 [-0.19, 1.11]		
	Irbesartan		25	17 (68.0)	0.92 (7.39)	-12.5	0.00	3.13	6.25	15.6			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 2													
< 45 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	82	73 (89.0)	85.74 (13.48)	43.8	78.13	90.63	93.75	100.0		
			Irbesartan	80	69 (86.3)	87.14 (13.91)	43.8	81.25	90.63	96.88	100.0		
		Week 24	Sparsentan	82	64 (78.0)	89.06 (12.17)	50.0	82.81	93.75	100.00	100.0		
			Irbesartan	80	48 (60.0)	87.83 (15.81)	40.6	82.81	95.31	100.00	100.0		
		Week 48	Sparsentan	82	61 (74.4)	89.09 (14.68)	0.0	87.50	90.63	96.88	100.0		
			Irbesartan	80	43 (53.8)	90.12 (12.81)	40.6	84.38	93.75	100.00	100.0		
		Week 70	Sparsentan	82	65 (79.3)	87.36 (14.46)	46.9	84.38	93.75	96.88	100.0		
			Irbesartan	80	45 (56.3)	87.36 (15.28)	37.5	84.38	90.63	100.00	100.0		
		Week 94	Sparsentan	82	60 (73.2)	89.22 (11.35)	56.3	84.38	93.75	98.44	100.0		
			Irbesartan	80	47 (58.8)	90.03 (12.48)	46.9	84.38	93.75	100.00	100.0		
		Week 110	Sparsentan	82	56 (68.3)	89.12 (10.56)	56.3	81.25	90.63	96.88	100.0		
			Irbesartan	80	44 (55.0)	89.56 (12.22)	46.9	85.94	93.75	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	82	64 (78.0)	3.22 (12.24)	-37.5	-3.13	3.13	9.38	31.3	0.31 [-0.07, 0.69]
			Irbesartan	80	48 (60.0)	-0.46 (11.22)	-37.5	-3.13	0.00	6.25	18.8		
	Week 48		Sparsentan	82	61 (74.4)	2.51 (17.36)	-100.0	0.00	3.13	6.25	56.3	0.13 [-0.26, 0.52]	
			Irbesartan	80	43 (53.8)	0.65 (9.46)	-25.0	-3.13	0.00	6.25	21.9		
	Week 70		Sparsentan	82	65 (79.3)	0.87 (12.16)	-43.8	-3.13	0.00	6.25	40.6	0.13 [-0.25, 0.51]	
			Irbesartan	80	45 (56.3)	-0.83 (14.81)	-62.5	-3.13	0.00	6.25	40.6		
Week 94	Sparsentan	82	60 (73.2)	2.66 (11.77)	-18.8	-3.13	0.00	7.81	56.3	0.05 [-0.33, 0.43]			
	Irbesartan	80	47 (58.8)	2.13 (9.26)	-21.9	0.00	0.00	6.25	37.5				
Week 110	Sparsentan	82	56 (68.3)	1.84 (10.47)	-15.6	-3.13	0.00	6.25	46.9	-0.02 [-0.42, 0.37]			
	Irbesartan	80	44 (55.0)	2.06 (9.95)	-15.6	-3.13	0.00	6.25	43.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
45 to < 60 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	45	43 (95.6)	91.57 (9.75)	62.5	87.50	93.75	100.00	100.0		
			Irbesartan	49	45 (91.8)	92.43 (8.41)	71.9	87.50	96.88	100.00	100.0		
		Week 24	Sparsentan	45	34 (75.6)	87.41 (14.13)	56.3	81.25	93.75	96.88	100.0		
			Irbesartan	49	32 (65.3)	93.36 (7.89)	75.0	85.94	96.88	100.00	100.0		
		Week 48	Sparsentan	45	34 (75.6)	92.83 (9.31)	68.8	87.50	96.88	100.00	100.0		
			Irbesartan	49	31 (63.3)	93.45 (7.92)	75.0	87.50	96.88	100.00	100.0		
		Week 70	Sparsentan	45	35 (77.8)	90.36 (12.76)	43.8	84.38	93.75	100.00	100.0		
			Irbesartan	49	33 (67.3)	94.51 (7.74)	71.9	93.75	96.88	100.00	100.0		
		Week 94	Sparsentan	45	32 (71.1)	90.63 (11.64)	56.3	85.94	93.75	100.00	100.0		
			Irbesartan	49	30 (61.2)	93.96 (7.34)	71.9	90.63	96.88	100.00	100.0		
		Week 110	Sparsentan	45	36 (80.0)	91.15 (11.99)	43.8	89.06	95.31	100.00	100.0		
			Irbesartan	49	28 (57.1)	92.63 (9.63)	68.8	87.50	96.88	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	45	34 (75.6)	-2.39 (14.02)	-37.5	-9.38	0.00	6.25	31.3	-0.26 [-0.75, 0.22]
				Irbesartan	49	32 (65.3)	0.49 (6.40)	-18.8	0.00	0.00	3.13	12.5	
	Week 48		Sparsentan	45	34 (75.6)	1.19 (8.57)	-15.6	-3.13	0.00	6.25	31.3	-0.05 [-0.54, 0.44]	
			Irbesartan	49	31 (63.3)	1.61 (8.34)	-15.6	0.00	0.00	6.25	25.0		
	Week 70		Sparsentan	45	35 (77.8)	-1.25 (12.37)	-31.3	-6.25	0.00	6.25	25.0	-0.29 [-0.76, 0.19]	
			Irbesartan	49	33 (67.3)	1.70 (7.58)	-21.9	0.00	0.00	3.13	15.6		
	Week 94		Sparsentan	45	32 (71.1)	0.98 (11.86)	-40.6	-3.13	1.56	6.25	28.1	-0.13 [-0.63, 0.37]	
			Irbesartan	49	30 (61.2)	2.19 (5.93)	-9.4	0.00	0.00	3.13	18.8		
Week 110	Sparsentan	45	36 (80.0)	0.43 (12.23)	-28.1	-6.25	0.00	4.69	34.4	-0.01 [-0.50, 0.48]			
	Irbesartan	49	28 (57.1)	0.56 (10.59)	-28.1	-1.56	0.00	6.25	18.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
60 to < 90 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	49	47 (95.9)	90.69 (12.66)	34.4	87.50	93.75	100.00	100.0	
			Irbesartan	48	44 (91.7)	87.93 (13.76)	46.9	81.25	92.19	100.00	100.0	
		Week 24	Sparsentan	49	38 (77.6)	90.46 (15.85)	9.4	87.50	93.75	100.00	100.0	
			Irbesartan	48	31 (64.6)	90.02 (12.69)	34.4	84.38	93.75	96.88	100.0	
		Week 48	Sparsentan	49	44 (89.8)	91.69 (9.31)	62.5	90.63	93.75	98.44	100.0	
			Irbesartan	48	32 (66.7)	90.43 (12.17)	50.0	84.38	95.31	100.00	100.0	
		Week 70	Sparsentan	49	38 (77.6)	92.19 (8.40)	68.8	90.63	93.75	100.00	100.0	
			Irbesartan	48	32 (66.7)	90.23 (16.61)	15.6	87.50	96.88	100.00	100.0	
		Week 94	Sparsentan	49	39 (79.6)	90.22 (10.47)	46.9	84.38	93.75	96.88	100.0	
			Irbesartan	48	34 (70.8)	90.17 (12.26)	43.8	87.50	93.75	100.00	100.0	
		Week 110	Sparsentan	49	36 (73.5)	90.45 (9.82)	62.5	87.50	93.75	96.88	100.0	
			Irbesartan	48	32 (66.7)	88.96 (11.69)	59.4	82.81	93.75	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	49	38 (77.6)	-0.25 (18.39)	-75.0	-6.25	0.00	6.25	43.8	-0.05 [-0.52, 0.43]
			Irbesartan	48	31 (64.6)	0.50 (13.74)	-53.1	-3.13	0.00	6.25	31.3	
		Week 48	Sparsentan	49	44 (89.8)	1.28 (10.83)	-37.5	-3.13	0.00	6.25	31.3	-0.05 [-0.51, 0.40]
			Irbesartan	48	32 (66.7)	1.86 (11.72)	-21.9	-3.13	0.00	9.38	25.0	
		Week 70	Sparsentan	49	38 (77.6)	0.99 (9.61)	-31.3	-3.13	0.00	3.13	34.4	-0.01 [-0.48, 0.46]
			Irbesartan	48	32 (66.7)	1.07 (17.31)	-71.9	-3.13	0.00	9.38	37.5	
		Week 94	Sparsentan	49	39 (79.6)	0.72 (9.19)	-18.8	-3.13	0.00	3.13	34.4	-0.15 [-0.61, 0.31]
			Irbesartan	48	34 (70.8)	2.30 (11.47)	-15.6	-6.25	0.00	9.38	40.6	
Week 110	Sparsentan	49	36 (73.5)	0.95 (11.92)	-21.9	-3.13	0.00	4.69	37.5	0.06 [-0.41, 0.54]		
	Irbesartan	48	32 (66.7)	0.20 (12.42)	-25.0	-6.25	0.00	6.25	40.6			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024



Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL: effect of kidney disease	Baseline	Sparsentan	26	24 (92.3)	86.46 (16.50)	28.1	84.38	90.63	98.44	100.0		
			Irbesartan	25	24 (96.0)	89.58 (14.96)	37.5	85.94	96.88	100.00	100.0		
		Week 24	Sparsentan	26	21 (80.8)	96.28 (3.91)	87.5	93.75	96.88	100.00	100.0		
			Irbesartan	25	20 (80.0)	83.28 (25.73)	0.0	78.13	93.75	96.88	100.0		
		Week 48	Sparsentan	26	23 (88.5)	91.98 (10.14)	56.3	87.50	93.75	100.00	100.0		
			Irbesartan	25	18 (72.0)	94.62 (7.64)	75.0	90.63	96.88	100.00	100.0		
		Week 70	Sparsentan	26	22 (84.6)	93.18 (7.00)	75.0	90.63	93.75	100.00	100.0		
			Irbesartan	25	20 (80.0)	94.38 (6.69)	81.3	89.06	96.88	100.00	100.0		
		Week 94	Sparsentan	26	20 (76.9)	94.53 (6.72)	78.1	93.75	96.88	100.00	100.0		
			Irbesartan	25	19 (76.0)	92.43 (12.21)	53.1	90.63	96.88	100.00	100.0		
		Week 110	Sparsentan	26	21 (80.8)	93.90 (5.46)	81.3	90.63	93.75	96.88	100.0		
			Irbesartan	25	17 (68.0)	93.01 (9.60)	68.8	90.63	100.00	100.00	100.0		
			KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	26	21 (80.8)	7.74 (11.08)	-12.5	3.13	3.13	12.50	34.4
		Irbesartan		25	20 (80.0)	-6.56 (23.65)	-100.0	-6.25	0.00	3.13	12.5		
	Week 48	Sparsentan		26	23 (88.5)	5.57 (17.50)	-40.6	-3.13	3.13	15.63	46.9	0.35 [-0.27, 0.98]	
		Irbesartan		25	18 (72.0)	0.69 (6.12)	-12.5	0.00	0.00	3.13	15.6		
	Week 70	Sparsentan		26	22 (84.6)	7.39 (16.27)	-15.6	-3.13	3.13	9.38	56.3	0.43 [-0.19, 1.04]	
		Irbesartan		25	20 (80.0)	1.72 (8.92)	-18.8	-1.56	3.13	4.69	21.9		
	Week 94	Sparsentan		26	20 (76.9)	6.09 (13.08)	-15.6	0.00	3.13	12.50	34.4	0.54 [-0.10, 1.18]	
		Irbesartan	25	19 (76.0)	0.16 (8.23)	-18.8	0.00	0.00	3.13	18.8			
Week 110	Sparsentan	26	21 (80.8)	5.36 (11.19)	-12.5	0.00	6.25	9.38	31.3	0.46 [-0.19, 1.11]			
	Irbesartan	25	17 (68.0)	0.92 (7.39)	-12.5	0.00	3.13	6.25	15.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline urine protein excretion														
<= 1.75 g/day	KDQOL: effect of kidney disease	Baseline	Sparsentan	98	93 (94.9)	87.40 (13.99)	34.4	84.38	90.63	96.88	100.0			
			Irbesartan	93	83 (89.2)	90.02 (13.11)	37.5	84.38	96.88	100.00	100.0			
		Week 24	Sparsentan	98	80 (81.6)	88.87 (12.68)	50.0	84.38	93.75	100.00	100.0			
			Irbesartan	93	55 (59.1)	90.68 (17.89)	0.0	87.50	96.88	100.00	100.0			
		Week 48	Sparsentan	98	82 (83.7)	90.47 (13.61)	0.0	87.50	93.75	100.00	100.0			
			Irbesartan	93	57 (61.3)	92.43 (10.68)	50.0	90.63	96.88	100.00	100.0			
		Week 70	Sparsentan	98	75 (76.5)	88.88 (13.08)	43.8	84.38	93.75	100.00	100.0			
			Irbesartan	93	61 (65.6)	93.24 (9.01)	56.3	87.50	96.88	100.00	100.0			
		Week 94	Sparsentan	98	76 (77.6)	89.39 (11.89)	46.9	82.81	93.75	100.00	100.0			
			Irbesartan	93	63 (67.7)	91.77 (11.26)	50.0	87.50	96.88	100.00	100.0			
		Week 110	Sparsentan	98	75 (76.5)	89.50 (11.78)	43.8	87.50	93.75	96.88	100.0			
			Irbesartan	93	57 (61.3)	91.61 (10.11)	59.4	87.50	93.75	100.00	100.0			
			KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	98	80 (81.6)	1.99 (15.14)	-37.5	-6.25	0.00	9.38	43.8	0.18 [-0.16, 0.53]
					Irbesartan	93	55 (59.1)	-0.85 (16.06)	-100.0	-3.13	0.00	6.25	18.8	
				Week 48	Sparsentan	98	82 (83.7)	1.91 (16.64)	-100.0	-3.13	3.13	6.25	56.3	0.11 [-0.23, 0.45]
					Irbesartan	93	57 (61.3)	0.38 (8.96)	-25.0	0.00	0.00	3.13	25.0	
				Week 70	Sparsentan	98	75 (76.5)	0.96 (12.51)	-31.3	-6.25	0.00	6.25	40.6	-0.00 [-0.34, 0.34]
					Irbesartan	93	61 (65.6)	0.97 (9.08)	-31.3	0.00	0.00	3.13	21.9	
		Week 94	Sparsentan	98	76 (77.6)	2.51 (13.26)	-40.6	-3.13	0.00	7.81	56.3	0.13 [-0.20, 0.47]		
			Irbesartan	93	63 (67.7)	0.99 (8.26)	-21.9	-3.13	0.00	6.25	18.8			
		Week 110	Sparsentan	98	75 (76.5)	1.71 (13.49)	-28.1	-3.13	0.00	6.25	46.9	0.04 [-0.30, 0.39]		
			Irbesartan	93	57 (61.3)	1.21 (7.69)	-25.0	0.00	0.00	6.25	15.6			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 1.75 g/day	KDQOL: effect of kidney disease	Baseline	Sparsentan	104	94 (90.4)	89.43 (12.19)	28.1	84.38	93.75	96.88	100.0		
			Irbesartan	109	99 (90.8)	88.07 (12.83)	46.9	84.38	90.63	96.88	100.0		
		Week 24	Sparsentan	104	77 (74.0)	91.19 (13.44)	9.4	87.50	96.88	100.00	100.0		
			Irbesartan	109	76 (69.7)	87.79 (14.10)	34.4	81.25	93.75	96.88	100.0		
		Week 48	Sparsentan	104	80 (76.9)	91.52 (9.49)	50.0	87.50	93.75	100.00	100.0		
			Irbesartan	109	67 (61.5)	91.04 (11.23)	40.6	84.38	93.75	100.00	100.0		
		Week 70	Sparsentan	104	85 (81.7)	90.92 (11.23)	46.9	87.50	93.75	100.00	100.0		
			Irbesartan	109	69 (63.3)	88.95 (15.98)	15.6	84.38	93.75	100.00	100.0		
		Week 94	Sparsentan	104	75 (72.1)	91.58 (9.35)	56.3	87.50	93.75	100.00	100.0		
			Irbesartan	109	67 (61.5)	90.90 (11.57)	43.8	87.50	93.75	100.00	100.0		
		Week 110	Sparsentan	104	74 (71.2)	91.72 (8.32)	71.9	87.50	93.75	100.00	100.0		
			Irbesartan	109	64 (58.7)	89.70 (12.05)	46.9	84.38	93.75	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	104	77 (74.0)	1.54 (13.76)	-75.0	-3.13	3.13	6.25	34.4	0.20 [-0.12, 0.52]
				Irbesartan	109	76 (69.7)	-0.99 (11.73)	-53.1	-3.13	0.00	3.13	31.3	
	Week 48		Sparsentan	104	80 (76.9)	2.77 (11.34)	-40.6	-3.13	0.00	6.25	46.9	0.08 [-0.24, 0.41]	
			Irbesartan	109	67 (61.5)	1.91 (9.70)	-21.9	-3.13	0.00	6.25	25.0		
	Week 70		Sparsentan	104	85 (81.7)	1.65 (12.49)	-43.8	-3.13	0.00	6.25	56.3	0.09 [-0.23, 0.41]	
			Irbesartan	109	69 (63.3)	0.41 (16.04)	-71.9	-3.13	0.00	6.25	40.6		
	Week 94		Sparsentan	104	75 (72.1)	2.00 (9.20)	-18.8	-3.13	0.00	6.25	34.4	-0.08 [-0.41, 0.25]	
			Irbesartan	109	67 (61.5)	2.75 (9.74)	-18.8	0.00	0.00	6.25	40.6		
Week 110	Sparsentan	104	74 (71.2)	1.86 (8.79)	-15.6	-3.13	0.00	6.25	31.3	0.09 [-0.25, 0.42]			
	Irbesartan	109	64 (58.7)	0.93 (12.41)	-28.1	-6.25	0.00	6.25	43.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline use of antihypertensives														
Yes	KDQOL: effect of kidney disease	Baseline	Sparsentan	90	79 (87.8)	89.79 (11.08)	50.0	84.38	93.75	96.88	100.0			
			Irbesartan	88	76 (86.4)	89.14 (11.91)	43.8	81.25	93.75	100.00	100.0			
		Week 24	Sparsentan	90	66 (73.3)	91.48 (10.89)	50.0	87.50	95.31	100.00	100.0			
			Irbesartan	88	49 (55.7)	89.48 (13.35)	50.0	84.38	93.75	100.00	100.0			
		Week 48	Sparsentan	90	65 (72.2)	91.59 (7.59)	75.0	87.50	93.75	100.00	100.0			
			Irbesartan	88	47 (53.4)	92.22 (10.05)	59.4	84.38	96.88	100.00	100.0			
		Week 70	Sparsentan	90	68 (75.6)	89.61 (11.81)	46.9	87.50	93.75	96.88	100.0			
			Irbesartan	88	49 (55.7)	91.26 (10.99)	56.3	87.50	96.88	100.00	100.0			
		Week 94	Sparsentan	90	63 (70.0)	89.98 (10.43)	56.3	84.38	93.75	100.00	100.0			
			Irbesartan	88	54 (61.4)	90.74 (11.50)	50.0	87.50	95.31	100.00	100.0			
		Week 110	Sparsentan	90	63 (70.0)	91.42 (9.31)	56.3	87.50	93.75	100.00	100.0			
			Irbesartan	88	51 (58.0)	89.95 (11.41)	59.4	84.38	93.75	100.00	100.0			
			KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	90	66 (73.3)	2.13 (11.36)	-37.5	-3.13	0.00	6.25	31.3	0.32 [-0.06, 0.69]
					Irbesartan	88	49 (55.7)	-1.34 (10.40)	-37.5	-3.13	0.00	6.25	18.8	
		Week 48		Sparsentan	90	65 (72.2)	1.68 (8.99)	-21.9	-3.13	0.00	6.25	31.3	-0.04 [-0.41, 0.34]	
				Irbesartan	88	47 (53.4)	1.99 (8.70)	-15.6	0.00	0.00	6.25	25.0		
		Week 70		Sparsentan	90	68 (75.6)	-0.74 (10.77)	-43.8	-3.13	0.00	6.25	21.9	-0.13 [-0.49, 0.24]	
				Irbesartan	88	49 (55.7)	0.57 (9.62)	-31.3	-3.13	0.00	6.25	18.8		
Week 94	Sparsentan	90		63 (70.0)	0.84 (10.09)	-21.9	-3.13	0.00	6.25	28.1	-0.04 [-0.40, 0.32]			
	Irbesartan	88		54 (61.4)	1.22 (8.31)	-21.9	0.00	0.00	6.25	18.8				
Week 110	Sparsentan	90		63 (70.0)	1.84 (10.13)	-21.9	-3.13	0.00	6.25	34.4	0.13 [-0.24, 0.50]			
	Irbesartan	88		51 (58.0)	0.55 (9.18)	-28.1	-3.13	0.00	6.25	18.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
No	KDQOL: effect of kidney disease	Baseline	Sparsentan	112	108 (96.4)	87.41 (14.39)	28.1	84.38	90.63	96.88	100.0		
			Irbesartan	114	106 (93.0)	88.83 (13.71)	37.5	84.38	93.75	100.00	100.0		
		Week 24	Sparsentan	112	91 (81.3)	88.94 (14.41)	9.4	84.38	93.75	100.00	100.0		
			Irbesartan	114	82 (71.9)	88.72 (17.17)	0.0	84.38	95.31	96.88	100.0		
		Week 48	Sparsentan	112	97 (86.6)	90.59 (13.85)	0.0	87.50	93.75	100.00	100.0		
			Irbesartan	114	77 (67.5)	91.36 (11.53)	40.6	87.50	96.88	100.00	100.0		
		Week 70	Sparsentan	112	92 (82.1)	90.22 (12.43)	43.8	87.50	93.75	100.00	100.0		
			Irbesartan	114	81 (71.1)	90.78 (14.59)	15.6	87.50	96.88	100.00	100.0		
		Week 94	Sparsentan	112	88 (78.6)	90.84 (10.98)	46.9	84.38	93.75	100.00	100.0		
			Irbesartan	114	76 (66.7)	91.74 (11.36)	43.8	87.50	93.75	100.00	100.0		
		Week 110	Sparsentan	112	86 (76.8)	90.01 (10.88)	43.8	84.38	93.75	96.88	100.0		
			Irbesartan	114	70 (61.4)	91.07 (11.06)	46.9	84.38	93.75	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	112	91 (81.3)	1.51 (16.37)	-75.0	-6.25	3.13	9.38	43.8	0.14 [-0.16, 0.44]
				Irbesartan	114	82 (71.9)	-0.69 (15.33)	-100.0	-3.13	0.00	3.13	31.3	
	Week 48		Sparsentan	112	97 (86.6)	2.77 (16.89)	-100.0	-3.13	3.13	6.25	56.3	0.14 [-0.16, 0.44]	
			Irbesartan	114	77 (67.5)	0.73 (9.76)	-25.0	-3.13	0.00	3.13	25.0		
	Week 70		Sparsentan	112	92 (82.1)	2.85 (13.43)	-31.3	-3.13	0.00	7.81	56.3	0.15 [-0.15, 0.45]	
			Irbesartan	114	81 (71.1)	0.73 (15.01)	-71.9	-3.13	0.00	6.25	40.6		
	Week 94		Sparsentan	112	88 (78.6)	3.27 (12.19)	-40.6	-3.13	3.13	6.25	56.3	0.08 [-0.23, 0.39]	
			Irbesartan	114	76 (66.7)	2.38 (9.59)	-12.5	-3.13	0.00	6.25	40.6		
Week 110	Sparsentan	112	86 (76.8)	1.74 (12.25)	-28.1	-3.13	0.00	6.25	46.9	0.03 [-0.29, 0.34]			
	Irbesartan	114	70 (61.4)	1.43 (11.28)	-25.0	-6.25	0.00	6.25	43.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Time since renal biopsy														
<= 5 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	113	104 (92.0)	88.07 (12.42)	34.4	84.38	92.19	96.88	100.0			
			Irbesartan	127	118 (92.9)	88.43 (13.48)	37.5	84.38	93.75	100.00	100.0			
		Week 24	Sparsentan	113	88 (77.9)	91.26 (11.47)	50.0	89.06	93.75	100.00	100.0			
			Irbesartan	127	86 (67.7)	87.61 (17.66)	0.0	84.38	93.75	96.88	100.0			
		Week 48	Sparsentan	113	92 (81.4)	91.78 (9.66)	50.0	87.50	93.75	100.00	100.0			
			Irbesartan	127	85 (66.9)	91.21 (12.04)	40.6	84.38	96.88	100.00	100.0			
		Week 70	Sparsentan	113	90 (79.6)	90.03 (11.26)	46.9	87.50	93.75	96.88	100.0			
			Irbesartan	127	86 (67.7)	91.64 (12.83)	15.6	87.50	96.88	100.00	100.0			
		Week 94	Sparsentan	113	86 (76.1)	90.08 (11.65)	46.9	84.38	93.75	100.00	100.0			
			Irbesartan	127	86 (67.7)	91.02 (11.72)	43.8	87.50	93.75	100.00	100.0			
		Week 110	Sparsentan	113	88 (77.9)	90.63 (9.48)	56.3	87.50	93.75	96.88	100.0			
			Irbesartan	127	80 (63.0)	90.43 (11.03)	59.4	84.38	93.75	100.00	100.0			
			KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	113	88 (77.9)	4.05 (14.26)	-37.5	-1.56	3.13	12.50	43.8	0.38 [0.08, 0.68]
					Irbesartan	127	86 (67.7)	-1.53 (15.24)	-100.0	-3.13	0.00	6.25	31.3	
		Week 48		Sparsentan	113	92 (81.4)	4.08 (10.86)	-40.6	0.00	3.13	7.81	34.4	0.26 [-0.04, 0.55]	
				Irbesartan	127	85 (66.9)	1.40 (9.92)	-25.0	0.00	0.00	6.25	25.0		
		Week 70		Sparsentan	113	90 (79.6)	1.63 (11.92)	-31.3	-3.13	0.00	6.25	34.4	0.06 [-0.24, 0.35]	
				Irbesartan	127	86 (67.7)	0.91 (12.54)	-71.9	0.00	0.00	6.25	28.1		
Week 94	Sparsentan	113		86 (76.1)	2.54 (11.98)	-40.6	-3.13	0.00	6.25	34.4	0.09 [-0.21, 0.39]			
	Irbesartan	127		86 (67.7)	1.60 (9.22)	-21.9	0.00	0.00	6.25	37.5				
Week 110	Sparsentan	113		88 (77.9)	2.52 (11.30)	-21.9	-3.13	0.00	6.25	37.5	0.16 [-0.14, 0.46]			
	Irbesartan	127		80 (63.0)	0.86 (9.54)	-25.0	-3.13	0.00	6.25	25.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 5 years	KDQOL: effect of kidney disease	Baseline	Sparsentan	89	83 (93.3)	88.86 (14.00)	28.1	84.38	93.75	100.00	100.0		
			Irbesartan	75	64 (85.3)	89.94 (11.97)	46.9	85.94	93.75	100.00	100.0		
		Week 24	Sparsentan	89	69 (77.5)	88.41 (14.80)	9.4	81.25	93.75	100.00	100.0		
			Irbesartan	75	45 (60.0)	91.67 (11.13)	50.0	87.50	96.88	100.00	100.0		
		Week 48	Sparsentan	89	70 (78.7)	89.96 (14.00)	0.0	87.50	93.75	100.00	100.0		
			Irbesartan	75	39 (52.0)	92.71 (8.15)	75.0	87.50	96.88	100.00	100.0		
		Week 70	Sparsentan	89	70 (78.7)	89.87 (13.26)	43.8	87.50	93.75	100.00	100.0		
			Irbesartan	75	44 (58.7)	89.63 (14.25)	37.5	84.38	95.31	100.00	100.0		
		Week 94	Sparsentan	89	65 (73.0)	91.01 (9.43)	59.4	84.38	93.75	100.00	100.0		
			Irbesartan	75	44 (58.7)	91.90 (10.81)	46.9	87.50	96.88	100.00	100.0		
		Week 110	Sparsentan	89	61 (68.5)	90.57 (11.32)	43.8	87.50	93.75	100.00	100.0		
			Irbesartan	75	41 (54.7)	90.93 (11.58)	46.9	87.50	93.75	100.00	100.0		
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	89	69 (77.5)	-1.13 (14.23)	-75.0	-6.25	0.00	6.25	31.3	-0.11 [-0.48, 0.27]
				Irbesartan	75	45 (60.0)	0.21 (10.02)	-37.5	-3.13	0.00	3.13	18.8	
	Week 48		Sparsentan	89	70 (78.7)	0.04 (17.55)	-100.0	-3.13	0.00	6.25	56.3	-0.05 [-0.44, 0.34]	
			Irbesartan	75	39 (52.0)	0.80 (8.10)	-15.6	-6.25	0.00	3.13	18.8		
	Week 70		Sparsentan	89	70 (78.7)	0.94 (13.21)	-43.8	-3.13	0.00	6.25	56.3	0.05 [-0.32, 0.43]	
			Irbesartan	75	44 (58.7)	0.21 (14.54)	-62.5	-3.13	0.00	3.13	40.6		
	Week 94		Sparsentan	89	65 (73.0)	1.88 (10.64)	-18.8	-3.13	0.00	6.25	56.3	-0.06 [-0.44, 0.32]	
			Irbesartan	75	44 (58.7)	2.49 (8.83)	-12.5	-1.56	0.00	6.25	40.6		
Week 110	Sparsentan	89	61 (68.5)	0.72 (11.46)	-28.1	-3.13	0.00	6.25	46.9	-0.06 [-0.46, 0.33]			
	Irbesartan	75	41 (54.7)	1.45 (12.06)	-28.1	-3.13	0.00	3.13	43.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: History of hypertension													
Yes	KDQOL: effect of kidney disease	Baseline	Sparsentan	155	141 (91.0)	87.61 (13.34)	28.1	84.38	93.75	96.88	100.0		
			Irbesartan	161	142 (88.2)	88.78 (12.57)	43.8	84.38	93.75	100.00	100.0		
		Week 24	Sparsentan	155	119 (76.8)	90.81 (11.28)	56.3	87.50	93.75	100.00	100.0		
			Irbesartan	161	100 (62.1)	90.88 (11.41)	50.0	84.38	96.88	100.00	100.0		
		Week 48	Sparsentan	155	121 (78.1)	90.37 (12.66)	0.0	87.50	93.75	100.00	100.0		
			Irbesartan	161	95 (59.0)	92.86 (9.21)	59.4	90.63	96.88	100.00	100.0		
		Week 70	Sparsentan	155	123 (79.4)	89.94 (11.77)	43.8	87.50	93.75	100.00	100.0		
			Irbesartan	161	100 (62.1)	91.84 (11.05)	40.6	87.50	96.88	100.00	100.0		
		Week 94	Sparsentan	155	114 (73.5)	89.86 (11.07)	46.9	84.38	93.75	100.00	100.0		
			Irbesartan	161	101 (62.7)	90.81 (12.15)	43.8	87.50	93.75	100.00	100.0		
	Week 110	Sparsentan	155	112 (72.3)	90.07 (10.94)	43.8	84.38	93.75	98.44	100.0			
		Irbesartan	161	92 (57.1)	91.13 (11.21)	46.9	87.50	95.31	100.00	100.0			
		KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	155	119 (76.8)	3.62 (12.08)	-37.5	-3.13	3.13	9.38	43.8	0.24 [-0.03, 0.51]
				Irbesartan	161	100 (62.1)	1.00 (9.60)	-37.5	-1.56	0.00	6.25	31.3	
	Week 48		Sparsentan	155	121 (78.1)	2.76 (14.44)	-100.0	0.00	3.13	6.25	46.9	0.00 [-0.27, 0.27]	
			Irbesartan	161	95 (59.0)	2.73 (8.89)	-21.9	0.00	0.00	6.25	25.0		
	Week 70		Sparsentan	155	123 (79.4)	2.18 (12.03)	-28.1	-3.13	0.00	6.25	56.3	-0.06 [-0.32, 0.21]	
			Irbesartan	161	100 (62.1)	2.84 (10.36)	-31.3	0.00	0.00	6.25	40.6		
	Week 94		Sparsentan	155	114 (73.5)	2.58 (9.88)	-21.9	-3.13	0.00	6.25	34.4	0.00 [-0.27, 0.27]	
			Irbesartan	161	101 (62.7)	2.57 (9.40)	-21.9	0.00	0.00	6.25	40.6		
Week 110	Sparsentan		155	112 (72.3)	2.04 (10.67)	-28.1	-3.13	0.00	6.25	37.5	-0.06 [-0.34, 0.21]		
	Irbesartan		161	92 (57.1)	2.72 (10.23)	-28.1	-1.56	0.00	6.25	43.8			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024



Table PT2KEFC\_FSHM: KDQOL: Course of effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL: effect of kidney disease	Baseline	Sparsentan	47	46 (97.9)	90.90 (12.22)	43.8	87.50	93.75	100.00	100.0	
			Irbesartan	41	40 (97.6)	89.61 (14.41)	37.5	87.50	95.31	100.00	100.0	
		Week 24	Sparsentan	47	38 (80.9)	87.50 (17.48)	9.4	81.25	93.75	100.00	100.0	
			Irbesartan	41	31 (75.6)	82.96 (24.55)	0.0	75.00	93.75	96.88	100.0	
		Week 48	Sparsentan	47	41 (87.2)	92.84 (8.28)	62.5	87.50	93.75	100.00	100.0	
			Irbesartan	41	29 (70.7)	87.82 (14.90)	40.6	81.25	93.75	100.00	100.0	
		Week 70	Sparsentan	47	37 (78.7)	90.03 (13.46)	50.0	87.50	93.75	100.00	100.0	
			Irbesartan	41	30 (73.2)	88.02 (18.95)	15.6	84.38	95.31	100.00	100.0	
		Week 94	Sparsentan	47	37 (78.7)	92.40 (9.48)	56.3	87.50	96.88	100.00	100.0	
			Irbesartan	41	29 (70.7)	93.10 (8.11)	71.9	90.63	93.75	100.00	100.0	
		Week 110	Sparsentan	47	37 (78.7)	92.23 (7.61)	75.0	87.50	93.75	100.00	100.0	
			Irbesartan	41	29 (70.7)	88.90 (11.10)	62.5	81.25	90.63	100.00	100.0	
	KDQOL: Change from baseline in effect of kidney disease	Week 24	Sparsentan	47	38 (80.9)	-4.03 (19.18)	-75.0	-12.50	0.00	3.13	34.4	0.16 [-0.32, 0.63]
			Irbesartan	41	31 (75.6)	-7.16 (21.26)	-100.0	-9.38	-3.13	3.13	15.6	
		Week 48	Sparsentan	47	41 (87.2)	1.07 (13.70)	-37.5	-3.13	0.00	6.25	56.3	0.40 [-0.08, 0.88]
			Irbesartan	41	29 (70.7)	-3.77 (9.28)	-25.0	-9.38	0.00	0.00	12.5	
		Week 70	Sparsentan	47	37 (78.7)	-1.52 (13.59)	-43.8	-6.25	0.00	6.25	34.4	0.32 [-0.17, 0.80]
			Irbesartan	41	30 (73.2)	-6.56 (18.38)	-71.9	-6.25	0.00	3.13	12.5	
		Week 94	Sparsentan	47	37 (78.7)	1.27 (15.26)	-40.6	-3.13	0.00	3.13	56.3	0.14 [-0.35, 0.62]
			Irbesartan	41	29 (70.7)	-0.43 (7.46)	-15.6	-6.25	0.00	3.13	18.8	
Week 110	Sparsentan	47	37 (78.7)	1.01 (13.38)	-21.9	-3.13	0.00	6.25	46.9	0.44 [-0.05, 0.93]		
	Irbesartan	41	29 (70.7)	-4.20 (9.35)	-25.0	-9.38	0.00	3.13	12.5			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Sex	Overall	Sparsentan								Interaction: 0.148
Male	Week 24	Sparsentan	139	109 (78.4)	-0.37 (1.02)	(-2.37, 1.63)	0.85 (1.50)	(-2.10, 3.80)	0.572	
		Irbesartan	143	93 (65.0)	-1.22 (1.10)	(-3.38, 0.94)				
	Week 48	Sparsentan	139	112 (80.6)	0.16 (1.00)	(-1.81, 2.13)	-0.29 (1.50)	(-3.24, 2.65)	0.847	
		Irbesartan	143	87 (60.8)	0.45 (1.11)	(-1.74, 2.63)				
	Week 70	Sparsentan	139	114 (82.0)	-0.30 (1.00)	(-2.27, 1.67)	-0.40 (1.49)	(-3.33, 2.52)	0.787	
		Irbesartan	143	92 (64.3)	0.10 (1.10)	(-2.06, 2.26)				
	Week 94	Sparsentan	139	105 (75.5)	0.16 (1.03)	(-1.87, 2.19)	-0.58 (1.50)	(-3.53, 2.37)	0.699	
		Irbesartan	143	96 (67.1)	0.74 (1.09)	(-1.39, 2.88)				
	Week 110	Sparsentan	139	103 (74.1)	0.43 (1.05)	(-1.64, 2.49)	0.20 (1.54)	(-2.84, 3.23)	0.899	
		Irbesartan	143	89 (62.2)	0.23 (1.13)	(-1.98, 2.45)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Female	Week 24	Sparsentan	63	48 (76.2)	5.84 (1.49)	(2.90, 8.77)	4.44 (2.25)	(0.00, 8.87)	0.050 *
		Irbesartan	59	38 (64.4)	1.40 (1.68)	(-1.90, 4.71)			
	Week 48	Sparsentan	63	50 (79.4)	4.96 (1.46)	(2.09, 7.83)	2.99 (2.24)	(-1.41, 7.38)	0.183
		Irbesartan	59	37 (62.7)	1.97 (1.69)	(-1.35, 5.30)			
	Week 70	Sparsentan	63	46 (73.0)	3.28 (1.52)	(0.30, 6.26)	0.73 (2.27)	(-3.73, 5.19)	0.748
		Irbesartan	59	38 (64.4)	2.55 (1.68)	(-0.75, 5.86)			
	Week 94	Sparsentan	63	46 (73.0)	4.57 (1.52)	(1.58, 7.56)	1.16 (2.33)	(-3.43, 5.75)	0.621
		Irbesartan	59	34 (57.6)	3.41 (1.76)	(-0.05, 6.87)			
	Week 110	Sparsentan	63	46 (73.0)	3.80 (1.53)	(0.79, 6.81)	0.96 (2.39)	(-3.73, 5.65)	0.687
		Irbesartan	59	32 (54.2)	2.84 (1.82)	(-0.75, 6.43)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.538
<= 45 years	Week 24	Sparsentan	96	71 (74.0)	1.21 (1.40)	(-1.54, 3.97)	1.94 (2.01)	(-2.02, 5.90)	0.336
		Irbesartan	99	67 (67.7)	-0.73 (1.44)	(-3.56, 2.11)			
	Week 48	Sparsentan	96	75 (78.1)	0.91 (1.36)	(-1.77, 3.59)	-0.25 (2.01)	(-4.20, 3.69)	0.900
		Irbesartan	99	62 (62.6)	1.16 (1.47)	(-1.73, 4.06)			
	Week 70	Sparsentan	96	72 (75.0)	1.13 (1.39)	(-1.60, 3.85)	-0.20 (2.03)	(-4.20, 3.79)	0.920
		Irbesartan	99	62 (62.6)	1.33 (1.49)	(-1.59, 4.25)			
	Week 94	Sparsentan	96	68 (70.8)	2.24 (1.42)	(-0.55, 5.03)	1.32 (2.05)	(-2.70, 5.34)	0.520
		Irbesartan	99	64 (64.6)	0.92 (1.47)	(-1.97, 3.82)			
	Week 110	Sparsentan	96	69 (71.9)	2.84 (1.43)	(0.03, 5.65)	2.45 (2.09)	(-1.66, 6.57)	0.242
		Irbesartan	99	59 (59.6)	0.39 (1.53)	(-2.63, 3.40)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 45 years	Week 24	Sparsentan	106	86 (81.1)	1.60 (1.01)	(-0.39, 3.58)	1.66 (1.55)	(-1.38, 4.71)	0.283
		Irbesartan	103	64 (62.1)	-0.07 (1.17)	(-2.37, 2.24)			
	Week 48	Sparsentan	106	87 (82.1)	2.16 (1.00)	(0.20, 4.13)	1.42 (1.54)	(-1.60, 4.45)	0.356
		Irbesartan	103	62 (60.2)	0.74 (1.17)	(-1.55, 3.04)			
	Week 70	Sparsentan	106	88 (83.0)	0.50 (1.00)	(-1.47, 2.47)	0.27 (1.52)	(-2.70, 3.25)	0.857
		Irbesartan	103	68 (66.0)	0.22 (1.13)	(-2.00, 2.45)			
	Week 94	Sparsentan	106	83 (78.3)	0.84 (1.03)	(-1.17, 2.86)	-1.00 (1.54)	(-4.03, 2.03)	0.518
		Irbesartan	103	66 (64.1)	1.84 (1.15)	(-0.42, 4.10)			
	Week 110	Sparsentan	106	80 (75.5)	0.18 (1.05)	(-1.88, 2.24)	-1.25 (1.59)	(-4.38, 1.87)	0.430
		Irbesartan	103	62 (60.2)	1.44 (1.19)	(-0.90, 3.78)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.495
<= 18 years	Week 24	Sparsentan	9	5 (55.6)	-1.59 (11.36)	(-28.17, 24.99)	-6.90 (19.01)	(-51.74, 37.95)	0.727
		Irbesartan	5	5 (100.0)	5.30 (14.27)	(-28.45, 39.06)			
	Week 48	Sparsentan	9	7 (77.8)	-4.77 (11.20)	(-31.20, 21.66)	-8.39 (19.02)	(-53.23, 36.46)	0.672
		Irbesartan	5	3 (60.0)	3.62 (14.41)	(-30.27, 37.50)			
	Week 70	Sparsentan	9	7 (77.8)	-4.97 (11.25)	(-31.44, 21.50)	-2.84 (19.04)	(-47.71, 42.02)	0.885
		Irbesartan	5	4 (80.0)	-2.13 (14.39)	(-36.00, 31.74)			
	Week 94	Sparsentan	9	5 (55.6)	0.36 (11.39)	(-26.24, 26.97)	-0.59 (19.21)	(-45.63, 44.45)	0.976
		Irbesartan	5	4 (80.0)	0.95 (14.50)	(-33.03, 34.94)			
	Week 110	Sparsentan	9	4 (44.4)	-0.16 (11.61)	(-27.00, 26.68)	-0.87 (19.68)	(-46.38, 44.65)	0.966
		Irbesartan	5	2 (40.0)	0.70 (15.02)	(-33.83, 35.24)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 18 to 40 years	Week 24	Sparsentan	102	79 (77.5)	2.06 (1.29)	(-0.47, 4.60)	3.46 (1.87)	(-0.21, 7.14)	0.065
		Irbesartan	109	71 (65.1)	-1.40 (1.36)	(-4.07, 1.26)			
	Week 48	Sparsentan	102	82 (80.4)	1.11 (1.27)	(-1.38, 3.59)	0.57 (1.87)	(-3.11, 4.24)	0.762
		Irbesartan	109	67 (61.5)	0.54 (1.38)	(-2.16, 3.24)			
	Week 70	Sparsentan	102	78 (76.5)	0.93 (1.29)	(-1.61, 3.47)	0.22 (1.89)	(-3.49, 3.93)	0.907
		Irbesartan	109	68 (62.4)	0.71 (1.38)	(-1.99, 3.42)			
	Week 94	Sparsentan	102	73 (71.6)	1.70 (1.33)	(-0.91, 4.31)	1.04 (1.92)	(-2.73, 4.80)	0.589
		Irbesartan	109	69 (63.3)	0.67 (1.38)	(-2.04, 3.37)			
	Week 110	Sparsentan	102	72 (70.6)	2.32 (1.35)	(-0.34, 4.98)	1.69 (1.96)	(-2.17, 5.55)	0.390
		Irbesartan	109	65 (59.6)	0.63 (1.42)	(-2.16, 3.42)			
> 40 years	Week 24	Sparsentan	91	73 (80.2)	1.36 (1.06)	(-0.73, 3.44)	1.16 (1.61)	(-2.02, 4.33)	0.474
		Irbesartan	88	55 (62.5)	0.20 (1.21)	(-2.19, 2.59)			
	Week 48	Sparsentan	91	73 (80.2)	2.30 (1.05)	(0.24, 4.37)	1.04 (1.61)	(-2.12, 4.20)	0.517
		Irbesartan	88	54 (61.4)	1.26 (1.21)	(-1.12, 3.65)			
	Week 70	Sparsentan	91	75 (82.4)	0.52 (1.04)	(-1.53, 2.58)	-0.61 (1.58)	(-3.72, 2.49)	0.699
		Irbesartan	88	58 (65.9)	1.14 (1.18)	(-1.19, 3.46)			
	Week 94	Sparsentan	91	73 (80.2)	0.92 (1.06)	(-1.16, 3.00)	-1.54 (1.60)	(-4.68, 1.59)	0.334
		Irbesartan	88	57 (64.8)	2.46 (1.19)	(0.12, 4.81)			
	Week 110	Sparsentan	91	73 (80.2)	0.20 (1.06)	(-1.89, 2.29)	-1.16 (1.63)	(-4.36, 2.04)	0.477
		Irbesartan	88	54 (61.4)	1.36 (1.23)	(-1.06, 3.78)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.296
North America	Week 24	Sparsentan	35	23 (65.7)	-2.17 (2.45)	(-7.02, 2.68)	0.18 (3.18)	(-6.10, 6.46)	0.954
		Irbesartan	46	35 (76.1)	-2.35 (2.00)	(-6.31, 1.61)			
	Week 48	Sparsentan	35	25 (71.4)	1.83 (2.38)	(-2.86, 6.53)	0.02 (3.16)	(-6.22, 6.26)	0.995
		Irbesartan	46	32 (69.6)	1.81 (2.05)	(-2.24, 5.87)			
	Week 70	Sparsentan	35	22 (62.9)	0.66 (2.49)	(-4.25, 5.58)	-0.16 (3.28)	(-6.63, 6.32)	0.962
		Irbesartan	46	30 (65.2)	0.82 (2.12)	(-3.36, 5.00)			
	Week 94	Sparsentan	35	23 (65.7)	0.70 (2.46)	(-4.17, 5.56)	-0.66 (3.27)	(-7.13, 5.80)	0.839
		Irbesartan	46	30 (65.2)	1.36 (2.14)	(-2.87, 5.59)			
	Week 110	Sparsentan	35	21 (60.0)	1.16 (2.57)	(-3.91, 6.23)	-0.87 (3.35)	(-7.48, 5.75)	0.796
		Irbesartan	46	31 (67.4)	2.03 (2.14)	(-2.21, 6.26)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Europe	Week 24	Sparsentan	98	73 (74.5)	3.32 (1.18)	(0.99, 5.64)	3.53 (1.75)	(0.10, 6.97)	0.044 *
		Irbesartan	115	62 (53.9)	-0.22 (1.29)	(-2.75, 2.31)			
	Week 48	Sparsentan	98	76 (77.6)	3.65 (1.16)	(1.36, 5.93)	3.38 (1.73)	(-0.02, 6.79)	0.051
		Irbesartan	115	59 (51.3)	0.26 (1.28)	(-2.25, 2.78)			
	Week 70	Sparsentan	98	75 (76.5)	1.42 (1.18)	(-0.90, 3.74)	1.45 (1.70)	(-1.90, 4.80)	0.395
		Irbesartan	115	69 (60.0)	-0.03 (1.22)	(-2.43, 2.37)			
	Week 94	Sparsentan	98	68 (69.4)	2.18 (1.22)	(-0.22, 4.57)	1.43 (1.73)	(-1.96, 4.83)	0.408
		Irbesartan	115	70 (60.9)	0.74 (1.22)	(-1.65, 3.14)			
	Week 110	Sparsentan	98	67 (68.4)	2.94 (1.24)	(0.49, 5.39)	2.65 (1.80)	(-0.89, 6.18)	0.142
		Irbesartan	115	61 (53.0)	0.29 (1.29)	(-2.24, 2.83)			
Asia Pacific	Week 24	Sparsentan	69	61 (88.4)	1.24 (1.39)	(-1.49, 3.98)	1.45 (2.32)	(-3.12, 6.01)	0.533
		Irbesartan	41	34 (82.9)	-0.20 (1.85)	(-3.84, 3.44)			
	Week 48	Sparsentan	69	61 (88.4)	-0.33 (1.38)	(-3.04, 2.39)	-0.42 (2.34)	(-5.02, 4.18)	0.858
		Irbesartan	41	33 (80.5)	0.09 (1.88)	(-3.60, 3.79)			
	Week 70	Sparsentan	69	63 (91.3)	0.69 (1.37)	(-2.00, 3.37)	-0.66 (2.39)	(-5.37, 4.05)	0.783
		Irbesartan	41	31 (75.6)	1.34 (1.95)	(-2.49, 5.18)			
	Week 94	Sparsentan	69	60 (87.0)	1.69 (1.39)	(-1.05, 4.43)	-0.12 (2.44)	(-4.92, 4.67)	0.960
		Irbesartan	41	30 (73.2)	1.81 (1.98)	(-2.09, 5.71)			
	Week 110	Sparsentan	69	61 (88.4)	0.67 (1.39)	(-2.06, 3.41)	0.88 (2.47)	(-3.97, 5.73)	0.722
		Irbesartan	41	29 (70.7)	-0.20 (2.02)	(-4.18, 3.77)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
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Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.826
< 27 kg/m**2	Week 24	Sparsentan	83	66 (79.5)	2.01 (1.45)	(-0.85, 4.86)	2.26 (2.06)	(-1.79, 6.31)	0.273
		Irbesartan	94	66 (70.2)	-0.25 (1.45)	(-3.10, 2.60)			
	Week 48	Sparsentan	83	66 (79.5)	1.36 (1.44)	(-1.48, 4.20)	0.31 (2.07)	(-3.76, 4.38)	0.883
		Irbesartan	94	60 (63.8)	1.05 (1.47)	(-1.85, 3.95)			
	Week 70	Sparsentan	83	64 (77.1)	0.70 (1.46)	(-2.18, 3.58)	-0.02 (2.10)	(-4.14, 4.11)	0.993
		Irbesartan	94	59 (62.8)	0.72 (1.49)	(-2.22, 3.66)			
	Week 94	Sparsentan	83	63 (75.9)	2.02 (1.48)	(-0.89, 4.92)	1.07 (2.10)	(-3.06, 5.21)	0.610
		Irbesartan	94	62 (66.0)	0.94 (1.49)	(-1.98, 3.87)			
	Week 110	Sparsentan	83	63 (75.9)	1.14 (1.49)	(-1.79, 4.08)	0.40 (2.15)	(-3.83, 4.63)	0.852
		Irbesartan	94	57 (60.6)	0.74 (1.55)	(-2.30, 3.78)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 27 kg/m**2	Week 24	Sparsentan	119	91 (76.5)	1.04 (1.03)	(-0.99, 3.07)	1.60 (1.61)	(-1.55, 4.76)	0.318
		Irbesartan	107	65 (60.7)	-0.57 (1.23)	(-2.97, 1.84)			
	Week 48	Sparsentan	119	96 (80.7)	1.63 (1.01)	(-0.35, 3.60)	0.40 (1.58)	(-2.71, 3.50)	
		Irbesartan	107	64 (59.8)	1.23 (1.22)	(-1.17, 3.63)			
	Week 70	Sparsentan	119	96 (80.7)	0.57 (1.01)	(-1.41, 2.55)	-0.58 (1.55)	(-3.62, 2.46)	
		Irbesartan	107	71 (66.4)	1.14 (1.17)	(-1.16, 3.45)			
	Week 94	Sparsentan	119	88 (73.9)	0.82 (1.05)	(-1.24, 2.88)	-1.41 (1.60)	(-4.55, 1.72)	
		Irbesartan	107	67 (62.6)	2.23 (1.20)	(-0.13, 4.60)			
	Week 110	Sparsentan	119	86 (72.3)	1.52 (1.07)	(-0.57, 3.62)	0.42 (1.64)	(-2.80, 3.64)	
		Irbesartan	107	63 (58.9)	1.10 (1.24)	(-1.34, 3.55)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
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Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan						Interaction:	0.614
eGFR Low and UP High	Week 24	Sparsentan	71	54 (76.1)	0.82 (1.33)	(-1.80, 3.44)	3.60 (2.03)	(-0.39, 7.60)	0.077
		Irbesartan	74	40 (54.1)	-2.78 (1.53)	(-5.79, 0.22)			
	Week 48	Sparsentan	71	51 (71.8)	1.80 (1.35)	(-0.85, 4.45)	1.81 (2.05)	(-2.23, 5.84)	0.379
		Irbesartan	74	37 (50.0)	-0.00 (1.54)	(-3.04, 3.03)			
	Week 70	Sparsentan	71	57 (80.3)	-0.56 (1.31)	(-3.14, 2.02)	-0.24 (2.04)	(-4.25, 3.77)	0.907
		Irbesartan	74	37 (50.0)	-0.32 (1.56)	(-3.39, 2.75)			
	Week 94	Sparsentan	71	51 (71.8)	0.46 (1.35)	(-2.20, 3.12)	0.33 (2.08)	(-3.77, 4.42)	0.876
		Irbesartan	74	37 (50.0)	0.13 (1.58)	(-2.97, 3.24)			
	Week 110	Sparsentan	71	53 (74.6)	0.25 (1.35)	(-2.42, 2.91)	0.09 (2.11)	(-4.06, 4.24)	0.966
		Irbesartan	74	36 (48.6)	0.16 (1.62)	(-3.03, 3.34)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
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A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
					Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
eGFR Low and UP Low	Week 24	Sparsentan	55	43 (78.2)	1.33 (1.61)	(-1.84, 4.50)	-2.88 (2.39)	(-7.57, 1.82)	0.229
		Irbesartan	55	37 (67.3)	4.21 (1.74)	(0.79, 7.64)			
	Week 48	Sparsentan	55	43 (78.2)	1.07 (1.60)	(-2.07, 4.22)	-1.57 (2.39)	(-6.27, 3.14)	0.513
		Irbesartan	55	35 (63.6)	2.64 (1.77)	(-0.85, 6.13)			
	Week 70	Sparsentan	55	42 (76.4)	1.27 (1.62)	(-1.93, 4.46)	-0.29 (2.35)	(-4.90, 4.33)	0.902
		Irbesartan	55	39 (70.9)	1.55 (1.69)	(-1.76, 4.87)			
	Week 94	Sparsentan	55	41 (74.5)	2.50 (1.65)	(-0.73, 5.74)	-0.74 (2.36)	(-5.39, 3.91)	0.755
		Irbesartan	55	39 (70.9)	3.24 (1.69)	(-0.08, 6.56)			
	Week 110	Sparsentan	55	37 (67.3)	1.81 (1.73)	(-1.59, 5.21)	-0.82 (2.50)	(-5.73, 4.09)	0.743
		Irbesartan	55	34 (61.8)	2.63 (1.80)	(-0.91, 6.17)			
eGFR High and UP High	Week 24	Sparsentan	37	26 (70.3)	2.60 (2.37)	(-2.08, 7.29)	3.17 (3.42)	(-3.60, 9.94)	0.356
		Irbesartan	36	26 (72.2)	-0.56 (2.47)	(-5.45, 4.32)			
	Week 48	Sparsentan	37	33 (89.2)	0.99 (2.20)	(-3.37, 5.34)	-1.17 (3.33)	(-7.77, 5.42)	0.725
		Irbesartan	36	24 (66.7)	2.16 (2.50)	(-2.79, 7.11)			
	Week 70	Sparsentan	37	31 (83.8)	1.34 (2.25)	(-3.11, 5.80)	1.27 (3.36)	(-5.39, 7.92)	0.707
		Irbesartan	36	25 (69.4)	0.08 (2.50)	(-4.87, 5.02)			
	Week 94	Sparsentan	37	27 (73.0)	2.67 (2.36)	(-1.98, 7.33)	0.29 (3.46)	(-6.55, 7.14)	0.933
		Irbesartan	36	24 (66.7)	2.38 (2.53)	(-2.63, 7.39)			
	Week 110	Sparsentan	37	26 (70.3)	2.94 (2.43)	(-1.87, 7.75)	4.33 (3.58)	(-2.75, 11.42)	0.228
		Irbesartan	36	22 (61.1)	-1.39 (2.63)	(-6.59, 3.80)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
eGFR High and UP Low	Week 24	Sparsentan	39	34 (87.2)	1.93 (1.74)	(-1.51, 5.36)	4.56 (2.58)	(-0.53, 9.65)	0.079
		Irbesartan	37	28 (75.7)	-2.64 (1.90)	(-6.39, 1.11)			
	Week 48	Sparsentan	39	35 (89.7)	2.32 (1.73)	(-1.08, 5.72)	2.01 (2.57)	(-3.05, 7.07)	0.435
		Irbesartan	37	28 (75.7)	0.31 (1.89)	(-3.42, 4.04)			
	Week 70	Sparsentan	39	30 (76.9)	1.38 (1.83)	(-2.22, 4.98)	-1.53 (2.63)	(-6.71, 3.65)	0.561
		Irbesartan	37	29 (78.4)	2.91 (1.88)	(-0.80, 6.61)			
	Week 94	Sparsentan	39	32 (82.1)	0.41 (1.79)	(-3.12, 3.93)	-0.79 (2.58)	(-5.88, 4.30)	0.760
		Irbesartan	37	30 (81.1)	1.20 (1.85)	(-2.46, 4.85)			
	Week 110	Sparsentan	39	33 (84.6)	1.08 (1.77)	(-2.41, 4.58)	-0.92 (2.59)	(-6.04, 4.19)	0.722
		Irbesartan	37	29 (78.4)	2.00 (1.89)	(-1.71, 5.72)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.040 #
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	98 (77.2)	0.68 (1.04)	(-1.37, 2.73)	0.29 (1.56)	(-2.77, 3.35)	0.854
		Irbesartan	129	80 (62.0)	0.39 (1.15)	(-1.88, 2.66)			
	Week 48	Sparsentan	127	95 (74.8)	1.65 (1.05)	(-0.41, 3.72)	0.75 (1.58)	(-2.35, 3.86)	0.634
		Irbesartan	129	74 (57.4)	0.90 (1.18)	(-1.42, 3.22)			
	Week 70	Sparsentan	127	100 (78.7)	-0.07 (1.03)	(-2.10, 1.97)	-0.39 (1.56)	(-3.44, 2.67)	0.805
		Irbesartan	129	78 (60.5)	0.32 (1.16)	(-1.96, 2.60)			
	Week 94	Sparsentan	127	92 (72.4)	1.34 (1.07)	(-0.76, 3.44)	-0.34 (1.59)	(-3.45, 2.78)	0.833
		Irbesartan	129	77 (59.7)	1.67 (1.17)	(-0.63, 3.97)			
	Week 110	Sparsentan	127	92 (72.4)	1.11 (1.08)	(-1.01, 3.22)	-0.41 (1.63)	(-3.61, 2.78)	0.800
		Irbesartan	129	72 (55.8)	1.52 (1.22)	(-0.87, 3.90)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	38 (77.6)	0.98 (1.83)	(-2.63, 4.59)	-0.28 (2.73)	(-5.67, 5.12)	0.919	
		Irbesartan	48	31 (64.6)	1.26 (2.02)	(-2.72, 5.24)				
	Week 48	Sparsentan	49	44 (89.8)	0.44 (1.75)	(-3.03, 3.91)	0.01 (2.66)	(-5.24, 5.25)	0.998	
		Irbesartan	48	32 (66.7)	0.43 (1.98)	(-3.47, 4.33)				
	Week 70	Sparsentan	49	38 (77.6)	0.29 (1.81)	(-3.29, 3.87)	-0.45 (2.70)	(-5.78, 4.89)	0.869	
		Irbesartan	48	32 (66.7)	0.73 (1.98)	(-3.18, 4.65)				
	Week 94	Sparsentan	49	39 (79.6)	-0.42 (1.82)	(-4.02, 3.18)	-1.65 (2.69)	(-6.97, 3.67)	0.542	
		Irbesartan	48	34 (70.8)	1.23 (1.97)	(-2.66, 5.11)				
	Week 110	Sparsentan	49	36 (73.5)	0.38 (1.89)	(-3.35, 4.10)	1.20 (2.77)	(-4.27, 6.66)	0.666	
		Irbesartan	48	32 (66.7)	-0.82 (2.01)	(-4.80, 3.16)				
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	21 (80.8)	7.02 (2.31)	(2.45, 11.58)	13.29 (3.31)	(6.76, 19.82)	<0.001 *	
		Irbesartan	25	20 (80.0)	-6.27 (2.37)	(-10.95, -1.60)				
	Week 48	Sparsentan	26	23 (88.5)	3.45 (2.22)	(-0.95, 7.85)	0.48 (3.35)	(-6.13, 7.09)	0.886	
		Irbesartan	25	18 (72.0)	2.97 (2.48)	(-1.92, 7.87)				
	Week 70	Sparsentan	26	22 (84.6)	4.82 (2.27)	(0.33, 9.32)	1.28 (3.31)	(-5.26, 7.82)	0.700	
		Irbesartan	25	20 (80.0)	3.55 (2.38)	(-1.16, 8.25)				
	Week 94	Sparsentan	26	20 (76.9)	5.49 (2.36)	(0.82, 10.15)	3.54 (3.40)	(-3.17, 10.26)	0.299	
		Irbesartan	25	19 (76.0)	1.94 (2.43)	(-2.86, 6.75)				
	Week 110	Sparsentan	26	21 (80.8)	4.69 (2.32)	(0.10, 9.27)	2.20 (3.46)	(-4.64, 9.04)	0.526	
		Irbesartan	25	17 (68.0)	2.49 (2.56)	(-2.58, 7.55)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.027 #
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	64 (78.0)	2.68 (1.37)	(-0.02, 5.38)	2.92 (2.10)	(-1.20, 7.04)	0.164
		Irbesartan	80	48 (60.0)	-0.25 (1.58)	(-3.36, 2.87)			
	Week 48	Sparsentan	82	61 (74.4)	1.92 (1.40)	(-0.82, 4.66)	1.43 (2.16)	(-2.81, 5.68)	0.507
		Irbesartan	80	43 (53.8)	0.49 (1.65)	(-2.75, 3.72)			
	Week 70	Sparsentan	82	65 (79.3)	0.37 (1.36)	(-2.30, 3.05)	1.37 (2.12)	(-2.79, 5.52)	0.519
		Irbesartan	80	45 (56.3)	-0.99 (1.62)	(-4.17, 2.18)			
	Week 94	Sparsentan	82	60 (73.2)	2.12 (1.41)	(-0.65, 4.89)	0.47 (2.13)	(-3.71, 4.65)	0.827
		Irbesartan	80	47 (58.8)	1.65 (1.59)	(-1.48, 4.78)			
	Week 110	Sparsentan	82	56 (68.3)	1.90 (1.46)	(-0.98, 4.77)	-0.00 (2.21)	(-4.34, 4.33)	0.998
		Irbesartan	80	44 (55.0)	1.90 (1.65)	(-1.34, 5.14)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	34 (75.6)	-3.08 (1.56)	(-6.16, 0.00)	-4.60 (2.26)	(-9.04, -0.15)	0.043 *
		Irbesartan	49	32 (65.3)	1.52 (1.62)	(-1.67, 4.70)			
	Week 48	Sparsentan	45	34 (75.6)	1.44 (1.55)	(-1.60, 4.49)	-0.37 (2.25)	(-4.80, 4.06)	0.869
		Irbesartan	49	31 (63.3)	1.81 (1.62)	(-1.39, 5.01)			
	Week 70	Sparsentan	45	35 (77.8)	-0.64 (1.54)	(-3.68, 2.40)	-2.96 (2.24)	(-7.37, 1.45)	0.187
		Irbesartan	49	33 (67.3)	2.32 (1.61)	(-0.85, 5.50)			
	Week 94	Sparsentan	45	32 (71.1)	-0.01 (1.59)	(-3.14, 3.13)	-1.85 (2.31)	(-6.39, 2.70)	0.424
		Irbesartan	49	30 (61.2)	1.84 (1.66)	(-1.44, 5.12)			
	Week 110	Sparsentan	45	36 (80.0)	-0.30 (1.55)	(-3.36, 2.75)	-1.36 (2.33)	(-5.95, 3.23)	0.560
		Irbesartan	49	28 (57.1)	1.06 (1.73)	(-2.35, 4.47)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	38 (77.6)	0.98 (1.83)	(-2.63, 4.59)	-0.28 (2.73)	(-5.67, 5.12)	0.919
		Irbesartan	48	31 (64.6)	1.26 (2.02)	(-2.72, 5.24)			
	Week 48	Sparsentan	49	44 (89.8)	0.44 (1.75)	(-3.03, 3.91)	0.01 (2.66)	(-5.24, 5.25)	0.998
		Irbesartan	48	32 (66.7)	0.43 (1.98)	(-3.47, 4.33)			
	Week 70	Sparsentan	49	38 (77.6)	0.29 (1.81)	(-3.29, 3.87)	-0.45 (2.70)	(-5.78, 4.89)	0.869
		Irbesartan	48	32 (66.7)	0.73 (1.98)	(-3.18, 4.65)			
	Week 94	Sparsentan	49	39 (79.6)	-0.42 (1.82)	(-4.02, 3.18)	-1.65 (2.69)	(-6.97, 3.67)	0.542
		Irbesartan	48	34 (70.8)	1.23 (1.97)	(-2.66, 5.11)			
	Week 110	Sparsentan	49	36 (73.5)	0.38 (1.89)	(-3.35, 4.10)	1.20 (2.77)	(-4.27, 6.66)	0.666
		Irbesartan	48	32 (66.7)	-0.82 (2.01)	(-4.80, 3.16)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	21 (80.8)	7.02 (2.31)	(2.45, 11.58)	13.29 (3.31)	(6.76, 19.82)	<0.001 *
		Irbesartan	25	20 (80.0)	-6.27 (2.37)	(-10.95, -1.60)			
	Week 48	Sparsentan	26	23 (88.5)	3.45 (2.22)	(-0.95, 7.85)	0.48 (3.35)	(-6.13, 7.09)	0.886
		Irbesartan	25	18 (72.0)	2.97 (2.48)	(-1.92, 7.87)			
	Week 70	Sparsentan	26	22 (84.6)	4.82 (2.27)	(0.33, 9.32)	1.28 (3.31)	(-5.26, 7.82)	0.700
		Irbesartan	25	20 (80.0)	3.55 (2.38)	(-1.16, 8.25)			
	Week 94	Sparsentan	26	20 (76.9)	5.49 (2.36)	(0.82, 10.15)	3.54 (3.40)	(-3.17, 10.26)	0.299
		Irbesartan	25	19 (76.0)	1.94 (2.43)	(-2.86, 6.75)			
	Week 110	Sparsentan	26	21 (80.8)	4.69 (2.32)	(0.10, 9.27)	2.20 (3.46)	(-4.64, 9.04)	0.526
		Irbesartan	25	17 (68.0)	2.49 (2.56)	(-2.58, 7.55)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline urine protein excretion		Overall	Sparsentan						Interaction:	0.165
<= 1.75 g/day	Week 24	Sparsentan	98	80 (81.6)	0.87 (1.21)	(-1.50, 3.25)	0.66 (1.89)	(-3.06, 4.38)	0.728	
		Irbesartan	93	55 (59.1)	0.22 (1.45)	(-2.64, 3.07)				
	Week 48	Sparsentan	98	82 (83.7)	1.36 (1.19)	(-0.98, 3.70)	0.39 (1.85)	(-3.25, 4.04)	0.832	
		Irbesartan	93	57 (61.3)	0.97 (1.42)	(-1.82, 3.76)				
	Week 70	Sparsentan	98	75 (76.5)	0.28 (1.24)	(-2.15, 2.71)	-1.58 (1.86)	(-5.23, 2.08)	0.397	
		Irbesartan	93	61 (65.6)	1.86 (1.38)	(-0.86, 4.57)				
	Week 94	Sparsentan	98	76 (77.6)	1.24 (1.24)	(-1.18, 3.67)	-0.23 (1.84)	(-3.84, 3.39)	0.903	
		Irbesartan	93	63 (67.7)	1.47 (1.36)	(-1.21, 4.14)				
	Week 110	Sparsentan	98	75 (76.5)	0.93 (1.25)	(-1.52, 3.38)	-0.75 (1.90)	(-4.48, 2.98)	0.694	
		Irbesartan	93	57 (61.3)	1.68 (1.43)	(-1.13, 4.48)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 1.75 g/day	Week 24	Sparsentan	104	77 (74.0)	2.28 (1.21)	(-0.10, 4.66)	3.56 (1.73)	(0.16, 6.96)	0.040 *
		Irbesartan	109	76 (69.7)	-1.28 (1.23)	(-3.70, 1.14)			
	Week 48	Sparsentan	104	80 (76.9)	1.94 (1.19)	(-0.40, 4.27)	1.13 (1.73)	(-2.27, 4.54)	0.513
		Irbesartan	109	67 (61.5)	0.80 (1.26)	(-1.67, 3.28)			
	Week 70	Sparsentan	104	85 (81.7)	1.33 (1.18)	(-0.99, 3.64)	1.64 (1.73)	(-1.76, 5.04)	0.343
		Irbesartan	109	69 (63.3)	-0.31 (1.26)	(-2.79, 2.17)			
	Week 94	Sparsentan	104	75 (72.1)	1.81 (1.22)	(-0.59, 4.21)	0.70 (1.78)	(-2.79, 4.19)	0.693
		Irbesartan	109	67 (61.5)	1.11 (1.28)	(-1.42, 3.63)			
	Week 110	Sparsentan	104	74 (71.2)	2.01 (1.25)	(-0.44, 4.46)	2.22 (1.82)	(-1.36, 5.80)	0.223
		Irbesartan	109	64 (58.7)	-0.21 (1.32)	(-2.81, 2.39)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline use of antihypertensives		Overall	Sparsentan						Interaction:	0.940
Yes	Week 24	Sparsentan	90	66 (73.3)	2.11 (1.09)	(-0.04, 4.25)	3.16 (1.67)	(-0.11, 6.44)	0.058	
		Irbesartan	88	49 (55.7)	-1.06 (1.26)	(-3.53, 1.41)				
	Week 48	Sparsentan	90	65 (72.2)	1.64 (1.09)	(-0.49, 3.78)	0.70 (1.66)	(-2.57, 3.97)	0.673	
		Irbesartan	88	47 (53.4)	0.94 (1.26)	(-1.53, 3.41)				
	Week 70	Sparsentan	90	68 (75.6)	-0.41 (1.08)	(-2.53, 1.71)	-0.59 (1.65)	(-3.84, 2.65)	0.720	
		Irbesartan	88	49 (55.7)	0.18 (1.25)	(-2.27, 2.63)				
	Week 94	Sparsentan	90	63 (70.0)	0.25 (1.10)	(-1.93, 2.42)	-0.55 (1.64)	(-3.77, 2.68)	0.738	
		Irbesartan	88	54 (61.4)	0.79 (1.21)	(-1.58, 3.17)				
	Week 110	Sparsentan	90	63 (70.0)	1.27 (1.12)	(-0.94, 3.47)	0.96 (1.68)	(-2.34, 4.26)	0.567	
		Irbesartan	88	51 (58.0)	0.31 (1.24)	(-2.13, 2.75)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
No	Week 24	Sparsentan	112	91 (81.3)	1.12 (1.23)	(-1.28, 3.53)	1.39 (1.78)	(-2.11, 4.89)	0.437
		Irbesartan	114	82 (71.9)	-0.26 (1.29)	(-2.80, 2.28)			
	Week 48	Sparsentan	112	97 (86.6)	1.71 (1.19)	(-0.62, 4.05)	0.85 (1.77)	(-2.63, 4.33)	0.631
		Irbesartan	114	77 (67.5)	0.86 (1.31)	(-1.72, 3.44)			
	Week 70	Sparsentan	112	92 (82.1)	1.81 (1.22)	(-0.59, 4.20)	0.71 (1.78)	(-2.78, 4.21)	0.689
		Irbesartan	114	81 (71.1)	1.09 (1.30)	(-1.45, 3.64)			
	Week 94	Sparsentan	112	88 (78.6)	2.53 (1.24)	(0.08, 4.97)	0.66 (1.83)	(-2.93, 4.24)	0.720
		Irbesartan	114	76 (66.7)	1.87 (1.34)	(-0.75, 4.50)			
	Week 110	Sparsentan	112	86 (76.8)	1.59 (1.26)	(-0.89, 4.08)	0.17 (1.88)	(-3.53, 3.87)	0.929
		Irbesartan	114	70 (61.4)	1.42 (1.40)	(-1.32, 4.16)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.080
<= 5 years	Week 24	Sparsentan	113	88 (77.9)	3.42 (1.10)	(1.25, 5.59)	4.76 (1.57)	(1.68, 7.84)	0.002 *
		Irbesartan	127	86 (67.7)	-1.34 (1.11)	(-3.52, 0.84)			
	Week 48	Sparsentan	113	92 (81.4)	3.43 (1.08)	(1.31, 5.54)	2.50 (1.54)	(-0.53, 5.54)	0.105
		Irbesartan	127	85 (66.9)	0.92 (1.10)	(-1.25, 3.09)			
	Week 70	Sparsentan	113	90 (79.6)	1.34 (1.09)	(-0.80, 3.48)	0.21 (1.55)	(-2.84, 3.26)	0.893
		Irbesartan	127	86 (67.7)	1.13 (1.11)	(-1.04, 3.30)			
	Week 94	Sparsentan	113	86 (76.1)	1.82 (1.11)	(-0.36, 4.00)	0.47 (1.57)	(-2.61, 3.55)	0.764
		Irbesartan	127	86 (67.7)	1.35 (1.11)	(-0.83, 3.52)			
	Week 110	Sparsentan	113	88 (77.9)	2.21 (1.11)	(0.03, 4.40)	1.25 (1.60)	(-1.90, 4.39)	0.436
		Irbesartan	127	80 (63.0)	0.97 (1.15)	(-1.30, 3.23)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 5 years	Week 24	Sparsentan	89	69 (77.5)	-0.86 (1.32)	(-3.46, 1.75)	-1.97 (2.13)	(-6.15, 2.21)	0.355
		Irbesartan	75	45 (60.0)	1.11 (1.66)	(-2.15, 4.38)			
	Week 48	Sparsentan	89	70 (78.7)	-0.55 (1.31)	(-3.13, 2.03)	-1.59 (2.17)	(-5.86, 2.69)	
		Irbesartan	75	39 (52.0)	1.04 (1.73)	(-2.36, 4.44)			
	Week 70	Sparsentan	89	70 (78.7)	0.25 (1.32)	(-2.34, 2.85)	0.29 (2.12)	(-3.88, 4.47)	
		Irbesartan	75	44 (58.7)	-0.04 (1.66)	(-3.30, 3.22)			
	Week 94	Sparsentan	89	65 (73.0)	1.30 (1.36)	(-1.37, 3.97)	-0.27 (2.15)	(-4.50, 3.97)	
		Irbesartan	75	44 (58.7)	1.57 (1.67)	(-1.71, 4.85)			
	Week 110	Sparsentan	89	61 (68.5)	0.53 (1.41)	(-2.24, 3.29)	-0.37 (2.23)	(-4.76, 4.01)	
		Irbesartan	75	41 (54.7)	0.90 (1.73)	(-2.49, 4.29)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.022 #
Yes	Week 24	Sparsentan	155	119 (76.8)	3.34 (0.84)	(1.70, 4.98)	1.70 (1.24)	(-0.73, 4.13)	0.170
		Irbesartan	161	100 (62.1)	1.64 (0.91)	(-0.14, 3.43)			
	Week 48	Sparsentan	155	121 (78.1)	2.20 (0.82)	(0.59, 3.82)	-0.90 (1.24)	(-3.33, 1.53)	0.468
		Irbesartan	161	95 (59.0)	3.10 (0.92)	(1.29, 4.91)			
	Week 70	Sparsentan	155	123 (79.4)	1.82 (0.82)	(0.20, 3.43)	-0.97 (1.22)	(-3.37, 1.43)	0.428
		Irbesartan	161	100 (62.1)	2.79 (0.91)	(1.01, 4.57)			
	Week 94	Sparsentan	155	114 (73.5)	2.05 (0.85)	(0.39, 3.72)	-0.19 (1.24)	(-2.63, 2.24)	0.876
		Irbesartan	161	101 (62.7)	2.25 (0.90)	(0.47, 4.02)			
	Week 110	Sparsentan	155	112 (72.3)	1.89 (0.86)	(0.19, 3.58)	-0.69 (1.28)	(-3.20, 1.82)	0.589
		Irbesartan	161	92 (57.1)	2.58 (0.95)	(0.72, 4.44)			

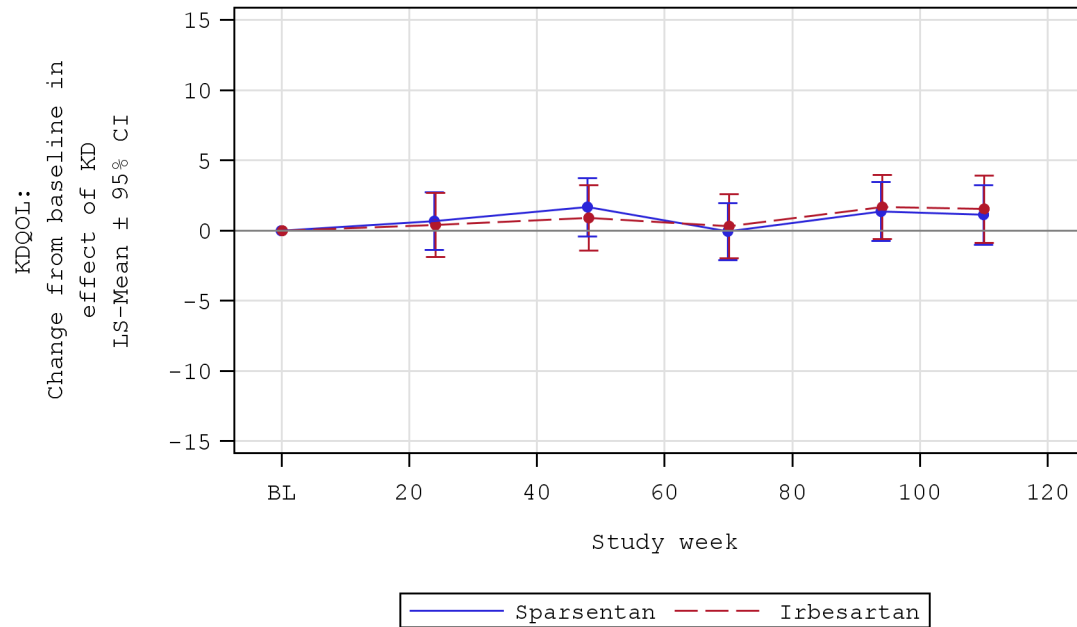
N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
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 Source Data: aqs, created on: 05MAR2024

Table PT2KEFC\_FSCM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in effect of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
No	Week 24	Sparsentan	47	38 (80.9)	-3.81 (2.36)	(-8.46, 0.84)	4.36 (3.53)	(-2.60, 11.32)	0.218
		Irbesartan	41	31 (75.6)	-8.17 (2.61)	(-13.33, -3.00)			
	Week 48	Sparsentan	47	41 (87.2)	-0.12 (2.29)	(-4.65, 4.41)	6.91 (3.50)	(0.00, 13.82)	0.050 *
		Irbesartan	41	29 (70.7)	-7.03 (2.63)	(-12.22, -1.84)			
	Week 70	Sparsentan	47	37 (78.7)	-2.14 (2.36)	(-6.81, 2.52)	4.11 (3.55)	(-2.88, 11.11)	0.248
		Irbesartan	41	30 (73.2)	-6.26 (2.63)	(-11.45, -1.07)			
	Week 94	Sparsentan	47	37 (78.7)	-0.03 (2.39)	(-4.75, 4.69)	1.92 (3.61)	(-5.20, 9.05)	0.595
		Irbesartan	41	29 (70.7)	-1.95 (2.69)	(-7.26, 3.35)			
	Week 110	Sparsentan	47	37 (78.7)	0.17 (2.41)	(-4.60, 4.94)	5.16 (3.66)	(-2.07, 12.38)	0.161
		Irbesartan	41	29 (70.7)	-4.99 (2.73)	(-10.38, 0.40)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

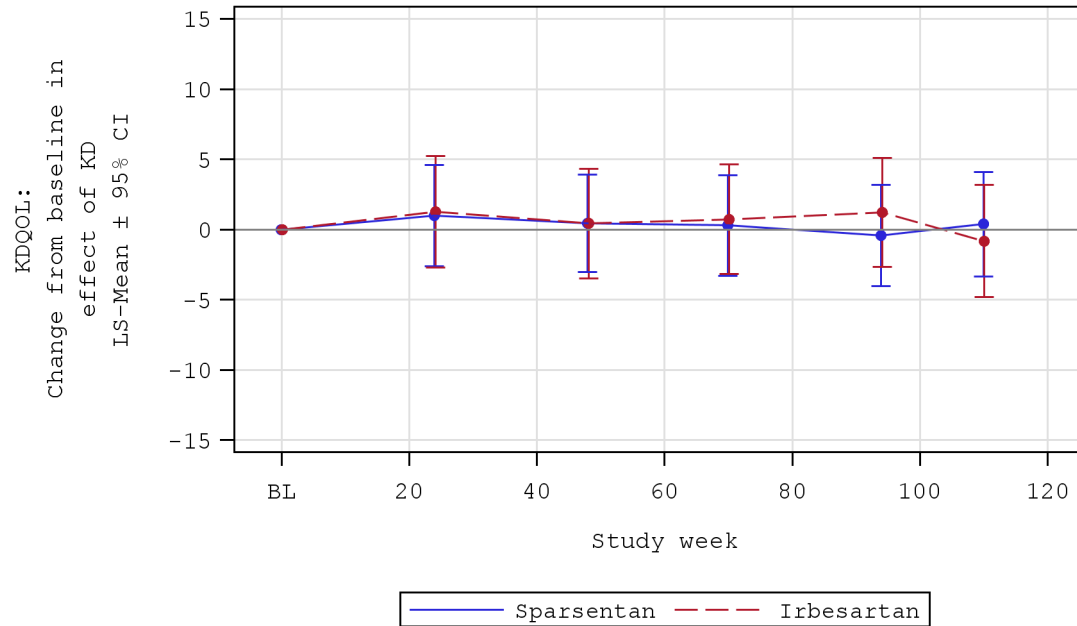
Figure PF2KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1: < 60 mL/min/1.73 m\*\*2



Sparsentan	98	95	100	92	92
Irbesartan	80	74	78	77	72

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KEFC\_FSCM

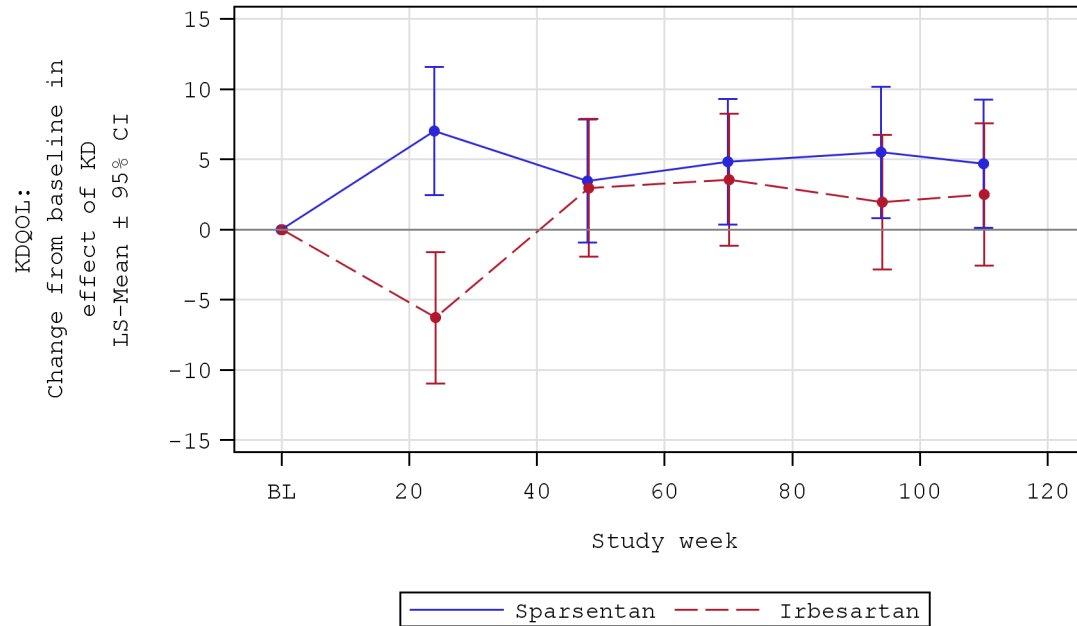
Figure PF2KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1: 60 to < 90 mL/min/1.73 m\*\*2



Sparsentan	38	44	38	39	36
Irbesartan	31	32	32	34	32

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KEFC\_FSCM

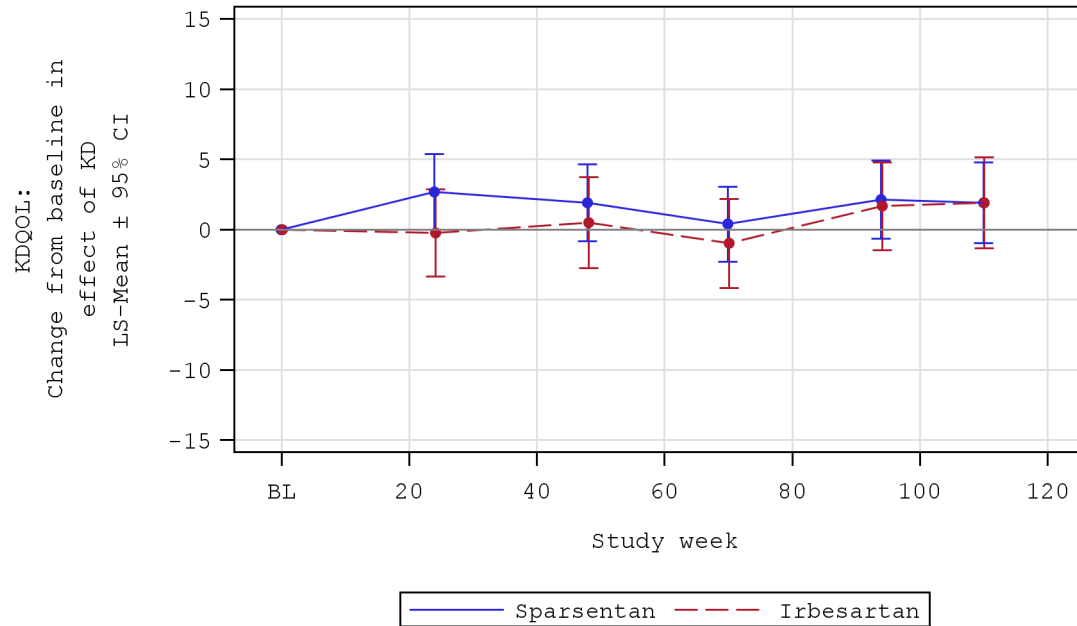
Figure PF2KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 1:  $\geq 90$  mL/min/1.73 m<sup>2</sup>



Sparsentan	21	23	22	20	21
Irbesartan	20	18	20	19	17

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low =  $30 < eGFR < 60$  ml/min/1.73m<sup>2</sup>, eGFR High  $\geq 60$  ml/min/1.73m<sup>2</sup>, UP Low =  $\leq 1.75$  g/day, UP High =  $> 1.75$  g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KEFC\_FSCM

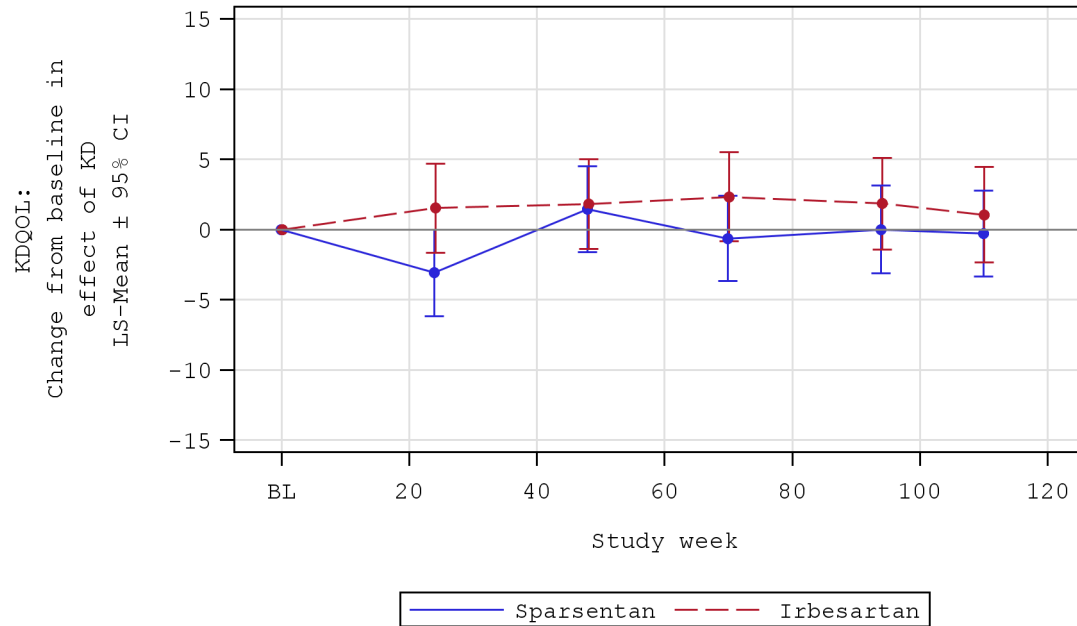
Figure PF2KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: < 45 mL/min/1.73 m\*\*2



Sparsentan	64	61	65	60	56
Irbesartan	48	43	45	47	44

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KEFC\_FSCM

Figure PF2KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: 45 to < 60 mL/min/1.73 m\*\*2

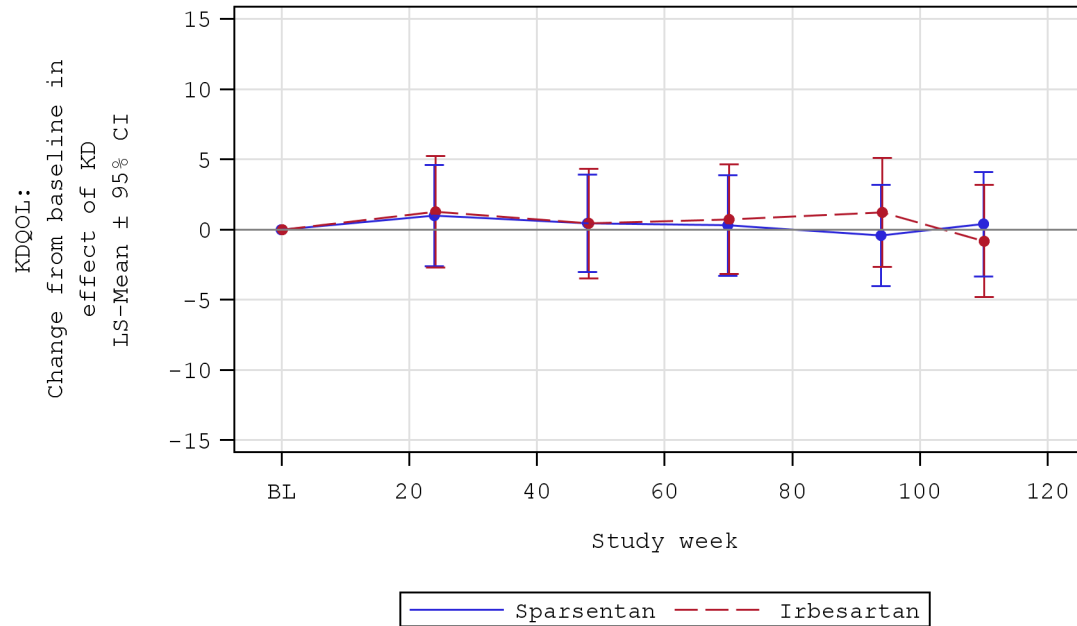


Sparsentan	34	34	35	32	36
Irbesartan	32	31	33	30	28

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KEFC\_FSCM



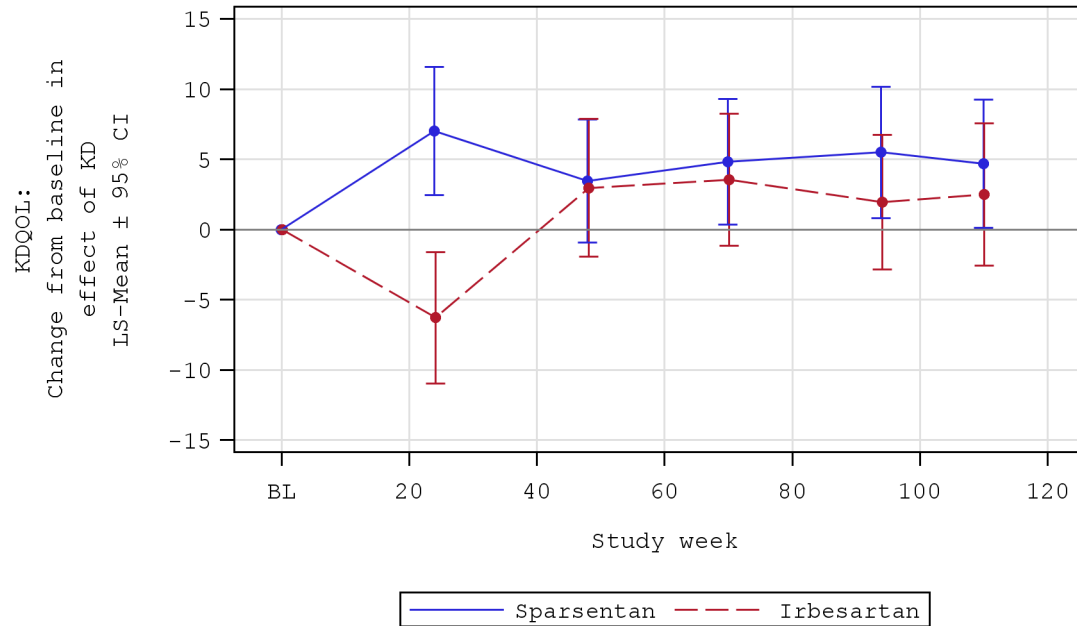
Figure PF2KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2: 60 to < 90 mL/min/1.73 m\*\*2



Sparsentan	38	44	38	39	36
Irbesartan	31	32	32	34	32

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KEFC\_FSCM

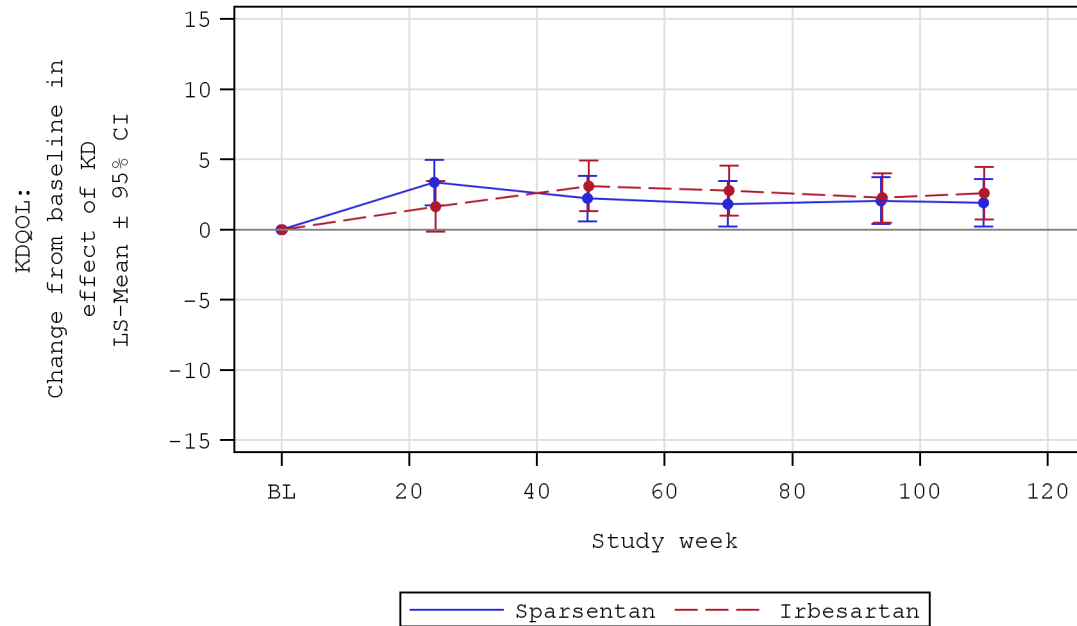
Figure PF2KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Baseline eGFR Group 2:  $\geq 90$  mL/min/1.73 m<sup>2</sup>



Sparsentan	21	23	22	20	21
Irbesartan	20	18	20	19	17

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m<sup>2</sup>, eGFR High  $\geq 60$  ml/min/1.73m<sup>2</sup>, UP Low =  $\leq 1.75$  g/day, UP High =  $> 1.75$  g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KEFC\_FSCM

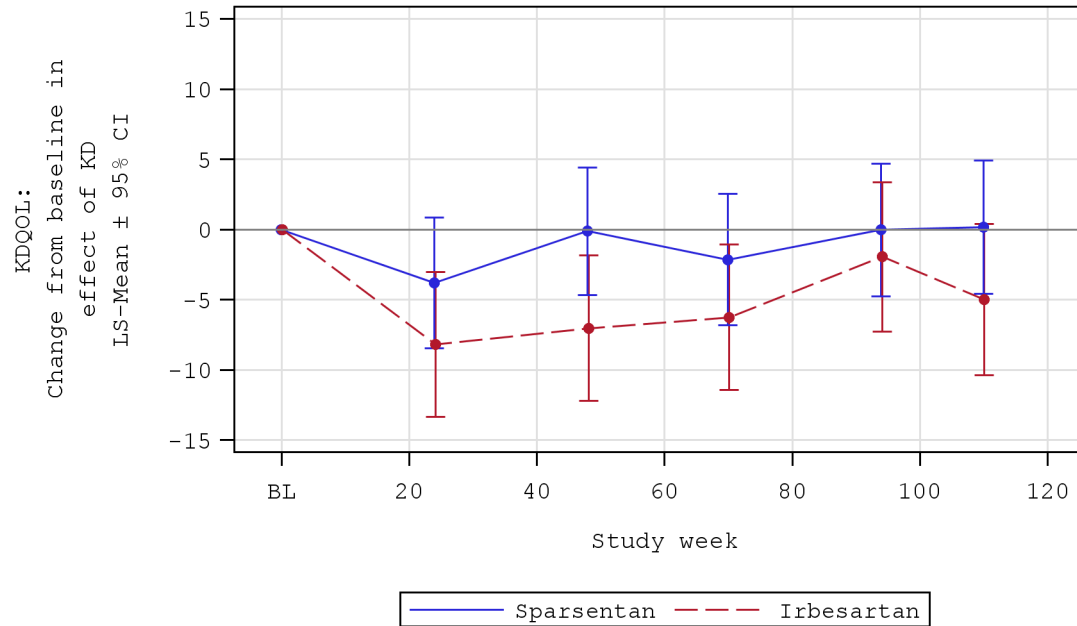
Figure PF2KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 History of hypertension: Yes



Sparsentan	119	121	123	114	112
Irbesartan	100	95	100	101	92

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KEFC\_FSCM

Figure PF2KEFC\_FSGM: KDQOL: Change from baseline in effect of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 History of hypertension: No



Sparsentan	38	41	37	37	37
Irbesartan	31	29	30	29	29

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KEFC\_FSCM

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.438
Male	Sparsentan	139	19 (13.7)	NE		1.853	(0.873, 3.932)	0.108
	Irbesartan	143	15 (10.5)	NE				
Female	Sparsentan	63	16 (25.4)	NE		0.935	(0.320, 2.734)	0.902
	Irbesartan	59	7 (11.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan							Interaction test: 0.776
<= 45 years	Sparsentan	96	19 (19.8)	NE		1.434	(0.667, 3.084)	0.356
	Irbesartan	99	12 (12.1)	NE				
> 45 years	Sparsentan	106	16 (15.1)	NE		1.749	(0.737, 4.151)	0.205
	Irbesartan	103	10 (9.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: <0.001 #
<= 18 years	Sparsentan	9	3 (33.3)	NE		0.263	(0.007, 9.271)	0.463
	Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Sparsentan	102	17 (16.7)	NE		2.171	(0.985, 4.783)	0.054
	Irbesartan	109	13 (11.9)	NE				
> 40 years	Sparsentan	91	15 (16.5)	NE		1.820	(0.704, 4.701)	0.216
	Irbesartan	88	8 (9.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.065
North America	Sparsentan	35	6 (17.1)	NE		1.532	(0.354, 6.639)	0.568
	Irbesartan	46	4 (8.7)	NE				
Europe	Sparsentan	98	14 (14.3)	NE		1.272	(0.585, 2.766)	0.545
	Irbesartan	115	16 (13.9)	NE				
Asia Pacific	Sparsentan	69	15 (21.7)	NE		7.401	(1.454, 37.680)	0.016 *
	Irbesartan	41	2 (4.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.676
< 27 kg/m**2	Sparsentan	83	18 (21.7)	NE		1.417	(0.619, 3.247)	0.410
	Irbesartan	94	10 (10.6)	NE				
≥ 27 kg/m**2	Sparsentan	119	17 (14.3)	NE		1.688	(0.749, 3.804)	0.206
	Irbesartan	107	12 (11.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.050
eGFR Low and UP High	Sparsentan	71	7 (9.9)	NE		1.428	(0.463, 4.410)	0.536
	Irbesartan	74	6 (8.1)	NE				
eGFR Low and UP Low	Sparsentan	55	13 (23.6)	NE		1.118	(0.379, 3.294)	0.840
	Irbesartan	55	6 (10.9)	NE				
eGFR High and UP High	Sparsentan	37	7 (18.9)	NE		0.731	(0.222, 2.407)	0.606
	Irbesartan	36	6 (16.7)	NE				
eGFR High and UP Low	Sparsentan	39	8 (20.5)	NE		5.973	(1.219, 29.260)	0.027 *
	Irbesartan	37	4 (10.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.794
< 60 mL/min/1.73 m**2	Sparsentan	127	20 (15.7)	NE		1.113	(0.525, 2.363)	0.780
	Irbesartan	129	12 (9.3)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	7 (14.3)	NE		1.807	(0.484, 6.753)	0.379
	Irbesartan	48	8 (16.7)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	8 (30.8)	NE		3.301	(0.665, 16.394)	0.144
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.909
< 45 mL/min/1.73 m**2	Sparsentan	82	15 (18.3)	NE		1.115	(0.447, 2.780)	0.815
	Irbesartan	80	8 (10.0)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	5 (11.1)	NE		0.739	(0.169, 3.224)	0.687
	Irbesartan	49	4 (8.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	7 (14.3)	NE		1.807	(0.484, 6.753)	0.379
	Irbesartan	48	8 (16.7)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	8 (30.8)	NE		3.301	(0.665, 16.394)	0.144
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.033 #
<= 1.75 g/day	Sparsentan	98	20 (20.4)	NE		2.881	(1.165, 7.124)	0.022 *
	Irbesartan	93	8 (8.6)	NE				
> 1.75 g/day	Sparsentan	104	15 (14.4)	NE		0.855	(0.395, 1.853)	0.692
	Irbesartan	109	14 (12.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.082
Yes	Sparsentan	90	14 (15.6)	NE		2.829	(1.094, 7.320)	0.032 *
	Irbesartan	88	7 (8.0)	NE				
No	Sparsentan	112	21 (18.8)	NE		1.023	(0.506, 2.068)	0.950
	Irbesartan	114	15 (13.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: <0.001 #
<= 5 years	Sparsentan	113	25 (22.1)	NE		4.053	(1.905, 8.623)	<0.001 *
	Irbesartan	127	13 (10.2)	NE				
> 5 years	Sparsentan	89	10 (11.2)	NE		0.396	(0.140, 1.119)	0.080
	Irbesartan	75	9 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFIT\_FSTM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

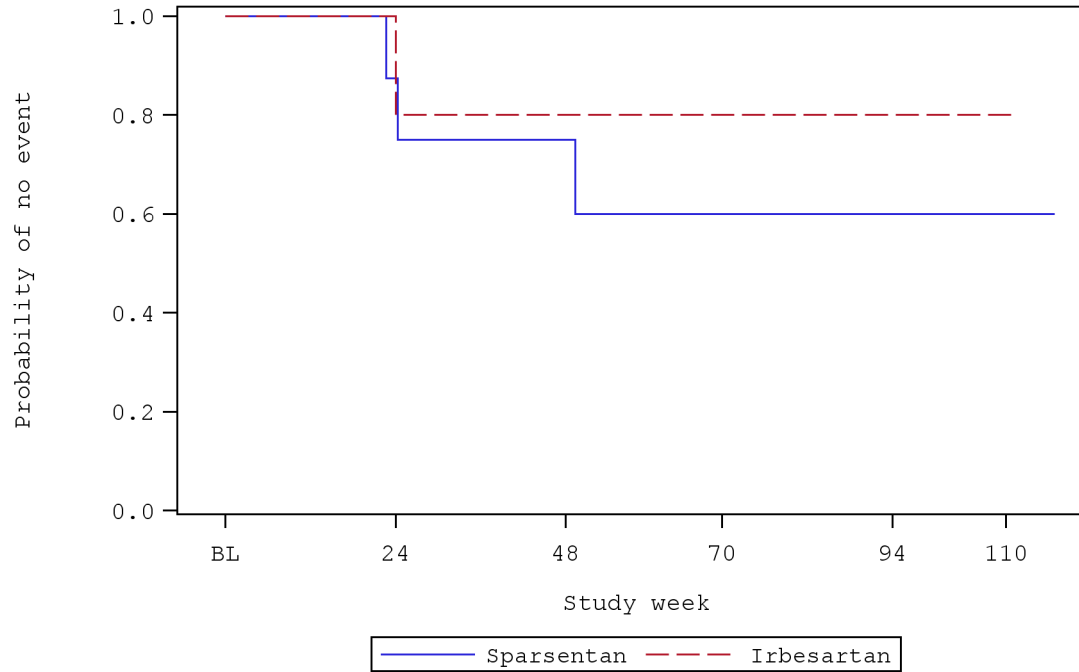
KDQOL: Time to increase in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.163
Yes	Sparsentan	155	30 (19.4)	NE		1.131	(0.625, 2.047)	0.685
	Irbesartan	161	20 (12.4)	NE				
No	Sparsentan	47	5 (10.6)	NE		10.943	(0.684, 175.145)	0.091
	Irbesartan	41	2 (4.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Figure PF2KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

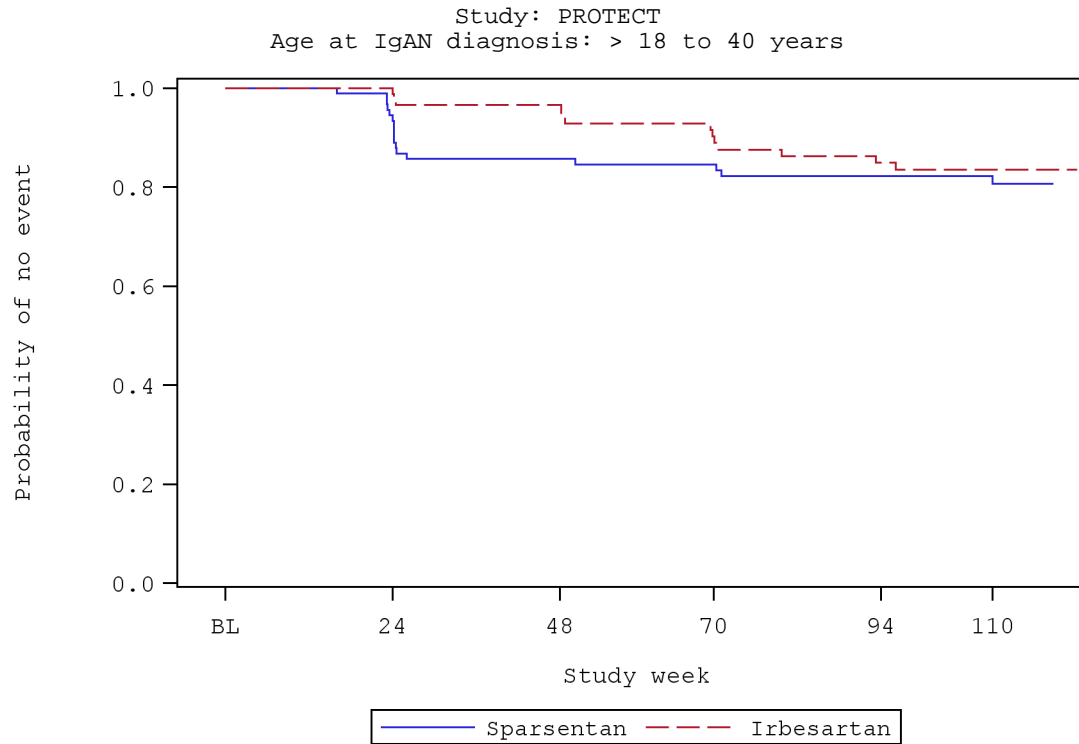
Study: PROTECT  
 Age at IgAN diagnosis: <= 18 years



Sparsentan	9	7	5	4	4	2
Irbesartan	5	5	3	3	2	1

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KEFIT\_FSTM

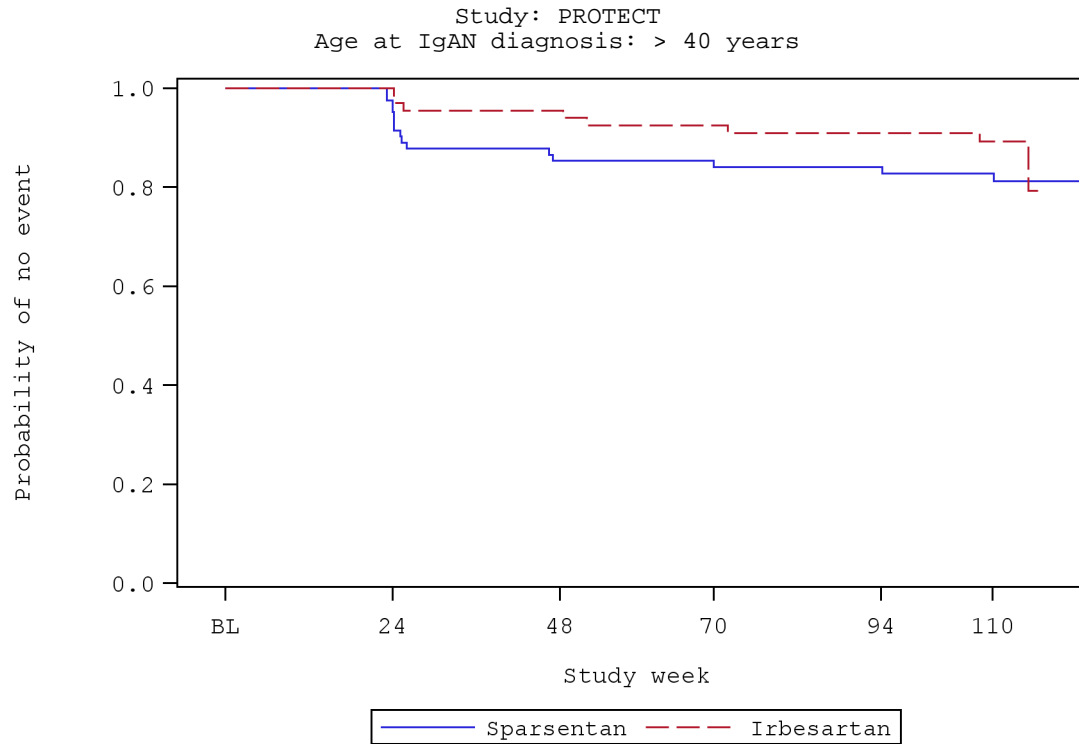
Figure PF2KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	102	86	77	74	64	54
Irbesartan	109	88	79	68	62	58

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KEFIT\_FSTM

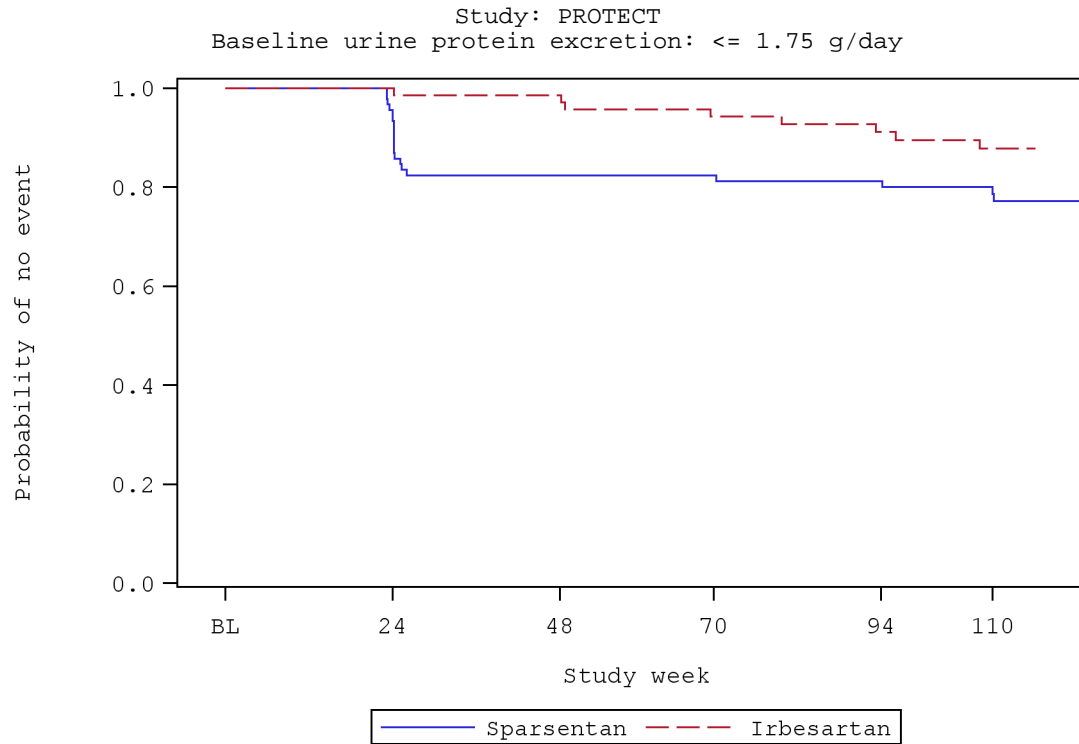
Figure PF2KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	91	80	69	69	65	56
Irbesartan	88	68	63	60	58	44

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KEFIT\_FSTM

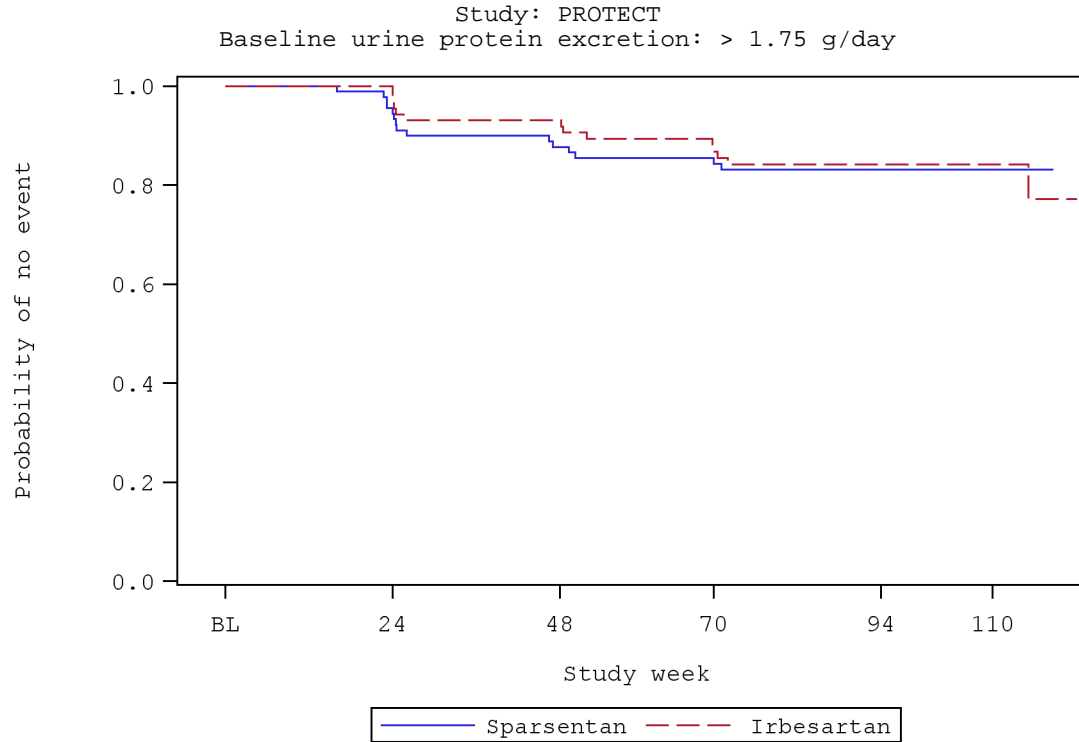
Figure PF2KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	98	87	73	73	68	56
Irbesartan	93	73	68	63	58	46

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KEFIT\_FSTM

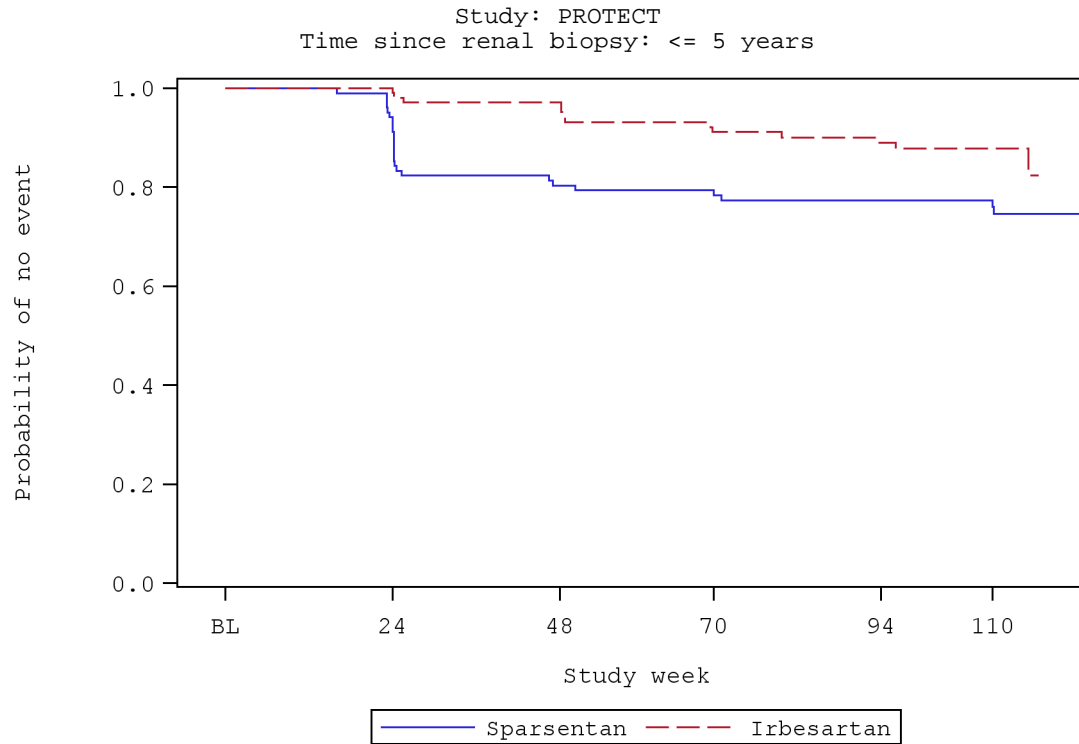
Figure PF2KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	104	86	78	74	65	56
Irbesartan	109	88	77	68	64	57

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KEFIT\_FSTM

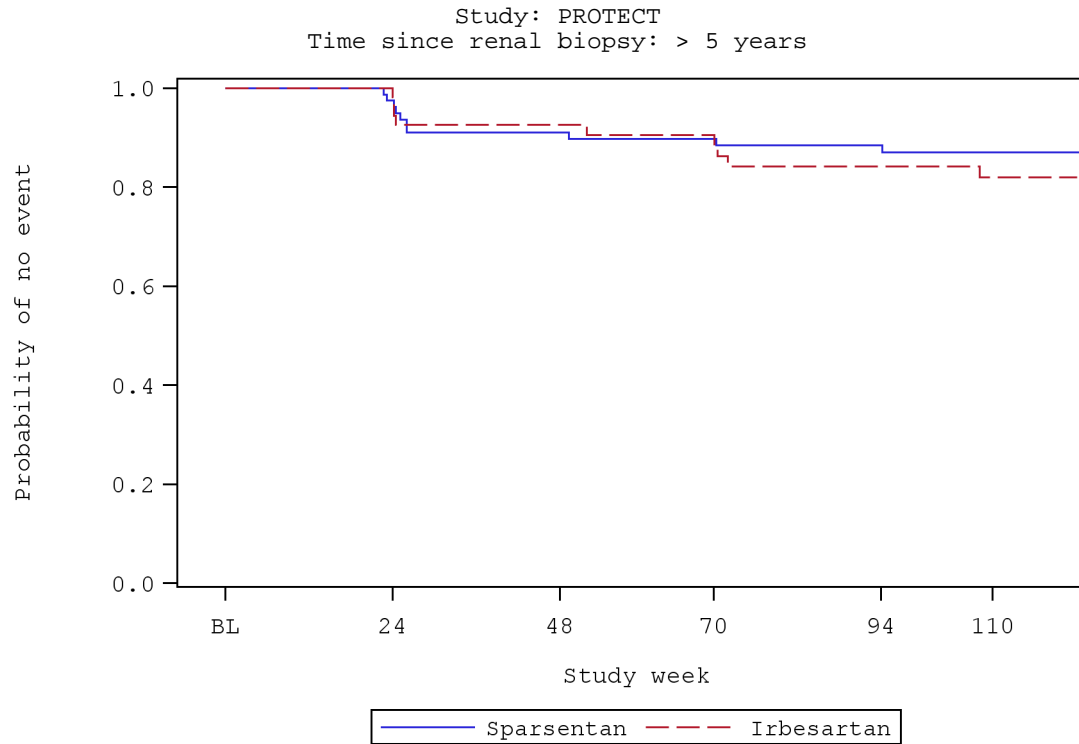
Figure PF2KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	113	96	81	79	70	58
Irbesartan	127	106	99	88	82	70

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KEFIT\_FSTM

Figure PF2KEFIT\_FSKM: KDQOL: Time to increase in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	89	77	70	68	63	54
Irbesartan	75	55	46	43	40	33

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KEFIT\_FSTM

Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.717
Male	Sparsentan	139	25 (18.0)	NE		0.933	(0.529, 1.648)	0.812
	Irbesartan	143	23 (16.1)	NE				
Female	Sparsentan	63	6 (9.5)	NE		0.712	(0.221, 2.294)	0.570
	Irbesartan	59	6 (10.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan							Interaction test: 0.770
<= 45 years	Sparsentan	96	16 (16.7)	NE		0.963	(0.476, 1.949)	0.916
	Irbesartan	99	15 (15.2)	NE				
> 45 years	Sparsentan	106	15 (14.2)	NE		0.870	(0.414, 1.829)	0.714
	Irbesartan	103	14 (13.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: 0.945
<= 18 years	Sparsentan	9	1 (11.1)	NE		NE		NE
	Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Sparsentan	102	16 (15.7)	NE		0.883	(0.439, 1.776)	0.728
	Irbesartan	109	16 (14.7)	NE				
> 40 years	Sparsentan	91	14 (15.4)	NE		0.876	(0.401, 1.912)	0.739
	Irbesartan	88	12 (13.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.605
North America	Sparsentan	35	5 (14.3)	NE		0.762	(0.242, 2.400)	0.643
	Irbesartan	46	8 (17.4)	NE				
Europe	Sparsentan	98	10 (10.2)	NE		0.588	(0.259, 1.338)	0.206
	Irbesartan	115	14 (12.2)	NE				
Asia Pacific	Sparsentan	69	16 (23.2)	NE		1.382	(0.552, 3.461)	0.490
	Irbesartan	41	7 (17.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan							Interaction test: 0.518
< 27 kg/m**2	Sparsentan	83	15 (18.1)	NE		1.026	(0.490, 2.148)	0.946
	Irbesartan	94	14 (14.9)	NE				
≥ 27 kg/m**2	Sparsentan	119	16 (13.4)	NE		0.815	(0.395, 1.678)	0.578
	Irbesartan	107	14 (13.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.563
eGFR Low and UP High	Sparsentan	71	12 (16.9)	NE		0.742	(0.337, 1.634)	0.459
	Irbesartan	74	13 (17.6)	NE				
eGFR Low and UP Low	Sparsentan	55	9 (16.4)	NE		1.889	(0.628, 5.685)	0.258
	Irbesartan	55	5 (9.1)	NE				
eGFR High and UP High	Sparsentan	37	5 (13.5)	NE		0.722	(0.207, 2.513)	0.609
	Irbesartan	36	5 (13.9)	NE				
eGFR High and UP Low	Sparsentan	39	5 (12.8)	NE		0.705	(0.212, 2.344)	0.568
	Irbesartan	37	6 (16.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.526
< 60 mL/min/1.73 m**2	Sparsentan	127	22 (17.3)	NE		1.042	(0.556, 1.956)	0.897
	Irbesartan	129	18 (14.0)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	6 (12.2)	NE		0.704	(0.223, 2.226)	0.550
	Irbesartan	48	6 (12.5)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	3 (11.5)	NE		0.622	(0.145, 2.658)	0.522
	Irbesartan	25	5 (20.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan							Interaction test: 0.012 #
< 45 mL/min/1.73 m**2	Sparsentan	82	11 (13.4)	NE		0.515	(0.238, 1.114)	0.092
	Irbesartan	80	16 (20.0)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	11 (24.4)	NE		6.080	(1.291, 28.628)	0.022 *
	Irbesartan	49	2 (4.1)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	6 (12.2)	NE		0.704	(0.223, 2.226)	0.550
	Irbesartan	48	6 (12.5)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	3 (11.5)	NE		0.622	(0.145, 2.658)	0.522
	Irbesartan	25	5 (20.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.534
<= 1.75 g/day	Sparsentan	98	17 (17.3)	NE		1.030	(0.498, 2.130)	0.937
	Irbesartan	93	13 (14.0)	NE				
> 1.75 g/day	Sparsentan	104	14 (13.5)	NE		0.709	(0.345, 1.457)	0.349
	Irbesartan	109	16 (14.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.655
Yes	Sparsentan	90	14 (15.6)	NE		0.736	(0.346, 1.563)	0.424
	Irbesartan	88	14 (15.9)	NE				
No	Sparsentan	112	17 (15.2)	NE		0.984	(0.490, 1.977)	0.964
	Irbesartan	114	15 (13.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.432
<= 5 years	Sparsentan	113	18 (15.9)	NE		0.772	(0.414, 1.441)	0.417
	Irbesartan	127	22 (17.3)	NE				
> 5 years	Sparsentan	89	13 (14.6)	NE		1.219	(0.476, 3.124)	0.679
	Irbesartan	75	7 (9.3)	NE				

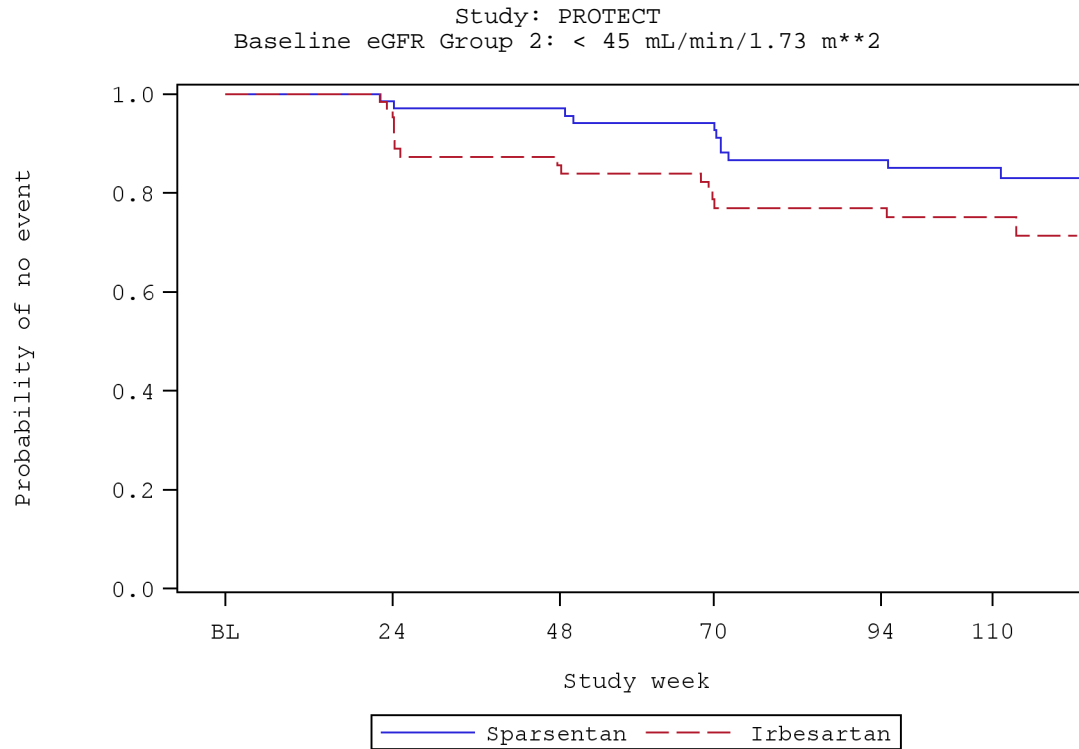
N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KEFDT\_FSTM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in effect of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							
							Interaction test:	0.260
Yes	Sparsentan	155	21 (13.5)	NE		1.044	(0.551, 1.981)	0.894
	Irbesartan	161	17 (10.6)	NE				
No	Sparsentan	47	10 (21.3)	NE		0.586	(0.249, 1.382)	0.222
	Irbesartan	41	12 (29.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

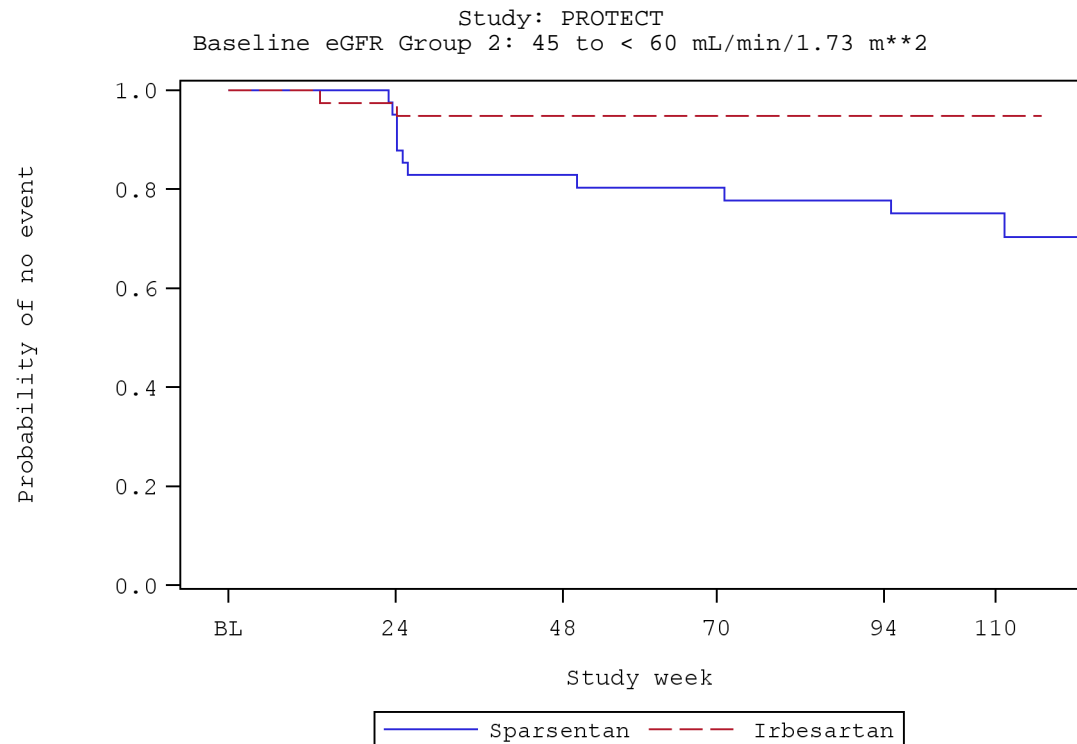
Figure PF2KEFDT\_FSKM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	82	68	66	63	55	48
Irbesartan	80	61	51	45	43	36

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KEFDT\_FSTM

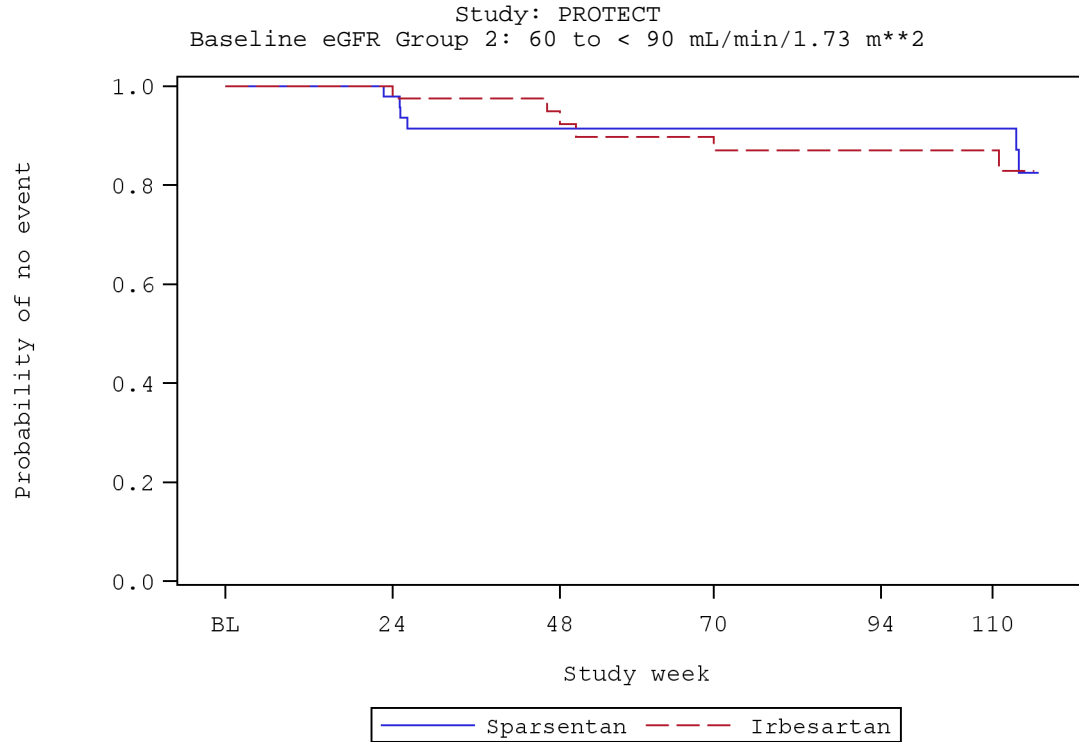
Figure PF2KEFDT\_FSKM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	45	39	33	32	29	22
Irbesartan	49	37	34	33	31	27

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KEFDT\_FSTM

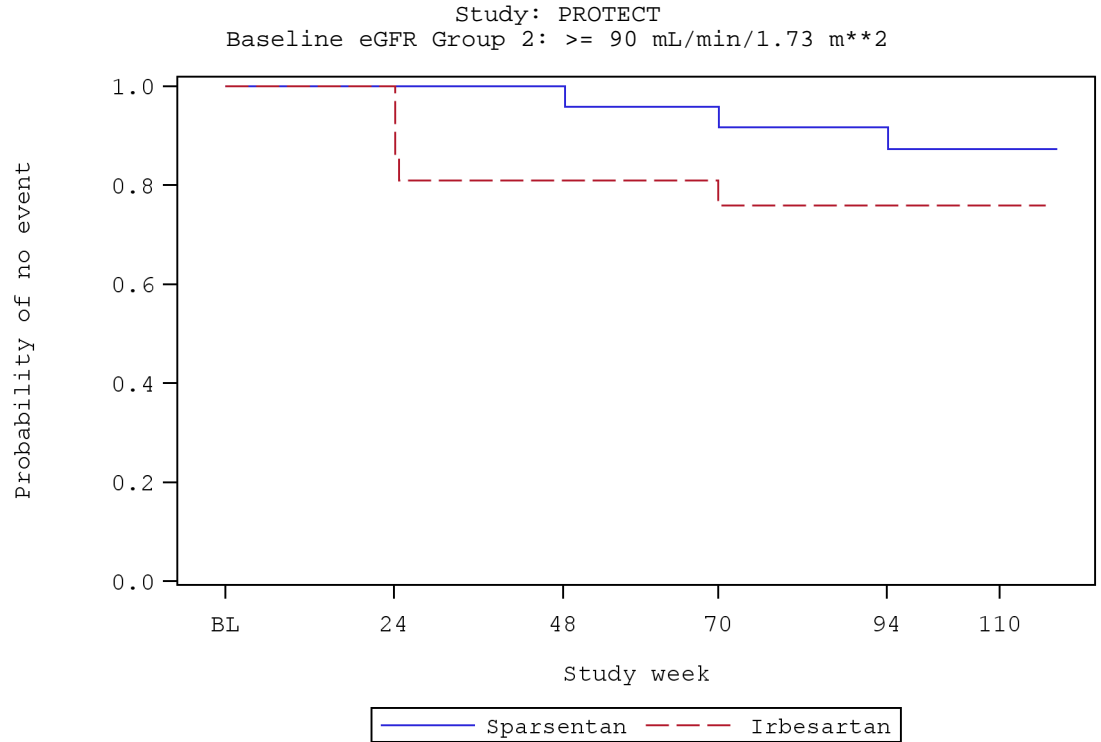
Figure PF2KEFDT\_FSKM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	49	46	43	42	39	34
Irbesartan	48	40	37	33	30	25

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KEFDT\_FSTM

Figure PF2KEFDT\_FSKM: KDQOL: Time to decrease in effect of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	26	24	24	23	21	19
Irbesartan	25	21	16	16	15	14

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KEFDT\_FSTM

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Sex														
Male	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	139	129 (92.8)	90.94 (8.24)	61.4	86.36	93.18	97.73	100.0			
			Irbesartan	143	131 (91.6)	89.63 (11.32)	38.6	86.36	93.18	97.73	100.0			
		Week 24	Sparsentan	139	109 (78.4)	89.51 (12.25)	4.5	84.09	93.18	97.73	100.0			
			Irbesartan	143	93 (65.0)	87.41 (16.42)	4.5	84.09	93.18	97.73	100.0			
		Week 48	Sparsentan	139	112 (80.6)	88.92 (15.05)	0.0	85.23	92.05	97.73	100.0			
			Irbesartan	143	87 (60.8)	89.58 (13.67)	36.4	86.36	93.18	100.00	100.0			
		Week 70	Sparsentan	139	114 (82.0)	88.98 (12.40)	38.6	86.36	93.18	97.73	100.0			
			Irbesartan	143	92 (64.3)	89.38 (14.25)	29.5	86.36	93.18	100.00	100.0			
		Week 94	Sparsentan	139	105 (75.5)	88.90 (10.74)	52.3	84.09	90.91	97.73	100.0			
			Irbesartan	143	96 (67.1)	87.57 (15.17)	25.0	81.82	92.05	97.73	100.0			
		Week 110	Sparsentan	139	103 (74.1)	88.83 (12.89)	6.8	84.09	93.18	97.73	100.0			
			Irbesartan	143	89 (62.2)	88.02 (15.04)	6.8	81.82	93.18	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	139	109 (78.4)	-1.48 (10.23)	-72.7	-4.55	0.00	2.27	25.0	0.01 [-0.26, 0.29]
					Irbesartan	143	93 (65.0)	-1.66 (14.94)	-90.9	-4.55	0.00	2.27	52.3	
		Week 48		Sparsentan	139	112 (80.6)	-1.75 (15.15)	-100.0	-4.55	0.00	4.55	27.3	-0.02 [-0.30, 0.26]	
				Irbesartan	143	87 (60.8)	-1.49 (10.19)	-52.3	-4.55	0.00	2.27	15.9		
		Week 70		Sparsentan	139	114 (82.0)	-2.17 (10.81)	-45.5	-4.55	0.00	4.55	20.5	-0.10 [-0.37, 0.17]	
				Irbesartan	143	92 (64.3)	-1.11 (10.40)	-59.1	-4.55	0.00	2.27	31.8		
	Week 94	Sparsentan		139	105 (75.5)	-1.90 (9.05)	-29.5	-4.55	0.00	4.55	20.5	0.05 [-0.22, 0.33]		
		Irbesartan		143	96 (67.1)	-2.44 (11.21)	-63.6	-4.55	0.00	2.27	20.5			
Week 110	Sparsentan	139		103 (74.1)	-2.29 (12.25)	-84.1	-6.82	0.00	2.27	18.2	-0.08 [-0.36, 0.21]			
	Irbesartan	143		89 (62.2)	-1.35 (12.55)	-81.8	-4.55	0.00	4.55	22.7				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Female	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	63	58 (92.1)	84.17 (15.05)	31.8	77.27	88.64	95.45	100.0			
			Irbesartan	59	51 (86.4)	88.19 (10.22)	50.0	81.82	90.91	95.45	100.0			
		Week 24	Sparsentan	63	48 (76.2)	88.30 (12.05)	54.5	81.82	92.05	97.73	100.0			
			Irbesartan	59	38 (64.4)	88.04 (17.36)	4.5	81.82	95.45	97.73	100.0			
		Week 48	Sparsentan	63	50 (79.4)	87.77 (12.68)	54.5	77.27	93.18	100.00	100.0			
			Irbesartan	59	37 (62.7)	86.24 (11.35)	54.5	79.55	88.64	93.18	100.0			
		Week 70	Sparsentan	63	46 (73.0)	84.39 (16.53)	27.3	79.55	89.77	97.73	100.0			
			Irbesartan	59	38 (64.4)	87.32 (13.26)	40.9	84.09	90.91	95.45	100.0			
		Week 94	Sparsentan	63	46 (73.0)	84.58 (14.81)	45.5	77.27	88.64	97.73	100.0			
			Irbesartan	59	34 (57.6)	88.90 (10.57)	59.1	84.09	93.18	97.73	100.0			
		Week 110	Sparsentan	63	46 (73.0)	85.92 (14.30)	43.2	77.27	90.91	97.73	100.0			
			Irbesartan	59	32 (54.2)	89.56 (8.73)	65.9	85.23	90.91	96.59	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	63	48 (76.2)	4.07 (13.54)	-25.0	-2.27	2.27	10.23	63.6	0.33 [-0.10, 0.76]
					Irbesartan	59	38 (64.4)	-1.02 (17.35)	-86.4	-2.27	0.00	4.55	29.5	
	Week 48	Sparsentan		63	50 (79.4)	3.55 (14.45)	-29.5	-2.27	0.00	11.36	59.1	0.42 [-0.01, 0.85]		
		Irbesartan		59	37 (62.7)	-1.84 (9.93)	-18.2	-9.09	-2.27	2.27	34.1			
	Week 70	Sparsentan		63	46 (73.0)	0.30 (16.38)	-61.4	-2.27	2.27	9.09	25.0	0.14 [-0.29, 0.57]		
		Irbesartan		59	38 (64.4)	-1.56 (9.15)	-27.3	-4.55	-2.27	4.55	20.5			
	Week 94	Sparsentan	63	46 (73.0)	1.28 (16.92)	-34.1	-6.82	0.00	9.09	68.2	0.19 [-0.25, 0.64]			
		Irbesartan	59	34 (57.6)	-1.54 (10.43)	-34.1	-6.82	0.00	2.27	20.5				
Week 110	Sparsentan	63	46 (73.0)	1.73 (18.76)	-34.1	-6.82	1.14	9.09	68.2	0.16 [-0.29, 0.61]				
	Irbesartan	59	32 (54.2)	-0.78 (8.61)	-18.2	-5.68	-1.14	2.27	20.5					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age														
<= 45 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	96	87 (90.6)	88.71 (12.19)	31.8	84.09	90.91	97.73	100.0			
			Irbesartan	99	92 (92.9)	89.30 (11.00)	52.3	86.36	92.05	97.73	100.0			
		Week 24	Sparsentan	96	71 (74.0)	90.08 (14.00)	4.5	86.36	93.18	97.73	100.0			
			Irbesartan	99	67 (67.7)	86.77 (18.91)	4.5	84.09	90.91	97.73	100.0			
		Week 48	Sparsentan	96	75 (78.1)	88.73 (15.03)	0.0	81.82	93.18	97.73	100.0			
			Irbesartan	99	62 (62.6)	88.42 (13.72)	36.4	84.09	90.91	100.00	100.0			
		Week 70	Sparsentan	96	72 (75.0)	88.57 (15.09)	27.3	84.09	93.18	100.00	100.0			
			Irbesartan	99	62 (62.6)	90.03 (12.48)	36.4	86.36	93.18	100.00	100.0			
		Week 94	Sparsentan	96	68 (70.8)	88.47 (12.29)	50.0	81.82	90.91	98.86	100.0			
			Irbesartan	99	64 (64.6)	86.65 (17.13)	25.0	79.55	93.18	100.00	100.0			
		Week 110	Sparsentan	96	69 (71.9)	89.69 (12.12)	43.2	81.82	93.18	100.00	100.0			
			Irbesartan	99	59 (59.6)	88.79 (11.77)	45.5	81.82	93.18	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	96	71 (74.0)	1.18 (13.52)	-72.7	-2.27	2.27	4.55	63.6	0.27 [-0.07, 0.60]
					Irbesartan	99	67 (67.7)	-2.85 (16.59)	-90.9	-6.82	0.00	2.27	15.9	
		Week 48		Sparsentan	96	75 (78.1)	-0.12 (15.45)	-90.9	-4.55	0.00	4.55	59.1	0.26 [-0.08, 0.59]	
				Irbesartan	99	62 (62.6)	-3.59 (10.67)	-52.3	-9.09	0.00	2.27	11.4		
		Week 70		Sparsentan	96	72 (75.0)	-0.19 (13.74)	-61.4	-3.41	2.27	6.82	22.7	0.14 [-0.20, 0.48]	
				Irbesartan	99	62 (62.6)	-1.87 (8.70)	-27.3	-4.55	0.00	2.27	25.0		
Week 94	Sparsentan	96		68 (70.8)	-0.10 (13.60)	-34.1	-4.55	0.00	4.55	68.2	0.30 [-0.04, 0.64]			
	Irbesartan	99		64 (64.6)	-4.15 (13.30)	-63.6	-7.95	0.00	2.27	20.5				
Week 110	Sparsentan	96		69 (71.9)	0.79 (15.11)	-45.5	-6.82	0.00	6.82	68.2	0.19 [-0.16, 0.54]			
	Irbesartan	99		59 (59.6)	-1.66 (9.19)	-27.3	-6.82	0.00	2.27	22.7				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 45 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	106	100 (94.3)	88.95 (10.37)	38.6	84.09	90.91	95.45	100.0			
			Irbesartan	103	90 (87.4)	89.14 (11.10)	38.6	84.09	92.05	97.73	100.0			
		Week 24	Sparsentan	106	86 (81.1)	88.37 (10.43)	50.0	84.09	90.91	95.45	100.0			
			Irbesartan	103	64 (62.1)	88.46 (13.96)	11.4	84.09	93.18	95.45	100.0			
		Week 48	Sparsentan	106	87 (82.1)	88.43 (13.80)	0.0	84.09	90.91	97.73	100.0			
			Irbesartan	103	62 (60.2)	88.75 (12.49)	43.2	84.09	93.18	97.73	100.0			
		Week 70	Sparsentan	106	88 (83.0)	86.91 (12.73)	38.6	80.68	90.91	95.45	100.0			
			Irbesartan	103	68 (66.0)	87.63 (15.16)	29.5	84.09	90.91	97.73	100.0			
		Week 94	Sparsentan	106	83 (78.3)	86.86 (12.21)	45.5	79.55	90.91	95.45	100.0			
			Irbesartan	103	66 (64.1)	89.15 (10.30)	54.5	81.82	92.05	97.73	100.0			
		Week 110	Sparsentan	106	80 (75.5)	86.42 (14.24)	6.8	78.41	89.77	95.45	100.0			
			Irbesartan	103	62 (60.2)	88.09 (15.29)	6.8	81.82	92.05	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	106	86 (81.1)	-0.58 (9.71)	-25.0	-4.55	0.00	4.55	36.4	-0.05 [-0.37, 0.28]
					Irbesartan	103	64 (62.1)	-0.04 (14.51)	-77.3	-4.55	0.00	4.55	52.3	
	Week 48	Sparsentan		106	87 (82.1)	-0.10 (14.87)	-100.0	-4.55	0.00	6.82	38.6	-0.04 [-0.37, 0.29]		
		Irbesartan		103	62 (60.2)	0.40 (9.09)	-27.3	-4.55	0.00	2.27	34.1			
	Week 70	Sparsentan		106	88 (83.0)	-2.51 (11.68)	-45.5	-6.82	0.00	3.41	25.0	-0.16 [-0.48, 0.16]		
		Irbesartan		103	68 (66.0)	-0.67 (11.12)	-59.1	-4.55	0.00	2.27	31.8			
	Week 94	Sparsentan	106	83 (78.3)	-1.62 (10.62)	-27.3	-6.82	-2.27	4.55	40.9	-0.14 [-0.46, 0.19]			
		Irbesartan	103	66 (64.1)	-0.31 (7.76)	-18.2	-4.55	0.00	4.55	20.5				
Week 110	Sparsentan	106	80 (75.5)	-2.64 (14.09)	-84.1	-6.82	0.00	2.27	36.4	-0.13 [-0.47, 0.20]				
	Irbesartan	103	62 (60.2)	-0.77 (13.58)	-81.8	-4.55	0.00	4.55	22.7					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age at IgAN diagnosis														
<= 18 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	9	9 (100.0)	85.86 (9.22)	75.0	77.27	84.09	93.18	100.0			
			Irbesartan	5	5 (100.0)	87.27 (19.85)	52.3	90.91	95.45	97.73	100.0			
		Week 24	Sparsentan	9	5 (55.6)	74.55 (39.48)	4.5	84.09	90.91	95.45	97.7			
			Irbesartan	5	5 (100.0)	85.45 (22.87)	45.5	88.64	93.18	100.00	100.0			
		Week 48	Sparsentan	9	7 (77.8)	86.04 (10.12)	72.7	75.00	88.64	95.45	100.0			
			Irbesartan	5	3 (60.0)	93.18 (2.27)	90.9	90.91	93.18	95.45	95.5			
		Week 70	Sparsentan	9	7 (77.8)	89.61 (5.99)	81.8	84.09	88.64	95.45	97.7			
			Irbesartan	5	4 (80.0)	77.84 (28.41)	36.4	60.23	87.50	95.45	100.0			
		Week 94	Sparsentan	9	5 (55.6)	84.55 (15.28)	68.2	68.18	90.91	95.45	100.0			
			Irbesartan	5	4 (80.0)	78.98 (23.65)	45.5	62.50	86.36	95.45	97.7			
		Week 110	Sparsentan	9	4 (44.4)	94.32 (8.40)	81.8	89.77	97.73	98.86	100.0			
			Irbesartan	5	2 (40.0)	61.36 (22.50)	45.5	45.45	61.36	77.27	77.3			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	9	5 (55.6)	-12.73 (33.81)	-72.7	-2.27	0.00	2.27	9.1	-0.45 [-1.71, 0.80]
					Irbesartan	5	5 (100.0)	-1.82 (4.66)	-6.8	-6.82	0.00	2.27	2.3	
				Week 48	Sparsentan	9	7 (77.8)	-2.27 (6.15)	-11.4	-9.09	0.00	2.27	4.5	-0.14 [-1.49, 1.22]
					Irbesartan	5	3 (60.0)	-1.52 (3.47)	-4.5	-4.55	-2.27	2.27	2.3	
				Week 70	Sparsentan	9	7 (77.8)	1.30 (5.23)	-4.5	-4.55	2.27	4.55	9.1	1.59 [0.19, 2.99]
					Irbesartan	5	4 (80.0)	-8.52 (7.74)	-15.9	-13.64	-10.23	-3.41	2.3	
		Week 94	Sparsentan	9	5 (55.6)	-4.55 (11.48)	-18.2	-13.64	-4.55	6.82	6.8	0.29 [-1.03, 1.62]		
			Irbesartan	5	4 (80.0)	-7.39 (6.53)	-15.9	-11.36	-6.82	-3.41	0.0			
		Week 110	Sparsentan	9	4 (44.4)	3.41 (7.07)	-2.3	-1.14	1.14	7.95	13.6	2.17 [0.08, 4.27]		
			Irbesartan	5	2 (40.0)	-12.50 (8.04)	-18.2	-18.18	-12.50	-6.82	-6.8			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 18 to 40 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	102	94 (92.2)	89.12 (12.10)	31.8	84.09	90.91	97.73	100.0			
			Irbesartan	109	99 (90.8)	89.69 (10.19)	52.3	86.36	93.18	97.73	100.0			
		Week 24	Sparsentan	102	79 (77.5)	90.91 (9.86)	54.5	86.36	93.18	97.73	100.0			
			Irbesartan	109	71 (65.1)	88.96 (14.96)	4.5	86.36	93.18	97.73	100.0			
		Week 48	Sparsentan	102	82 (80.4)	88.91 (14.70)	0.0	81.82	94.32	97.73	100.0			
			Irbesartan	109	67 (61.5)	87.96 (13.85)	36.4	84.09	90.91	100.00	100.0			
		Week 70	Sparsentan	102	78 (76.5)	87.82 (15.22)	27.3	81.82	93.18	100.00	100.0			
			Irbesartan	109	68 (62.4)	90.71 (10.03)	54.5	86.36	93.18	97.73	100.0			
		Week 94	Sparsentan	102	73 (71.6)	88.48 (11.83)	50.0	81.82	90.91	97.73	100.0			
			Irbesartan	109	69 (63.3)	87.58 (15.84)	25.0	81.82	93.18	97.73	100.0			
		Week 110	Sparsentan	102	72 (70.6)	89.05 (12.29)	43.2	81.82	93.18	98.86	100.0			
			Irbesartan	109	65 (59.6)	89.97 (10.64)	52.3	84.09	93.18	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	102	79 (77.5)	2.04 (9.95)	-18.2	-2.27	2.27	4.55	63.6	0.30 [-0.02, 0.63]
					Irbesartan	109	71 (65.1)	-1.31 (12.13)	-86.4	-4.55	0.00	2.27	15.9	
			Week 48	Sparsentan	102	82 (80.4)	0.08 (14.97)	-90.9	-4.55	0.00	4.55	59.1	0.31 [-0.01, 0.64]	
				Irbesartan	109	67 (61.5)	-4.07 (10.80)	-52.3	-11.36	0.00	2.27	11.4		
			Week 70	Sparsentan	102	78 (76.5)	-0.82 (13.74)	-61.4	-4.55	0.00	6.82	22.7	0.07 [-0.25, 0.40]	
				Irbesartan	109	68 (62.4)	-1.64 (8.20)	-27.3	-4.55	0.00	2.27	25.0		
			Week 94	Sparsentan	102	73 (71.6)	0.09 (12.91)	-34.1	-4.55	0.00	4.55	68.2	0.29 [-0.04, 0.62]	
				Irbesartan	109	69 (63.3)	-3.62 (12.97)	-63.6	-6.82	0.00	2.27	20.5		
		Week 110	Sparsentan	102	72 (70.6)	0.54 (14.88)	-45.5	-6.82	0.00	6.82	68.2	0.15 [-0.19, 0.48]		
			Irbesartan	109	65 (59.6)	-1.29 (9.17)	-27.3	-4.55	0.00	2.27	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 40 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	91	84 (92.3)	88.85 (10.44)	38.6	85.23	90.91	95.45	100.0			
			Irbesartan	88	78 (88.6)	88.75 (11.49)	38.6	84.09	90.91	97.73	100.0			
		Week 24	Sparsentan	91	73 (80.2)	88.23 (10.47)	50.0	81.82	90.91	95.45	100.0			
			Irbesartan	88	55 (62.5)	86.03 (18.22)	4.5	81.82	90.91	95.45	100.0			
		Week 48	Sparsentan	91	73 (80.2)	88.42 (14.38)	0.0	84.09	90.91	97.73	100.0			
			Irbesartan	88	54 (61.4)	89.10 (12.45)	43.2	84.09	93.18	97.73	100.0			
		Week 70	Sparsentan	91	75 (82.4)	87.30 (12.88)	38.6	81.82	93.18	95.45	100.0			
			Irbesartan	88	58 (65.9)	87.26 (16.26)	29.5	84.09	90.91	97.73	100.0			
		Week 94	Sparsentan	91	73 (80.2)	86.89 (12.53)	45.5	79.55	90.91	95.45	100.0			
			Irbesartan	88	57 (64.8)	88.96 (10.74)	54.5	81.82	90.91	97.73	100.0			
		Week 110	Sparsentan	91	73 (80.2)	86.49 (14.47)	6.8	79.55	90.91	95.45	100.0			
			Irbesartan	88	54 (61.4)	87.58 (15.60)	6.8	81.82	90.91	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	91	73 (80.2)	-0.87 (10.18)	-25.0	-4.55	-2.27	2.27	36.4	0.05 [-0.30, 0.40]
					Irbesartan	88	55 (62.5)	-1.65 (19.85)	-90.9	-4.55	0.00	4.55	52.3	
	Week 48	Sparsentan		91	73 (80.2)	-0.12 (15.93)	-100.0	-2.27	0.00	6.82	38.6	-0.12 [-0.47, 0.23]		
		Irbesartan		88	54 (61.4)	1.47 (8.51)	-20.5	-2.27	0.00	4.55	34.1			
	Week 70	Sparsentan		91	75 (82.4)	-2.39 (11.98)	-45.5	-6.82	0.00	2.27	25.0	-0.18 [-0.52, 0.17]		
		Irbesartan		88	58 (65.9)	-0.27 (11.84)	-59.1	-4.55	0.00	4.55	31.8			
	Week 94	Sparsentan	91	73 (80.2)	-1.71 (11.18)	-27.3	-6.82	-2.27	4.55	40.9	-0.16 [-0.51, 0.19]			
		Irbesartan	88	57 (64.8)	-0.12 (7.89)	-18.2	-4.55	0.00	4.55	20.5				
Week 110	Sparsentan	91	73 (80.2)	-2.86 (14.57)	-84.1	-6.82	0.00	2.27	36.4	-0.15 [-0.51, 0.20]				
	Irbesartan	88	54 (61.4)	-0.67 (14.04)	-81.8	-2.27	0.00	4.55	22.7					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Geographic region														
North America	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	35	31 (88.6)	91.50 (7.63)	75.0	86.36	93.18	97.73	100.0			
		Week 24	Irbesartan	46	43 (93.5)	90.91 (10.80)	50.0	86.36	93.18	100.00	100.0			
			Sparsentan	35	23 (65.7)	90.71 (19.71)	4.5	90.91	97.73	100.00	100.0			
		Week 48	Sparsentan	46	35 (76.1)	89.03 (17.09)	4.5	84.09	93.18	100.00	100.0			
			Sparsentan	35	25 (71.4)	91.45 (10.56)	54.5	88.64	95.45	97.73	100.0			
		Week 70	Irbesartan	46	32 (69.6)	88.57 (13.99)	36.4	84.09	93.18	97.73	100.0			
			Sparsentan	35	22 (62.9)	93.39 (7.55)	72.7	90.91	95.45	100.00	100.0			
		Week 94	Irbesartan	46	30 (65.2)	90.38 (14.52)	40.9	88.64	95.45	100.00	100.0			
			Sparsentan	35	23 (65.7)	90.91 (10.68)	50.0	88.64	90.91	97.73	100.0			
		Week 110	Irbesartan	46	30 (65.2)	91.21 (10.76)	63.6	81.82	95.45	100.00	100.0			
			Sparsentan	35	21 (60.0)	92.75 (8.02)	75.0	90.91	95.45	97.73	100.0			
			Irbesartan	Sparsentan	46	31 (67.4)	91.13 (9.14)	65.9	84.09	93.18	97.73	100.0		
				Sparsentan	35	23 (65.7)	-2.77 (15.83)	-72.7	-2.27	0.00	2.27	11.4	-0.07 [-0.59, 0.46]	
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Irbesartan	46	35 (76.1)	-1.69 (16.41)	-86.4	-4.55	0.00	2.27	29.5	
					Sparsentan	35	25 (71.4)	-1.00 (8.14)	-29.5	-2.27	-2.27	2.27	18.2	0.10 [-0.42, 0.62]
				Week 48	Irbesartan	46	32 (69.6)	-2.13 (13.50)	-52.3	-7.95	0.00	2.27	34.1	
					Sparsentan	35	22 (62.9)	0.10 (5.18)	-11.4	-2.27	0.00	2.27	9.1	0.21 [-0.34, 0.76]
				Week 70	Irbesartan	46	30 (65.2)	-1.36 (8.17)	-27.3	-4.55	0.00	2.27	18.2	
Sparsentan	35				23 (65.7)	-2.57 (9.10)	-34.1	-4.55	-2.27	0.00	20.5	-0.25 [-0.80, 0.30]		
Week 94	Irbesartan			46	30 (65.2)	-0.38 (8.48)	-18.2	-4.55	0.00	2.27	20.5			
	Sparsentan			35	21 (60.0)	-1.41 (8.11)	-22.7	-2.27	0.00	2.27	18.2	-0.14 [-0.69, 0.42]		
Week 110	Irbesartan			46	31 (67.4)	-0.22 (9.15)	-18.2	-4.55	0.00	4.55	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Europe	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	98	89 (90.8)	88.13 (12.99)	31.8	84.09	90.91	97.73	100.0			
			Irbesartan	115	98 (85.2)	88.59 (11.45)	38.6	84.09	90.91	95.45	100.0			
		Week 24	Sparsentan	98	73 (74.5)	89.35 (10.00)	59.1	84.09	93.18	97.73	100.0			
			Irbesartan	115	62 (53.9)	86.44 (18.46)	4.5	84.09	92.05	97.73	100.0			
		Week 48	Sparsentan	98	76 (77.6)	87.41 (14.86)	0.0	80.68	90.91	97.73	100.0			
			Irbesartan	115	59 (51.3)	86.98 (14.10)	43.2	79.55	90.91	100.00	100.0			
		Week 70	Sparsentan	98	75 (76.5)	88.00 (11.67)	50.0	81.82	90.91	95.45	100.0			
			Irbesartan	115	69 (60.0)	86.69 (15.74)	29.5	84.09	90.91	95.45	100.0			
		Week 94	Sparsentan	98	68 (69.4)	87.67 (11.75)	45.5	79.55	90.91	97.73	100.0			
			Irbesartan	115	70 (60.9)	85.29 (16.56)	25.0	79.55	89.77	97.73	100.0			
		Week 110	Sparsentan	98	67 (68.4)	87.62 (15.19)	6.8	81.82	93.18	97.73	100.0			
			Irbesartan	115	61 (53.0)	86.62 (16.74)	6.8	81.82	90.91	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	98	73 (74.5)	1.46 (12.31)	-25.0	-4.55	0.00	4.55	63.6	0.23 [-0.11, 0.57]
					Irbesartan	115	62 (53.9)	-2.02 (18.03)	-90.9	-4.55	0.00	4.55	52.3	
	Week 48	Sparsentan		98	76 (77.6)	0.33 (16.79)	-100.0	-4.55	0.00	4.55	59.1	0.22 [-0.12, 0.57]		
		Irbesartan		115	59 (51.3)	-2.81 (9.15)	-27.3	-6.82	0.00	2.27	11.4			
	Week 70	Sparsentan		98	75 (76.5)	-0.21 (10.88)	-45.5	-4.55	0.00	4.55	25.0	0.16 [-0.16, 0.49]		
		Irbesartan		115	69 (60.0)	-2.08 (11.72)	-59.1	-4.55	0.00	2.27	31.8			
	Week 94	Sparsentan	98	68 (69.4)	0.94 (13.07)	-29.5	-4.55	0.00	4.55	68.2	0.35 [0.01, 0.69]			
		Irbesartan	115	70 (60.9)	-3.60 (12.82)	-63.6	-6.82	0.00	2.27	18.2				
Week 110	Sparsentan	98	67 (68.4)	0.24 (16.73)	-84.1	-4.55	0.00	4.55	68.2	0.11 [-0.24, 0.45]				
	Irbesartan	115	61 (53.0)	-1.42 (13.96)	-81.8	-6.82	0.00	4.55	22.7					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Asia Pacific	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	69	67 (97.1)	88.57 (9.98)	50.0	84.09	88.64	97.73	100.0			
			Irbesartan	41	41 (100.0)	88.97 (10.22)	52.3	84.09	93.18	95.45	100.0			
		Week 24	Sparsentan	69	61 (88.4)	88.30 (10.98)	50.0	84.09	90.91	97.73	100.0			
			Irbesartan	41	34 (82.9)	88.24 (12.40)	50.0	86.36	93.18	95.45	100.0			
		Week 48	Sparsentan	69	61 (88.4)	88.82 (15.02)	0.0	84.09	93.18	97.73	100.0			
			Irbesartan	41	33 (80.5)	91.46 (9.64)	54.5	88.64	93.18	97.73	100.0			
		Week 70	Sparsentan	69	63 (91.3)	85.25 (17.03)	27.3	79.55	93.18	95.45	100.0			
			Irbesartan	41	31 (75.6)	91.86 (6.96)	70.5	88.64	90.91	97.73	100.0			
		Week 94	Sparsentan	69	60 (87.0)	86.21 (13.23)	52.3	78.41	89.77	96.59	100.0			
			Irbesartan	41	30 (73.2)	90.76 (8.87)	65.9	84.09	93.18	97.73	100.0			
		Week 110	Sparsentan	69	61 (88.4)	86.62 (12.45)	50.0	79.55	90.91	95.45	100.0			
			Irbesartan	41	29 (70.7)	89.34 (9.55)	59.1	84.09	90.91	95.45	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	69	61 (88.4)	-0.15 (8.36)	-22.7	-4.55	0.00	4.55	13.6	0.01 [-0.41, 0.43]
					Irbesartan	41	34 (82.9)	-0.27 (8.93)	-25.0	-4.55	0.00	2.27	20.5	
	Week 48	Sparsentan		69	61 (88.4)	-0.30 (15.23)	-90.9	-4.55	0.00	9.09	27.3	-0.11 [-0.53, 0.32]		
		Irbesartan		41	33 (80.5)	1.10 (7.14)	-15.9	-2.27	0.00	4.55	20.5			
	Week 70	Sparsentan		69	63 (91.3)	-3.50 (15.92)	-61.4	-6.82	0.00	6.82	22.7	-0.31 [-0.74, 0.12]		
		Irbesartan		41	31 (75.6)	0.73 (7.05)	-13.6	-4.55	0.00	4.55	20.5			
	Week 94	Sparsentan	69	60 (87.0)	-2.42 (11.66)	-27.3	-7.95	-1.14	3.41	25.0	-0.16 [-0.60, 0.28]			
		Irbesartan	41	30 (73.2)	-0.76 (7.93)	-18.2	-4.55	0.00	2.27	20.5				
Week 110	Sparsentan	69	61 (88.4)	-2.35 (13.93)	-45.5	-6.82	0.00	4.55	29.5	-0.04 [-0.49, 0.40]				
	Irbesartan	41	29 (70.7)	-1.80 (8.29)	-18.2	-4.55	0.00	2.27	20.5					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline BMI														
< 27 kg/m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	83	75 (90.4)	89.09 (9.68)	47.7	84.09	88.64	97.73	100.0			
			Irbesartan	94	86 (91.5)	88.85 (10.97)	38.6	86.36	90.91	95.45	100.0			
		Week 24	Sparsentan	83	66 (79.5)	88.22 (14.76)	4.5	84.09	92.05	97.73	100.0			
			Irbesartan	94	66 (70.2)	86.95 (15.74)	4.5	81.82	92.05	97.73	100.0			
		Week 48	Sparsentan	83	66 (79.5)	89.39 (10.68)	65.9	81.82	93.18	97.73	100.0			
			Irbesartan	94	60 (63.8)	85.53 (14.82)	36.4	78.41	89.77	96.59	100.0			
		Week 70	Sparsentan	83	64 (77.1)	87.71 (13.31)	27.3	81.82	93.18	96.59	100.0			
			Irbesartan	94	59 (62.8)	87.37 (14.38)	36.4	84.09	90.91	95.45	100.0			
		Week 94	Sparsentan	83	63 (75.9)	88.02 (12.25)	45.5	84.09	90.91	97.73	100.0			
			Irbesartan	94	62 (66.0)	86.25 (15.71)	25.0	79.55	90.91	97.73	100.0			
		Week 110	Sparsentan	83	63 (75.9)	88.28 (11.05)	50.0	81.82	90.91	97.73	100.0			
			Irbesartan	94	57 (60.6)	86.28 (12.88)	45.5	77.27	88.64	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	83	66 (79.5)	-1.41 (12.36)	-72.7	-6.82	0.00	4.55	20.5	-0.01 [-0.35, 0.33]
					Irbesartan	94	66 (70.2)	-1.24 (14.50)	-86.4	-4.55	0.00	4.55	52.3	
				Week 48	Sparsentan	83	66 (79.5)	0.45 (9.47)	-18.2	-4.55	0.00	4.55	25.0	0.39 [0.04, 0.75]
					Irbesartan	94	60 (63.8)	-3.60 (11.14)	-52.3	-6.82	0.00	2.27	13.6	
				Week 70	Sparsentan	83	64 (77.1)	-1.92 (12.63)	-61.4	-6.82	0.00	4.55	25.0	0.05 [-0.30, 0.40]
					Irbesartan	94	59 (62.8)	-2.47 (8.15)	-27.3	-4.55	0.00	2.27	15.9	
				Week 94	Sparsentan	83	63 (75.9)	-1.77 (10.55)	-29.5	-6.82	0.00	4.55	20.5	0.20 [-0.15, 0.55]
					Irbesartan	94	62 (66.0)	-4.03 (12.42)	-63.6	-6.82	0.00	2.27	13.6	
		Week 110	Sparsentan	83	63 (75.9)	-1.84 (10.64)	-34.1	-6.82	0.00	4.55	20.5	0.10 [-0.26, 0.45]		
			Irbesartan	94	57 (60.6)	-2.79 (9.21)	-27.3	-9.09	0.00	2.27	18.2			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
>= 27 kg/m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	119	112 (94.1)	88.68 (12.19)	31.8	84.09	90.91	97.73	100.0			
			Irbesartan	107	95 (88.8)	89.88 (10.71)	47.7	84.09	93.18	97.73	100.0			
		Week 24	Sparsentan	119	91 (76.5)	89.81 (9.90)	50.0	84.09	93.18	97.73	100.0			
			Irbesartan	107	65 (60.7)	88.25 (17.59)	4.5	86.36	93.18	97.73	100.0			
		Week 48	Sparsentan	119	96 (80.7)	88.00 (16.41)	0.0	84.09	92.05	97.73	100.0			
			Irbesartan	107	64 (59.8)	91.44 (10.50)	43.2	88.64	94.32	100.00	100.0			
		Week 70	Sparsentan	119	96 (80.7)	87.62 (14.21)	38.6	82.95	93.18	97.73	100.0			
			Irbesartan	107	71 (66.4)	89.95 (13.57)	29.5	86.36	93.18	100.00	100.0			
		Week 94	Sparsentan	119	88 (73.9)	87.27 (12.29)	50.0	79.55	90.91	97.73	100.0			
			Irbesartan	107	67 (62.6)	90.23 (10.56)	54.5	84.09	95.45	97.73	100.0			
		Week 110	Sparsentan	119	86 (72.3)	87.68 (14.89)	6.8	81.82	93.18	97.73	100.0			
			Irbesartan	107	63 (58.9)	90.95 (13.36)	6.8	86.36	93.18	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	119	91 (76.5)	1.40 (10.90)	-22.7	-4.55	2.27	4.55	63.6	0.23 [-0.09, 0.55]
					Irbesartan	107	65 (60.7)	-1.71 (16.77)	-90.9	-4.55	0.00	2.27	29.5	
			Week 48	Sparsentan	119	96 (80.7)	-0.50 (18.01)	-100.0	-3.41	0.00	4.55	59.1	-0.05 [-0.37, 0.26]	
				Irbesartan	107	64 (59.8)	0.28 (8.62)	-18.2	-3.41	1.14	4.55	34.1		
			Week 70	Sparsentan	119	96 (80.7)	-1.16 (12.74)	-50.0	-4.55	0.00	4.55	22.7	-0.08 [-0.38, 0.23]	
				Irbesartan	107	71 (66.4)	-0.22 (11.29)	-59.1	-4.55	0.00	4.55	31.8		
			Week 94	Sparsentan	119	88 (73.9)	-0.34 (13.02)	-34.1	-4.55	0.00	4.55	68.2	-0.01 [-0.33, 0.31]	
				Irbesartan	107	67 (62.6)	-0.20 (8.91)	-34.1	-4.55	0.00	4.55	20.5		
		Week 110	Sparsentan	119	86 (72.3)	-0.48 (17.00)	-84.1	-4.55	0.00	4.55	68.2	-0.05 [-0.38, 0.27]		
			Irbesartan	107	63 (58.9)	0.32 (13.38)	-81.8	-2.27	0.00	4.55	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Randomization strata														
eGFR Low and UP High	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	71	63 (88.7)	90.98 (8.33)	63.6	86.36	93.18	97.73	100.0			
			Irbesartan	74	63 (85.1)	86.51 (13.18)	38.6	81.82	88.64	95.45	100.0			
		Week 24	Sparsentan	71	54 (76.1)	89.77 (9.86)	50.0	86.36	93.18	97.73	100.0			
			Irbesartan	74	40 (54.1)	86.25 (13.71)	45.5	82.95	88.64	95.45	100.0			
		Week 48	Sparsentan	71	51 (71.8)	90.55 (10.07)	65.9	84.09	93.18	100.00	100.0			
			Irbesartan	74	37 (50.0)	83.97 (17.24)	36.4	77.27	90.91	97.73	100.0			
		Week 70	Sparsentan	71	57 (80.3)	87.60 (13.97)	38.6	86.36	93.18	95.45	100.0			
			Irbesartan	74	37 (50.0)	85.57 (16.84)	36.4	84.09	90.91	97.73	100.0			
		Week 94	Sparsentan	71	51 (71.8)	87.97 (11.84)	52.3	84.09	90.91	95.45	100.0			
			Irbesartan	74	37 (50.0)	84.52 (15.59)	36.4	79.55	88.64	95.45	100.0			
		Week 110	Sparsentan	71	53 (74.6)	88.64 (11.79)	43.2	81.82	93.18	97.73	100.0			
			Irbesartan	74	36 (48.6)	85.80 (14.83)	45.5	77.27	90.91	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	71	54 (76.1)	-1.81 (8.13)	-25.0	-4.55	-2.27	2.27	25.0	-0.11 [-0.52, 0.30]
					Irbesartan	74	40 (54.1)	-0.80 (10.85)	-22.7	-6.82	-1.14	2.27	52.3	
				Week 48	Sparsentan	71	51 (71.8)	-0.53 (9.05)	-20.5	-6.82	0.00	4.55	27.3	0.36 [-0.07, 0.79]
					Irbesartan	74	37 (50.0)	-4.36 (12.46)	-52.3	-9.09	0.00	2.27	11.4	
				Week 70	Sparsentan	71	57 (80.3)	-3.47 (11.80)	-45.5	-6.82	0.00	2.27	18.2	-0.16 [-0.57, 0.26]
					Irbesartan	74	37 (50.0)	-1.72 (10.08)	-22.7	-9.09	0.00	2.27	31.8	
				Week 94	Sparsentan	71	51 (71.8)	-2.99 (7.50)	-20.5	-6.82	-2.27	0.00	13.6	-0.02 [-0.44, 0.40]
					Irbesartan	74	37 (50.0)	-2.83 (8.94)	-22.7	-6.82	0.00	2.27	18.2	
		Week 110	Sparsentan	71	53 (74.6)	-3.00 (9.04)	-27.3	-6.82	-2.27	2.27	18.2	-0.33 [-0.76, 0.09]		
			Irbesartan	74	36 (48.6)	0.06 (9.32)	-18.2	-6.82	0.00	4.55	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR Low and UP Low	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	55	52 (94.5)	86.28 (14.66)	31.8	81.82	88.64	96.59	100.0			
			Irbesartan	55	50 (90.9)	90.45 (7.33)	75.0	84.09	92.05	95.45	100.0			
		Week 24	Sparsentan	55	43 (78.2)	87.95 (10.62)	63.6	79.55	90.91	97.73	100.0			
			Irbesartan	55	37 (67.3)	91.58 (8.95)	68.2	88.64	95.45	97.73	100.0			
		Week 48	Sparsentan	55	43 (78.2)	84.25 (21.53)	0.0	79.55	90.91	95.45	100.0			
			Irbesartan	55	35 (63.6)	89.55 (9.99)	61.4	84.09	90.91	100.00	100.0			
		Week 70	Sparsentan	55	42 (76.4)	88.47 (11.99)	54.5	79.55	93.18	97.73	100.0			
			Irbesartan	55	39 (70.9)	89.10 (10.48)	54.5	84.09	90.91	97.73	100.0			
		Week 94	Sparsentan	55	41 (74.5)	86.97 (14.00)	45.5	79.55	90.91	97.73	100.0			
			Irbesartan	55	39 (70.9)	89.74 (9.93)	59.1	84.09	90.91	97.73	100.0			
		Week 110	Sparsentan	55	37 (67.3)	84.28 (17.84)	6.8	77.27	86.36	97.73	100.0			
			Irbesartan	55	34 (61.8)	88.84 (10.96)	59.1	81.82	92.05	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	55	43 (78.2)	2.54 (13.73)	-18.2	-6.82	0.00	6.82	63.6	0.12 [-0.32, 0.56]
					Irbesartan	55	37 (67.3)	1.23 (7.62)	-18.2	-2.27	0.00	4.55	20.5	
	Week 48	Sparsentan		55	43 (78.2)	-1.69 (25.72)	-100.0	-4.55	0.00	9.09	59.1	-0.06 [-0.50, 0.39]		
		Irbesartan		55	35 (63.6)	-0.58 (7.80)	-15.9	-4.55	0.00	4.55	20.5			
	Week 70	Sparsentan		55	42 (76.4)	1.57 (10.58)	-22.7	-4.55	2.27	6.82	25.0	0.23 [-0.20, 0.67]		
		Irbesartan		55	39 (70.9)	-0.70 (8.60)	-27.3	-4.55	0.00	4.55	20.5			
	Week 94	Sparsentan	55	41 (74.5)	1.94 (17.06)	-34.1	-4.55	0.00	6.82	68.2	0.21 [-0.23, 0.65]			
		Irbesartan	55	39 (70.9)	-0.99 (9.41)	-34.1	-4.55	0.00	4.55	20.5				
Week 110	Sparsentan	55	37 (67.3)	-0.80 (22.33)	-84.1	-6.82	0.00	6.82	68.2	0.04 [-0.43, 0.50]				
	Irbesartan	55	34 (61.8)	-1.47 (9.38)	-27.3	-4.55	0.00	2.27	20.5					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP High	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	37	35 (94.6)	88.05 (9.38)	65.9	84.09	88.64	97.73	100.0			
			Irbesartan	36	33 (91.7)	89.60 (10.68)	50.0	84.09	93.18	97.73	100.0			
		Week 24	Sparsentan	37	26 (70.3)	87.85 (19.12)	4.5	84.09	90.91	100.00	100.0			
			Irbesartan	36	26 (72.2)	88.29 (17.41)	11.4	86.36	93.18	97.73	100.0			
		Week 48	Sparsentan	37	33 (89.2)	88.43 (11.37)	61.4	81.82	90.91	97.73	100.0			
			Irbesartan	36	24 (66.7)	88.07 (12.07)	47.7	84.09	90.91	95.45	100.0			
		Week 70	Sparsentan	37	31 (83.8)	85.70 (17.76)	27.3	81.82	90.91	97.73	100.0			
			Irbesartan	36	25 (69.4)	86.36 (18.16)	29.5	86.36	90.91	95.45	100.0			
		Week 94	Sparsentan	37	27 (73.0)	87.12 (10.64)	63.6	79.55	90.91	95.45	100.0			
			Irbesartan	36	24 (66.7)	88.54 (14.96)	29.5	84.09	93.18	97.73	100.0			
		Week 110	Sparsentan	37	26 (70.3)	91.43 (9.19)	59.1	88.64	93.18	97.73	100.0			
			Irbesartan	36	22 (61.1)	87.19 (19.72)	6.8	86.36	92.05	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	37	26 (70.3)	-0.52 (16.88)	-72.7	-2.27	1.14	9.09	22.7	0.01 [-0.54, 0.55]
					Irbesartan	36	26 (72.2)	-0.61 (17.82)	-77.3	-4.55	1.14	9.09	29.5	
			Week 48	Sparsentan	37	33 (89.2)	0.41 (9.53)	-22.7	-4.55	0.00	4.55	20.5	0.15 [-0.37, 0.68]	
				Irbesartan	36	24 (66.7)	-1.23 (12.08)	-27.3	-6.82	0.00	2.27	34.1		
			Week 70	Sparsentan	37	31 (83.8)	-2.57 (16.66)	-61.4	-4.55	0.00	4.55	22.7	0.05 [-0.48, 0.57]	
				Irbesartan	36	25 (69.4)	-3.27 (14.12)	-59.1	-4.55	-2.27	2.27	15.9		
			Week 94	Sparsentan	37	27 (73.0)	-1.35 (11.59)	-29.5	-6.82	0.00	4.55	25.0	0.04 [-0.51, 0.59]	
				Irbesartan	36	24 (66.7)	-1.80 (12.25)	-45.5	-5.68	0.00	3.41	20.5		
		Week 110	Sparsentan	37	26 (70.3)	2.71 (11.82)	-31.8	-2.27	2.27	6.82	34.1	0.41 [-0.16, 0.99]		
			Irbesartan	36	22 (61.1)	-3.93 (19.98)	-81.8	-4.55	-1.14	4.55	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP Low	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	39	37 (94.9)	89.56 (11.19)	50.0	86.36	90.91	97.73	100.0			
			Irbesartan	37	36 (97.3)	91.92 (10.83)	52.3	89.77	95.45	97.73	100.0			
		Week 24	Sparsentan	39	34 (87.2)	90.64 (10.79)	54.5	86.36	94.32	97.73	100.0			
			Irbesartan	37	28 (75.7)	83.60 (25.14)	4.5	79.55	94.32	97.73	100.0			
		Week 48	Sparsentan	39	35 (89.7)	91.10 (9.66)	65.9	88.64	93.18	97.73	100.0			
			Irbesartan	37	28 (75.7)	93.91 (8.46)	65.9	92.05	96.59	100.00	100.0			
		Week 70	Sparsentan	39	30 (76.9)	88.64 (11.59)	59.1	84.09	92.05	97.73	100.0			
			Irbesartan	37	29 (78.4)	94.51 (6.83)	68.2	90.91	95.45	100.00	100.0			
		Week 94	Sparsentan	39	32 (82.1)	88.14 (12.21)	52.3	82.95	90.91	97.73	100.0			
			Irbesartan	37	30 (81.1)	89.24 (15.90)	25.0	86.36	95.45	100.00	100.0			
		Week 110	Sparsentan	39	33 (84.6)	88.15 (12.21)	52.3	79.55	90.91	97.73	100.0			
			Irbesartan	37	29 (78.4)	92.16 (8.00)	75.0	88.64	95.45	100.00	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	39	34 (87.2)	1.07 (7.59)	-22.7	0.00	2.27	4.55	13.6	0.46 [-0.05, 0.96]
					Irbesartan	37	28 (75.7)	-6.82 (24.31)	-90.9	-4.55	0.00	2.27	15.9	
			Week 48	Sparsentan	39	35 (89.7)	1.95 (6.89)	-11.4	-2.27	2.27	6.82	15.9	0.22 [-0.28, 0.72]	
				Irbesartan	37	28 (75.7)	0.49 (6.13)	-15.9	-2.27	1.14	2.27	15.9		
			Week 70	Sparsentan	39	30 (76.9)	-0.76 (11.88)	-38.6	-4.55	2.27	6.82	13.6	-0.12 [-0.63, 0.39]	
				Irbesartan	37	29 (78.4)	0.39 (7.29)	-13.6	-2.27	0.00	4.55	25.0		
			Week 94	Sparsentan	39	32 (82.1)	-0.99 (10.16)	-25.0	-6.82	0.00	5.68	15.9	0.19 [-0.31, 0.69]	
				Irbesartan	37	30 (81.1)	-3.33 (14.07)	-63.6	-6.82	0.00	2.27	13.6		
		Week 110	Sparsentan	39	33 (84.6)	-1.17 (12.97)	-45.5	-6.82	0.00	4.55	27.3	-0.07 [-0.57, 0.43]		
			Irbesartan	37	29 (78.4)	-0.39 (7.56)	-20.5	-4.55	0.00	2.27	11.4			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 1														
< 60 mL/min/1.73 m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	127	116 (91.3)	88.99 (11.76)	31.8	84.09	90.91	97.73	100.0			
			Irbesartan	129	114 (88.4)	88.62 (11.07)	38.6	84.09	90.91	95.45	100.0			
		Week 24	Sparsentan	127	98 (77.2)	88.82 (10.06)	50.0	84.09	90.91	95.45	100.0			
			Irbesartan	129	80 (62.0)	87.78 (15.06)	4.5	84.09	92.05	97.73	100.0			
		Week 48	Sparsentan	127	95 (74.8)	88.18 (16.14)	0.0	84.09	90.91	97.73	100.0			
			Irbesartan	129	74 (57.4)	87.62 (13.97)	36.4	79.55	93.18	97.73	100.0			
		Week 70	Sparsentan	127	100 (78.7)	87.68 (13.33)	38.6	81.82	93.18	95.45	100.0			
			Irbesartan	129	78 (60.5)	88.00 (13.71)	36.4	84.09	90.91	97.73	100.0			
		Week 94	Sparsentan	127	92 (72.4)	87.48 (12.23)	45.5	80.68	90.91	97.73	100.0			
			Irbesartan	129	77 (59.7)	87.66 (13.47)	36.4	81.82	90.91	97.73	100.0			
		Week 110	Sparsentan	127	92 (72.4)	86.51 (14.93)	6.8	78.41	90.91	97.73	100.0			
			Irbesartan	129	72 (55.8)	87.72 (13.00)	45.5	81.82	90.91	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	127	98 (77.2)	-0.09 (11.06)	-25.0	-4.55	-1.14	4.55	63.6	0.08 [-0.22, 0.37]
					Irbesartan	129	80 (62.0)	-1.02 (13.75)	-90.9	-4.55	0.00	2.27	52.3	
				Week 48	Sparsentan	127	95 (74.8)	-0.72 (18.19)	-100.0	-4.55	0.00	4.55	59.1	0.07 [-0.23, 0.37]
					Irbesartan	129	74 (57.4)	-1.78 (10.37)	-52.3	-4.55	0.00	2.27	20.5	
				Week 70	Sparsentan	127	100 (78.7)	-1.77 (12.06)	-45.5	-4.55	0.00	4.55	25.0	-0.08 [-0.38, 0.22]
					Irbesartan	129	78 (60.5)	-0.90 (9.01)	-27.3	-4.55	0.00	2.27	31.8	
		Week 94	Sparsentan	127	92 (72.4)	-0.96 (12.69)	-27.3	-6.82	-2.27	2.27	68.2	0.05 [-0.25, 0.36]		
			Irbesartan	129	77 (59.7)	-1.56 (9.24)	-34.1	-4.55	0.00	4.55	20.5			
		Week 110	Sparsentan	127	92 (72.4)	-2.59 (16.34)	-84.1	-7.95	0.00	2.27	68.2	-0.15 [-0.46, 0.16]		
			Irbesartan	129	72 (55.8)	-0.51 (9.27)	-27.3	-4.55	0.00	4.55	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024



Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
60 to < 90 mL/min/1.73 m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	49	47 (95.9)	88.83 (9.65)	50.0	84.09	88.64	97.73	100.0			
			Irbesartan	48	44 (91.7)	89.51 (10.53)	50.0	86.36	93.18	97.73	100.0			
		Week 24	Sparsentan	49	38 (77.6)	88.52 (17.64)	4.5	84.09	94.32	97.73	100.0			
			Irbesartan	48	31 (64.6)	88.12 (16.30)	11.4	84.09	93.18	97.73	100.0			
		Week 48	Sparsentan	49	44 (89.8)	88.58 (11.60)	54.5	82.95	92.05	97.73	100.0			
			Irbesartan	48	32 (66.7)	87.57 (13.33)	47.7	84.09	90.91	100.00	100.0			
		Week 70	Sparsentan	49	38 (77.6)	88.94 (12.12)	40.9	84.09	92.05	97.73	100.0			
			Irbesartan	48	32 (66.7)	86.22 (17.13)	29.5	86.36	90.91	97.73	100.0			
		Week 94	Sparsentan	49	39 (79.6)	87.88 (12.60)	50.0	86.36	90.91	97.73	100.0			
			Irbesartan	48	34 (70.8)	85.16 (16.58)	25.0	81.82	88.64	95.45	100.0			
		Week 110	Sparsentan	49	36 (73.5)	90.15 (9.93)	59.1	87.50	93.18	97.73	100.0			
			Irbesartan	48	32 (66.7)	87.43 (16.67)	6.8	84.09	90.91	96.59	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	49	38 (77.6)	-0.84 (14.37)	-72.7	-2.27	2.27	4.55	13.6	-0.02 [-0.49, 0.46]
					Irbesartan	48	31 (64.6)	-0.59 (16.21)	-77.3	-4.55	2.27	4.55	29.5	
	Week 48	Sparsentan		49	44 (89.8)	-0.15 (8.90)	-29.5	-4.55	0.00	4.55	18.2	0.23 [-0.23, 0.69]		
		Irbesartan		48	32 (66.7)	-2.41 (10.89)	-27.3	-6.82	-2.27	2.27	34.1			
	Week 70	Sparsentan		49	38 (77.6)	-0.18 (10.95)	-50.0	-2.27	2.27	4.55	13.6	0.39 [-0.08, 0.87]		
		Irbesartan		48	32 (66.7)	-4.76 (12.33)	-59.1	-7.95	-2.27	1.14	13.6			
	Week 94	Sparsentan	49	39 (79.6)	-1.28 (10.28)	-34.1	-2.27	0.00	4.55	15.9	0.35 [-0.12, 0.81]			
		Irbesartan	48	34 (70.8)	-5.61 (14.65)	-63.6	-9.09	-3.41	0.00	20.5				
Week 110	Sparsentan	49	36 (73.5)	1.01 (11.11)	-31.8	-5.68	0.00	6.82	27.3	0.36 [-0.12, 0.84]				
	Irbesartan	48	32 (66.7)	-3.98 (16.29)	-81.8	-6.82	-2.27	2.27	22.7					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
>= 90 mL/min/1.73 m**2	KDQOL: symptom/ problems of kidney disease	Baseline	Sparsentan	26	24 (92.3)	88.16 (11.84)	61.4	81.82	89.77	97.73	100.0			
			Irbesartan	25	24 (96.0)	91.57 (11.77)	52.3	87.50	96.59	100.00	100.0			
		Week 24	Sparsentan	26	21 (80.8)	91.77 (8.87)	70.5	88.64	90.91	100.00	100.0			
			Irbesartan	25	20 (80.0)	86.02 (22.97)	4.5	85.23	95.45	97.73	100.0			
		Week 48	Sparsentan	26	23 (88.5)	90.12 (11.13)	61.4	81.82	93.18	100.00	100.0			
			Irbesartan	25	18 (72.0)	94.32 (5.69)	81.8	88.64	94.32	100.00	100.0			
		Week 70	Sparsentan	26	22 (84.6)	85.33 (18.46)	27.3	79.55	89.77	100.00	100.0			
			Irbesartan	25	20 (80.0)	95.91 (3.58)	88.6	94.32	95.45	100.00	100.0			
		Week 94	Sparsentan	26	20 (76.9)	87.50 (12.13)	63.6	77.27	92.05	98.86	100.0			
			Irbesartan	25	19 (76.0)	93.90 (9.99)	70.5	95.45	97.73	100.00	100.0			
		Week 110	Sparsentan	26	21 (80.8)	90.37 (10.46)	68.2	84.09	93.18	100.00	100.0			
			Irbesartan	25	17 (68.0)	93.32 (8.78)	75.0	88.64	97.73	100.00	100.0			
			KDQOL: Change from baseline in symptom/ problems of kidney disease	Week 24	Sparsentan	26	21 (80.8)	3.57 (7.56)	-9.1	0.00	2.27	9.09	22.7	0.52 [-0.10, 1.14]
					Irbesartan	25	20 (80.0)	-4.66 (21.30)	-86.4	-3.41	-1.14	2.27	15.9	
				Week 48	Sparsentan	26	23 (88.5)	2.47 (9.52)	-18.2	-2.27	0.00	11.36	20.5	0.22 [-0.40, 0.83]
					Irbesartan	25	18 (72.0)	0.63 (7.05)	-15.9	-2.27	2.27	6.82	13.6	
				Week 70	Sparsentan	26	22 (84.6)	-2.27 (17.66)	-61.4	-9.09	0.00	6.82	22.7	-0.38 [-0.99, 0.23]
					Irbesartan	25	20 (80.0)	3.07 (7.95)	-9.1	-2.27	1.14	6.82	25.0	
				Week 94	Sparsentan	26	20 (76.9)	-0.11 (12.61)	-29.5	-4.55	1.14	7.95	25.0	-0.13 [-0.76, 0.50]
				Irbesartan	25	19 (76.0)	1.32 (8.55)	-25.0	0.00	0.00	2.27	13.6		
		Week 110	Sparsentan	26	21 (80.8)	2.16 (11.05)	-18.2	-4.55	2.27	6.82	34.1	0.10 [-0.53, 0.74]		
			Irbesartan	25	17 (68.0)	1.07 (9.61)	-20.5	0.00	0.00	6.82	15.9			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline eGFR Group 2														
< 45 mL/min/1.73 m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	82	73 (89.0)	87.58 (11.81)	31.8	81.82	90.91	95.45	100.0			
			Irbesartan	80	69 (86.3)	87.09 (12.66)	38.6	81.82	90.91	95.45	100.0			
		Week 24	Sparsentan	82	64 (78.0)	88.81 (9.58)	50.0	84.09	90.91	95.45	100.0			
			Irbesartan	80	48 (60.0)	87.12 (13.59)	45.5	82.95	90.91	96.59	100.0			
		Week 48	Sparsentan	82	61 (74.4)	86.74 (14.89)	0.0	79.55	90.91	95.45	100.0			
			Irbesartan	80	43 (53.8)	84.73 (16.75)	36.4	72.73	90.91	97.73	100.0			
		Week 70	Sparsentan	82	65 (79.3)	86.75 (13.46)	38.6	79.55	93.18	95.45	100.0			
			Irbesartan	80	45 (56.3)	84.49 (16.51)	36.4	77.27	88.64	97.73	100.0			
		Week 94	Sparsentan	82	60 (73.2)	86.52 (13.02)	45.5	78.41	90.91	95.45	100.0			
			Irbesartan	80	47 (58.8)	84.48 (15.25)	36.4	77.27	88.64	97.73	100.0			
		Week 110	Sparsentan	82	56 (68.3)	85.59 (16.52)	6.8	77.27	90.91	96.59	100.0			
			Irbesartan	80	44 (55.0)	84.81 (14.80)	45.5	78.41	88.64	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	82	64 (78.0)	0.67 (11.57)	-22.7	-4.55	-1.14	4.55	63.6	0.10 [-0.27, 0.48]
					Irbesartan	80	48 (60.0)	-0.47 (10.77)	-22.7	-5.68	0.00	2.27	52.3	
				Week 48	Sparsentan	82	61 (74.4)	-0.89 (17.08)	-90.9	-6.82	-2.27	4.55	59.1	0.14 [-0.25, 0.53]
					Irbesartan	80	43 (53.8)	-3.07 (12.15)	-52.3	-9.09	0.00	2.27	20.5	
				Week 70	Sparsentan	82	65 (79.3)	-1.05 (11.39)	-40.9	-4.55	0.00	4.55	25.0	0.10 [-0.28, 0.48]
					Irbesartan	80	45 (56.3)	-2.17 (10.70)	-27.3	-9.09	0.00	2.27	31.8	
		Week 94	Sparsentan	82	60 (73.2)	-1.10 (12.81)	-25.0	-6.82	-2.27	3.41	68.2	0.14 [-0.24, 0.52]		
			Irbesartan	80	47 (58.8)	-2.80 (11.00)	-34.1	-9.09	0.00	2.27	20.5			
		Week 110	Sparsentan	82	56 (68.3)	-2.64 (17.71)	-84.1	-7.95	-1.14	2.27	68.2	-0.11 [-0.50, 0.29]		
			Irbesartan	80	44 (55.0)	-0.98 (11.04)	-27.3	-7.95	0.00	4.55	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
45 to < 60 mL/min/1.73 m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	45	43 (95.6)	91.38 (11.41)	38.6	86.36	95.45	100.00	100.0			
			Irbesartan	49	45 (91.8)	90.96 (7.59)	68.2	86.36	93.18	97.73	100.0			
		Week 24	Sparsentan	45	34 (75.6)	88.84 (11.05)	59.1	81.82	93.18	95.45	100.0			
			Irbesartan	49	32 (65.3)	88.78 (17.21)	4.5	86.36	94.32	97.73	100.0			
		Week 48	Sparsentan	45	34 (75.6)	90.78 (18.12)	0.0	90.91	97.73	100.00	100.0			
			Irbesartan	49	31 (63.3)	91.64 (7.29)	77.3	86.36	93.18	97.73	100.0			
		Week 70	Sparsentan	45	35 (77.8)	89.42 (13.09)	50.0	81.82	95.45	100.00	100.0			
			Irbesartan	49	33 (67.3)	92.77 (6.05)	79.5	88.64	93.18	97.73	100.0			
		Week 94	Sparsentan	45	32 (71.1)	89.28 (10.57)	59.1	82.95	92.05	97.73	100.0			
			Irbesartan	49	30 (61.2)	92.65 (8.02)	68.2	88.64	95.45	97.73	100.0			
		Week 110	Sparsentan	45	36 (80.0)	87.94 (12.15)	52.3	79.55	93.18	97.73	100.0			
			Irbesartan	49	28 (57.1)	92.29 (7.73)	75.0	87.50	94.32	98.86	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	45	34 (75.6)	-1.54 (10.04)	-25.0	-4.55	-1.14	2.27	36.4	0.02 [-0.46, 0.50]
					Irbesartan	49	32 (65.3)	-1.85 (17.46)	-90.9	-3.41	0.00	4.55	15.9	
				Week 48	Sparsentan	45	34 (75.6)	-0.40 (20.29)	-100.0	-2.27	0.00	4.55	38.6	-0.03 [-0.51, 0.46]
					Irbesartan	49	31 (63.3)	0.00 (7.02)	-13.6	-2.27	0.00	4.55	15.9	
				Week 70	Sparsentan	45	35 (77.8)	-3.12 (13.30)	-45.5	-6.82	0.00	4.55	22.7	-0.38 [-0.86, 0.10]
					Irbesartan	49	33 (67.3)	0.83 (5.71)	-13.6	-2.27	2.27	4.55	13.6	
				Week 94	Sparsentan	45	32 (71.1)	-0.71 (12.68)	-27.3	-4.55	-1.14	2.27	40.9	-0.11 [-0.61, 0.39]
				Irbesartan	49	30 (61.2)	0.38 (5.03)	-11.4	-2.27	1.14	4.55	6.8		
		Week 110	Sparsentan	45	36 (80.0)	-2.53 (14.19)	-45.5	-6.82	0.00	2.27	36.4	-0.25 [-0.74, 0.25]		
			Irbesartan	49	28 (57.1)	0.24 (5.54)	-13.6	-2.27	0.00	3.41	11.4			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
60 to < 90 mL/min/1.73 m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	49	47 (95.9)	88.83 (9.65)	50.0	84.09	88.64	97.73	100.0			
			Irbesartan	48	44 (91.7)	89.51 (10.53)	50.0	86.36	93.18	97.73	100.0			
		Week 24	Sparsentan	49	38 (77.6)	88.52 (17.64)	4.5	84.09	94.32	97.73	100.0			
			Irbesartan	48	31 (64.6)	88.12 (16.30)	11.4	84.09	93.18	97.73	100.0			
		Week 48	Sparsentan	49	44 (89.8)	88.58 (11.60)	54.5	82.95	92.05	97.73	100.0			
			Irbesartan	48	32 (66.7)	87.57 (13.33)	47.7	84.09	90.91	100.00	100.0			
		Week 70	Sparsentan	49	38 (77.6)	88.94 (12.12)	40.9	84.09	92.05	97.73	100.0			
			Irbesartan	48	32 (66.7)	86.22 (17.13)	29.5	86.36	90.91	97.73	100.0			
		Week 94	Sparsentan	49	39 (79.6)	87.88 (12.60)	50.0	86.36	90.91	97.73	100.0			
			Irbesartan	48	34 (70.8)	85.16 (16.58)	25.0	81.82	88.64	95.45	100.0			
		Week 110	Sparsentan	49	36 (73.5)	90.15 (9.93)	59.1	87.50	93.18	97.73	100.0			
			Irbesartan	48	32 (66.7)	87.43 (16.67)	6.8	84.09	90.91	96.59	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	49	38 (77.6)	-0.84 (14.37)	-72.7	-2.27	2.27	4.55	13.6	-0.02 [-0.49, 0.46]
					Irbesartan	48	31 (64.6)	-0.59 (16.21)	-77.3	-4.55	2.27	4.55	29.5	
			Week 48	Sparsentan	49	44 (89.8)	-0.15 (8.90)	-29.5	-4.55	0.00	4.55	18.2	0.23 [-0.23, 0.69]	
				Irbesartan	48	32 (66.7)	-2.41 (10.89)	-27.3	-6.82	-2.27	2.27	34.1		
			Week 70	Sparsentan	49	38 (77.6)	-0.18 (10.95)	-50.0	-2.27	2.27	4.55	13.6	0.39 [-0.08, 0.87]	
				Irbesartan	48	32 (66.7)	-4.76 (12.33)	-59.1	-7.95	-2.27	1.14	13.6		
			Week 94	Sparsentan	49	39 (79.6)	-1.28 (10.28)	-34.1	-2.27	0.00	4.55	15.9	0.35 [-0.12, 0.81]	
				Irbesartan	48	34 (70.8)	-5.61 (14.65)	-63.6	-9.09	-3.41	0.00	20.5		
		Week 110	Sparsentan	49	36 (73.5)	1.01 (11.11)	-31.8	-5.68	0.00	6.82	27.3	0.36 [-0.12, 0.84]		
			Irbesartan	48	32 (66.7)	-3.98 (16.29)	-81.8	-6.82	-2.27	2.27	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
>= 90 mL/min/1.73 m**2	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	26	24 (92.3)	88.16 (11.84)	61.4	81.82	89.77	97.73	100.0			
			Irbesartan	25	24 (96.0)	91.57 (11.77)	52.3	87.50	96.59	100.00	100.0			
		Week 24	Sparsentan	26	21 (80.8)	91.77 (8.87)	70.5	88.64	90.91	100.00	100.0			
			Irbesartan	25	20 (80.0)	86.02 (22.97)	4.5	85.23	95.45	97.73	100.0			
		Week 48	Sparsentan	26	23 (88.5)	90.12 (11.13)	61.4	81.82	93.18	100.00	100.0			
			Irbesartan	25	18 (72.0)	94.32 (5.69)	81.8	88.64	94.32	100.00	100.0			
		Week 70	Sparsentan	26	22 (84.6)	85.33 (18.46)	27.3	79.55	89.77	100.00	100.0			
			Irbesartan	25	20 (80.0)	95.91 (3.58)	88.6	94.32	95.45	100.00	100.0			
		Week 94	Sparsentan	26	20 (76.9)	87.50 (12.13)	63.6	77.27	92.05	98.86	100.0			
			Irbesartan	25	19 (76.0)	93.90 (9.99)	70.5	95.45	97.73	100.00	100.0			
		Week 110	Sparsentan	26	21 (80.8)	90.37 (10.46)	68.2	84.09	93.18	100.00	100.0			
			Irbesartan	25	17 (68.0)	93.32 (8.78)	75.0	88.64	97.73	100.00	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	26	21 (80.8)	3.57 (7.56)	-9.1	0.00	2.27	9.09	22.7	0.52 [-0.10, 1.14]
					Irbesartan	25	20 (80.0)	-4.66 (21.30)	-86.4	-3.41	-1.14	2.27	15.9	
			Week 48	Sparsentan	26	23 (88.5)	2.47 (9.52)	-18.2	-2.27	0.00	11.36	20.5	0.22 [-0.40, 0.83]	
				Irbesartan	25	18 (72.0)	0.63 (7.05)	-15.9	-2.27	2.27	6.82	13.6		
			Week 70	Sparsentan	26	22 (84.6)	-2.27 (17.66)	-61.4	-9.09	0.00	6.82	22.7	-0.38 [-0.99, 0.23]	
				Irbesartan	25	20 (80.0)	3.07 (7.95)	-9.1	-2.27	1.14	6.82	25.0		
			Week 94	Sparsentan	26	20 (76.9)	-0.11 (12.61)	-29.5	-4.55	1.14	7.95	25.0	-0.13 [-0.76, 0.50]	
				Irbesartan	25	19 (76.0)	1.32 (8.55)	-25.0	0.00	0.00	2.27	13.6		
		Week 110	Sparsentan	26	21 (80.8)	2.16 (11.05)	-18.2	-4.55	2.27	6.82	34.1	0.10 [-0.53, 0.74]		
			Irbesartan	25	17 (68.0)	1.07 (9.61)	-20.5	0.00	0.00	6.82	15.9			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline urine protein excretion														
<= 1.75 g/day	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	98	93 (94.9)	87.85 (13.45)	31.8	84.09	90.91	97.73	100.0			
			Irbesartan	93	83 (89.2)	90.47 (9.47)	52.3	86.36	93.18	97.73	100.0			
		Week 24	Sparsentan	98	80 (81.6)	88.55 (10.53)	54.5	81.82	90.91	97.73	100.0			
			Irbesartan	93	55 (59.1)	87.89 (19.21)	4.5	84.09	95.45	97.73	100.0			
		Week 48	Sparsentan	98	82 (83.7)	87.97 (17.09)	0.0	81.82	93.18	97.73	100.0			
			Irbesartan	93	57 (61.3)	91.71 (9.54)	65.9	88.64	93.18	100.00	100.0			
		Week 70	Sparsentan	98	75 (76.5)	87.24 (13.54)	40.9	79.55	90.91	97.73	100.0			
			Irbesartan	93	61 (65.6)	91.54 (9.38)	61.4	88.64	95.45	100.00	100.0			
		Week 94	Sparsentan	98	76 (77.6)	87.35 (12.63)	45.5	79.55	90.91	97.73	100.0			
			Irbesartan	93	63 (67.7)	88.28 (14.79)	25.0	81.82	93.18	97.73	100.0			
		Week 110	Sparsentan	98	75 (76.5)	87.06 (13.40)	43.2	77.27	90.91	97.73	100.0			
			Irbesartan	93	57 (61.3)	89.87 (10.86)	52.3	84.09	93.18	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	98	80 (81.6)	0.94 (11.43)	-22.7	-5.68	0.00	4.55	63.6	0.22 [-0.13, 0.56]
					Irbesartan	93	55 (59.1)	-2.27 (18.41)	-90.9	-2.27	0.00	4.55	20.5	
				Week 48	Sparsentan	98	82 (83.7)	0.19 (19.15)	-100.0	-2.27	0.00	6.82	59.1	0.01 [-0.33, 0.35]
					Irbesartan	93	57 (61.3)	0.04 (6.84)	-18.2	-2.27	0.00	2.27	20.5	
				Week 70	Sparsentan	98	75 (76.5)	-0.97 (12.79)	-50.0	-4.55	0.00	4.55	25.0	-0.08 [-0.41, 0.26]
					Irbesartan	93	61 (65.6)	-0.15 (7.51)	-15.9	-4.55	0.00	4.55	25.0	
		Week 94	Sparsentan	98	76 (77.6)	0.39 (13.85)	-27.3	-6.82	0.00	5.68	68.2	0.24 [-0.09, 0.58]		
			Irbesartan	93	63 (67.7)	-2.78 (11.87)	-63.6	-6.82	0.00	2.27	20.5			
		Week 110	Sparsentan	98	75 (76.5)	-0.42 (15.81)	-45.5	-9.09	0.00	6.82	68.2	0.05 [-0.29, 0.40]		
			Irbesartan	93	57 (61.3)	-1.12 (7.62)	-20.5	-4.55	0.00	2.27	20.5			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 1.75 g/day	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	104	94 (90.4)	89.82 (8.42)	65.9	84.09	90.91	97.73	100.0			
			Irbesartan	109	99 (90.8)	88.18 (12.11)	38.6	84.09	90.91	97.73	100.0			
		Week 24	Sparsentan	104	77 (74.0)	89.76 (13.69)	4.5	88.64	93.18	97.73	100.0			
			Irbesartan	109	76 (69.7)	87.38 (14.62)	11.4	84.09	90.91	95.45	100.0			
		Week 48	Sparsentan	104	80 (76.9)	89.18 (10.89)	54.5	82.95	90.91	97.73	100.0			
			Irbesartan	109	67 (61.5)	85.92 (15.01)	36.4	79.55	90.91	97.73	100.0			
		Week 70	Sparsentan	104	85 (81.7)	88.02 (14.13)	27.3	86.36	93.18	97.73	100.0			
			Irbesartan	109	69 (63.3)	86.33 (16.69)	29.5	84.09	90.91	97.73	100.0			
		Week 94	Sparsentan	104	75 (72.1)	87.82 (11.91)	50.0	84.09	90.91	95.45	100.0			
			Irbesartan	109	67 (61.5)	87.58 (13.50)	29.5	81.82	90.91	97.73	100.0			
		Week 110	Sparsentan	104	74 (71.2)	88.82 (13.35)	6.8	81.82	93.18	97.73	100.0			
			Irbesartan	109	64 (58.7)	87.14 (15.68)	6.8	81.82	92.05	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	104	77 (74.0)	-0.53 (11.77)	-72.7	-4.55	0.00	4.55	25.0	0.03 [-0.29, 0.35]
					Irbesartan	109	76 (69.7)	-0.90 (13.33)	-77.3	-5.68	0.00	2.27	52.3	
			Week 48	Sparsentan	104	80 (76.9)	-0.43 (9.38)	-29.5	-4.55	0.00	4.55	27.3	0.24 [-0.09, 0.57]	
				Irbesartan	109	67 (61.5)	-2.99 (12.05)	-52.3	-9.09	0.00	2.27	34.1		
			Week 70	Sparsentan	104	85 (81.7)	-1.90 (12.60)	-61.4	-4.55	0.00	4.55	22.7	0.03 [-0.29, 0.34]	
				Irbesartan	109	69 (63.3)	-2.21 (11.77)	-59.1	-4.55	0.00	2.27	31.8		
			Week 94	Sparsentan	104	75 (72.1)	-2.27 (9.77)	-34.1	-4.55	-2.27	2.27	25.0	-0.06 [-0.39, 0.27]	
				Irbesartan	109	67 (61.5)	-1.66 (10.13)	-45.5	-6.82	0.00	4.55	20.5		
		Week 110	Sparsentan	104	74 (71.2)	-1.69 (13.40)	-84.1	-4.55	0.00	2.27	34.1	-0.03 [-0.36, 0.30]		
			Irbesartan	109	64 (58.7)	-1.28 (14.32)	-81.8	-4.55	0.00	4.55	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024



Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline use of antihypertensives														
Yes	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	90	79 (87.8)	88.38 (10.79)	38.6	84.09	90.91	95.45	100.0			
		Week 24	Irbesartan	88	76 (86.4)	88.88 (10.04)	38.6	84.09	90.91	95.45	100.0			
			Sparsentan	90	66 (73.3)	88.81 (10.77)	50.0	84.09	90.91	97.73	100.0			
		Week 48	Irbesartan	88	49 (55.7)	89.42 (9.27)	56.8	86.36	93.18	95.45	100.0			
			Sparsentan	90	65 (72.2)	89.13 (10.79)	54.5	84.09	90.91	97.73	100.0			
		Week 70	Irbesartan	88	47 (53.4)	88.88 (11.49)	47.7	86.36	93.18	95.45	100.0			
			Sparsentan	90	68 (75.6)	86.80 (13.84)	38.6	81.82	90.91	95.45	100.0			
		Week 94	Irbesartan	88	49 (55.7)	88.50 (12.00)	40.9	84.09	90.91	97.73	100.0			
			Sparsentan	90	63 (70.0)	86.15 (13.39)	45.5	79.55	90.91	95.45	100.0			
		Week 110	Irbesartan	88	54 (61.4)	87.54 (11.80)	36.4	81.82	90.91	97.73	100.0			
			Sparsentan	90	63 (70.0)	88.17 (14.02)	6.8	81.82	93.18	97.73	100.0			
				Irbesartan	88	51 (58.0)	88.95 (10.53)	52.3	84.09	90.91	97.73	100.0		
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	90	66 (73.3)	0.48 (10.40)	-25.0	-4.55	0.00	4.55	36.4	-0.01 [-0.38, 0.36]
		Week 48		Irbesartan	88	49 (55.7)	0.56 (9.99)	-22.7	-4.55	0.00	4.55	52.3		
				Sparsentan	90	65 (72.2)	1.08 (10.71)	-29.5	-2.27	0.00	4.55	38.6	0.19 [-0.18, 0.57]	
		Week 70		Irbesartan	88	47 (53.4)	-0.87 (9.50)	-27.3	-4.55	0.00	4.55	15.9		
				Sparsentan	90	68 (75.6)	-2.47 (12.35)	-45.5	-7.95	0.00	4.55	25.0	-0.08 [-0.45, 0.28]	
		Week 94		Irbesartan	88	49 (55.7)	-1.58 (8.24)	-22.7	-4.55	0.00	2.27	18.2		
Sparsentan	90			63 (70.0)	-1.55 (11.86)	-34.1	-4.55	-2.27	4.55	40.9	0.08 [-0.28, 0.44]			
Week 110	Irbesartan	88		54 (61.4)	-2.40 (8.82)	-22.7	-6.82	0.00	2.27	20.5				
	Sparsentan	90		63 (70.0)	-0.11 (14.79)	-84.1	-4.55	0.00	4.55	36.4	-0.02 [-0.39, 0.35]			
		Irbesartan		88	51 (58.0)	0.13 (7.75)	-18.2	-2.27	0.00	4.55	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
No	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	112	108 (96.4)	89.18 (11.57)	31.8	84.09	90.91	97.73	100.0		
			Irbesartan	114	106 (93.0)	89.47 (11.71)	47.7	84.09	93.18	97.73	100.0		
		Week 24	Sparsentan	112	91 (81.3)	89.39 (13.13)	4.5	86.36	93.18	97.73	100.0		
			Irbesartan	114	82 (71.9)	86.50 (19.75)	4.5	84.09	93.18	97.73	100.0		
		Week 48	Sparsentan	112	97 (86.6)	88.19 (16.33)	0.0	81.82	93.18	97.73	100.0		
			Irbesartan	114	77 (67.5)	88.40 (14.01)	36.4	84.09	90.91	100.00	100.0		
		Week 70	Sparsentan	112	92 (82.1)	88.29 (13.84)	27.3	81.82	93.18	97.73	100.0		
			Irbesartan	114	81 (71.1)	88.95 (15.07)	29.5	86.36	93.18	100.00	100.0		
		Week 94	Sparsentan	112	88 (78.6)	88.61 (11.30)	52.3	81.82	90.91	97.73	100.0		
			Irbesartan	114	76 (66.7)	88.19 (15.58)	25.0	82.95	95.45	98.86	100.0		
		Week 110	Sparsentan	112	86 (76.8)	87.76 (12.94)	43.2	79.55	92.05	97.73	100.0		
			Irbesartan	114	70 (61.4)	88.05 (15.58)	6.8	81.82	93.18	97.73	100.0		
		KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	112	91 (81.3)	0.02 (12.43)	-72.7	-4.55	0.00	4.55	63.6	0.18 [-0.12, 0.48]
				Irbesartan	114	82 (71.9)	-2.69 (18.11)	-90.9	-4.55	0.00	4.55	29.5	
	Week 48		Sparsentan	112	97 (86.6)	-0.91 (17.43)	-100.0	-4.55	0.00	4.55	59.1	0.08 [-0.22, 0.38]	
			Irbesartan	114	77 (67.5)	-2.04 (10.44)	-52.3	-4.55	0.00	2.27	34.1		
	Week 70		Sparsentan	112	92 (82.1)	-0.72 (12.90)	-61.4	-4.55	0.00	6.82	22.7	0.03 [-0.27, 0.33]	
			Irbesartan	114	81 (71.1)	-1.04 (11.00)	-59.1	-4.55	0.00	4.55	31.8		
	Week 94		Sparsentan	112	88 (78.6)	-0.49 (12.20)	-29.5	-6.82	0.00	4.55	68.2	0.13 [-0.18, 0.44]	
			Irbesartan	114	76 (66.7)	-2.06 (12.34)	-63.6	-4.55	0.00	2.27	20.5		
Week 110	Sparsentan		112	86 (76.8)	-1.74 (14.55)	-45.5	-6.82	0.00	2.27	68.2	0.03 [-0.29, 0.35]		
	Irbesartan		114	70 (61.4)	-2.18 (13.73)	-81.8	-6.82	0.00	4.55	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Time since renal biopsy														
<= 5 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	113	104 (92.0)	88.96 (11.04)	38.6	84.09	90.91	97.73	100.0			
			Irbesartan	127	118 (92.9)	88.66 (12.00)	38.6	84.09	90.91	97.73	100.0			
		Week 24	Sparsentan	113	88 (77.9)	90.55 (10.12)	50.0	86.36	93.18	97.73	100.0			
			Irbesartan	127	86 (67.7)	86.05 (19.07)	4.5	81.82	93.18	97.73	100.0			
		Week 48	Sparsentan	113	92 (81.4)	90.04 (13.52)	0.0	86.36	93.18	97.73	100.0			
			Irbesartan	127	85 (66.9)	88.32 (14.37)	36.4	86.36	93.18	100.00	100.0			
		Week 70	Sparsentan	113	90 (79.6)	87.55 (14.62)	27.3	81.82	93.18	97.73	100.0			
			Irbesartan	127	86 (67.7)	88.64 (14.91)	29.5	86.36	93.18	97.73	100.0			
		Week 94	Sparsentan	113	86 (76.1)	87.63 (12.97)	45.5	81.82	90.91	97.73	100.0			
			Irbesartan	127	86 (67.7)	88.37 (13.55)	29.5	81.82	93.18	97.73	100.0			
		Week 110	Sparsentan	113	88 (77.9)	89.46 (10.54)	50.0	82.95	92.05	97.73	100.0			
			Irbesartan	127	80 (63.0)	88.15 (14.42)	6.8	81.82	93.18	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	113	88 (77.9)	1.60 (8.50)	-22.7	-2.27	1.14	4.55	36.4	0.25 [-0.05, 0.54]
					Irbesartan	127	86 (67.7)	-1.98 (18.82)	-90.9	-4.55	0.00	4.55	52.3	
				Week 48	Sparsentan	113	92 (81.4)	1.24 (14.56)	-100.0	-2.27	0.00	6.82	38.6	0.18 [-0.12, 0.48]
					Irbesartan	127	85 (66.9)	-1.10 (10.89)	-52.3	-4.55	0.00	2.27	34.1	
				Week 70	Sparsentan	113	90 (79.6)	-1.99 (13.49)	-61.4	-4.55	0.00	4.55	25.0	-0.07 [-0.37, 0.22]
					Irbesartan	127	86 (67.7)	-1.11 (10.73)	-59.1	-4.55	0.00	4.55	25.0	
		Week 94	Sparsentan	113	86 (76.1)	-0.90 (11.51)	-34.1	-4.55	0.00	4.55	40.9	0.07 [-0.23, 0.37]		
			Irbesartan	127	86 (67.7)	-1.61 (9.91)	-45.5	-6.82	0.00	4.55	20.5			
		Week 110	Sparsentan	113	88 (77.9)	0.31 (11.52)	-45.5	-4.55	0.00	4.55	36.4	0.11 [-0.19, 0.42]		
			Irbesartan	127	80 (63.0)	-1.05 (12.74)	-81.8	-4.55	0.00	4.55	22.7			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
> 5 years	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	89	83 (93.3)	88.69 (11.52)	31.8	84.09	90.91	97.73	100.0			
			Irbesartan	75	64 (85.3)	90.27 (8.91)	52.3	86.36	93.18	96.59	100.0			
		Week 24	Sparsentan	89	69 (77.5)	87.35 (14.23)	4.5	81.82	90.91	97.73	100.0			
			Irbesartan	75	45 (60.0)	90.56 (10.07)	45.5	86.36	93.18	97.73	100.0			
		Week 48	Sparsentan	89	70 (78.7)	86.62 (15.22)	0.0	79.55	90.91	97.73	100.0			
			Irbesartan	75	39 (52.0)	89.16 (9.77)	61.4	84.09	90.91	95.45	100.0			
		Week 70	Sparsentan	89	70 (78.7)	87.79 (12.81)	40.9	81.82	93.18	95.45	100.0			
			Irbesartan	75	44 (58.7)	89.05 (12.01)	36.4	84.09	90.91	97.73	100.0			
		Week 94	Sparsentan	89	65 (73.0)	87.52 (11.29)	63.6	77.27	90.91	95.45	100.0			
			Irbesartan	75	44 (58.7)	87.04 (15.20)	25.0	81.82	88.64	97.73	100.0			
		Week 110	Sparsentan	89	61 (68.5)	85.73 (16.45)	6.8	77.27	93.18	97.73	100.0			
			Irbesartan	75	41 (54.7)	88.97 (12.11)	45.5	81.82	90.91	100.00	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	89	69 (77.5)	-1.55 (14.48)	-72.7	-6.82	-2.27	4.55	63.6	-0.09 [-0.46, 0.29]
				Irbesartan	75	45 (60.0)	-0.51 (5.85)	-18.2	-4.55	0.00	2.27	11.4		
	Week 48	Sparsentan		89	70 (78.7)	-1.88 (15.69)	-90.9	-6.82	-2.27	4.55	59.1	0.06 [-0.33, 0.45]		
		Irbesartan		75	39 (52.0)	-2.68 (8.04)	-27.3	-6.82	-2.27	2.27	11.4			
	Week 70	Sparsentan		89	70 (78.7)	-0.78 (11.57)	-50.0	-4.55	0.00	4.55	22.7	0.07 [-0.31, 0.45]		
		Irbesartan		75	44 (58.7)	-1.50 (8.57)	-15.9	-4.55	-1.14	2.27	31.8			
	Week 94	Sparsentan	89	65 (73.0)	-0.98 (12.78)	-27.3	-6.82	-2.27	2.27	68.2	0.19 [-0.20, 0.57]			
		Irbesartan	75	44 (58.7)	-3.36 (12.86)	-63.6	-6.82	0.00	2.27	18.2				
Week 110	Sparsentan	89	61 (68.5)	-3.02 (18.13)	-84.1	-9.09	-2.27	2.27	68.2	-0.10 [-0.50, 0.30]				
	Irbesartan	75	41 (54.7)	-1.50 (9.13)	-18.2	-4.55	0.00	2.27	22.7					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: History of hypertension														
Yes	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	155	141 (91.0)	88.31 (10.78)	38.6	84.09	90.91	95.45	100.0			
			Irbesartan	161	142 (88.2)	89.34 (10.98)	38.6	84.09	93.18	97.73	100.0			
		Week 24	Sparsentan	155	119 (76.8)	88.92 (10.87)	50.0	84.09	93.18	97.73	100.0			
			Irbesartan	161	100 (62.1)	89.30 (13.32)	4.5	86.36	93.18	97.73	100.0			
		Week 48	Sparsentan	155	121 (78.1)	87.94 (13.44)	0.0	81.82	90.91	97.73	100.0			
			Irbesartan	161	95 (59.0)	88.66 (13.46)	36.4	84.09	93.18	97.73	100.0			
		Week 70	Sparsentan	155	123 (79.4)	87.84 (12.72)	38.6	81.82	93.18	97.73	100.0			
			Irbesartan	161	100 (62.1)	88.91 (13.98)	36.4	85.23	93.18	97.73	100.0			
		Week 94	Sparsentan	155	114 (73.5)	86.86 (12.33)	45.5	79.55	90.91	95.45	100.0			
			Irbesartan	161	101 (62.7)	87.56 (14.67)	25.0	81.82	90.91	97.73	100.0			
		Week 110	Sparsentan	155	112 (72.3)	87.60 (13.00)	6.8	81.82	90.91	97.73	100.0			
			Irbesartan	161	92 (57.1)	89.70 (11.16)	45.5	84.09	93.18	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	155	119 (76.8)	0.67 (8.86)	-25.0	-4.55	0.00	4.55	36.4	0.05 [-0.21, 0.32]
		Irbesartan			161	100 (62.1)	0.09 (12.56)	-90.9	-4.55	0.00	4.55	52.3		
		Week 48		Sparsentan	155	121 (78.1)	-0.11 (13.02)	-90.9	-4.55	0.00	4.55	38.6	0.12 [-0.15, 0.39]	
				Irbesartan	161	95 (59.0)	-1.60 (10.71)	-52.3	-4.55	0.00	2.27	34.1		
		Week 70		Sparsentan	155	123 (79.4)	-0.76 (11.01)	-45.5	-4.55	0.00	4.55	25.0	-0.03 [-0.29, 0.23]	
				Irbesartan	161	100 (62.1)	-0.45 (8.81)	-27.3	-4.55	0.00	2.27	31.8		
Week 94	Sparsentan	155		114 (73.5)	-0.96 (10.67)	-34.1	-6.82	-1.14	4.55	40.9	0.13 [-0.14, 0.39]			
	Irbesartan	161		101 (62.7)	-2.32 (10.86)	-63.6	-6.82	0.00	2.27	20.5				
Week 110	Sparsentan	155		112 (72.3)	-0.89 (13.49)	-84.1	-4.55	0.00	4.55	36.4	-0.11 [-0.38, 0.17]			
	Irbesartan	161		92 (57.1)	0.32 (8.08)	-18.2	-4.55	0.00	4.55	22.7				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSHM: KDQOL: Course of symptom/problems of kidney disease by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
No	KDQOL: symptom/problems of kidney disease	Baseline	Sparsentan	47	46 (97.9)	90.46 (12.47)	31.8	86.36	94.32	100.00	100.0			
			Irbesartan	41	40 (97.6)	88.81 (11.29)	52.3	86.36	90.91	97.73	100.0			
		Week 24	Sparsentan	47	38 (80.9)	89.83 (15.70)	4.5	88.64	93.18	97.73	100.0			
			Irbesartan	41	31 (75.6)	82.11 (23.93)	4.5	72.73	90.91	97.73	100.0			
		Week 48	Sparsentan	47	41 (87.2)	90.41 (16.74)	0.0	88.64	95.45	100.00	100.0			
			Irbesartan	41	29 (70.7)	88.32 (11.91)	54.5	84.09	90.91	97.73	100.0			
		Week 70	Sparsentan	47	37 (78.7)	87.04 (17.15)	27.3	84.09	93.18	95.45	100.0			
			Irbesartan	41	30 (73.2)	88.33 (14.07)	29.5	86.36	90.91	97.73	100.0			
		Week 94	Sparsentan	47	37 (78.7)	89.80 (11.84)	63.6	86.36	93.18	100.00	100.0			
			Irbesartan	41	29 (70.7)	89.18 (11.99)	59.1	81.82	93.18	100.00	100.0			
		Week 110	Sparsentan	47	37 (78.7)	88.94 (14.52)	43.2	86.36	93.18	100.00	100.0			
			Irbesartan	41	29 (70.7)	84.40 (19.23)	6.8	77.27	88.64	97.73	100.0			
			KDQOL: Change from baseline in symptom/problems of kidney disease	Week 24	Sparsentan	47	38 (80.9)	-1.20 (17.69)	-72.7	-6.82	0.00	4.55	63.6	0.27 [-0.21, 0.74]
					Irbesartan	41	31 (75.6)	-6.52 (22.39)	-86.4	-9.09	0.00	4.55	20.5	
	Week 48	Sparsentan		47	41 (87.2)	-0.11 (20.22)	-100.0	-2.27	0.00	6.82	59.1	0.09 [-0.39, 0.57]		
		Irbesartan		41	29 (70.7)	-1.57 (7.80)	-20.5	-4.55	0.00	2.27	20.5			
	Week 70	Sparsentan		47	37 (78.7)	-3.81 (17.02)	-61.4	-4.55	0.00	4.55	22.7	0.00 [-0.48, 0.49]		
		Irbesartan		41	30 (73.2)	-3.86 (13.12)	-59.1	-4.55	-2.27	0.00	20.5			
	Week 94	Sparsentan	47	37 (78.7)	-0.86 (15.68)	-27.3	-4.55	0.00	2.27	68.2	0.07 [-0.42, 0.55]			
		Irbesartan	41	29 (70.7)	-1.80 (11.56)	-34.1	-6.82	0.00	2.27	20.5				
Week 110	Sparsentan	47	37 (78.7)	-1.54 (17.84)	-45.5	-9.09	0.00	4.55	68.2	0.25 [-0.24, 0.74]				
	Irbesartan	41	29 (70.7)	-6.03 (18.26)	-81.8	-13.64	-2.27	2.27	20.5					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Time	Treatment	KDQOL: Change from baseline in symptom/problems of kidney disease		Repeated measures analysis				
					Change from Baseline		Treatment Difference		
			N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Sex	Overall	Sparsentan							Interaction: 0.334
Male	Week 24	Sparsentan	139	109 (78.4)	-1.49 (1.13)	(-3.70, 0.72)	0.59 (1.66)	(-2.67, 3.85)	0.723
		Irbesartan	143	93 (65.0)	-2.08 (1.22)	(-4.47, 0.31)			
	Week 48	Sparsentan	139	112 (80.6)	-2.11 (1.11)	(-4.28, 0.07)	-0.19 (1.67)	(-3.46, 3.08)	0.909
		Irbesartan	143	87 (60.8)	-1.92 (1.24)	(-4.36, 0.52)			
	Week 70	Sparsentan	139	114 (82.0)	-2.08 (1.10)	(-4.24, 0.09)	-0.73 (1.65)	(-3.97, 2.50)	0.657
		Irbesartan	143	92 (64.3)	-1.34 (1.22)	(-3.74, 1.05)			
	Week 94	Sparsentan	139	105 (75.5)	-2.05 (1.14)	(-4.29, 0.19)	0.60 (1.66)	(-2.66, 3.86)	0.719
		Irbesartan	143	96 (67.1)	-2.65 (1.20)	(-5.01, -0.29)			
	Week 110	Sparsentan	139	103 (74.1)	-2.15 (1.16)	(-4.43, 0.12)	-0.33 (1.71)	(-3.68, 3.02)	0.845
		Irbesartan	143	89 (62.2)	-1.82 (1.25)	(-4.27, 0.63)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Time	Treatment	KDQOL: Change from baseline in symptom/problems of kidney disease		Repeated measures analysis				
					Change from Baseline		Treatment Difference		
			N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Female	Week 24	Sparsentan	63	48 (76.2)	2.51 (1.76)	(-0.95, 5.96)	2.18 (2.66)	(-3.05, 7.41)	0.412
		Irbesartan	59	38 (64.4)	0.32 (1.98)	(-3.57, 4.22)			
	Week 48	Sparsentan	63	50 (79.4)	2.14 (1.72)	(-1.24, 5.52)	3.20 (2.64)	(-1.99, 8.40)	0.226
		Irbesartan	59	37 (62.7)	-1.06 (1.99)	(-4.98, 2.85)			
	Week 70	Sparsentan	63	46 (73.0)	-0.25 (1.78)	(-3.75, 3.25)	0.60 (2.68)	(-4.67, 5.86)	0.824
		Irbesartan	59	38 (64.4)	-0.85 (1.98)	(-4.74, 3.05)			
	Week 94	Sparsentan	63	46 (73.0)	0.08 (1.79)	(-3.45, 3.60)	0.74 (2.76)	(-4.70, 6.17)	0.790
		Irbesartan	59	34 (57.6)	-0.66 (2.08)	(-4.75, 3.42)			
	Week 110	Sparsentan	63	46 (73.0)	0.66 (1.80)	(-2.89, 4.21)	-0.20 (2.83)	(-5.77, 5.37)	0.944
		Irbesartan	59	32 (54.2)	0.86 (2.16)	(-3.39, 5.10)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024



Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Age	Overall		Sparsentan							Interaction: 0.045 #
<= 45 years	Week 24		Sparsentan	96	71 (74.0)	0.69 (1.49)	(-2.24, 3.62)	3.56 (2.14)	(-0.65, 7.76)	0.097
			Irbesartan	99	67 (67.7)	-2.87 (1.54)	(-5.88, 0.15)			
	Week 48		Sparsentan	96	75 (78.1)	-1.10 (1.45)	(-3.94, 1.75)	1.77 (2.14)	(-2.44, 5.98)	0.408
			Irbesartan	99	62 (62.6)	-2.87 (1.58)	(-5.97, 0.23)			
	Week 70		Sparsentan	96	72 (75.0)	-0.82 (1.48)	(-3.72, 2.08)	0.96 (2.17)	(-3.30, 5.22)	0.659
			Irbesartan	99	62 (62.6)	-1.78 (1.59)	(-4.90, 1.34)			
	Week 94		Sparsentan	96	68 (70.8)	-0.60 (1.52)	(-3.57, 2.38)	3.68 (2.18)	(-0.60, 7.97)	0.092
			Irbesartan	99	64 (64.6)	-4.28 (1.57)	(-7.36, -1.20)			
	Week 110		Sparsentan	96	69 (71.9)	0.36 (1.52)	(-2.62, 3.34)	1.90 (2.23)	(-2.47, 6.28)	0.394
			Irbesartan	99	59 (59.6)	-1.54 (1.63)	(-4.75, 1.67)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Time	Treatment	KDQOL: Change from baseline in symptom/problems of kidney disease		Repeated measures analysis				
					Change from Baseline		Treatment Difference		
			N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 45 years	Week 24	Sparsentan	106	86 (81.1)	-1.00 (1.23)	(-3.42, 1.42)	-1.03 (1.89)	(-4.74, 2.68)	0.586
		Irbesartan	103	64 (62.1)	0.03 (1.43)	(-2.77, 2.84)			
	Week 48	Sparsentan	106	87 (82.1)	-0.42 (1.22)	(-2.82, 1.97)	0.34 (1.88)	(-3.35, 4.03)	0.856
		Irbesartan	103	62 (60.2)	-0.76 (1.43)	(-3.56, 2.04)			
	Week 70	Sparsentan	106	88 (83.0)	-1.88 (1.22)	(-4.28, 0.52)	-1.02 (1.85)	(-4.64, 2.61)	0.582
		Irbesartan	103	68 (66.0)	-0.86 (1.38)	(-3.58, 1.85)			
	Week 94	Sparsentan	106	83 (78.3)	-1.99 (1.25)	(-4.44, 0.47)	-1.64 (1.88)	(-5.33, 2.06)	0.384
		Irbesartan	103	66 (64.1)	-0.35 (1.40)	(-3.11, 2.40)			
	Week 110	Sparsentan	106	80 (75.5)	-2.51 (1.28)	(-5.02, 0.00)	-1.52 (1.94)	(-5.32, 2.29)	0.434
		Irbesartan	103	62 (60.2)	-1.00 (1.45)	(-3.85, 1.86)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Age at IgAN diagnosis	Overall		Sparsentan							Interaction: 0.252
<= 18 years	Week 24		Sparsentan	9	5 (55.6)	-6.74 (9.23)	(-28.15, 14.67)	-6.64 (15.33)	(-42.75, 29.48)	0.678
			Irbesartan	5	5 (100.0)	-0.10 (11.52)	(-27.42, 27.22)			
	Week 48		Sparsentan	9	7 (77.8)	-11.32 (8.98)	(-32.48, 9.85)	-9.14 (15.33)	(-45.24, 26.96)	0.569
			Irbesartan	5	3 (60.0)	-2.18 (11.73)	(-29.67, 25.32)			
	Week 70		Sparsentan	9	7 (77.8)	-7.40 (9.04)	(-28.62, 13.83)	-2.15 (15.37)	(-38.30, 33.99)	0.892
			Irbesartan	5	4 (80.0)	-5.24 (11.71)	(-32.74, 22.25)			
	Week 94		Sparsentan	9	5 (55.6)	-12.30 (9.25)	(-33.75, 9.15)	-8.21 (15.64)	(-44.64, 28.22)	0.615
			Irbesartan	5	4 (80.0)	-4.09 (11.88)	(-31.76, 23.59)			
	Week 110		Sparsentan	9	4 (44.4)	-3.92 (9.58)	(-25.74, 17.91)	1.40 (16.36)	(-35.84, 38.64)	0.934
			Irbesartan	5	2 (40.0)	-5.32 (12.63)	(-33.87, 23.23)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
> 18 to 40 years	Week 24	Sparsentan	102	79 (77.5)	1.66 (1.27)	(-0.84, 4.15)	3.08 (1.84)	(-0.54, 6.70)	0.095	
		Irbesartan	109	71 (65.1)	-1.42 (1.34)	(-4.05, 1.20)				
	Week 48	Sparsentan	102	82 (80.4)	-0.43 (1.25)	(-2.87, 2.02)	3.12 (1.85)	(-0.52, 6.75)	0.092	
		Irbesartan	109	67 (61.5)	-3.55 (1.36)	(-6.22, -0.87)				
	Week 70	Sparsentan	102	78 (76.5)	-1.26 (1.27)	(-3.76, 1.25)	0.25 (1.87)	(-3.42, 3.92)	0.894	
		Irbesartan	109	68 (62.4)	-1.51 (1.36)	(-4.19, 1.17)				
	Week 94	Sparsentan	102	73 (71.6)	-0.45 (1.31)	(-3.04, 2.13)	3.35 (1.89)	(-0.37, 7.07)	0.077	
		Irbesartan	109	69 (63.3)	-3.80 (1.36)	(-6.47, -1.14)				
	Week 110	Sparsentan	102	72 (70.6)	0.02 (1.33)	(-2.60, 2.64)	1.22 (1.94)	(-2.59, 5.02)	0.531	
		Irbesartan	109	65 (59.6)	-1.20 (1.40)	(-3.95, 1.55)				
> 40 years	Week 24	Sparsentan	91	73 (80.2)	-1.22 (1.46)	(-4.09, 1.66)	0.75 (2.23)	(-3.63, 5.13)	0.738	
		Irbesartan	88	55 (62.5)	-1.97 (1.68)	(-5.26, 1.33)				
	Week 48	Sparsentan	91	73 (80.2)	-0.38 (1.45)	(-3.23, 2.48)	-0.57 (2.22)	(-4.93, 3.80)	0.798	
		Irbesartan	88	54 (61.4)	0.19 (1.68)	(-3.11, 3.49)				
	Week 70	Sparsentan	91	75 (82.4)	-1.52 (1.44)	(-4.36, 1.32)	-0.69 (2.19)	(-4.98, 3.61)	0.753	
		Irbesartan	88	58 (65.9)	-0.83 (1.64)	(-4.05, 2.39)				
	Week 94	Sparsentan	91	73 (80.2)	-1.86 (1.46)	(-4.73, 1.01)	-1.56 (2.20)	(-5.89, 2.77)	0.479	
		Irbesartan	88	57 (64.8)	-0.30 (1.65)	(-3.54, 2.94)				
	Week 110	Sparsentan	91	73 (80.2)	-2.46 (1.47)	(-5.35, 0.42)	-1.32 (2.25)	(-5.74, 3.10)	0.557	
		Irbesartan	88	54 (61.4)	-1.14 (1.70)	(-4.49, 2.21)				

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
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Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Geographic region	Overall		Sparsentan							Interaction: 0.095
North America	Week 24		Sparsentan	35	23 (65.7)	-2.87 (2.49)	(-7.79, 2.05)	0.18 (3.23)	(-6.21, 6.57)	0.956
			Irbesartan	46	35 (76.1)	-3.05 (2.05)	(-7.10, 1.00)			
	Week 48		Sparsentan	35	25 (71.4)	-3.17 (2.41)	(-7.94, 1.60)	-1.10 (3.20)	(-7.43, 5.23)	0.732
			Irbesartan	46	32 (69.6)	-2.07 (2.08)	(-6.19, 2.05)			
	Week 70		Sparsentan	35	22 (62.9)	0.66 (2.50)	(-4.28, 5.61)	2.91 (3.30)	(-3.62, 9.44)	0.379
			Irbesartan	46	30 (65.2)	-2.25 (2.14)	(-6.49, 1.99)			
	Week 94		Sparsentan	35	23 (65.7)	-1.81 (2.49)	(-6.74, 3.11)	0.35 (3.32)	(-6.21, 6.91)	0.917
			Irbesartan	46	30 (65.2)	-2.16 (2.18)	(-6.46, 2.14)			
	Week 110		Sparsentan	35	21 (60.0)	-0.60 (2.60)	(-5.74, 4.55)	0.30 (3.41)	(-6.45, 7.04)	0.931
			Irbesartan	46	31 (67.4)	-0.89 (2.19)	(-5.23, 3.44)			

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A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Europe	Week 24	Sparsentan	98	73 (74.5)	1.01 (1.49)	(-1.92, 3.95)	2.76 (2.21)	(-1.58, 7.10)	0.213	
		Irbesartan	115	62 (53.9)	-1.75 (1.63)	(-4.94, 1.45)				
	Week 48	Sparsentan	98	76 (77.6)	0.05 (1.47)	(-2.82, 2.93)	3.30 (2.20)	(-1.02, 7.62)	0.134	
		Irbesartan	115	59 (51.3)	-3.24 (1.64)	(-6.46, -0.03)				
	Week 70	Sparsentan	98	75 (76.5)	0.20 (1.48)	(-2.71, 3.12)	2.33 (2.14)	(-1.88, 6.54)	0.277	
		Irbesartan	115	69 (60.0)	-2.13 (1.54)	(-5.16, 0.90)				
	Week 94	Sparsentan	98	68 (69.4)	0.21 (1.54)	(-2.83, 3.24)	3.62 (2.18)	(-0.66, 7.90)	0.097	
		Irbesartan	115	70 (60.9)	-3.41 (1.53)	(-6.43, -0.40)				
	Week 110	Sparsentan	98	67 (68.4)	0.02 (1.57)	(-3.07, 3.10)	1.73 (2.27)	(-2.72, 6.19)	0.445	
		Irbesartan	115	61 (53.0)	-1.72 (1.63)	(-4.92, 1.49)				
Asia Pacific	Week 24	Sparsentan	69	61 (88.4)	-0.29 (1.43)	(-3.10, 2.51)	0.28 (2.39)	(-4.41, 4.97)	0.906	
		Irbesartan	41	34 (82.9)	-0.58 (1.91)	(-4.33, 3.18)				
	Week 48	Sparsentan	69	61 (88.4)	-0.50 (1.42)	(-3.30, 2.30)	-1.54 (2.41)	(-6.28, 3.19)	0.521	
		Irbesartan	41	33 (80.5)	1.04 (1.94)	(-2.77, 4.85)				
	Week 70	Sparsentan	69	63 (91.3)	-3.63 (1.41)	(-6.39, -0.87)	-4.58 (2.46)	(-9.41, 0.24)	0.063	
		Irbesartan	41	31 (75.6)	0.96 (2.00)	(-2.99, 4.90)				
	Week 94	Sparsentan	69	60 (87.0)	-2.63 (1.44)	(-5.45, 0.19)	-2.18 (2.50)	(-7.10, 2.73)	0.383	
		Irbesartan	41	30 (73.2)	-0.45 (2.04)	(-4.46, 3.56)				
	Week 110	Sparsentan	69	61 (88.4)	-2.44 (1.43)	(-5.26, 0.37)	-1.04 (2.52)	(-6.00, 3.92)	0.680	
		Irbesartan	41	29 (70.7)	-1.40 (2.07)	(-5.48, 2.67)				

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Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline BMI	Overall		Sparsentan							Interaction: 0.265
< 27 kg/m**2	Week 24		Sparsentan	83	66 (79.5)	-1.51 (1.37)	(-4.20, 1.18)	0.07 (1.94)	(-3.74, 3.89)	0.969
			Irbesartan	94	66 (70.2)	-1.58 (1.37)	(-4.28, 1.11)			
	Week 48		Sparsentan	83	66 (79.5)	-0.28 (1.36)	(-2.96, 2.40)	3.59 (1.97)	(-0.27, 7.45)	0.069
			Irbesartan	94	60 (63.8)	-3.87 (1.41)	(-6.65, -1.09)			
	Week 70		Sparsentan	83	64 (77.1)	-2.07 (1.39)	(-4.79, 0.65)	0.68 (1.99)	(-3.24, 4.60)	0.733
			Irbesartan	94	59 (62.8)	-2.75 (1.43)	(-5.56, 0.06)			
	Week 94		Sparsentan	83	63 (75.9)	-2.00 (1.40)	(-4.74, 0.75)	2.39 (1.99)	(-1.51, 6.30)	0.228
			Irbesartan	94	62 (66.0)	-4.39 (1.41)	(-7.16, -1.62)			
	Week 110		Sparsentan	83	63 (75.9)	-1.62 (1.41)	(-4.38, 1.15)	1.78 (2.03)	(-2.22, 5.77)	0.383
			Irbesartan	94	57 (60.6)	-3.39 (1.47)	(-6.28, -0.51)			

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Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
>= 27 kg/m**2	Week 24	Sparsentan	119	91 (76.5)	0.76 (1.30)	(-1.78, 3.31)	2.16 (2.01)	(-1.79, 6.12)	0.283	
		Irbesartan	107	65 (60.7)	-1.40 (1.54)	(-4.42, 1.62)				
	Week 48	Sparsentan	119	96 (80.7)	-1.07 (1.26)	(-3.55, 1.40)	-1.52 (1.98)	(-5.42, 2.37)	0.443	
		Irbesartan	107	64 (59.8)	0.45 (1.53)	(-2.55, 3.46)				
	Week 70	Sparsentan	119	96 (80.7)	-1.06 (1.26)	(-3.54, 1.42)	-1.27 (1.94)	(-5.08, 2.55)	0.515	
		Irbesartan	107	71 (66.4)	0.20 (1.47)	(-2.69, 3.09)				
	Week 94	Sparsentan	119	88 (73.9)	-1.02 (1.31)	(-3.60, 1.56)	-1.26 (2.00)	(-5.19, 2.68)	0.530	
		Irbesartan	107	67 (62.6)	0.24 (1.51)	(-2.73, 3.20)				
	Week 110	Sparsentan	119	86 (72.3)	-1.05 (1.34)	(-3.68, 1.57)	-2.08 (2.06)	(-6.12, 1.96)	0.313	
		Irbesartan	107	63 (58.9)	1.03 (1.56)	(-2.04, 4.09)				

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Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Randomization strata	Overall		Sparsentan							Interaction: 0.749
eGFR Low and UP High	Week 24		Sparsentan	71	54 (76.1)	-1.15 (1.32)	(-3.74, 1.45)	0.49 (2.02)	(-3.49, 4.46)	0.810
			Irbesartan	74	40 (54.1)	-1.63 (1.52)	(-4.62, 1.35)			
	Week 48		Sparsentan	71	51 (71.8)	-0.63 (1.33)	(-3.26, 1.99)	4.32 (2.04)	(0.30, 8.34)	0.035 *
			Irbesartan	74	37 (50.0)	-4.96 (1.54)	(-7.99, -1.93)			
	Week 70		Sparsentan	71	57 (80.3)	-3.18 (1.29)	(-5.72, -0.63)	0.23 (2.03)	(-3.77, 4.22)	0.912
			Irbesartan	74	37 (50.0)	-3.40 (1.56)	(-6.46, -0.34)			
	Week 94		Sparsentan	71	51 (71.8)	-3.12 (1.34)	(-5.75, -0.48)	0.55 (2.07)	(-3.52, 4.63)	0.790
			Irbesartan	74	37 (50.0)	-3.67 (1.57)	(-6.76, -0.58)			
	Week 110		Sparsentan	71	53 (74.6)	-2.85 (1.34)	(-5.48, -0.22)	-1.94 (2.10)	(-6.08, 2.20)	0.356
			Irbesartan	74	36 (48.6)	-0.90 (1.61)	(-4.07, 2.26)			

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Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
					Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
eGFR Low and UP Low	Week 24	Sparsentan	55	43 (78.2)	0.26 (1.91)	(-3.50, 4.02)	-2.68 (2.82)	(-8.24, 2.87)	0.343
		Irbesartan	55	37 (67.3)	2.94 (2.07)	(-1.12, 7.01)			
	Week 48	Sparsentan	55	43 (78.2)	-3.16 (1.90)	(-6.91, 0.58)	-3.73 (2.84)	(-9.32, 1.86)	
		Irbesartan	55	35 (63.6)	0.57 (2.10)	(-3.56, 4.70)			
	Week 70	Sparsentan	55	42 (76.4)	1.37 (1.93)	(-2.42, 5.17)	0.85 (2.79)	(-4.63, 6.33)	
		Irbesartan	55	39 (70.9)	0.52 (2.00)	(-3.42, 4.46)			
	Week 94	Sparsentan	55	41 (74.5)	0.11 (1.96)	(-3.74, 3.96)	-0.33 (2.82)	(-5.87, 5.21)	
		Irbesartan	55	39 (70.9)	0.44 (2.01)	(-3.51, 4.40)			
	Week 110	Sparsentan	55	37 (67.3)	-2.29 (2.06)	(-6.34, 1.76)	-2.55 (2.99)	(-8.43, 3.32)	
		Irbesartan	55	34 (61.8)	0.27 (2.15)	(-3.96, 4.49)			
eGFR High and UP High	Week 24	Sparsentan	37	26 (70.3)	-0.40 (2.82)	(-5.96, 5.16)	0.01 (4.03)	(-7.94, 7.97)	0.998
		Irbesartan	36	26 (72.2)	-0.42 (2.88)	(-6.11, 5.28)			
	Week 48	Sparsentan	37	33 (89.2)	-0.98 (2.56)	(-6.04, 4.08)	2.12 (3.91)	(-5.59, 9.83)	
		Irbesartan	36	24 (66.7)	-3.10 (2.95)	(-8.92, 2.72)			
	Week 70	Sparsentan	37	31 (83.8)	-2.81 (2.63)	(-8.01, 2.39)	-0.45 (3.94)	(-8.23, 7.33)	
		Irbesartan	36	25 (69.4)	-2.36 (2.93)	(-8.14, 3.43)			
	Week 94	Sparsentan	37	27 (73.0)	-1.17 (2.79)	(-6.67, 4.33)	0.11 (4.08)	(-7.94, 8.15)	
		Irbesartan	36	24 (66.7)	-1.28 (2.97)	(-7.15, 4.59)			
	Week 110	Sparsentan	37	26 (70.3)	2.21 (2.87)	(-3.45, 7.87)	6.68 (4.23)	(-1.67, 15.02)	
		Irbesartan	36	22 (61.1)	-4.46 (3.11)	(-10.59, 1.66)			

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KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
eGFR High and UP Low	Week 24	Sparsentan	39	34 (87.2)	0.53 (1.94)	(-3.28, 4.35)	7.43 (2.87)	(1.78, 13.09)	0.010	*
		Irbesartan	37	28 (75.7)	-6.90 (2.12)	(-11.08, -2.72)				
	Week 48	Sparsentan	39	35 (89.7)	1.35 (1.91)	(-2.42, 5.11)	-0.52 (2.86)	(-6.15, 5.11)	0.857	
		Irbesartan	37	28 (75.7)	1.87 (2.12)	(-2.31, 6.04)				
	Week 70	Sparsentan	39	30 (76.9)	-1.61 (2.04)	(-5.64, 2.41)	-3.16 (2.93)	(-8.93, 2.62)	0.283	
		Irbesartan	37	29 (78.4)	1.54 (2.09)	(-2.58, 5.67)				
	Week 94	Sparsentan	39	32 (82.1)	-1.61 (1.99)	(-5.52, 2.31)	1.14 (2.87)	(-4.51, 6.79)	0.691	
		Irbesartan	37	30 (81.1)	-2.75 (2.06)	(-6.80, 1.31)				
	Week 110	Sparsentan	39	33 (84.6)	-1.71 (1.97)	(-5.58, 2.16)	-2.06 (2.88)	(-7.72, 3.61)	0.476	
		Irbesartan	37	29 (78.4)	0.35 (2.09)	(-3.78, 4.47)				

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						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline eGFR Group 1	Overall		Sparsentan							Interaction: 0.493
< 60 mL/min/1.73 m**2	Week 24		Sparsentan	127	98 (77.2)	-0.35 (1.19)	(-2.69, 1.98)	0.84 (1.78)	(-2.64, 4.33)	0.635
			Irbesartan	129	80 (62.0)	-1.20 (1.32)	(-3.78, 1.39)			
	Week 48		Sparsentan	127	95 (74.8)	-1.10 (1.20)	(-3.45, 1.26)	0.87 (1.80)	(-2.67, 4.41)	0.629
			Irbesartan	129	74 (57.4)	-1.97 (1.35)	(-4.61, 0.68)			
	Week 70		Sparsentan	127	100 (78.7)	-1.33 (1.18)	(-3.65, 0.98)	0.14 (1.77)	(-3.35, 3.62)	0.939
			Irbesartan	129	78 (60.5)	-1.47 (1.32)	(-4.07, 1.13)			
	Week 94		Sparsentan	127	92 (72.4)	-1.36 (1.22)	(-3.76, 1.03)	0.59 (1.81)	(-2.96, 4.15)	0.743
			Irbesartan	129	77 (59.7)	-1.96 (1.33)	(-4.58, 0.66)			
	Week 110		Sparsentan	127	92 (72.4)	-2.47 (1.23)	(-4.88, -0.05)	-1.74 (1.86)	(-5.38, 1.90)	0.349
			Irbesartan	129	72 (55.8)	-0.73 (1.39)	(-3.45, 1.99)			

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	38 (77.6)	-1.55 (2.10)	(-5.68, 2.58)	-1.30 (3.13)	(-7.47, 4.87)	0.678	
		Irbesartan	48	31 (64.6)	-0.25 (2.32)	(-4.81, 4.32)				
	Week 48	Sparsentan	49	44 (89.8)	-1.37 (1.98)	(-5.28, 2.54)	2.08 (3.02)	(-3.87, 8.03)	0.491	
		Irbesartan	48	32 (66.7)	-3.45 (2.26)	(-7.91, 1.01)				
	Week 70	Sparsentan	49	38 (77.6)	-0.87 (2.08)	(-4.97, 3.22)	2.74 (3.09)	(-3.34, 8.83)	0.375	
		Irbesartan	48	32 (66.7)	-3.61 (2.27)	(-8.09, 0.86)				
	Week 94	Sparsentan	49	39 (79.6)	-1.69 (2.08)	(-5.79, 2.41)	3.47 (3.06)	(-2.57, 9.51)	0.259	
		Irbesartan	48	34 (70.8)	-5.16 (2.24)	(-9.58, -0.75)				
	Week 110	Sparsentan	49	36 (73.5)	0.32 (2.16)	(-3.95, 4.58)	4.53 (3.17)	(-1.71, 10.77)	0.154	
		Irbesartan	48	32 (66.7)	-4.21 (2.30)	(-8.75, 0.33)				
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	21 (80.8)	2.80 (2.45)	(-2.03, 7.63)	7.05 (3.51)	(0.13, 13.96)	0.046 *	
		Irbesartan	25	20 (80.0)	-4.25 (2.51)	(-9.19, 0.70)				
	Week 48	Sparsentan	26	23 (88.5)	1.38 (2.34)	(-3.24, 6.00)	-1.06 (3.55)	(-8.07, 5.95)	0.766	
		Irbesartan	25	18 (72.0)	2.44 (2.65)	(-2.79, 7.67)				
	Week 70	Sparsentan	26	22 (84.6)	-3.30 (2.40)	(-8.03, 1.43)	-7.71 (3.49)	(-14.59, -0.83)	0.028 *	
		Irbesartan	25	20 (80.0)	4.41 (2.52)	(-0.56, 9.37)				
	Week 94	Sparsentan	26	20 (76.9)	-1.08 (2.51)	(-6.03, 3.88)	-3.66 (3.61)	(-10.78, 3.47)	0.313	
		Irbesartan	25	19 (76.0)	2.58 (2.58)	(-2.51, 7.67)				
	Week 110	Sparsentan	26	21 (80.8)	1.40 (2.45)	(-3.44, 6.23)	-0.65 (3.67)	(-7.90, 6.59)	0.859	
		Irbesartan	25	17 (68.0)	2.05 (2.72)	(-3.32, 7.42)				

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
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Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline eGFR Group 2	Overall		Sparsentan							Interaction: 0.252
< 45 mL/min/1.73 m**2	Week 24		Sparsentan	82	64 (78.0)	1.05 (1.50)	(-1.90, 4.00)	1.69 (2.29)	(-2.80, 6.19)	0.460
			Irbesartan	80	48 (60.0)	-0.64 (1.73)	(-4.04, 2.75)			
	Week 48		Sparsentan	82	61 (74.4)	-1.27 (1.52)	(-4.26, 1.72)	2.41 (2.35)	(-2.20, 7.03)	0.304
			Irbesartan	80	43 (53.8)	-3.68 (1.79)	(-7.19, -0.17)			
	Week 70		Sparsentan	82	65 (79.3)	-0.91 (1.49)	(-3.84, 2.02)	2.39 (2.30)	(-2.14, 6.92)	0.300
			Irbesartan	80	45 (56.3)	-3.30 (1.76)	(-6.75, 0.15)			
	Week 94		Sparsentan	82	60 (73.2)	-1.21 (1.54)	(-4.24, 1.81)	2.43 (2.32)	(-2.12, 6.99)	0.295
			Irbesartan	80	47 (58.8)	-3.65 (1.74)	(-7.06, -0.24)			
	Week 110		Sparsentan	82	56 (68.3)	-2.00 (1.60)	(-5.14, 1.14)	-0.26 (2.41)	(-5.00, 4.47)	0.913
			Irbesartan	80	44 (55.0)	-1.74 (1.80)	(-5.28, 1.80)			

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Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	34 (75.6)	-3.09 (1.93)	(-6.89, 0.70)	-0.99 (2.77)	(-6.45, 4.47)	0.721	
		Irbesartan	49	32 (65.3)	-2.10 (1.99)	(-6.02, 1.81)				
	Week 48	Sparsentan	45	34 (75.6)	-0.76 (1.91)	(-4.52, 3.00)	-1.43 (2.78)	(-6.90, 4.04)	0.608	
		Irbesartan	49	31 (63.3)	0.67 (2.01)	(-3.29, 4.62)				
	Week 70	Sparsentan	45	35 (77.8)	-2.16 (1.90)	(-5.90, 1.58)	-3.59 (2.74)	(-8.99, 1.81)	0.192	
		Irbesartan	49	33 (67.3)	1.43 (1.97)	(-2.45, 5.31)				
	Week 94	Sparsentan	45	32 (71.1)	-1.78 (1.97)	(-5.66, 2.11)	-2.72 (2.85)	(-8.34, 2.89)	0.341	
		Irbesartan	49	30 (61.2)	0.94 (2.05)	(-3.09, 4.98)				
	Week 110	Sparsentan	45	36 (80.0)	-3.40 (1.89)	(-7.13, 0.32)	-4.51 (2.86)	(-10.14, 1.11)	0.116	
		Irbesartan	49	28 (57.1)	1.11 (2.13)	(-3.09, 5.31)				
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	38 (77.6)	-1.55 (2.10)	(-5.68, 2.58)	-1.30 (3.13)	(-7.47, 4.87)	0.678	
		Irbesartan	48	31 (64.6)	-0.25 (2.32)	(-4.81, 4.32)				
	Week 48	Sparsentan	49	44 (89.8)	-1.37 (1.98)	(-5.28, 2.54)	2.08 (3.02)	(-3.87, 8.03)	0.491	
		Irbesartan	48	32 (66.7)	-3.45 (2.26)	(-7.91, 1.01)				
	Week 70	Sparsentan	49	38 (77.6)	-0.87 (2.08)	(-4.97, 3.22)	2.74 (3.09)	(-3.34, 8.83)	0.375	
		Irbesartan	48	32 (66.7)	-3.61 (2.27)	(-8.09, 0.86)				
	Week 94	Sparsentan	49	39 (79.6)	-1.69 (2.08)	(-5.79, 2.41)	3.47 (3.06)	(-2.57, 9.51)	0.259	
		Irbesartan	48	34 (70.8)	-5.16 (2.24)	(-9.58, -0.75)				
	Week 110	Sparsentan	49	36 (73.5)	0.32 (2.16)	(-3.95, 4.58)	4.53 (3.17)	(-1.71, 10.77)	0.154	
		Irbesartan	48	32 (66.7)	-4.21 (2.30)	(-8.75, 0.33)				

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Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	21 (80.8)	2.80 (2.45)	(-2.03, 7.63)	7.05 (3.51)	(0.13, 13.96)	0.046	*
		Irbesartan	25	20 (80.0)	-4.25 (2.51)	(-9.19, 0.70)				
	Week 48	Sparsentan	26	23 (88.5)	1.38 (2.34)	(-3.24, 6.00)	-1.06 (3.55)	(-8.07, 5.95)	0.766	
		Irbesartan	25	18 (72.0)	2.44 (2.65)	(-2.79, 7.67)				
	Week 70	Sparsentan	26	22 (84.6)	-3.30 (2.40)	(-8.03, 1.43)	-7.71 (3.49)	(-14.59, -0.83)	0.028	*
		Irbesartan	25	20 (80.0)	4.41 (2.52)	(-0.56, 9.37)				
	Week 94	Sparsentan	26	20 (76.9)	-1.08 (2.51)	(-6.03, 3.88)	-3.66 (3.61)	(-10.78, 3.47)	0.313	
		Irbesartan	25	19 (76.0)	2.58 (2.58)	(-2.51, 7.67)				
	Week 110	Sparsentan	26	21 (80.8)	1.40 (2.45)	(-3.44, 6.23)	-0.65 (3.67)	(-7.90, 6.59)	0.859	
		Irbesartan	25	17 (68.0)	2.05 (2.72)	(-3.32, 7.42)				

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Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline urine protein excretion		Overall	Sparsentan							Interaction: 0.510
<= 1.75 g/day	Week 24	Sparsentan	98	80 (81.6)	-0.17 (1.35)	(-2.81, 2.48)	1.81 (2.11)	(-2.33, 5.94)	0.391	
		Irbesartan	93	55 (59.1)	-1.98 (1.62)	(-5.15, 1.20)				
	Week 48	Sparsentan	98	82 (83.7)	-0.89 (1.33)	(-3.49, 1.72)	-1.95 (2.07)	(-6.01, 2.11)	0.346	
		Irbesartan	93	57 (61.3)	1.06 (1.58)	(-2.04, 4.17)				
	Week 70	Sparsentan	98	75 (76.5)	-1.09 (1.38)	(-3.80, 1.62)	-1.62 (2.07)	(-5.70, 2.45)	0.434	
		Irbesartan	93	61 (65.6)	0.53 (1.54)	(-2.49, 3.56)				
	Week 94	Sparsentan	98	76 (77.6)	-0.52 (1.38)	(-3.22, 2.19)	1.52 (2.05)	(-2.51, 5.56)	0.459	
		Irbesartan	93	63 (67.7)	-2.04 (1.52)	(-5.03, 0.95)				
	Week 110	Sparsentan	98	75 (76.5)	-1.03 (1.39)	(-3.76, 1.71)	-0.91 (2.12)	(-5.07, 3.26)	0.670	
		Irbesartan	93	57 (61.3)	-0.12 (1.60)	(-3.25, 3.01)				

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Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
> 1.75 g/day	Week 24		Sparsentan	104	77 (74.0)	-0.37 (1.35)	(-3.02, 2.28)	0.92 (1.92)	(-2.86, 4.70)	0.633
			Irbesartan	109	76 (69.7)	-1.29 (1.37)	(-3.97, 1.40)			
	Week 48		Sparsentan	104	80 (76.9)	-0.70 (1.32)	(-3.29, 1.89)	3.26 (1.94)	(-0.55, 7.08)	0.094
			Irbesartan	109	67 (61.5)	-3.96 (1.42)	(-6.76, -1.17)			
	Week 70		Sparsentan	104	85 (81.7)	-1.87 (1.30)	(-4.41, 0.68)	0.77 (1.92)	(-3.01, 4.55)	0.690
			Irbesartan	109	69 (63.3)	-2.63 (1.42)	(-5.42, 0.15)			
	Week 94		Sparsentan	104	75 (72.1)	-2.36 (1.36)	(-5.04, 0.32)	-0.27 (1.99)	(-4.17, 3.63)	0.893
			Irbesartan	109	67 (61.5)	-2.09 (1.44)	(-4.92, 0.74)			
	Week 110		Sparsentan	104	74 (71.2)	-1.59 (1.38)	(-4.31, 1.13)	0.43 (2.03)	(-3.56, 4.41)	0.833
			Irbesartan	109	64 (58.7)	-2.02 (1.48)	(-4.93, 0.89)			

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						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline use of antihypertensives	Overall		Sparsentan							Interaction: 0.512
Yes	Week 24		Sparsentan	90	66 (73.3)	0.11 (1.22)	(-2.29, 2.50)	-0.43 (1.87)	(-4.10, 3.23)	0.816
			Irbesartan	88	49 (55.7)	0.54 (1.41)	(-2.23, 3.32)			
	Week 48		Sparsentan	90	65 (72.2)	0.43 (1.22)	(-1.96, 2.83)	1.23 (1.88)	(-2.46, 4.92)	0.513
			Irbesartan	88	47 (53.4)	-0.80 (1.43)	(-3.60, 2.01)			
	Week 70		Sparsentan	90	68 (75.6)	-2.01 (1.20)	(-4.38, 0.35)	-0.92 (1.85)	(-4.57, 2.72)	0.619
			Irbesartan	88	49 (55.7)	-1.09 (1.41)	(-3.86, 1.68)			
	Week 94		Sparsentan	90	63 (70.0)	-2.03 (1.24)	(-4.47, 0.41)	-0.27 (1.84)	(-3.87, 3.34)	0.884
			Irbesartan	88	54 (61.4)	-1.77 (1.35)	(-4.42, 0.89)			
	Week 110		Sparsentan	90	63 (70.0)	-0.35 (1.25)	(-2.81, 2.11)	-0.67 (1.87)	(-4.35, 3.01)	0.721
			Irbesartan	88	51 (58.0)	0.32 (1.39)	(-2.41, 3.05)			

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Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
No	Week 24	Sparsentan	112	91 (81.3)	-0.42 (1.38)	(-3.13, 2.29)	2.40 (2.01)	(-1.54, 6.34)	0.233	
			114	82 (71.9)	-2.81 (1.46)	(-5.67, 0.04)				
	Week 48	Sparsentan	112	97 (86.6)	-1.52 (1.34)	(-4.15, 1.11)	0.79 (2.00)	(-3.13, 4.71)	0.693	
			114	77 (67.5)	-2.31 (1.48)	(-5.22, 0.60)				
	Week 70	Sparsentan	112	92 (82.1)	-0.97 (1.37)	(-3.66, 1.73)	0.32 (2.00)	(-3.62, 4.25)	0.875	
			114	81 (71.1)	-1.29 (1.46)	(-4.15, 1.58)				
	Week 94	Sparsentan	112	88 (78.6)	-0.83 (1.40)	(-3.58, 1.92)	1.57 (2.06)	(-2.47, 5.61)	0.445	
			114	76 (66.7)	-2.40 (1.51)	(-5.36, 0.56)				
	Week 110	Sparsentan	112	86 (76.8)	-1.91 (1.42)	(-4.70, 0.89)	0.28 (2.12)	(-3.89, 4.45)	0.894	
			114	70 (61.4)	-2.19 (1.57)	(-5.28, 0.90)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.117
<= 5 years	Week 24	Sparsentan	113	88 (77.9)	1.04 (1.28)	(-1.47, 3.56)	3.61 (1.82)	(0.03, 7.19)	0.048 *
		Irbesartan	127	86 (67.7)	-2.57 (1.30)	(-5.11, -0.02)			
	Week 48	Sparsentan	113	92 (81.4)	0.99 (1.25)	(-1.47, 3.44)	2.47 (1.80)	(-1.06, 6.00)	0.170
		Irbesartan	127	85 (66.9)	-1.49 (1.29)	(-4.02, 1.05)			
	Week 70	Sparsentan	113	90 (79.6)	-1.49 (1.26)	(-3.97, 0.99)	-0.05 (1.81)	(-3.59, 3.50)	0.979
		Irbesartan	127	86 (67.7)	-1.44 (1.29)	(-3.98, 1.09)			
	Week 94	Sparsentan	113	86 (76.1)	-1.21 (1.29)	(-3.74, 1.33)	0.60 (1.83)	(-2.99, 4.18)	0.744
		Irbesartan	127	86 (67.7)	-1.80 (1.29)	(-4.34, 0.73)			
	Week 110	Sparsentan	113	88 (77.9)	0.37 (1.29)	(-2.16, 2.90)	1.67 (1.86)	(-1.98, 5.32)	0.369
		Irbesartan	127	80 (63.0)	-1.30 (1.34)	(-3.94, 1.33)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
> 5 years	Week 24	Sparsentan	89	69 (77.5)	-1.63 (1.43)	(-4.45, 1.19)	-1.98 (2.31)	(-6.52, 2.55)	0.390	
			75	45 (60.0)	0.36 (1.80)	(-3.18, 3.89)				
	Week 48	Sparsentan	89	70 (78.7)	-3.02 (1.42)	(-5.82, -0.23)	-0.73 (2.36)	(-5.36, 3.91)	0.758	
			75	39 (52.0)	-2.30 (1.87)	(-5.98, 1.38)				
	Week 70	Sparsentan	89	70 (78.7)	-1.35 (1.43)	(-4.16, 1.46)	-0.36 (2.30)	(-4.89, 4.17)	0.876	
			75	44 (58.7)	-0.99 (1.80)	(-4.52, 2.55)				
	Week 94	Sparsentan	89	65 (73.0)	-1.44 (1.47)	(-4.33, 1.46)	1.57 (2.34)	(-3.03, 6.16)	0.503	
			75	44 (58.7)	-3.00 (1.81)	(-6.56, 0.55)				
	Week 110	Sparsentan	89	61 (68.5)	-3.40 (1.52)	(-6.39, -0.40)	-2.33 (2.42)	(-7.08, 2.43)	0.336	
			75	41 (54.7)	-1.07 (1.87)	(-4.74, 2.61)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
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 Source Data: aqs, created on: 05MAR2024

Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
History of hypertension	Overall		Sparsentan							Interaction: 0.101
Yes	Week 24		Sparsentan	155	119 (76.8)	0.49 (0.95)	(-1.37, 2.36)	0.44 (1.41)	(-2.32, 3.20)	0.753
			Irbesartan	161	100 (62.1)	0.05 (1.04)	(-1.98, 2.08)			
	Week 48		Sparsentan	155	121 (78.1)	-0.59 (0.94)	(-2.43, 1.26)	0.85 (1.41)	(-1.92, 3.62)	0.547
			Irbesartan	161	95 (59.0)	-1.44 (1.05)	(-3.50, 0.63)			
	Week 70		Sparsentan	155	123 (79.4)	-0.62 (0.94)	(-2.46, 1.21)	0.02 (1.40)	(-2.72, 2.76)	0.989
			Irbesartan	161	100 (62.1)	-0.64 (1.03)	(-2.67, 1.39)			
	Week 94		Sparsentan	155	114 (73.5)	-1.40 (0.97)	(-3.30, 0.49)	0.92 (1.41)	(-1.85, 3.70)	0.513
			Irbesartan	161	101 (62.7)	-2.33 (1.03)	(-4.35, -0.30)			
	Week 110		Sparsentan	155	112 (72.3)	-0.95 (0.98)	(-2.88, 0.98)	-1.34 (1.46)	(-4.20, 1.52)	0.359
			Irbesartan	161	92 (57.1)	0.39 (1.08)	(-1.73, 2.51)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs, created on: 05MAR2024

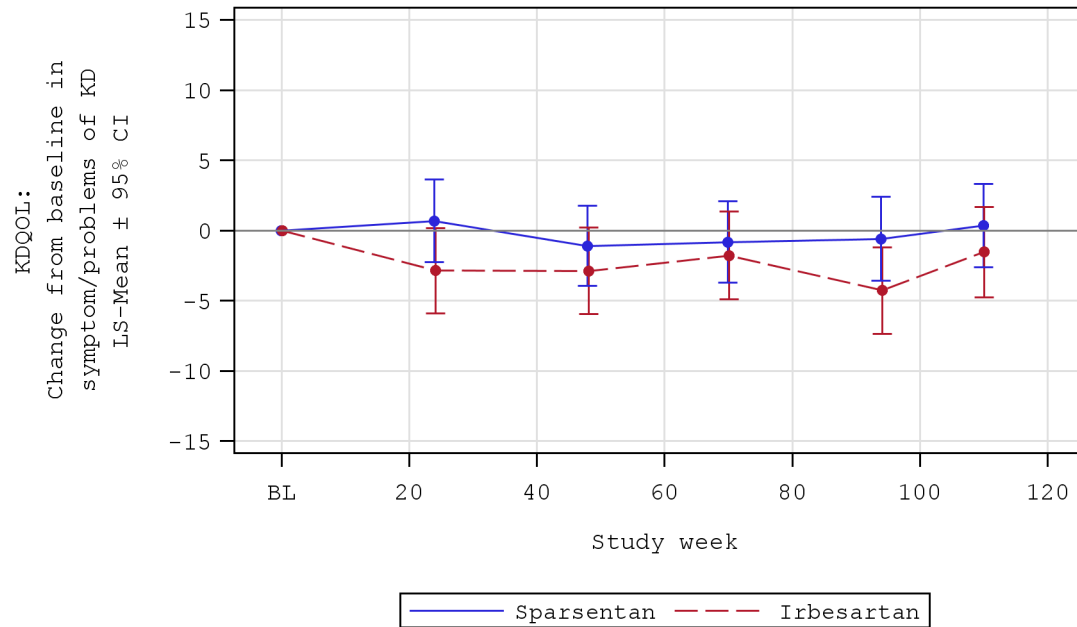
Table PT2KSYC\_FSCM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
Full Analysis Set

KDQOL: Change from baseline in symptom/problems of kidney disease	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
No	Week 24	Sparsentan	Irbesartan	47	38 (80.9)	-1.81 (2.62)	(-6.98, 3.35)	5.68 (3.92)	(-2.05, 13.41)	0.149
				41	31 (75.6)	-7.49 (2.91)	(-13.23, -1.76)			
	Week 48	Sparsentan	Irbesartan	47	41 (87.2)	-0.95 (2.54)	(-5.96, 4.06)	2.76 (3.90)	(-4.92, 10.45)	0.479
				41	29 (70.7)	-3.72 (2.95)	(-9.52, 2.09)			
	Week 70	Sparsentan	Irbesartan	47	37 (78.7)	-3.70 (2.64)	(-8.89, 1.50)	0.17 (3.95)	(-7.62, 7.96)	0.965
				41	30 (73.2)	-3.87 (2.93)	(-9.65, 1.92)			
	Week 94	Sparsentan	Irbesartan	47	37 (78.7)	-0.69 (2.66)	(-5.93, 4.55)	1.62 (4.02)	(-6.30, 9.54)	0.688
				41	29 (70.7)	-2.31 (3.00)	(-8.22, 3.61)			
	Week 110	Sparsentan	Irbesartan	47	37 (78.7)	-1.49 (2.68)	(-6.77, 3.79)	5.35 (4.06)	(-2.66, 13.35)	0.189
				41	29 (70.7)	-6.84 (3.04)	(-12.83, -0.86)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
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Source Data: aqs, created on: 05MAR2024



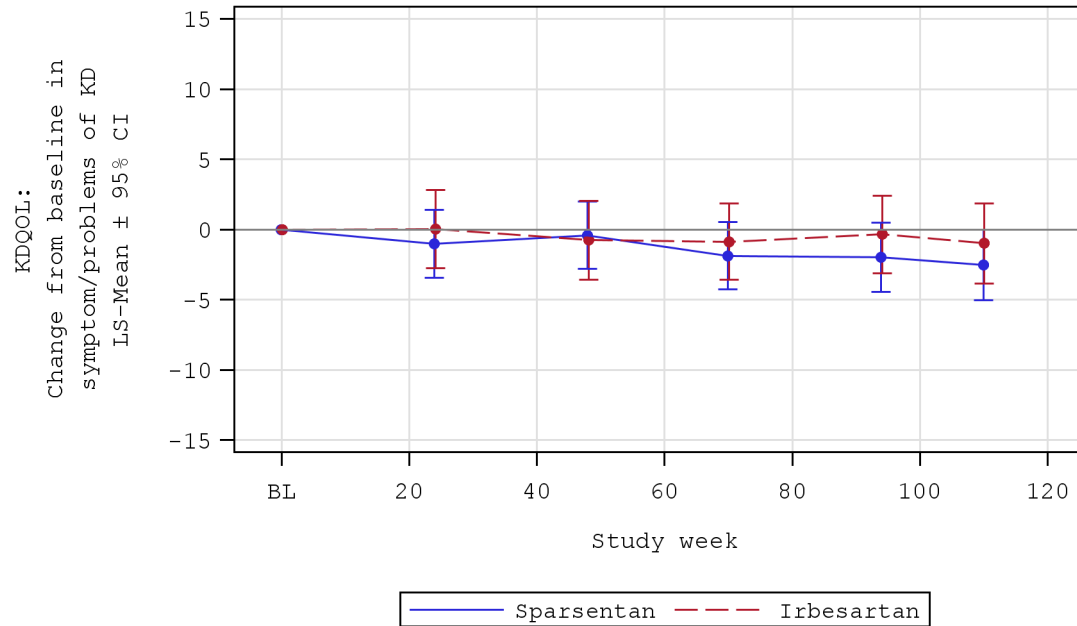
Figure PF2KSYC\_FSGM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Age: <= 45 years



Sparsentan	71	75	72	68	69
Irbesartan	67	62	62	64	59

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KSYC\_FSCM

Figure PF2KSYC\_FSGM: KDQOL: Change from baseline in symptom/problems of kidney disease by subgroup  
 Full Analysis Set  
 Study: PROTECT  
 Age: > 45 years



Sparsentan	86	87	88	83	80
Irbesartan	64	62	68	66	62

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Reference table: PT2KSYC\_FSCM

Table PT2KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.125
Male	Sparsentan	139	9 (6.5)	NE		2.500	(0.730, 8.555)	0.144
	Irbesartan	143	7 (4.9)	NE				
Female	Sparsentan	63	11 (17.5)	NE		0.733	(0.236, 2.277)	0.591
	Irbesartan	59	6 (10.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.130
<= 45 years	Sparsentan	96	11 (11.5)	NE		2.831	(0.703, 11.395)	0.143
	Irbesartan	99	3 (3.0)	NE				
> 45 years	Sparsentan	106	9 (8.5)	NE		0.741	(0.271, 2.028)	0.559
	Irbesartan	103	10 (9.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	0 (0.0) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	12 (11.8)	NE		1.873	(0.557, 6.294)	0.310
	Irbesartan	109	4 (3.7)	NE				
> 40 years	Sparsentan	91	8 (8.8)	NE		0.790	(0.263, 2.373)	0.675
	Irbesartan	88	9 (10.2)	NE				

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 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.692
North America	Sparsentan	35	2 (5.7)	NE		0.883	(0.121, 6.425)	0.902
	Irbesartan	46	4 (8.7)	NE				
Europe	Sparsentan	98	10 (10.2)	NE		1.703	(0.456, 6.355)	0.428
	Irbesartan	115	5 (4.3)	NE				
Asia Pacific	Sparsentan	69	8 (11.6)	NE		0.789	(0.223, 2.795)	0.714
	Irbesartan	41	4 (9.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

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 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.118
< 27 kg/m**2	Sparsentan	83	7 (8.4)	NE		4.117	(0.671, 25.245)	0.126
	Irbesartan	94	3 (3.2)	NE				
≥ 27 kg/m**2	Sparsentan	119	13 (10.9)	NE		0.973	(0.384, 2.468)	0.954
	Irbesartan	107	10 (9.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.247
eGFR Low and UP High	Sparsentan	71	3 (4.2)	NE		16.002	(0.623, 411.105)	0.094
	Irbesartan	74	3 (4.1)	NE				
eGFR Low and UP Low	Sparsentan	55	9 (16.4)	NE		0.953	(0.246, 3.692)	0.945
	Irbesartan	55	4 (7.3)	NE				
eGFR High and UP High	Sparsentan	37	4 (10.8)	NE		1.267	(0.253, 6.358)	0.774
	Irbesartan	36	4 (11.1)	NE				
eGFR High and UP Low	Sparsentan	39	4 (10.3)	NE		0.805	(0.132, 4.908)	0.814
	Irbesartan	37	2 (5.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
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 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.863
< 60 mL/min/1.73 m**2	Sparsentan	127	12 (9.4)	NE		1.302	(0.471, 3.597)	0.611
	Irbesartan	129	8 (6.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	3 (6.1)	NE		1.368	(0.162, 11.554)	0.774
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	5 (19.2)	NE		0.641	(0.142, 2.886)	0.562
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.696
< 45 mL/min/1.73 m**2	Sparsentan	82	8 (9.8)	NE		1.883	(0.522, 6.796)	0.334
	Irbesartan	80	5 (6.3)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	4 (8.9)	NE		0.512	(0.068, 3.843)	0.515
	Irbesartan	49	3 (6.1)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	3 (6.1)	NE		1.368	(0.162, 11.554)	0.774
	Irbesartan	48	2 (4.2)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	5 (19.2)	NE		0.641	(0.142, 2.886)	0.562
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.472
<= 1.75 g/day	Sparsentan	98	14 (14.3)	NE		2.246	(0.591, 8.539)	0.235
	Irbesartan	93	3 (3.2)	NE				
> 1.75 g/day	Sparsentan	104	6 (5.8)	NE		1.291	(0.389, 4.285)	0.676
	Irbesartan	109	10 (9.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan						Interaction test:	0.724
Yes	Sparsentan	90	9 (10.0)	NE		2.709	(0.680, 10.790)	0.157
	Irbesartan	88	5 (5.7)	NE				
No	Sparsentan	112	11 (9.8)	NE		0.851	(0.314, 2.307)	0.752
	Irbesartan	114	8 (7.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.445
<= 5 years	Sparsentan	113	13 (11.5)	NE		1.405	(0.556, 3.552)	0.472
	Irbesartan	127	10 (7.9)	NE				
> 5 years	Sparsentan	89	7 (7.9)	NE		0.798	(0.126, 5.030)	0.810
	Irbesartan	75	3 (4.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KSYIT\_FSTM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to increase in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.786
Yes	Sparsentan	155	15 (9.7)	NE		1.044	(0.412, 2.647)	0.928
	Irbesartan	161	11 (6.8)	NE				
No	Sparsentan	47	5 (10.6)	NE		1.752	(0.316, 9.718)	0.521
	Irbesartan	41	2 (4.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Figure PF2KSYIT\_FSKM: KDQOL: Time to increase in symptom/problems of kidney disease by at least 15 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_ An increase reflects an improvement of the status of the patient.  
Reference table: PT2KSYIT\_FSTM

Table PT2KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan							Interaction test: 0.839
Male	Sparsentan	139	31 (22.3)	NE		1.221	(0.709, 2.104)	0.471
	Irbesartan	143	23 (16.1)	NE				
Female	Sparsentan	63	14 (22.2)	NE		1.161	(0.503, 2.682)	0.726
	Irbesartan	59	10 (16.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan							Interaction test: 0.024 #
<= 45 years	Sparsentan	96	17 (17.7)	NE		0.665	(0.349, 1.267)	0.215
	Irbesartan	99	21 (21.2)	NE				
> 45 years	Sparsentan	106	28 (26.4)	NE		2.030	(1.025, 4.024)	0.042 *
	Irbesartan	103	12 (11.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	0.117
<= 18 years	Sparsentan	9	2 (22.2)	NE		0.286	(0.019, 4.375)	0.369
	Irbesartan	5	2 (40.0)	95.3	(70.0, NE)			
> 18 to 40 years	Sparsentan	102	19 (18.6)	NE		0.771	(0.410, 1.449)	0.420
	Irbesartan	109	21 (19.3)	NE				
> 40 years	Sparsentan	91	24 (26.4)	NE		1.922	(0.915, 4.036)	0.084
	Irbesartan	88	10 (11.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.409
North America	Sparsentan	35	5 (14.3)	115.7	(115.7, NE)	0.737	(0.232, 2.346)	0.606
	Irbesartan	46	8 (17.4)	NE				
Europe	Sparsentan	98	20 (20.4)	NE		0.967	(0.514, 1.821)	0.918
	Irbesartan	115	19 (16.5)	NE				
Asia Pacific	Sparsentan	69	20 (29.0)	NE		1.929	(0.764, 4.870)	0.165
	Irbesartan	41	6 (14.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.197
< 27 kg/m**2	Sparsentan	83	21 (25.3)	NE		0.941	(0.516, 1.717)	0.843
	Irbesartan	94	22 (23.4)	NE				
≥ 27 kg/m**2	Sparsentan	119	24 (20.2)	NE		1.785	(0.849, 3.750)	0.126
	Irbesartan	107	10 (9.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
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 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.799
eGFR Low and UP High	Sparsentan	71	16 (22.5)	NE		1.010	(0.478, 2.132)	0.980
	Irbesartan	74	14 (18.9)	NE				
eGFR Low and UP Low	Sparsentan	55	13 (23.6)	NE		1.703	(0.699, 4.148)	0.241
	Irbesartan	55	8 (14.5)	NE				
eGFR High and UP High	Sparsentan	37	8 (21.6)	115.7	(115.7, NE)	1.269	(0.413, 3.898)	0.677
	Irbesartan	36	5 (13.9)	NE				
eGFR High and UP Low	Sparsentan	39	8 (20.5)	NE		1.241	(0.426, 3.620)	0.692
	Irbesartan	37	6 (16.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.569
< 60 mL/min/1.73 m**2	Sparsentan	127	29 (22.8)	NE		1.220	(0.694, 2.147)	0.490
	Irbesartan	129	21 (16.3)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	9 (18.4)	115.7	(115.7, NE)	0.856	(0.333, 2.200)	0.747
	Irbesartan	48	9 (18.8)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	7 (26.9)	NE		1.885	(0.480, 7.404)	0.364
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.054
< 45 mL/min/1.73 m**2	Sparsentan	82	18 (22.0)	NE		0.773	(0.405, 1.479)	0.437
	Irbesartan	80	19 (23.8)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	11 (24.4)	NE		6.308	(1.379, 28.862)	0.018 *
	Irbesartan	49	2 (4.1)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	9 (18.4)	115.7	(115.7, NE)	0.856	(0.333, 2.200)	0.747
	Irbesartan	48	9 (18.8)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	7 (26.9)	NE		1.885	(0.480, 7.404)	0.364
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
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Table PT2KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan						Interaction test:	0.580
<= 1.75 g/day	Sparsentan	98	23 (23.5)	NE		1.354	(0.681, 2.689)	0.387
	Irbesartan	93	13 (14.0)	NE				
> 1.75 g/day	Sparsentan	104	22 (21.2)	NE		1.069	(0.577, 1.981)	0.832
	Irbesartan	109	20 (18.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan						Interaction test:	0.460
Yes	Sparsentan	90	20 (22.2)	NE		1.513	(0.737, 3.106)	0.260
	Irbesartan	88	12 (13.6)	NE				
No	Sparsentan	112	25 (22.3)	NE		1.018	(0.567, 1.826)	0.953
	Irbesartan	114	21 (18.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment  
 was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.965
<= 5 years	Sparsentan	113	23 (20.4)	NE		1.113	(0.611, 2.028)	0.726
	Irbesartan	127	20 (15.7)	NE				
> 5 years	Sparsentan	89	22 (24.7)	NE		1.133	(0.562, 2.285)	0.726
	Irbesartan	75	13 (17.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

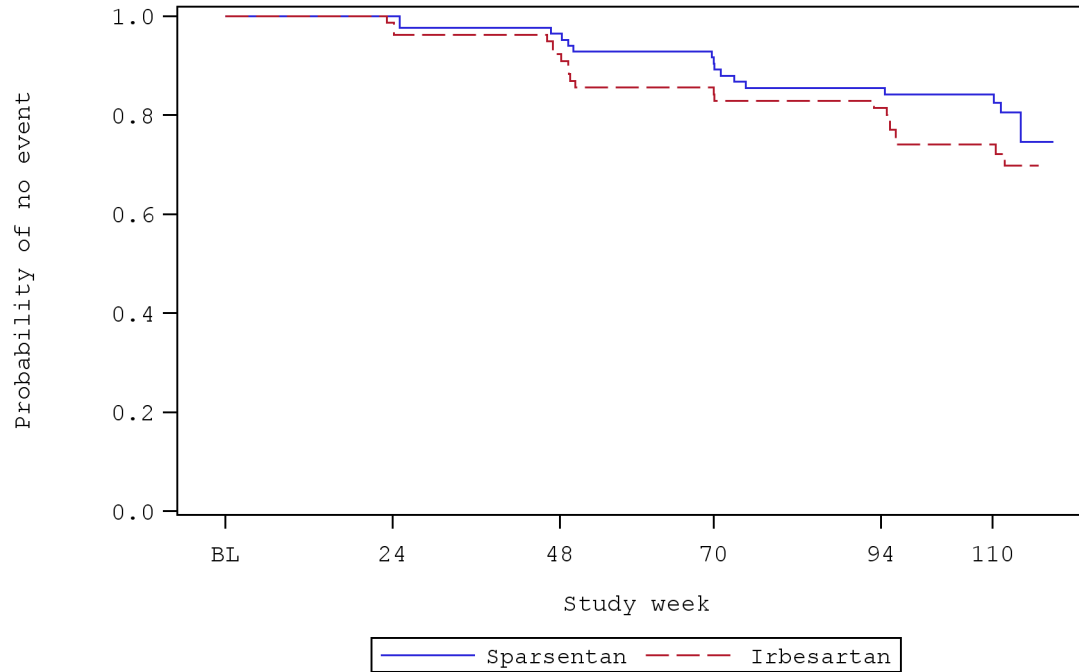
Table PT2KSYDT\_FSTM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan						Interaction test:	0.490
Yes	Sparsentan	155	31 (20.0)	NE		1.239	(0.717, 2.140)	0.443
	Irbesartan	161	22 (13.7)	NE				
No	Sparsentan	47	14 (29.8)	NE		0.888	(0.393, 2.007)	0.775
	Irbesartan	41	11 (26.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Figure PF2KSYDT\_FSKM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set

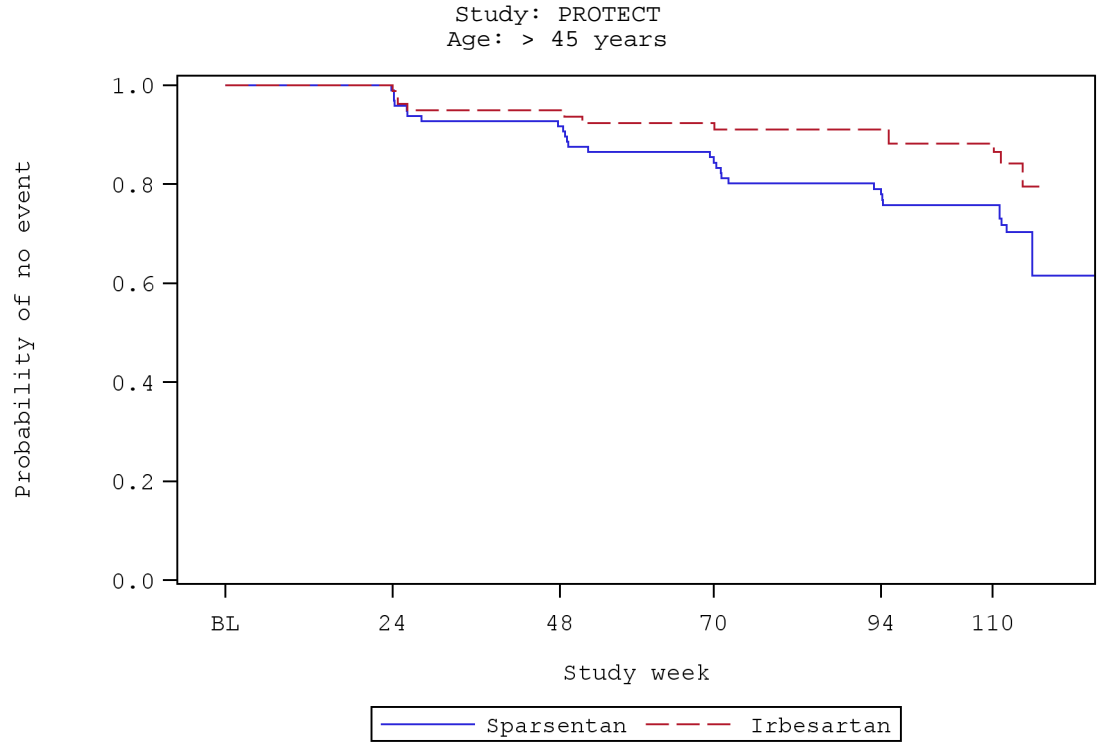
Study: PROTECT  
 Age: <= 45 years



Sparsentan	96	85	81	76	65	53
Irbesartan	99	79	70	62	56	46

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KSYDT\_FSTM

Figure PF2KSYDT\_FSKM: KDQOL: Time to decrease in symptom/problems of kidney disease by at least 15 points by subgroup  
 Full Analysis Set



Sparsentan	106	96	88	81	72	61
Irbesartan	103	82	74	69	66	52

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KSYDT\_FSTM

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Sex														
Male	KDQOL-SF12: PCS	Baseline	Sparsentan	139	130 (93.5)	51.62 (7.25)	25.5	47.38	53.46	56.74	62.3			
			Irbesartan	143	132 (92.3)	52.00 (7.33)	21.6	49.77	54.43	56.42	64.3			
		Week 24	Sparsentan	139	110 (79.1)	52.25 (6.99)	16.3	49.67	53.52	56.66	64.0			
			Irbesartan	143	100 (69.9)	51.71 (7.37)	20.8	48.17	54.17	56.22	64.3			
		Week 48	Sparsentan	139	115 (82.7)	51.97 (7.34)	25.4	47.31	53.93	56.71	66.1			
			Irbesartan	143	90 (62.9)	52.53 (7.88)	18.0	50.59	54.95	57.20	64.5			
		Week 70	Sparsentan	139	117 (84.2)	51.33 (7.30)	28.0	46.39	53.18	56.42	66.8			
			Irbesartan	143	97 (67.8)	51.81 (7.62)	21.7	49.01	54.24	56.71	63.0			
		Week 94	Sparsentan	139	108 (77.7)	51.71 (7.53)	23.8	48.01	54.39	56.83	63.5			
			Irbesartan	143	99 (69.2)	52.02 (7.24)	20.0	49.60	54.08	56.42	62.7			
		Week 110	Sparsentan	139	104 (74.8)	51.66 (7.11)	22.8	47.42	53.21	56.46	63.2			
			Irbesartan	143	89 (62.2)	51.90 (7.59)	25.4	49.07	53.50	56.71	62.6			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	139	110 (79.1)	0.62 (5.47)	-13.4	-2.71	0.18	3.69	15.5	0.14 [-0.13, 0.41]
					Irbesartan	143	100 (69.9)	-0.14 (5.63)	-20.0	-2.39	0.00	1.92	16.9	
		Week 48		Sparsentan	139	115 (82.7)	0.43 (6.43)	-18.7	-3.27	-0.08	4.02	20.6	0.04 [-0.24, 0.31]	
				Irbesartan	143	90 (62.9)	0.16 (8.54)	-35.7	-3.45	0.29	3.64	20.4		
		Week 70		Sparsentan	139	117 (84.2)	-0.45 (6.85)	-25.7	-3.10	-0.04	2.98	15.6	0.04 [-0.23, 0.31]	
				Irbesartan	143	97 (67.8)	-0.75 (7.82)	-28.0	-3.16	0.00	2.69	17.8		
Week 94	Sparsentan	139		108 (77.7)	0.11 (7.08)	-19.6	-3.22	-0.27	5.10	20.7	0.04 [-0.23, 0.31]			
	Irbesartan	143		99 (69.2)	-0.17 (6.49)	-20.1	-3.49	0.41	3.17	15.4				
Week 110	Sparsentan	139		104 (74.8)	-0.17 (7.24)	-20.5	-2.88	-0.17	4.77	16.4	0.00 [-0.28, 0.28]			
	Irbesartan	143		89 (62.2)	-0.19 (7.01)	-18.5	-3.49	0.00	4.22	15.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Female	KDQOL-SF12: PCS	Baseline	Sparsentan	63	58 (92.1)	49.83 (9.66)	12.4	44.71	52.36	56.15	63.5		
			Irbesartan	59	51 (86.4)	50.77 (6.10)	29.6	48.31	51.04	55.59	61.1		
	Week 24	Sparsentan	63	49 (77.8)	50.38 (8.06)	27.3	45.43	52.50	56.24	62.9			
		Irbesartan	59	38 (64.4)	51.59 (6.04)	39.0	46.70	53.46	55.88	60.2			
	Week 48	Sparsentan	63	50 (79.4)	50.37 (7.18)	29.8	46.58	51.30	55.57	61.7			
		Irbesartan	59	37 (62.7)	49.54 (6.70)	33.7	45.30	50.13	55.33	60.0			
	Week 70	Sparsentan	63	46 (73.0)	48.77 (8.67)	21.9	42.16	51.61	55.86	59.2			
		Irbesartan	59	38 (64.4)	49.44 (8.93)	22.3	47.10	51.36	56.95	60.2			
	Week 94	Sparsentan	63	47 (74.6)	48.82 (7.52)	27.9	41.40	50.23	54.80	59.6			
		Irbesartan	59	34 (57.6)	50.58 (6.70)	37.0	43.33	52.45	56.69	59.9			
	Week 110	Sparsentan	63	46 (73.0)	48.94 (8.08)	32.6	41.42	50.68	55.31	64.2			
		Irbesartan	59	32 (54.2)	49.47 (6.89)	39.0	43.26	49.88	55.37	60.0			
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	63	49 (77.8)	-0.10 (8.69)	-24.8	-5.43	0.00	4.08	20.4	-0.00 [-0.42, 0.42]	
			Irbesartan	59	38 (64.4)	-0.10 (5.02)	-12.8	-2.61	0.14	3.78	10.9		
			Week 48	Sparsentan	63	50 (79.4)	0.29 (7.96)	-22.3	-3.28	0.16	2.67	24.7	0.20 [-0.23, 0.63]
			Irbesartan	59	37 (62.7)	-1.24 (7.23)	-18.8	-5.09	0.00	2.18	16.4		
			Week 70	Sparsentan	63	46 (73.0)	-0.57 (9.59)	-30.2	-5.20	0.17	3.73	31.3	0.19 [-0.24, 0.62]
			Irbesartan	59	38 (64.4)	-2.16 (6.85)	-27.2	-6.02	-1.18	1.91	9.7		
			Week 94	Sparsentan	63	47 (74.6)	-1.09 (7.75)	-24.1	-4.58	-0.14	2.40	15.9	-0.04 [-0.48, 0.41]
			Irbesartan	59	34 (57.6)	-0.83 (6.69)	-14.7	-5.11	-1.58	3.53	18.0		
Week 110			Sparsentan	63	46 (73.0)	-1.52 (7.87)	-17.4	-6.37	-0.89	2.13	16.1	0.08 [-0.37, 0.53]	
Irbesartan			59	32 (54.2)	-2.12 (6.13)	-11.9	-7.60	-1.66	0.97	15.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age														
<= 45 years	KDQOL-SF12: PCS	Baseline	Sparsentan	96	88 (91.7)	52.37 (7.89)	12.4	49.42	54.38	57.08	63.5			
			Irbesartan	99	92 (92.9)	52.20 (7.28)	21.6	50.06	54.38	56.39	64.3			
		Week 24	Sparsentan	96	72 (75.0)	53.39 (6.68)	27.3	51.19	54.47	57.49	64.0			
			Irbesartan	99	70 (70.7)	52.86 (6.92)	20.8	52.09	54.79	56.40	64.3			
		Week 48	Sparsentan	96	76 (79.2)	53.07 (6.85)	29.8	48.45	54.70	57.34	66.1			
			Irbesartan	99	62 (62.6)	52.18 (7.81)	18.0	48.88	54.95	56.71	63.5			
		Week 70	Sparsentan	96	75 (78.1)	51.68 (7.84)	21.9	48.98	54.09	56.44	66.8			
			Irbesartan	99	64 (64.6)	52.94 (7.05)	34.6	50.89	55.87	57.63	63.0			
		Week 94	Sparsentan	96	70 (72.9)	52.49 (7.11)	27.9	50.22	54.70	56.95	63.5			
			Irbesartan	99	64 (64.6)	51.82 (7.29)	30.4	47.74	54.15	56.95	62.7			
		Week 110	Sparsentan	96	70 (72.9)	52.63 (7.00)	34.7	50.22	54.82	57.49	63.2			
			Irbesartan	99	59 (59.6)	52.09 (7.54)	25.4	46.91	54.80	57.58	62.6			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	96	72 (75.0)	0.53 (5.86)	-24.8	-3.15	0.75	4.08	14.7	0.14 [-0.19, 0.47]
		Irbesartan			99	70 (70.7)	-0.23 (4.91)	-15.6	-2.69	-0.41	1.65	13.3		
Week 48	Sparsentan	96		76 (79.2)	0.27 (6.85)	-22.3	-2.78	0.00	3.76	24.7	0.17 [-0.16, 0.51]			
	Irbesartan	99		62 (62.6)	-1.03 (8.22)	-35.7	-2.77	0.04	2.42	16.4				
Week 70	Sparsentan	96		75 (78.1)	-0.46 (7.74)	-30.2	-2.95	0.08	3.37	31.3	0.00 [-0.33, 0.34]			
	Irbesartan	99		64 (64.6)	-0.48 (6.73)	-24.4	-2.58	0.33	3.05	11.2				
Week 94	Sparsentan	96		70 (72.9)	-0.43 (6.22)	-24.1	-3.46	0.03	3.24	11.9	0.09 [-0.25, 0.43]			
	Irbesartan	99		64 (64.6)	-1.01 (7.23)	-20.1	-4.17	0.00	2.98	18.0				
Week 110	Sparsentan	96		70 (72.9)	-0.78 (6.58)	-20.5	-2.86	-0.04	2.33	12.2	-0.08 [-0.42, 0.27]			
	Irbesartan	99		59 (59.6)	-0.27 (6.91)	-18.5	-3.49	0.00	4.56	15.0				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024



Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 45 years	KDQOL-SF12: PCS	Baseline	Sparsentan	106	100 (94.3)	49.92 (8.11)	25.5	45.69	52.32	56.15	61.8		
			Irbesartan	103	91 (88.3)	51.12 (6.73)	27.9	48.76	52.91	56.15	59.6		
		Week 24	Sparsentan	106	87 (82.1)	50.25 (7.63)	16.3	44.66	52.35	56.15	62.9		
			Irbesartan	103	68 (66.0)	50.47 (6.94)	31.2	45.43	52.51	56.15	61.9		
		Week 48	Sparsentan	106	89 (84.0)	50.14 (7.45)	25.4	46.31	52.59	55.09	62.1		
			Irbesartan	103	65 (63.1)	51.17 (7.53)	24.9	47.87	53.71	56.48	64.5		
		Week 70	Sparsentan	106	88 (83.0)	49.69 (7.64)	28.0	43.41	51.73	56.15	60.9		
			Irbesartan	103	71 (68.9)	49.53 (8.59)	21.7	46.63	52.09	55.35	59.6		
		Week 94	Sparsentan	106	85 (80.2)	49.47 (7.80)	23.8	42.20	52.40	55.40	61.7		
			Irbesartan	103	69 (67.0)	51.50 (7.00)	20.0	49.27	53.46	56.15	60.9		
	Week 110	Sparsentan	106	80 (75.5)	49.24 (7.60)	22.8	43.65	50.21	54.95	64.2			
		Irbesartan	103	62 (60.2)	50.47 (7.36)	28.1	44.94	51.97	55.93	62.0			
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	106	87 (82.1)	0.30 (7.20)	-16.1	-3.48	-0.27	3.53	20.4	0.05 [-0.27, 0.37]
	Irbesartan			103	68 (66.0)	-0.03 (5.99)	-20.0	-2.36	0.29	3.47	16.9		
	Week 48		Sparsentan	106	89 (84.0)	0.49 (6.99)	-15.4	-3.60	-0.29	3.93	20.6	-0.00 [-0.32, 0.32]	
			Irbesartan	103	65 (63.1)	0.50 (8.13)	-20.8	-4.24	0.29	3.64	20.4		
	Week 70		Sparsentan	106	88 (83.0)	-0.50 (7.69)	-20.6	-4.44	-0.09	3.07	21.2	0.16 [-0.16, 0.47]	
			Irbesartan	103	71 (68.9)	-1.74 (8.24)	-28.0	-5.89	-0.50	2.42	17.8		
	Week 94		Sparsentan	106	85 (80.2)	-0.11 (8.09)	-20.0	-4.03	-0.27	5.11	20.7	-0.05 [-0.37, 0.26]	
			Irbesartan	103	69 (67.0)	0.28 (5.78)	-13.0	-2.83	0.29	4.53	13.6		
Week 110	Sparsentan		106	80 (75.5)	-0.42 (8.15)	-17.9	-4.89	-0.65	4.62	16.4	0.09 [-0.24, 0.42]		
	Irbesartan		103	62 (60.2)	-1.11 (6.77)	-17.3	-4.93	-1.04	2.61	15.6			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age at IgAN diagnosis														
<= 18 years	KDQOL-SF12: PCS	Baseline	Sparsentan	9	9 (100.0)	49.16 (14.77)	12.4	53.62	54.80	55.47	58.8			
			Irbesartan	5	5 (100.0)	50.57 (8.50)	39.0	44.78	53.18	56.15	59.7			
		Week 24	Sparsentan	9	5 (55.6)	56.34 (5.69)	47.0	55.09	58.85	59.89	60.9			
			Irbesartan	5	5 (100.0)	51.63 (6.38)	44.9	46.02	51.35	56.15	59.7			
		Week 48	Sparsentan	9	7 (77.8)	50.92 (9.91)	36.9	37.10	54.23	58.84	60.8			
			Irbesartan	5	3 (60.0)	55.44 (0.79)	54.6	54.59	55.59	56.15	56.1			
		Week 70	Sparsentan	9	7 (77.8)	51.64 (7.11)	40.9	43.65	53.37	57.76	59.4			
			Irbesartan	5	4 (80.0)	52.79 (8.96)	40.5	46.31	54.96	59.27	60.8			
		Week 94	Sparsentan	9	5 (55.6)	45.61 (7.92)	36.5	39.66	44.96	51.36	55.6			
			Irbesartan	5	4 (80.0)	53.17 (7.73)	46.3	46.91	51.85	59.44	62.7			
		Week 110	Sparsentan	9	4 (44.4)	50.36 (6.59)	40.9	46.52	52.20	54.19	56.1			
			Irbesartan	5	2 (40.0)	48.22 (4.59)	45.0	44.97	48.22	51.46	51.5			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	9	5 (55.6)	4.11 (3.63)	-2.1	3.76	6.13	6.27	6.5	0.87 [-0.43, 2.16]
				Irbesartan	5	5 (100.0)	1.06 (3.42)	-1.8	0.00	0.00	0.12	7.0		
Week 48	Sparsentan	9		7 (77.8)	3.35 (9.92)	-3.6	-2.97	0.60	3.75	24.7	-0.28 [-1.63, 1.08]			
	Irbesartan	5		3 (60.0)	5.99 (8.38)	0.0	0.00	2.40	15.57	15.6				
Week 70	Sparsentan	9		7 (77.8)	4.07 (12.15)	-3.3	-2.10	0.40	2.25	31.3	0.47 [-0.77, 1.72]			
	Irbesartan	5		4 (80.0)	-0.67 (2.68)	-4.3	-2.67	0.01	1.33	1.6				
Week 94	Sparsentan	9	5 (55.6)	-6.76 (7.79)	-19.0	-10.13	-2.27	-1.61	-0.8	-1.01 [-2.41, 0.38]				
	Irbesartan	5	4 (80.0)	-0.28 (3.77)	-5.6	-2.82	0.74	2.25	3.0					
Week 110	Sparsentan	9	4 (44.4)	-1.65 (1.50)	-2.9	-2.77	-2.07	-0.53	0.4	-0.60 [-2.34, 1.13]				
	Irbesartan	5	2 (40.0)	-0.76 (1.36)	-1.7	-1.73	-0.76	0.20	0.2					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 18 to 40 years	KDQOL-SF12: PCS	Baseline	Sparsentan	102	94 (92.2)	52.86 (7.28)	25.5	49.93	54.52	57.30	63.5	
			Irbesartan	109	100 (91.7)	52.36 (6.99)	21.6	50.06	54.33	56.34	64.3	
		Week 24	Sparsentan	102	80 (78.4)	53.14 (6.62)	27.3	50.98	54.47	57.49	64.0	
			Irbesartan	109	73 (67.0)	52.87 (6.54)	20.8	51.90	54.53	56.71	64.3	
		Week 48	Sparsentan	102	83 (81.4)	53.40 (6.87)	29.8	50.03	54.80	57.76	66.1	
			Irbesartan	109	69 (63.3)	52.20 (7.45)	18.0	49.48	54.80	56.71	63.5	
		Week 70	Sparsentan	102	80 (78.4)	51.73 (8.11)	21.9	49.29	54.80	56.53	66.8	
			Irbesartan	109	71 (65.1)	52.43 (7.44)	22.3	48.59	55.28	57.22	63.0	
		Week 94	Sparsentan	102	76 (74.5)	52.84 (7.17)	27.9	51.07	54.80	57.33	63.5	
			Irbesartan	109	69 (63.3)	51.72 (6.71)	34.4	47.89	53.47	56.62	61.3	
	Week 110	Sparsentan	102	73 (71.6)	52.42 (7.72)	22.8	50.22	54.84	57.76	63.2		
		Irbesartan	109	65 (59.6)	52.38 (7.39)	25.4	49.07	55.01	57.58	62.0		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	102	80 (78.4)	0.43 (6.07)	-24.8	-2.87	0.22	4.04	14.7	0.13 [-0.19, 0.45]
			Irbesartan	109	73 (67.0)	-0.27 (4.50)	-15.0	-2.38	0.00	1.65	13.3	
		Week 48	Sparsentan	102	83 (81.4)	0.20 (6.16)	-22.3	-2.88	0.00	3.69	15.6	0.15 [-0.17, 0.46]
			Irbesartan	109	69 (63.3)	-0.84 (8.19)	-35.7	-2.98	0.56	2.22	16.4	
		Week 70	Sparsentan	102	80 (78.4)	-0.95 (7.45)	-30.2	-3.95	0.17	3.21	15.6	-0.01 [-0.33, 0.31]
			Irbesartan	109	71 (65.1)	-0.84 (7.33)	-27.2	-2.69	0.67	2.69	15.7	
		Week 94	Sparsentan	102	76 (74.5)	-0.00 (6.16)	-24.1	-3.24	0.30	3.47	11.9	0.18 [-0.15, 0.51]
			Irbesartan	109	69 (63.3)	-1.16 (6.82)	-18.6	-4.31	0.00	2.45	18.0	
Week 110		Sparsentan	102	73 (71.6)	-0.97 (6.47)	-20.5	-3.20	-0.52	2.13	12.2	-0.08 [-0.42, 0.25]	
		Irbesartan	109	65 (59.6)	-0.42 (6.78)	-18.5	-3.25	0.00	3.57	15.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 PCS = physical component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 40 years	KDQOL-SF12: PCS	Baseline	Sparsentan	91	85 (93.4)	49.29 (7.67)	25.8	44.69	50.89	54.80	60.2	
			Irbesartan	88	78 (88.6)	50.84 (6.96)	27.9	48.47	52.28	56.15	59.6	
		Week 24	Sparsentan	91	74 (81.3)	49.77 (7.79)	16.3	43.35	52.21	56.15	62.9	
			Irbesartan	88	60 (68.2)	50.24 (7.44)	31.2	44.37	53.20	56.15	61.9	
		Week 48	Sparsentan	91	75 (82.4)	49.42 (7.04)	25.4	45.70	51.29	54.80	59.6	
			Irbesartan	88	55 (62.5)	50.79 (8.04)	24.9	45.30	52.76	56.97	64.5	
		Week 70	Sparsentan	91	76 (83.5)	49.32 (7.35)	28.0	43.41	51.43	55.87	59.6	
			Irbesartan	88	60 (68.2)	49.51 (8.52)	21.7	46.86	51.57	54.80	59.6	
		Week 94	Sparsentan	91	74 (81.3)	49.13 (7.57)	23.8	42.20	50.72	55.33	61.7	
			Irbesartan	88	60 (68.2)	51.47 (7.62)	20.0	49.44	53.46	56.18	61.3	
	Week 110	Sparsentan	91	73 (80.2)	49.25 (7.06)	32.6	43.40	49.86	54.80	64.2		
		Irbesartan	88	54 (61.4)	50.02 (7.50)	28.1	44.69	51.33	55.09	62.6		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	91	74 (81.3)	0.12 (7.28)	-16.1	-3.94	-0.73	3.49	20.4	0.03 [-0.32, 0.37]
			Irbesartan	88	60 (68.2)	-0.06 (6.58)	-20.0	-3.29	0.00	3.78	16.9	
		Week 48	Sparsentan	91	75 (82.4)	0.32 (7.40)	-15.4	-4.33	-0.54	5.05	20.6	0.02 [-0.33, 0.37]
			Irbesartan	88	55 (62.5)	0.15 (8.14)	-20.8	-4.31	0.00	4.04	20.4	
		Week 70	Sparsentan	91	76 (83.5)	-0.41 (7.43)	-20.4	-4.10	-0.33	3.31	21.2	0.15 [-0.19, 0.48]
			Irbesartan	88	60 (68.2)	-1.54 (8.10)	-28.0	-6.07	-0.27	2.43	17.8	
		Week 94	Sparsentan	91	74 (81.3)	-0.07 (8.16)	-19.6	-4.39	-0.14	5.11	20.7	-0.09 [-0.43, 0.25]
			Irbesartan	88	60 (68.2)	0.61 (6.27)	-20.1	-2.67	0.35	4.97	13.6	
Week 110		Sparsentan	91	73 (80.2)	-0.15 (8.49)	-17.9	-5.45	0.00	5.40	16.4	0.11 [-0.24, 0.46]	
		Irbesartan	88	54 (61.4)	-1.03 (7.04)	-17.3	-5.80	-1.11	3.16	15.6		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Geographic region													
North America	KDQOL-SF12: PCS	Baseline	Sparsentan	35	32 (91.4)	52.20 (9.84)	12.4	50.86	54.80	57.76	62.3		
			Irbesartan	46	43 (93.5)	50.66 (8.22)	27.9	47.65	54.24	56.15	61.1		
		Week 24	Sparsentan	35	23 (65.7)	54.24 (6.48)	35.4	52.64	56.15	57.49	64.0		
			Irbesartan	46	37 (80.4)	51.74 (6.97)	31.2	48.45	54.51	56.22	60.2		
		Week 48	Sparsentan	35	25 (71.4)	53.35 (6.94)	37.1	50.24	55.59	57.76	66.0		
			Irbesartan	46	33 (71.7)	52.26 (8.67)	18.0	47.21	55.64	57.47	64.5		
		Week 70	Sparsentan	35	23 (65.7)	52.71 (6.17)	42.2	48.98	53.17	57.20	63.2		
			Irbesartan	46	31 (67.4)	53.82 (6.76)	23.5	52.64	55.86	57.31	59.9		
		Week 94	Sparsentan	35	24 (68.6)	54.43 (4.00)	44.8	51.80	56.02	57.12	61.3		
			Irbesartan	46	30 (65.2)	52.63 (6.78)	29.2	47.92	55.12	56.95	59.9		
		Week 110	Sparsentan	35	21 (60.0)	52.83 (6.87)	35.4	49.10	54.80	57.76	63.2		
			Irbesartan	46	31 (67.4)	52.87 (5.82)	35.0	47.77	54.51	57.58	60.5		
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	35	23 (65.7)	-1.34 (5.27)	-16.1	-3.22	-0.27	2.88	6.1	-0.41 [-0.94, 0.12]
			Irbesartan	46	37 (80.4)	0.99 (5.94)	-20.0	-1.35	0.80	3.27	13.3		
	Week 48		Sparsentan	35	25 (71.4)	0.40 (6.91)	-15.4	-1.90	-0.52	2.67	24.7	-0.14 [-0.66, 0.38]	
			Irbesartan	46	33 (71.7)	1.64 (10.41)	-35.7	-1.06	1.35	6.73	20.4		
	Week 70		Sparsentan	35	23 (65.7)	0.10 (7.78)	-9.4	-2.89	-0.47	1.61	31.3	-0.16 [-0.70, 0.38]	
			Irbesartan	46	31 (67.4)	1.19 (5.88)	-13.6	-1.74	1.35	2.69	17.8		
Week 94	Sparsentan	35	24 (68.6)	0.51 (5.69)	-7.3	-3.14	-0.32	3.20	20.7	-0.20 [-0.74, 0.33]			
	Irbesartan	46	30 (65.2)	1.75 (6.41)	-10.7	-1.82	1.05	5.11	18.0				
Week 110	Sparsentan	35	21 (60.0)	-2.48 (5.35)	-20.5	-3.51	-1.35	0.56	3.9	-0.76 [-1.33, -0.19]			
	Irbesartan	46	31 (67.4)	2.12 (6.46)	-10.2	-1.42	1.35	6.00	15.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Europe	KDQOL-SF12: PCS	Baseline	Sparsentan	98	89 (90.8)	51.21 (7.64)	25.5	48.20	53.46	56.62	62.3		
			Irbesartan	115	99 (86.1)	52.29 (6.36)	29.2	49.79	53.95	56.71	62.0		
		Week 24	Sparsentan	98	75 (76.5)	52.04 (7.44)	16.3	50.24	53.44	56.71	62.9		
			Irbesartan	115	65 (56.5)	51.69 (6.45)	33.8	46.99	52.94	56.15	64.3		
		Week 48	Sparsentan	98	77 (78.6)	51.47 (7.46)	25.4	47.31	54.23	56.15	62.1		
			Irbesartan	115	61 (53.0)	51.74 (7.75)	24.9	48.01	54.63	57.49	63.5		
		Week 70	Sparsentan	98	76 (77.6)	50.75 (7.42)	28.0	46.33	52.50	56.15	63.2		
			Irbesartan	115	72 (62.6)	49.95 (8.98)	21.7	44.37	52.41	56.43	63.0		
		Week 94	Sparsentan	98	71 (72.4)	50.39 (7.84)	23.8	43.49	53.18	56.15	61.7		
			Irbesartan	115	72 (62.6)	51.30 (7.34)	20.0	47.72	53.21	56.18	62.7		
	Week 110	Sparsentan	98	68 (69.4)	51.46 (7.96)	22.8	47.18	52.77	56.59	64.2			
		Irbesartan	115	61 (53.0)	50.85 (8.69)	25.4	44.97	53.46	56.71	62.6			
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	98	75 (76.5)	1.11 (6.03)	-13.4	-2.71	0.29	4.30	14.7	0.41 [0.07, 0.75]
	Irbesartan			115	65 (56.5)	-1.15 (4.85)	-15.0	-3.09	-0.83	1.33	10.9		
	Week 48		Sparsentan	98	77 (78.6)	0.68 (6.33)	-18.7	-3.10	0.56	4.31	17.5	0.26 [-0.08, 0.59]	
			Irbesartan	115	61 (53.0)	-1.08 (7.43)	-24.7	-4.24	0.00	2.40	16.0		
	Week 70		Sparsentan	98	76 (77.6)	-0.37 (7.63)	-25.7	-4.01	0.55	3.93	21.2	0.29 [-0.04, 0.61]	
			Irbesartan	115	72 (62.6)	-2.63 (8.17)	-28.0	-7.08	-1.08	1.75	15.7		
	Week 94		Sparsentan	98	71 (72.4)	-0.17 (7.70)	-20.0	-3.04	0.27	4.86	15.9	0.19 [-0.14, 0.52]	
			Irbesartan	115	72 (62.6)	-1.50 (6.08)	-18.6	-4.29	-0.61	2.19	13.6		
Week 110	Sparsentan		98	68 (69.4)	0.69 (7.30)	-17.8	-1.91	0.41	5.45	16.1	0.35 [-0.00, 0.69]		
	Irbesartan		115	61 (53.0)	-1.77 (6.92)	-18.5	-4.79	-1.06	2.69	11.1			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Asia Pacific	KDQOL-SF12: PCS	Baseline	Sparsentan	69	67 (97.1)	50.34 (7.79)	32.4	44.69	52.08	56.15	63.5	
			Irbesartan	41	41 (100.0)	51.18 (7.18)	21.6	49.48	51.57	55.86	64.3	
		Week 24	Sparsentan	69	61 (88.4)	50.25 (7.37)	27.3	45.37	52.40	56.15	62.0	
			Irbesartan	41	36 (87.8)	51.61 (8.14)	20.8	49.25	54.51	56.33	59.4	
		Week 48	Sparsentan	69	63 (91.3)	50.77 (7.24)	29.8	46.56	52.88	56.15	66.1	
			Irbesartan	41	33 (80.5)	50.93 (6.47)	33.7	48.88	53.17	55.09	58.8	
		Week 70	Sparsentan	69	64 (92.8)	49.68 (8.60)	21.9	44.11	53.05	56.31	66.8	
			Irbesartan	41	32 (78.0)	51.24 (6.41)	30.0	48.52	52.16	55.66	58.8	
		Week 94	Sparsentan	69	60 (87.0)	49.92 (8.13)	27.9	42.66	52.71	56.28	63.5	
			Irbesartan	41	31 (75.6)	51.52 (7.03)	30.4	49.06	53.46	56.15	60.9	
	Week 110	Sparsentan	69	61 (88.4)	49.43 (7.02)	34.7	43.07	50.91	55.31	59.7		
		Irbesartan	41	29 (70.7)	50.41 (6.02)	40.0	45.69	50.55	55.09	61.5		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	69	61 (88.4)	0.19 (7.63)	-24.8	-3.94	0.00	4.90	20.4	-0.05 [-0.46, 0.36]
			Irbesartan	41	36 (87.8)	0.56 (5.77)	-15.6	-2.46	0.00	3.77	16.9	
		Week 48	Sparsentan	69	63 (91.3)	0.03 (7.64)	-22.3	-3.58	-0.86	4.02	20.6	0.08 [-0.34, 0.51]
			Irbesartan	41	33 (80.5)	-0.59 (6.82)	-18.8	-4.39	0.00	3.24	15.5	
		Week 70	Sparsentan	69	64 (92.8)	-0.82 (7.83)	-30.2	-4.44	-0.23	3.31	15.6	-0.10 [-0.52, 0.32]
			Irbesartan	41	32 (78.0)	-0.06 (7.00)	-25.1	-3.10	0.73	3.71	15.2	
		Week 94	Sparsentan	69	60 (87.0)	-0.65 (7.43)	-24.1	-4.30	-0.41	5.51	13.7	-0.13 [-0.57, 0.30]
			Irbesartan	41	31 (75.6)	0.33 (7.23)	-20.1	-5.11	1.10	5.07	12.0	
Week 110		Sparsentan	69	61 (88.4)	-1.37 (8.04)	-17.9	-5.47	-1.34	4.70	16.4	0.01 [-0.43, 0.45]	
		Irbesartan	41	29 (70.7)	-1.45 (6.33)	-11.3	-5.71	-1.09	1.73	15.5		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline BMI														
< 27 kg/m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	83	75 (90.4)	51.42 (8.74)	12.4	47.38	53.46	57.49	63.5			
			Irbesartan	94	86 (91.5)	53.13 (5.19)	39.0	49.84	54.52	56.42	61.5			
		Week 24	Sparsentan	83	66 (79.5)	52.61 (6.55)	37.6	49.19	53.85	57.49	64.0			
			Irbesartan	94	69 (73.4)	52.59 (5.57)	36.8	50.19	54.51	56.15	60.8			
		Week 48	Sparsentan	83	68 (81.9)	51.99 (6.87)	33.2	47.22	53.23	57.09	66.1			
			Irbesartan	94	60 (63.8)	51.48 (7.72)	18.0	47.16	54.52	56.84	63.5			
		Week 70	Sparsentan	83	66 (79.5)	50.70 (8.08)	30.4	44.81	53.05	56.15	66.8			
			Irbesartan	94	62 (66.0)	51.66 (7.58)	26.4	47.78	54.38	57.20	60.8			
		Week 94	Sparsentan	83	64 (77.1)	51.48 (7.81)	31.0	45.82	53.46	57.46	63.5			
			Irbesartan	94	64 (68.1)	52.49 (5.29)	38.9	48.29	54.23	56.35	62.7			
		Week 110	Sparsentan	83	63 (75.9)	51.36 (7.74)	32.6	44.33	53.46	57.76	64.2			
			Irbesartan	94	57 (60.6)	52.54 (6.45)	36.2	49.31	54.80	57.55	61.5			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	83	66 (79.5)	0.67 (6.29)	-11.9	-3.48	-0.07	5.98	20.4	0.24 [-0.10, 0.58]
		Irbesartan			94	69 (73.4)	-0.62 (4.08)	-15.0	-2.61	-0.51	1.35	10.4		
Week 48	Sparsentan	83		68 (81.9)	0.35 (6.64)	-13.9	-4.13	-0.30	4.34	24.7	0.29 [-0.06, 0.64]			
	Irbesartan	94		60 (63.8)	-1.71 (7.67)	-35.7	-4.06	0.00	2.04	15.6				
Week 70	Sparsentan	83		66 (79.5)	-1.02 (8.60)	-25.7	-5.11	-1.09	2.90	31.3	0.08 [-0.26, 0.43]			
	Irbesartan	94		62 (66.0)	-1.69 (7.54)	-28.0	-2.69	-0.13	1.89	15.7				
Week 94	Sparsentan	83		64 (77.1)	-0.66 (7.26)	-20.0	-3.59	-0.64	3.59	20.7	0.09 [-0.25, 0.44]			
	Irbesartan	94		64 (68.1)	-1.27 (5.54)	-18.6	-3.76	0.14	2.21	9.1				
Week 110	Sparsentan	83		63 (75.9)	-1.39 (7.72)	-17.9	-6.37	-0.52	2.22	15.6	-0.08 [-0.44, 0.28]			
	Irbesartan	94		57 (60.6)	-0.84 (5.55)	-18.3	-3.45	-0.54	1.95	11.3				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024



Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 27 kg/m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	119	113 (95.0)	50.83 (7.64)	25.5	46.66	52.40	56.15	62.3		
			Irbesartan	107	96 (89.7)	50.47 (8.07)	21.6	47.36	52.90	56.10	64.3		
	Week 24	Sparsentan	119	93 (78.2)	51.01 (7.85)	16.3	47.24	52.70	56.42	62.9			
		Irbesartan	107	69 (64.5)	50.77 (8.14)	20.8	45.88	53.46	56.15	64.3			
	Week 48	Sparsentan	119	97 (81.5)	51.14 (7.62)	25.4	46.91	53.71	56.15	62.1			
		Irbesartan	107	67 (62.6)	51.83 (7.64)	24.9	48.88	54.80	56.71	64.5			
	Week 70	Sparsentan	119	97 (81.5)	50.54 (7.59)	21.9	46.03	52.61	56.19	61.1			
		Irbesartan	107	73 (68.2)	50.70 (8.46)	21.7	48.04	52.62	56.15	63.0			
	Week 94	Sparsentan	119	91 (76.5)	50.38 (7.50)	23.8	45.84	52.91	56.15	59.3			
		Irbesartan	107	68 (63.6)	51.11 (8.26)	20.0	48.08	53.46	56.43	61.3			
	Week 110	Sparsentan	119	87 (73.1)	50.44 (7.34)	22.8	46.04	52.50	55.92	63.1			
		Irbesartan	107	63 (58.9)	50.51 (7.57)	28.1	46.13	51.57	56.15	62.6			
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	119	93 (78.2)	0.21 (6.86)	-24.8	-3.22	0.21	3.49	15.5	-0.02 [-0.33, 0.29]
				Irbesartan	107	69 (64.5)	0.36 (6.54)	-20.0	-2.55	0.29	3.64	16.9	
	Week 48		Sparsentan	119	97 (81.5)	0.42 (7.12)	-22.3	-3.10	0.00	3.60	20.6	-0.08 [-0.40, 0.23]	
			Irbesartan	107	67 (62.6)	1.06 (8.45)	-20.8	-4.31	1.09	5.40	20.4		
	Week 70		Sparsentan	119	97 (81.5)	-0.12 (7.02)	-30.2	-2.69	0.27	3.37	15.8	0.08 [-0.23, 0.38]	
			Irbesartan	107	73 (68.2)	-0.68 (7.61)	-27.2	-5.05	0.51	2.69	17.8		
	Week 94		Sparsentan	119	91 (76.5)	0.03 (7.33)	-24.1	-3.52	0.27	4.51	15.9	-0.08 [-0.39, 0.24]	
			Irbesartan	107	68 (63.6)	0.59 (7.29)	-20.1	-3.36	0.61	5.09	18.0		
Week 110	Sparsentan		119	87 (73.1)	-0.01 (7.21)	-20.5	-2.86	-0.41	4.50	16.4	0.05 [-0.28, 0.37]		
	Irbesartan		107	63 (58.9)	-0.36 (7.71)	-18.5	-5.13	0.00	4.84	15.6			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Randomization strata														
eGFR Low and UP High	KDQOL-SF12: PCS	Baseline	Sparsentan	71	64 (90.1)	51.45 (7.23)	29.7	46.69	53.18	56.41	63.5			
			Irbesartan	74	64 (86.5)	50.34 (7.86)	29.2	44.67	52.50	56.15	60.1			
		Week 24	Sparsentan	71	54 (76.1)	51.20 (7.62)	16.3	47.24	52.97	56.66	60.9			
			Irbesartan	74	44 (59.5)	50.24 (7.51)	33.8	43.77	53.04	56.18	61.9			
		Week 48	Sparsentan	71	53 (74.6)	51.36 (7.35)	25.4	47.92	53.24	57.22	61.9			
			Irbesartan	74	38 (51.4)	49.94 (9.76)	18.0	45.30	52.63	56.48	64.5			
		Week 70	Sparsentan	71	58 (81.7)	50.24 (8.02)	28.0	42.96	52.12	56.40	61.1			
			Irbesartan	74	41 (55.4)	49.86 (8.38)	21.7	47.36	52.11	55.86	59.6			
		Week 94	Sparsentan	71	52 (73.2)	51.03 (7.58)	23.8	46.47	53.04	56.93	59.6			
			Irbesartan	74	38 (51.4)	49.91 (7.97)	20.0	47.59	52.89	54.53	57.8			
		Week 110	Sparsentan	71	53 (74.6)	50.91 (7.25)	32.6	44.40	52.62	56.15	64.2			
			Irbesartan	74	36 (48.6)	49.95 (7.78)	25.4	45.33	52.32	55.47	58.9			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	71	54 (76.1)	-0.38 (6.55)	-13.4	-3.70	-0.27	3.49	20.4	0.01 [-0.39, 0.41]
				Irbesartan	74	44 (59.5)	-0.46 (5.53)	-15.6	-2.69	0.20	3.29	12.3		
	Week 48	Sparsentan		71	53 (74.6)	0.02 (7.21)	-18.7	-3.60	-0.29	3.75	20.6	0.05 [-0.36, 0.47]		
		Irbesartan		74	38 (51.4)	-0.47 (11.00)	-35.7	-5.82	0.00	5.13	20.4			
	Week 70	Sparsentan		71	58 (81.7)	-1.13 (6.36)	-20.6	-4.50	-0.08	2.30	15.6	-0.05 [-0.45, 0.35]		
		Irbesartan		74	41 (55.4)	-0.78 (8.38)	-28.0	-4.57	-0.29	2.69	17.8			
	Week 94	Sparsentan	71	52 (73.2)	-0.56 (5.87)	-15.2	-3.58	-0.85	3.72	13.7	-0.09 [-0.51, 0.32]			
		Irbesartan	74	38 (51.4)	0.06 (7.55)	-20.1	-3.49	0.00	3.53	18.0				
Week 110	Sparsentan	71	53 (74.6)	-0.91 (7.41)	-17.9	-2.90	-0.74	2.13	16.4	-0.07 [-0.49, 0.36]				
	Irbesartan	74	36 (48.6)	-0.43 (7.21)	-18.3	-4.20	-0.94	4.45	15.6					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR Low and UP Low	KDQOL-SF12: PCS	Baseline	Sparsentan	55	52 (94.5)	49.20 (8.16)	25.8	45.51	50.70	54.97	61.1	
			Irbesartan	55	50 (90.9)	52.29 (4.45)	39.6	50.07	52.68	56.15	62.0	
		Week 24	Sparsentan	55	44 (80.0)	50.64 (7.32)	31.2	44.35	52.97	55.89	64.0	
			Irbesartan	55	39 (70.9)	51.58 (6.48)	32.9	46.99	52.62	56.22	64.3	
		Week 48	Sparsentan	55	43 (78.2)	51.31 (7.15)	36.0	46.31	53.71	55.95	66.0	
			Irbesartan	55	36 (65.5)	52.06 (6.11)	37.7	47.94	53.58	56.61	63.5	
		Week 70	Sparsentan	55	42 (76.4)	50.79 (7.47)	30.4	48.80	53.27	56.15	63.2	
			Irbesartan	55	40 (72.7)	50.11 (7.96)	22.3	46.68	52.04	56.15	60.3	
		Week 94	Sparsentan	55	42 (76.4)	50.49 (7.11)	31.0	46.35	53.18	55.90	61.3	
			Irbesartan	55	40 (72.7)	51.67 (7.11)	34.4	47.90	54.43	56.79	60.8	
	Week 110	Sparsentan	55	38 (69.1)	49.90 (7.40)	34.1	43.40	52.16	54.86	63.2		
		Irbesartan	55	34 (61.8)	50.93 (8.28)	36.1	41.84	54.56	57.79	62.0		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	55	44 (80.0)	1.24 (6.33)	-14.9	-3.17	1.14	4.45	14.7	0.32 [-0.12, 0.75]
			Irbesartan	55	39 (70.9)	-0.73 (6.05)	-20.0	-3.09	0.00	2.22	16.9	
		Week 48	Sparsentan	55	43 (78.2)	1.61 (5.51)	-11.5	-1.54	0.69	6.14	14.8	0.25 [-0.19, 0.70]
			Irbesartan	55	36 (65.5)	0.11 (6.40)	-11.9	-2.78	0.55	2.59	16.0	
		Week 70	Sparsentan	55	42 (76.4)	0.63 (6.88)	-20.4	-2.95	0.96	4.22	21.2	0.32 [-0.11, 0.76]
			Irbesartan	55	40 (72.7)	-1.71 (7.67)	-27.2	-5.77	-0.13	3.52	15.2	
		Week 94	Sparsentan	55	42 (76.4)	1.24 (7.35)	-19.6	-2.77	0.19	6.33	20.7	0.25 [-0.18, 0.69]
			Irbesartan	55	40 (72.7)	-0.48 (6.04)	-13.0	-4.29	0.81	3.69	12.0	
Week 110		Sparsentan	55	38 (69.1)	0.11 (7.90)	-15.3	-4.31	0.42	5.95	14.2	0.14 [-0.32, 0.60]	
		Irbesartan	55	34 (61.8)	-0.98 (7.69)	-17.3	-8.29	-0.01	3.57	15.5		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP High	KDQOL-SF12: PCS	Baseline	Sparsentan	37	35 (94.6)	52.49 (9.30)	12.4	50.75	54.80	56.71	62.9	
			Irbesartan	36	33 (91.7)	51.69 (7.00)	32.5	48.47	53.46	56.15	64.3	
		Week 24	Sparsentan	37	27 (73.0)	53.52 (8.37)	27.3	50.86	57.20	57.78	61.8	
			Irbesartan	36	26 (72.2)	53.12 (4.22)	43.0	51.40	54.51	56.15	58.9	
		Week 48	Sparsentan	37	34 (91.9)	51.81 (7.99)	29.8	47.31	53.55	57.49	62.1	
			Irbesartan	36	25 (69.4)	51.76 (5.43)	40.3	49.48	53.17	55.32	60.8	
		Week 70	Sparsentan	37	31 (83.8)	52.34 (8.04)	21.9	50.55	55.21	57.20	63.2	
			Irbesartan	36	25 (69.4)	51.72 (7.76)	23.5	51.57	53.47	56.15	59.9	
		Week 94	Sparsentan	37	27 (73.0)	51.28 (8.01)	27.9	45.84	54.59	56.95	61.7	
			Irbesartan	36	25 (69.4)	52.72 (5.44)	39.8	48.24	54.12	56.15	61.3	
	Week 110	Sparsentan	37	26 (70.3)	54.40 (5.75)	34.7	50.91	56.22	58.64	59.4		
		Irbesartan	36	22 (61.1)	51.79 (6.14)	32.1	49.25	52.52	55.09	60.5		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	37	27 (73.0)	-0.24 (7.30)	-24.8	-1.42	1.57	4.02	9.9	-0.25 [-0.79, 0.29]
			Irbesartan	36	26 (72.2)	1.32 (5.10)	-5.9	-1.86	0.00	2.69	13.3	
		Week 48	Sparsentan	37	34 (91.9)	-0.62 (8.07)	-22.3	-3.76	-1.15	3.39	24.7	-0.05 [-0.57, 0.46]
			Irbesartan	36	25 (69.4)	-0.22 (6.59)	-15.8	-3.90	0.00	4.31	13.1	
		Week 70	Sparsentan	37	31 (83.8)	0.42 (9.53)	-30.2	-3.34	1.06	4.13	31.3	0.12 [-0.41, 0.65]
			Irbesartan	36	25 (69.4)	-0.59 (7.05)	-20.4	-2.75	0.67	2.61	11.2	
		Week 94	Sparsentan	37	27 (73.0)	-1.83 (8.09)	-24.1	-5.12	-0.53	2.96	8.2	-0.30 [-0.85, 0.25]
			Irbesartan	36	25 (69.4)	0.44 (6.93)	-13.4	-3.49	0.00	4.04	15.4	
Week 110		Sparsentan	37	26 (70.3)	1.07 (6.12)	-17.4	-1.94	0.53	5.40	13.3	0.20 [-0.37, 0.77]	
		Irbesartan	36	22 (61.1)	-0.21 (6.78)	-18.5	-1.73	-0.65	3.46	13.8		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
eGFR High and UP Low	KDQOL-SF12: PCS	Baseline	Sparsentan	39	37 (94.9)	51.70 (8.03)	25.5	46.73	53.46	57.30	62.3			
			Irbesartan	37	36 (97.3)	53.11 (8.14)	21.6	51.72	55.73	56.96	60.8			
		Week 24	Sparsentan	39	34 (87.2)	52.30 (6.02)	39.4	49.19	52.57	56.42	62.9			
			Irbesartan	37	29 (78.4)	52.70 (8.63)	20.8	52.62	54.80	56.71	60.8			
		Week 48	Sparsentan	39	35 (89.7)	51.58 (7.05)	32.1	46.58	54.61	56.45	66.1			
			Irbesartan	37	28 (75.7)	53.40 (7.79)	33.7	52.73	56.28	58.02	63.4			
		Week 70	Sparsentan	39	32 (82.1)	49.36 (7.48)	31.6	44.91	51.47	54.80	66.8			
			Irbesartan	37	29 (78.4)	53.89 (7.61)	30.0	52.29	56.66	58.82	63.0			
		Week 94	Sparsentan	39	34 (87.2)	50.61 (8.28)	30.9	44.79	53.24	56.15	63.5			
			Irbesartan	37	30 (81.1)	52.94 (7.08)	29.2	49.81	54.71	57.22	62.7			
		Week 110	Sparsentan	39	33 (84.6)	48.93 (8.49)	22.8	43.24	50.22	54.84	59.7			
			Irbesartan	37	29 (78.4)	52.87 (6.97)	35.0	48.68	56.15	57.76	62.6			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	39	34 (87.2)	1.07 (6.59)	-8.4	-3.69	-0.79	5.98	14.3	0.21 [-0.29, 0.70]
		Irbesartan			37	29 (78.4)	-0.14 (4.78)	-12.8	-2.16	-0.32	2.42	10.4		
		Week 48		Sparsentan	39	35 (89.7)	0.43 (6.84)	-13.6	-3.28	0.02	4.02	17.5	0.12 [-0.38, 0.62]	
				Irbesartan	37	28 (75.7)	-0.42 (7.38)	-20.8	-1.89	0.95	3.14	12.5		
		Week 70		Sparsentan	39	32 (82.1)	-1.64 (8.90)	-25.7	-7.93	-1.60	4.43	15.8	-0.03 [-0.54, 0.47]	
				Irbesartan	37	29 (78.4)	-1.36 (6.92)	-25.1	-2.48	0.51	1.89	10.0		
		Week 94		Sparsentan	39	34 (87.2)	-0.37 (8.42)	-18.4	-7.35	0.21	4.04	15.9	0.13 [-0.36, 0.62]	
				Irbesartan	37	30 (81.1)	-1.32 (5.52)	-14.7	-4.04	0.33	2.45	6.6		
Week 110	Sparsentan	39		33 (84.6)	-2.17 (7.84)	-20.5	-6.76	-2.52	1.07	16.1	-0.16 [-0.66, 0.34]			
	Irbesartan	37		29 (78.4)	-1.06 (5.45)	-13.7	-4.48	0.00	1.61	8.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 1													
< 60 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	127	117 (92.1)	50.36 (7.64)	25.8	46.21	52.35	56.15	63.5		
			Irbesartan	129	115 (89.1)	51.35 (6.58)	27.9	48.45	52.89	56.15	62.0		
		Sparsentan	127	99 (78.0)	50.77 (7.40)	16.3	45.43	52.77	56.15	64.0			
		Irbesartan	129	85 (65.9)	50.78 (7.10)	31.2	45.08	52.62	56.15	64.3			
	Week 24	Sparsentan	127	97 (76.4)	51.64 (7.08)	25.4	47.92	53.46	56.15	66.0			
		Irbesartan	129	76 (58.9)	51.51 (8.31)	18.0	47.94	54.51	56.84	64.5			
	Week 48	Sparsentan	127	101 (79.5)	50.48 (7.77)	28.0	44.05	52.35	56.42	63.2			
		Irbesartan	129	83 (64.3)	50.13 (7.99)	21.7	47.36	52.22	56.13	60.3			
	Week 70	Sparsentan	127	94 (74.0)	50.53 (7.52)	23.8	44.96	53.04	56.15	61.3			
		Irbesartan	129	79 (61.2)	51.02 (7.65)	20.0	47.89	53.46	56.15	61.3			
	Week 94	Sparsentan	127	93 (73.2)	50.57 (7.34)	32.6	44.05	52.62	55.92	64.2			
		Irbesartan	129	72 (55.8)	50.32 (8.06)	25.4	44.81	51.97	56.40	62.6			
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	127	99 (78.0)	0.25 (6.51)	-14.9	-3.48	0.00	3.69	20.4	0.09 [-0.20, 0.38]
				Irbesartan	129	85 (65.9)	-0.30 (5.62)	-20.0	-2.69	0.00	2.96	16.9	
			Week 48	Sparsentan	127	97 (76.4)	1.12 (6.29)	-18.7	-2.83	0.00	4.71	20.6	0.10 [-0.20, 0.40]
				Irbesartan	129	76 (58.9)	0.37 (8.77)	-35.7	-4.23	0.81	3.66	20.4	
			Week 70	Sparsentan	127	101 (79.5)	-0.29 (6.60)	-20.6	-2.95	0.38	2.98	21.2	0.12 [-0.17, 0.41]
				Irbesartan	129	83 (64.3)	-1.12 (7.91)	-28.0	-4.76	-0.26	3.49	17.8	
		Week 94	Sparsentan	127	94 (74.0)	0.14 (6.79)	-19.6	-3.46	-0.14	4.86	20.7	0.02 [-0.28, 0.32]	
			Irbesartan	129	79 (61.2)	-0.02 (6.06)	-18.6	-3.47	0.41	3.77	13.6		
		Week 110	Sparsentan	127	93 (73.2)	-0.38 (7.54)	-17.9	-2.90	0.00	4.53	16.4	0.06 [-0.25, 0.37]	
			Irbesartan	129	72 (55.8)	-0.82 (7.09)	-18.3	-5.03	-0.29	4.05	15.6		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 PCS = physical component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	49	47 (95.9)	53.00 (6.69)	37.4	48.78	53.79	57.76	61.8		
			Irbesartan	48	44 (91.7)	52.03 (7.49)	29.6	49.15	54.95	56.30	64.3		
		Week 24	Sparsentan	49	39 (79.6)	53.54 (6.86)	27.3	50.55	56.15	57.49	62.9		
			Irbesartan	48	33 (68.8)	53.37 (5.41)	34.9	52.40	54.53	56.15	60.8		
		Week 48	Sparsentan	49	45 (91.8)	51.72 (8.05)	29.8	46.56	53.83	57.49	66.1		
			Irbesartan	48	33 (68.8)	50.68 (6.74)	33.7	45.97	52.37	56.15	58.6		
		Week 70	Sparsentan	49	40 (81.6)	52.29 (7.02)	21.9	50.42	53.41	56.15	66.8		
			Irbesartan	48	32 (66.7)	50.79 (8.76)	23.5	47.65	53.48	57.07	59.6		
		Week 94	Sparsentan	49	41 (83.7)	51.67 (7.62)	27.9	45.88	54.26	57.35	63.5		
			Irbesartan	48	35 (72.9)	50.98 (6.51)	30.4	47.55	52.60	56.15	59.9		
	Week 110	Sparsentan	49	36 (73.5)	52.30 (6.30)	34.7	47.36	53.92	57.83	59.4			
		Irbesartan	48	32 (66.7)	51.75 (6.92)	32.1	47.17	53.34	57.67	60.9			
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	49	39 (79.6)	0.64 (6.99)	-24.8	-2.13	0.00	4.30	14.3	0.05 [-0.41, 0.51]
			Irbesartan	48	33 (68.8)	0.30 (6.11)	-15.6	-1.89	0.00	3.24	13.3		
	Week 48		Sparsentan	49	45 (91.8)	-1.08 (7.58)	-22.3	-3.76	-0.88	3.45	17.5	0.10 [-0.35, 0.55]	
			Irbesartan	48	33 (68.8)	-1.84 (7.76)	-20.8	-4.32	-1.06	1.61	16.4		
	Week 70		Sparsentan	49	40 (81.6)	-0.20 (7.76)	-30.2	-3.69	-0.31	3.88	15.8	0.36 [-0.11, 0.82]	
			Irbesartan	48	32 (66.7)	-2.97 (7.82)	-25.1	-7.96	-1.08	1.61	9.7		
	Week 94		Sparsentan	49	41 (83.7)	-0.79 (8.77)	-24.1	-3.31	0.56	3.49	15.9	0.09 [-0.36, 0.54]	
			Irbesartan	48	35 (72.9)	-1.58 (8.56)	-20.1	-6.39	-1.62	3.17	18.0		
Week 110	Sparsentan		49	36 (73.5)	-0.09 (7.28)	-17.4	-3.89	-0.60	3.85	16.1	0.14 [-0.34, 0.62]		
	Irbesartan		48	32 (66.7)	-1.13 (7.54)	-18.5	-5.19	-1.03	3.04	15.0			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	26	24 (92.3)	50.73 (11.70)	12.4	49.48	54.33	56.81	62.9		
			Irbesartan	25	24 (96.0)	52.47 (8.27)	21.6	51.67	55.35	56.15	61.1		
	Week 24	Sparsentan	26	21 (80.8)	52.48 (7.66)	35.4	51.29	56.15	57.49	59.7			
		Irbesartan	25	20 (80.0)	52.72 (8.49)	20.8	51.95	54.58	57.35	59.7			
	Week 48	Sparsentan	26	23 (88.5)	50.41 (6.97)	32.1	46.58	51.29	56.15	58.6			
		Irbesartan	25	18 (72.0)	54.09 (5.97)	40.3	53.17	56.28	58.02	60.8			
	Week 70	Sparsentan	26	22 (84.6)	48.12 (8.65)	31.6	42.16	50.41	56.22	58.1			
		Irbesartan	25	20 (80.0)	55.94 (5.29)	43.3	55.34	57.48	59.28	63.0			
	Week 94	Sparsentan	26	20 (76.9)	50.55 (8.35)	30.9	44.80	53.91	56.15	61.0			
		Irbesartan	25	19 (76.0)	55.52 (4.33)	47.4	51.84	56.44	57.49	62.7			
	Week 110	Sparsentan	26	21 (80.8)	49.45 (9.78)	22.8	46.04	52.38	57.20	59.7			
		Irbesartan	25	17 (68.0)	54.34 (4.69)	43.1	51.02	56.15	56.71	61.3			
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	26	21 (80.8)	0.68 (6.68)	-16.1	-2.65	0.98	2.98	14.1	0.15 [-0.47, 0.76]
			Irbesartan	25	20 (80.0)	-0.11 (3.39)	-7.0	-1.42	-0.16	0.96	10.0		
	Week 48		Sparsentan	26	23 (88.5)	0.19 (7.81)	-11.7	-4.37	-0.08	2.28	24.7	0.02 [-0.60, 0.63]	
			Irbesartan	25	18 (72.0)	0.07 (6.08)	-11.6	-2.98	1.33	2.42	13.1		
	Week 70		Sparsentan	26	22 (84.6)	-1.89 (11.57)	-25.7	-8.46	-2.63	3.78	31.3	-0.40 [-1.01, 0.21]	
			Irbesartan	25	20 (80.0)	1.69 (4.40)	-7.0	-0.57	1.48	3.63	11.2		
	Week 94		Sparsentan	26	20 (76.9)	-0.98 (6.39)	-15.9	-4.89	-0.64	3.17	9.8	-0.31 [-0.94, 0.32]	
			Irbesartan	25	19 (76.0)	0.60 (3.20)	-5.5	-2.02	0.81	3.01	6.3		
Week 110	Sparsentan		26	21 (80.8)	-2.35 (7.32)	-20.5	-5.65	-1.80	2.03	12.2	-0.50 [-1.15, 0.15]		
	Irbesartan		25	17 (68.0)	0.62 (3.66)	-5.7	-1.09	0.00	1.61	8.4			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024



Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 2													
< 45 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	82	73 (89.0)	49.80 (7.82)	29.2	44.56	51.81	55.95	60.8		
			Irbesartan	80	70 (87.5)	50.48 (6.75)	29.2	46.31	52.11	56.06	60.1		
		Sparsentan	82	64 (78.0)	50.57 (7.38)	16.3	46.16	52.64	56.15	60.9			
		Irbesartan	80	51 (63.8)	49.80 (6.61)	33.8	43.80	52.09	55.88	61.9			
	Week 24	Sparsentan	82	63 (76.8)	51.48 (7.52)	25.4	46.85	53.71	56.71	61.9			
		Irbesartan	80	45 (56.3)	50.17 (9.31)	18.0	47.10	52.50	55.86	64.5			
	Week 48	Sparsentan	82	65 (79.3)	50.39 (8.09)	28.0	44.11	53.17	56.62	61.1			
		Irbesartan	80	50 (62.5)	48.63 (8.90)	21.7	44.28	51.23	55.28	60.3			
	Week 70	Sparsentan	82	61 (74.4)	49.78 (7.93)	23.8	43.05	52.91	56.15	59.1			
		Irbesartan	80	48 (60.0)	49.65 (7.93)	20.0	46.20	52.23	55.67	58.8			
	Week 94	Sparsentan	82	57 (69.5)	50.75 (6.59)	34.1	47.39	52.50	55.33	63.1			
		Irbesartan	80	44 (55.0)	48.81 (8.17)	25.4	44.43	51.33	55.32	60.0			
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	82	64 (78.0)	0.42 (5.94)	-13.4	-3.24	-0.12	3.92	15.5	0.09 [-0.27, 0.46]
			Irbesartan	80	51 (63.8)	-0.13 (5.73)	-15.0	-2.38	0.00	3.57	16.9		
	Week 48		Sparsentan	82	63 (76.8)	1.48 (6.23)	-18.7	-2.43	0.00	4.71	20.6	0.22 [-0.17, 0.60]	
			Irbesartan	80	45 (56.3)	-0.30 (10.46)	-35.7	-4.31	-0.27	3.28	20.4		
	Week 70		Sparsentan	82	65 (79.3)	0.11 (6.71)	-20.6	-2.89	0.29	2.98	21.2	0.22 [-0.15, 0.59]	
			Irbesartan	80	50 (62.5)	-1.62 (8.97)	-28.0	-5.69	-0.82	3.64	17.8		
Week 94	Sparsentan	82	61 (74.4)	-0.45 (7.13)	-19.6	-3.24	-1.22	2.69	20.7	0.05 [-0.33, 0.42]			
	Irbesartan	80	48 (60.0)	-0.78 (6.99)	-18.6	-4.69	0.29	3.27	13.6				
Week 110	Sparsentan	82	57 (69.5)	0.22 (7.02)	-17.9	-2.69	0.00	3.91	16.4	0.31 [-0.09, 0.70]			
	Irbesartan	80	44 (55.0)	-2.06 (7.86)	-18.3	-7.95	-0.98	1.89	15.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
45 to < 60 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	45	44 (97.8)	51.30 (7.34)	25.8	46.86	52.52	56.15	63.5	
		Week 24	Irbesartan	49	45 (91.8)	52.71 (6.14)	27.9	50.34	53.18	56.71	62.0	
			Sparsentan	45	35 (77.8)	51.14 (7.54)	31.2	45.43	53.03	56.48	64.0	
		Week 48	Irbesartan	49	34 (69.4)	52.25 (7.65)	31.2	47.89	54.95	57.03	64.3	
			Sparsentan	45	34 (75.6)	51.92 (6.27)	33.2	49.81	53.19	55.09	66.0	
		Week 70	Irbesartan	49	31 (63.3)	53.46 (6.23)	39.0	49.95	54.80	57.76	63.4	
			Sparsentan	45	36 (80.0)	50.65 (7.27)	35.2	43.90	51.99	56.17	63.2	
		Week 94	Irbesartan	49	33 (67.3)	52.40 (5.78)	34.6	50.07	53.71	56.15	59.3	
			Sparsentan	45	33 (73.3)	51.90 (6.58)	34.1	49.92	53.17	56.15	61.3	
		Week 110	Irbesartan	49	31 (63.3)	53.13 (6.80)	29.2	52.50	54.53	56.96	61.3	
	Sparsentan		45	36 (80.0)	50.27 (8.48)	32.6	42.27	54.13	56.15	64.2		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	45	35 (77.8)	-0.06 (7.53)	-14.9	-4.25	0.27	2.88	20.4	0.08 [-0.40, 0.55]
			Irbesartan	49	34 (69.4)	-0.56 (5.51)	-20.0	-3.09	0.00	2.22	10.1	
		Week 48	Sparsentan	45	34 (75.6)	0.46 (6.45)	-13.9	-3.27	-0.31	4.84	17.0	-0.15 [-0.63, 0.34]
			Irbesartan	49	31 (63.3)	1.34 (5.47)	-11.3	0.00	2.10	4.04	12.5	
		Week 70	Sparsentan	45	36 (80.0)	-1.00 (6.43)	-16.7	-5.16	0.46	3.04	11.0	-0.10 [-0.57, 0.37]
			Irbesartan	49	33 (67.3)	-0.37 (6.02)	-24.4	-2.07	0.00	2.61	9.1	
		Week 94	Sparsentan	45	33 (73.3)	1.24 (6.06)	-12.6	-3.46	1.89	6.13	14.8	0.01 [-0.48, 0.50]
			Irbesartan	49	31 (63.3)	1.17 (4.07)	-7.0	-1.82	1.29	4.70	9.1	
		Week 110	Sparsentan	45	36 (80.0)	-1.33 (8.31)	-17.8	-7.16	0.00	4.97	15.6	-0.35 [-0.84, 0.15]
Irbesartan			49	28 (57.1)	1.14 (5.23)	-8.9	-2.57	1.75	4.59	11.3		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	49	47 (95.9)	53.00 (6.69)	37.4	48.78	53.79	57.76	61.8		
			Irbesartan	48	44 (91.7)	52.03 (7.49)	29.6	49.15	54.95	56.30	64.3		
	Week 24	Sparsentan	49	39 (79.6)	53.54 (6.86)	27.3	50.55	56.15	57.49	62.9			
		Irbesartan	48	33 (68.8)	53.37 (5.41)	34.9	52.40	54.53	56.15	60.8			
	Week 48	Sparsentan	49	45 (91.8)	51.72 (8.05)	29.8	46.56	53.83	57.49	66.1			
		Irbesartan	48	33 (68.8)	50.68 (6.74)	33.7	45.97	52.37	56.15	58.6			
	Week 70	Sparsentan	49	40 (81.6)	52.29 (7.02)	21.9	50.42	53.41	56.15	66.8			
		Irbesartan	48	32 (66.7)	50.79 (8.76)	23.5	47.65	53.48	57.07	59.6			
	Week 94	Sparsentan	49	41 (83.7)	51.67 (7.62)	27.9	45.88	54.26	57.35	63.5			
		Irbesartan	48	35 (72.9)	50.98 (6.51)	30.4	47.55	52.60	56.15	59.9			
	Week 110	Sparsentan	49	36 (73.5)	52.30 (6.30)	34.7	47.36	53.92	57.83	59.4			
		Irbesartan	48	32 (66.7)	51.75 (6.92)	32.1	47.17	53.34	57.67	60.9			
	KDQOL-SF12: change from baseline in PCS		Week 24	Sparsentan	49	39 (79.6)	0.64 (6.99)	-24.8	-2.13	0.00	4.30	14.3	0.05 [-0.41, 0.51]
				Irbesartan	48	33 (68.8)	0.30 (6.11)	-15.6	-1.89	0.00	3.24	13.3	
			Week 48	Sparsentan	49	45 (91.8)	-1.08 (7.58)	-22.3	-3.76	-0.88	3.45	17.5	0.10 [-0.35, 0.55]
				Irbesartan	48	33 (68.8)	-1.84 (7.76)	-20.8	-4.32	-1.06	1.61	16.4	
			Week 70	Sparsentan	49	40 (81.6)	-0.20 (7.76)	-30.2	-3.69	-0.31	3.88	15.8	0.36 [-0.11, 0.82]
				Irbesartan	48	32 (66.7)	-2.97 (7.82)	-25.1	-7.96	-1.08	1.61	9.7	
			Week 94	Sparsentan	49	41 (83.7)	-0.79 (8.77)	-24.1	-3.31	0.56	3.49	15.9	0.09 [-0.36, 0.54]
				Irbesartan	48	35 (72.9)	-1.58 (8.56)	-20.1	-6.39	-1.62	3.17	18.0	
Week 110			Sparsentan	49	36 (73.5)	-0.09 (7.28)	-17.4	-3.89	-0.60	3.85	16.1	0.14 [-0.34, 0.62]	
			Irbesartan	48	32 (66.7)	-1.13 (7.54)	-18.5	-5.19	-1.03	3.04	15.0		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL-SF12: PCS	Baseline	Sparsentan	26	24 (92.3)	50.73 (11.70)	12.4	49.48	54.33	56.81	62.9		
			Irbesartan	25	24 (96.0)	52.47 (8.27)	21.6	51.67	55.35	56.15	61.1		
	Week 24	Sparsentan	26	21 (80.8)	52.48 (7.66)	35.4	51.29	56.15	57.49	59.7			
		Irbesartan	25	20 (80.0)	52.72 (8.49)	20.8	51.95	54.58	57.35	59.7			
	Week 48	Sparsentan	26	23 (88.5)	50.41 (6.97)	32.1	46.58	51.29	56.15	58.6			
		Irbesartan	25	18 (72.0)	54.09 (5.97)	40.3	53.17	56.28	58.02	60.8			
	Week 70	Sparsentan	26	22 (84.6)	48.12 (8.65)	31.6	42.16	50.41	56.22	58.1			
		Irbesartan	25	20 (80.0)	55.94 (5.29)	43.3	55.34	57.48	59.28	63.0			
	Week 94	Sparsentan	26	20 (76.9)	50.55 (8.35)	30.9	44.80	53.91	56.15	61.0			
		Irbesartan	25	19 (76.0)	55.52 (4.33)	47.4	51.84	56.44	57.49	62.7			
	Week 110	Sparsentan	26	21 (80.8)	49.45 (9.78)	22.8	46.04	52.38	57.20	59.7			
		Irbesartan	25	17 (68.0)	54.34 (4.69)	43.1	51.02	56.15	56.71	61.3			
		KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	26	21 (80.8)	0.68 (6.68)	-16.1	-2.65	0.98	2.98	14.1	0.15 [-0.47, 0.76]
			Irbesartan	25	20 (80.0)	-0.11 (3.39)	-7.0	-1.42	-0.16	0.96	10.0		
	Week 48		Sparsentan	26	23 (88.5)	0.19 (7.81)	-11.7	-4.37	-0.08	2.28	24.7	0.02 [-0.60, 0.63]	
			Irbesartan	25	18 (72.0)	0.07 (6.08)	-11.6	-2.98	1.33	2.42	13.1		
	Week 70		Sparsentan	26	22 (84.6)	-1.89 (11.57)	-25.7	-8.46	-2.63	3.78	31.3	-0.40 [-1.01, 0.21]	
			Irbesartan	25	20 (80.0)	1.69 (4.40)	-7.0	-0.57	1.48	3.63	11.2		
	Week 94		Sparsentan	26	20 (76.9)	-0.98 (6.39)	-15.9	-4.89	-0.64	3.17	9.8	-0.31 [-0.94, 0.32]	
			Irbesartan	25	19 (76.0)	0.60 (3.20)	-5.5	-2.02	0.81	3.01	6.3		
Week 110	Sparsentan		26	21 (80.8)	-2.35 (7.32)	-20.5	-5.65	-1.80	2.03	12.2	-0.50 [-1.15, 0.15]		
	Irbesartan		25	17 (68.0)	0.62 (3.66)	-5.7	-1.09	0.00	1.61	8.4			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline urine protein excretion														
<= 1.75 g/day	KDQOL-SF12: PCS	Baseline	Sparsentan	98	93 (94.9)	50.35 (8.28)	25.5	46.10	52.40	56.74	62.3			
			Irbesartan	93	83 (89.2)	51.93 (6.71)	21.6	49.79	53.43	56.15	60.8			
		Week 24	Sparsentan	98	81 (82.7)	50.96 (7.37)	27.3	45.00	52.70	56.15	64.0			
			Irbesartan	93	59 (63.4)	52.35 (7.13)	20.8	50.61	54.51	56.71	60.8			
		Week 48	Sparsentan	98	83 (84.7)	51.41 (7.33)	29.8	46.85	53.39	56.15	66.1			
			Irbesartan	93	57 (61.3)	52.48 (6.32)	33.7	49.68	54.55	57.20	63.4			
		Week 70	Sparsentan	98	77 (78.6)	49.52 (8.26)	21.9	44.05	51.63	56.15	66.8			
			Irbesartan	93	62 (66.7)	52.24 (6.82)	30.0	48.45	53.59	57.22	63.0			
		Week 94	Sparsentan	98	79 (80.6)	49.89 (8.21)	27.9	41.68	52.91	56.15	63.5			
			Irbesartan	93	64 (68.8)	51.87 (6.69)	34.4	47.49	54.15	56.69	62.7			
		Week 110	Sparsentan	98	76 (77.6)	49.14 (8.38)	22.8	42.51	51.19	54.98	63.2			
			Irbesartan	93	57 (61.3)	51.27 (8.30)	25.4	44.94	53.68	57.76	62.6			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	98	81 (82.7)	0.82 (7.48)	-24.8	-3.69	0.00	4.90	20.4	0.16 [-0.18, 0.49]
		Irbesartan			93	59 (63.4)	-0.24 (5.70)	-20.0	-2.16	0.00	2.31	16.9		
Week 48	Sparsentan	98		83 (84.7)	0.88 (6.62)	-22.3	-2.38	0.60	4.84	17.5	0.15 [-0.18, 0.49]			
	Irbesartan	93		57 (61.3)	-0.14 (6.55)	-20.8	-2.24	0.29	2.42	16.4				
Week 70	Sparsentan	98		77 (78.6)	-0.97 (8.31)	-30.2	-4.50	-0.27	3.78	21.2	-0.03 [-0.36, 0.31]			
	Irbesartan	93		62 (66.7)	-0.75 (6.36)	-25.1	-3.77	0.14	2.69	15.2				
Week 94	Sparsentan	98		79 (80.6)	-0.10 (8.37)	-24.1	-4.03	0.19	5.11	20.7	0.06 [-0.27, 0.39]			
	Irbesartan	93		64 (68.8)	-0.57 (6.05)	-14.7	-3.75	0.35	2.71	18.0				
Week 110	Sparsentan	98		76 (77.6)	-1.42 (8.21)	-20.5	-6.74	-0.85	4.22	16.1	-0.04 [-0.38, 0.31]			
	Irbesartan	93		57 (61.3)	-1.14 (7.29)	-18.5	-5.71	0.00	2.15	15.5				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 1.75 g/day	KDQOL-SF12: PCS	Baseline	Sparsentan	104	95 (91.3)	51.77 (7.87)	12.4	48.84	53.50	56.68	63.5	
			Irbesartan	109	100 (91.7)	51.44 (7.29)	27.9	48.39	53.46	56.34	64.3	
	Week 24	Sparsentan	104	78 (75.0)	52.41 (7.32)	16.3	50.30	54.26	57.49	61.8		
		Irbesartan	109	79 (72.5)	51.18 (6.92)	31.2	45.96	53.46	56.15	64.3		
	Week 48	Sparsentan	104	82 (78.8)	51.56 (7.33)	25.4	47.13	53.46	57.22	62.1		
		Irbesartan	109	70 (64.2)	51.00 (8.57)	18.0	46.17	54.55	56.71	64.5		
	Week 70	Sparsentan	104	86 (82.7)	51.57 (7.21)	28.0	47.70	54.10	56.40	63.2		
		Irbesartan	109	73 (67.0)	50.21 (8.91)	21.7	46.63	53.46	56.15	60.3		
	Week 94	Sparsentan	104	76 (73.1)	51.81 (6.87)	23.8	49.11	54.02	56.49	61.7		
		Irbesartan	109	69 (63.3)	51.45 (7.52)	20.0	48.24	53.46	56.15	61.3		
	Week 110	Sparsentan	104	74 (71.2)	52.56 (6.06)	32.6	49.10	53.46	57.20	64.2		
		Irbesartan	109	64 (58.7)	51.25 (6.69)	28.1	48.22	52.78	56.13	62.0		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	104	78 (75.0)	-0.03 (5.59)	-16.1	-2.18	0.14	3.10	15.5	0.00 [-0.31, 0.32]
			Irbesartan	109	79 (72.5)	-0.05 (5.30)	-15.6	-2.69	0.00	3.22	13.3	
		Week 48	Sparsentan	104	82 (78.8)	-0.11 (7.18)	-18.7	-3.58	-0.58	3.45	24.7	0.03 [-0.29, 0.35]
			Irbesartan	109	70 (64.2)	-0.34 (9.35)	-35.7	-5.29	0.18	5.13	20.4	
		Week 70	Sparsentan	104	86 (82.7)	-0.05 (7.11)	-20.6	-2.77	0.17	2.90	31.3	0.18 [-0.13, 0.50]
			Irbesartan	109	73 (67.0)	-1.48 (8.48)	-28.0	-4.57	-0.29	2.61	17.8	
		Week 94	Sparsentan	104	76 (73.1)	-0.41 (6.01)	-20.0	-3.50	-0.52	3.32	13.7	-0.04 [-0.37, 0.28]
			Irbesartan	109	69 (63.3)	-0.12 (6.97)	-20.1	-3.49	0.00	4.04	15.4	
Week 110		Sparsentan	104	74 (71.2)	0.27 (6.50)	-17.9	-2.69	-0.04	3.64	16.4	0.09 [-0.25, 0.42]	
		Irbesartan	109	64 (58.7)	-0.30 (6.41)	-18.3	-3.47	-0.88	4.45	15.6		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline use of antihypertensives														
Yes	KDQOL-SF12: PCS	Baseline	Sparsentan	90	80 (88.9)	50.28 (8.52)	25.8	46.20	52.90	56.15	62.3			
			Irbesartan	88	77 (87.5)	50.39 (7.25)	27.9	46.31	51.28	55.86	64.3			
		Week 24	Sparsentan	90	68 (75.6)	51.03 (8.35)	16.3	45.53	52.84	57.08	64.0			
			Irbesartan	88	52 (59.1)	50.60 (6.50)	31.2	45.81	52.77	56.01	61.9			
		Week 48	Sparsentan	90	68 (75.6)	50.68 (8.14)	25.4	45.19	53.35	56.15	66.0			
			Irbesartan	88	48 (54.5)	50.84 (7.08)	32.8	46.16	51.91	56.15	64.5			
		Week 70	Sparsentan	90	69 (76.7)	49.51 (8.14)	28.0	42.96	51.57	56.15	63.2			
			Irbesartan	88	51 (58.0)	49.41 (8.02)	26.4	44.28	51.67	55.28	59.6			
		Week 94	Sparsentan	90	66 (73.3)	49.88 (7.62)	23.8	43.05	51.60	56.15	61.3			
			Irbesartan	88	57 (64.8)	50.26 (6.69)	29.2	47.89	52.79	54.53	58.6			
		Week 110	Sparsentan	90	64 (71.1)	50.89 (7.12)	32.6	44.82	52.25	56.22	63.2			
			Irbesartan	88	51 (58.0)	49.37 (8.28)	25.4	43.10	51.26	56.15	61.2			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	90	68 (75.6)	0.38 (7.03)	-16.1	-3.41	-0.64	3.89	20.4	0.03 [-0.33, 0.39]
		Irbesartan			88	52 (59.1)	0.17 (6.38)	-15.0	-2.93	0.00	3.50	16.9		
Week 48	Sparsentan	90		68 (75.6)	0.56 (7.03)	-15.4	-3.52	-0.03	4.78	20.6	-0.05 [-0.42, 0.32]			
	Irbesartan	88		48 (54.5)	0.93 (9.19)	-24.7	-4.07	-0.06	6.95	20.4				
Week 70	Sparsentan	90		69 (76.7)	-1.09 (6.82)	-20.6	-3.34	0.29	2.69	15.6	0.08 [-0.29, 0.44]			
	Irbesartan	88		51 (58.0)	-1.65 (8.27)	-28.0	-5.65	-0.29	2.44	15.2				
Week 94	Sparsentan	90		66 (73.3)	0.08 (7.73)	-19.6	-3.64	-0.23	4.99	20.7	-0.05 [-0.40, 0.31]			
	Irbesartan	88		57 (64.8)	0.42 (7.26)	-18.6	-3.49	1.06	4.87	18.0				
Week 110	Sparsentan	90		64 (71.1)	0.24 (7.37)	-16.4	-2.82	0.00	4.86	16.4	0.22 [-0.15, 0.59]			
	Irbesartan	88		51 (58.0)	-1.45 (8.12)	-18.5	-6.59	-1.01	3.77	15.5				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
No	KDQOL-SF12: PCS	Baseline	Sparsentan	112	108 (96.4)	51.65 (7.74)	12.4	47.39	53.45	56.86	63.5			
			Irbesartan	114	106 (93.0)	52.58 (6.73)	21.6	50.34	54.51	56.37	62.0			
		Week 24	Sparsentan	112	91 (81.3)	52.15 (6.54)	27.3	50.15	53.46	56.66	62.0			
			Irbesartan	114	86 (75.4)	52.33 (7.26)	20.8	51.35	54.66	56.22	64.3			
		Week 48	Sparsentan	112	97 (86.6)	52.05 (6.65)	29.8	48.31	53.46	56.44	66.1			
			Irbesartan	114	79 (69.3)	52.16 (7.98)	18.0	49.48	54.80	57.49	63.5			
		Week 70	Sparsentan	112	94 (83.9)	51.41 (7.43)	21.9	46.39	53.27	56.42	66.8			
			Irbesartan	114	84 (73.7)	52.20 (7.93)	21.7	48.73	54.80	57.20	63.0			
		Week 94	Sparsentan	112	89 (79.5)	51.54 (7.58)	27.9	48.14	54.53	56.56	63.5			
			Irbesartan	114	76 (66.7)	52.70 (7.28)	20.0	49.03	54.80	57.36	62.7			
		Week 110	Sparsentan	112	86 (76.8)	50.78 (7.81)	22.8	46.35	53.46	56.42	64.2			
			Irbesartan	114	70 (61.4)	52.64 (6.52)	28.1	49.25	53.91	57.55	62.6			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	112	91 (81.3)	0.42 (6.32)	-24.8	-3.25	0.29	4.08	14.3	0.13 [-0.17, 0.42]
		Irbesartan			114	86 (75.4)	-0.31 (4.84)	-20.0	-2.13	0.00	1.71	10.9		
		Week 48		Sparsentan	112	97 (86.6)	0.27 (6.85)	-22.3	-3.11	-0.08	3.45	24.7	0.17 [-0.12, 0.47]	
				Irbesartan	114	79 (69.3)	-0.97 (7.48)	-35.7	-4.22	0.54	2.22	18.9		
		Week 70		Sparsentan	112	94 (83.9)	-0.04 (8.28)	-30.2	-4.04	-0.23	3.73	31.3	0.10 [-0.19, 0.40]	
				Irbesartan	114	84 (73.7)	-0.84 (7.14)	-27.2	-3.31	0.00	2.65	17.8		
		Week 94		Sparsentan	112	89 (79.5)	-0.50 (6.97)	-24.1	-3.31	-0.14	3.46	15.9	0.06 [-0.24, 0.37]	
				Irbesartan	114	76 (66.7)	-0.91 (5.90)	-20.1	-3.63	0.00	2.57	13.6		
Week 110	Sparsentan	112		86 (76.8)	-1.20 (7.47)	-20.5	-4.78	-0.63	2.69	16.1	-0.16 [-0.47, 0.16]			
	Irbesartan	114		70 (61.4)	-0.15 (5.69)	-13.7	-4.22	0.00	3.46	15.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024



Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Time since renal biopsy														
<= 5 years	KDQOL-SF12: PCS	Baseline	Sparsentan	113	105 (92.9)	49.79 (7.95)	25.8	45.62	51.57	56.15	63.5			
			Irbesartan	127	118 (92.9)	51.56 (7.62)	21.6	48.83	53.95	56.42	64.3			
		Week 24	Sparsentan	113	90 (79.6)	51.43 (7.53)	16.3	47.50	53.04	56.48	64.0			
			Irbesartan	127	91 (71.7)	51.22 (7.54)	20.8	46.70	54.36	56.15	59.7			
		Week 48	Sparsentan	113	93 (82.3)	50.51 (7.28)	25.4	46.11	52.88	55.87	66.0			
			Irbesartan	127	86 (67.7)	51.71 (8.36)	18.0	47.21	54.72	57.02	64.5			
		Week 70	Sparsentan	113	91 (80.5)	50.10 (7.78)	28.0	44.11	52.40	56.15	63.2			
			Irbesartan	127	88 (69.3)	51.65 (8.42)	21.7	48.93	54.38	57.20	63.0			
		Week 94	Sparsentan	113	88 (77.9)	50.56 (7.67)	23.8	44.90	53.04	56.15	61.7			
			Irbesartan	127	88 (69.3)	51.87 (7.55)	20.0	49.16	54.15	56.53	62.7			
		Week 110	Sparsentan	113	88 (77.9)	50.96 (6.49)	35.4	46.19	52.37	56.03	64.2			
			Irbesartan	127	80 (63.0)	51.41 (7.84)	25.4	46.22	53.59	57.10	62.6			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	113	90 (79.6)	1.38 (6.75)	-16.1	-2.69	0.75	4.84	20.4	0.27 [-0.02, 0.56]
		Irbesartan			127	91 (71.7)	-0.34 (5.99)	-20.0	-2.77	-0.88	3.22	16.9		
Week 48	Sparsentan	113		93 (82.3)	0.57 (7.15)	-18.7	-3.28	-0.06	4.84	20.6	0.06 [-0.23, 0.36]			
	Irbesartan	127		86 (67.7)	0.06 (8.94)	-35.7	-4.22	0.93	4.27	20.4				
Week 70	Sparsentan	113		91 (80.5)	-0.08 (7.43)	-25.7	-3.10	0.27	3.78	21.2	0.07 [-0.23, 0.36]			
	Irbesartan	127		88 (69.3)	-0.59 (7.85)	-28.0	-3.31	0.00	3.57	17.8				
Week 94	Sparsentan	113		88 (77.9)	0.66 (7.52)	-19.6	-3.28	0.27	5.05	20.7	0.07 [-0.23, 0.37]			
	Irbesartan	127		88 (69.3)	0.16 (6.80)	-20.1	-3.48	0.89	4.27	18.0				
Week 110	Sparsentan	113		88 (77.9)	0.60 (7.59)	-20.5	-3.17	0.00	5.13	16.4	0.09 [-0.21, 0.40]			
	Irbesartan	127		80 (63.0)	-0.10 (7.32)	-18.5	-4.44	-0.01	4.57	15.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 5 years	KDQOL-SF12: PCS	Baseline	Sparsentan	89	83 (93.3)	52.69 (8.01)	12.4	49.93	54.80	57.25	62.3	
			Irbesartan	75	65 (86.7)	51.85 (5.81)	34.9	49.51	53.43	56.11	62.0	
		Week 24	Sparsentan	89	69 (77.5)	51.99 (7.18)	27.3	49.67	53.46	57.20	62.9	
			Irbesartan	75	47 (62.7)	52.58 (5.82)	39.0	47.89	53.46	57.05	64.3	
		Week 48	Sparsentan	89	72 (80.9)	52.75 (7.20)	29.8	50.07	53.89	57.49	66.1	
			Irbesartan	75	41 (54.7)	51.57 (6.00)	37.7	48.01	53.46	56.15	60.8	
		Week 70	Sparsentan	89	72 (80.9)	51.24 (7.76)	21.9	45.42	53.71	56.43	66.8	
			Irbesartan	75	47 (62.7)	50.21 (7.32)	22.3	46.26	52.11	56.13	59.7	
		Week 94	Sparsentan	89	67 (75.3)	51.20 (7.60)	27.9	46.35	53.46	56.95	63.5	
			Irbesartan	75	45 (60.0)	51.23 (6.22)	37.7	47.29	53.17	56.15	61.3	
	Week 110	Sparsentan	89	62 (69.7)	50.64 (8.79)	22.8	43.24	53.18	56.48	63.1		
		Irbesartan	75	41 (54.7)	50.97 (6.75)	36.1	46.91	51.90	56.48	62.0		
	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	89	69 (77.5)	-0.88 (6.25)	-24.8	-4.03	-0.27	2.69	14.1	-0.21 [-0.58, 0.16]
			Irbesartan	75	47 (62.7)	0.28 (4.25)	-12.7	-0.83	0.56	2.31	10.8	
		Week 48	Sparsentan	89	72 (80.9)	0.16 (6.61)	-22.3	-3.04	-0.29	3.49	24.7	0.16 [-0.22, 0.55]
			Irbesartan	75	41 (54.7)	-0.89 (6.36)	-12.5	-3.45	0.00	1.35	15.6	
		Week 70	Sparsentan	89	72 (80.9)	-0.99 (8.03)	-30.2	-4.89	-0.38	2.46	31.3	0.16 [-0.21, 0.52]
			Irbesartan	75	47 (62.7)	-2.18 (6.96)	-27.2	-4.76	-0.93	1.61	15.7	
		Week 94	Sparsentan	89	67 (75.3)	-1.45 (6.83)	-24.1	-3.53	-1.42	2.69	10.2	-0.02 [-0.40, 0.36]
			Irbesartan	75	45 (60.0)	-1.31 (5.90)	-14.7	-5.11	0.00	1.56	13.6	
Week 110		Sparsentan	89	62 (69.7)	-2.27 (6.93)	-17.9	-5.45	-1.18	2.13	8.6	-0.06 [-0.46, 0.33]	
		Irbesartan	75	41 (54.7)	-1.86 (5.63)	-17.3	-4.65	-0.81	1.61	9.7		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: History of hypertension														
Yes	KDQOL-SF12: PCS	Baseline	Sparsentan	155	142 (91.6)	50.13 (8.54)	12.4	45.76	52.34	56.15	62.3			
			Irbesartan	161	143 (88.8)	51.59 (6.91)	27.9	48.76	53.24	56.15	64.3			
		Week 24	Sparsentan	155	121 (78.1)	51.12 (7.51)	16.3	46.96	52.77	56.15	64.0			
			Irbesartan	161	106 (65.8)	51.43 (6.90)	31.2	46.58	53.70	56.15	64.3			
		Week 48	Sparsentan	155	124 (80.0)	50.83 (7.21)	25.4	46.57	53.18	56.06	66.0			
			Irbesartan	161	97 (60.2)	51.62 (7.94)	18.0	48.01	54.80	56.71	64.5			
		Week 70	Sparsentan	155	125 (80.6)	50.23 (7.43)	28.0	44.05	52.35	56.15	63.2			
			Irbesartan	161	104 (64.6)	50.18 (8.61)	21.7	46.86	52.35	56.15	63.0			
		Week 94	Sparsentan	155	118 (76.1)	50.36 (7.10)	23.8	44.96	52.56	56.15	61.7			
			Irbesartan	161	103 (64.0)	50.93 (7.33)	20.0	47.89	53.17	56.15	61.3			
		Week 110	Sparsentan	155	113 (72.9)	50.56 (7.52)	22.8	44.33	52.40	56.15	64.2			
			Irbesartan	161	92 (57.1)	50.89 (7.83)	25.4	45.91	52.79	56.68	62.6			
			KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	155	121 (78.1)	0.63 (6.50)	-16.1	-3.24	0.21	4.07	20.4	0.15 [-0.11, 0.41]
		Irbesartan			161	106 (65.8)	-0.28 (5.84)	-20.0	-2.69	0.00	2.42	16.9		
Week 48	Sparsentan	155		124 (80.0)	0.59 (6.93)	-18.7	-3.56	-0.14	4.12	24.7	0.08 [-0.19, 0.35]			
	Irbesartan	161		97 (60.2)	-0.02 (8.54)	-35.7	-3.90	0.08	3.04	20.4				
Week 70	Sparsentan	155		125 (80.6)	-0.14 (7.13)	-20.4	-3.80	0.08	3.18	31.3	0.21 [-0.05, 0.47]			
	Irbesartan	161		104 (64.6)	-1.76 (8.11)	-28.0	-5.77	-0.28	2.43	17.8				
Week 94	Sparsentan	155		118 (76.1)	-0.03 (7.01)	-20.0	-3.31	-0.23	4.51	20.7	0.07 [-0.20, 0.33]			
	Irbesartan	161		103 (64.0)	-0.51 (6.85)	-20.1	-4.03	0.00	3.17	18.0				
Week 110	Sparsentan	155		113 (72.9)	-0.20 (7.43)	-17.9	-3.20	0.00	4.53	16.4	0.04 [-0.23, 0.32]			
	Irbesartan	161		92 (57.1)	-0.50 (7.14)	-18.5	-4.20	-0.01	3.99	15.6				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSHM: KDQOL-SF12: Course of PCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
No	KDQOL-SF12: PCS	Baseline	Sparsentan	47	46 (97.9)	53.97 (5.64)	37.4	51.80	54.80	57.49	63.5			
			Irbesartan	41	40 (97.6)	51.92 (7.46)	21.6	49.37	54.25	56.26	60.1			
		Week 24	Sparsentan	47	38 (80.9)	53.44 (6.65)	27.3	51.61	56.15	57.49	60.9			
			Irbesartan	41	32 (78.0)	52.50 (7.41)	20.8	51.26	54.58	56.44	59.7			
		Week 48	Sparsentan	47	41 (87.2)	53.48 (7.33)	29.8	50.07	55.09	57.76	66.1			
			Irbesartan	41	30 (73.2)	51.81 (6.77)	37.9	47.87	53.44	56.71	63.5			
		Week 70	Sparsentan	47	38 (80.9)	51.85 (8.79)	21.9	47.79	54.21	56.62	66.8			
			Irbesartan	41	31 (75.6)	54.38 (4.59)	42.8	52.43	55.88	57.22	60.8			
		Week 94	Sparsentan	47	37 (78.7)	52.33 (9.03)	27.9	51.57	55.09	57.49	63.5			
			Irbesartan	41	30 (73.2)	54.14 (5.74)	40.6	52.79	55.79	57.70	62.7			
		Week 110	Sparsentan	47	37 (78.7)	51.62 (7.47)	34.7	49.73	53.50	56.71	59.7			
			Irbesartan	41	29 (70.7)	52.43 (6.13)	39.8	49.31	53.22	56.71	62.0			
		KDQOL-SF12: change from baseline in PCS	KDQOL-SF12: change from baseline in PCS	Week 24	Sparsentan	47	38 (80.9)	-0.32 (6.99)	-24.8	-3.69	-0.13	2.70	14.3	-0.12 [-0.59, 0.35]
					Irbesartan	41	32 (78.0)	0.36 (3.94)	-8.9	-1.83	0.00	2.03	10.4	
				Week 48	Sparsentan	47	41 (87.2)	-0.22 (6.87)	-22.3	-2.69	-0.04	2.28	15.6	0.11 [-0.36, 0.58]
					Irbesartan	41	30 (73.2)	-0.98 (6.99)	-20.8	-4.66	0.55	2.22	15.6	
				Week 70	Sparsentan	47	38 (80.9)	-1.62 (9.31)	-30.2	-4.04	-0.11	3.17	15.6	-0.33 [-0.81, 0.15]
					Irbesartan	41	31 (75.6)	0.90 (4.88)	-14.2	-1.27	1.06	3.78	15.7	
				Week 94	Sparsentan	47	37 (78.7)	-0.95 (8.16)	-24.1	-3.87	-0.14	3.46	12.9	-0.17 [-0.65, 0.31]
					Irbesartan	41	30 (73.2)	0.26 (5.33)	-14.7	-3.00	1.45	3.53	7.8	
Week 110	Sparsentan			47	37 (78.7)	-1.77 (7.45)	-20.5	-5.45	-1.34	2.03	14.6	-0.07 [-0.55, 0.42]		
	Irbesartan			41	29 (70.7)	-1.31 (5.76)	-13.7	-5.13	-1.06	1.61	11.3			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.

95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

PCS = physical component summary.

Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.833
Male	Week 24	Sparsentan	139	110 (79.1)	0.65 (0.58)	(-0.49, 1.80)	0.96 (0.84)	(-0.70, 2.61)	0.258
		Irbesartan	143	100 (69.9)	-0.30 (0.61)	(-1.50, 0.90)			
	Week 48	Sparsentan	139	115 (82.7)	0.24 (0.57)	(-0.88, 1.36)	-0.07 (0.85)	(-1.74, 1.60)	0.938
		Irbesartan	143	90 (62.9)	0.30 (0.63)	(-0.93, 1.54)			
	Week 70	Sparsentan	139	117 (84.2)	-0.35 (0.57)	(-1.47, 0.76)	0.31 (0.84)	(-1.34, 1.96)	0.711
		Irbesartan	143	97 (67.8)	-0.66 (0.62)	(-1.88, 0.55)			
	Week 94	Sparsentan	139	108 (77.7)	-0.13 (0.59)	(-1.28, 1.02)	0.00 (0.85)	(-1.67, 1.67)	1.000
		Irbesartan	143	99 (69.2)	-0.13 (0.62)	(-1.34, 1.08)			
	Week 110	Sparsentan	139	104 (74.8)	-0.18 (0.60)	(-1.36, 1.00)	-0.22 (0.88)	(-1.95, 1.52)	0.807
		Irbesartan	143	89 (62.2)	0.03 (0.65)	(-1.24, 1.30)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Female	Week 24	Sparsentan	63	49 (77.8)	-0.17 (0.92)	(-1.99, 1.64)	-0.83 (1.40)	(-3.59, 1.92)	0.552
		Irbesartan	59	38 (64.4)	0.66 (1.05)	(-1.41, 2.73)			
	Week 48	Sparsentan	63	50 (79.4)	-0.35 (0.91)	(-2.14, 1.45)	0.55 (1.39)	(-2.19, 3.30)	0.692
		Irbesartan	59	37 (62.7)	-0.90 (1.05)	(-2.97, 1.17)			
	Week 70	Sparsentan	63	46 (73.0)	-1.13 (0.94)	(-2.98, 0.73)	0.44 (1.41)	(-2.35, 3.22)	0.758
		Irbesartan	59	38 (64.4)	-1.56 (1.05)	(-3.62, 0.50)			
	Week 94	Sparsentan	63	47 (74.6)	-1.16 (0.94)	(-3.01, 0.69)	-0.23 (1.45)	(-3.07, 2.62)	0.874
		Irbesartan	59	34 (57.6)	-0.93 (1.10)	(-3.09, 1.22)			
	Week 110	Sparsentan	63	46 (73.0)	-1.51 (0.95)	(-3.39, 0.37)	0.14 (1.49)	(-2.79, 3.07)	0.924
		Irbesartan	59	32 (54.2)	-1.65 (1.14)	(-3.89, 0.59)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.463
<= 45 years	Week 24	Sparsentan	96	72 (75.0)	0.49 (0.71)	(-0.91, 1.88)	0.66 (1.01)	(-1.33, 2.65)	0.514
		Irbesartan	99	70 (70.7)	-0.17 (0.72)	(-1.59, 1.24)			
	Week 48	Sparsentan	96	76 (79.2)	-0.10 (0.69)	(-1.46, 1.26)	0.86 (1.02)	(-1.14, 2.87)	0.398
		Irbesartan	99	62 (62.6)	-0.96 (0.75)	(-2.44, 0.51)			
	Week 70	Sparsentan	96	75 (78.1)	-0.78 (0.70)	(-2.14, 0.59)	-0.09 (1.02)	(-2.10, 1.92)	0.930
		Irbesartan	99	64 (64.6)	-0.69 (0.75)	(-2.16, 0.78)			
	Week 94	Sparsentan	96	70 (72.9)	-0.38 (0.72)	(-1.79, 1.02)	1.06 (1.04)	(-0.98, 3.10)	0.307
		Irbesartan	99	64 (64.6)	-1.44 (0.75)	(-2.92, 0.03)			
	Week 110	Sparsentan	96	70 (72.9)	-0.58 (0.72)	(-2.00, 0.84)	-0.04 (1.07)	(-2.13, 2.05)	0.971
		Irbesartan	99	59 (59.6)	-0.54 (0.78)	(-2.08, 1.00)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 45 years	Week 24	Sparsentan	106	87 (82.1)	0.25 (0.68)	(-1.09, 1.58)	0.13 (1.03)	(-1.89, 2.16)	0.897
		Irbesartan	103	68 (66.0)	0.11 (0.77)	(-1.41, 1.63)			
	Week 48	Sparsentan	106	89 (84.0)	0.13 (0.67)	(-1.19, 1.46)	-0.68 (1.03)	(-2.71, 1.34)	
		Irbesartan	103	65 (63.1)	0.82 (0.78)	(-0.71, 2.34)			
	Week 70	Sparsentan	106	88 (83.0)	-0.46 (0.68)	(-1.80, 0.87)	0.66 (1.02)	(-1.34, 2.66)	
		Irbesartan	103	71 (68.9)	-1.13 (0.76)	(-2.61, 0.36)			
	Week 94	Sparsentan	106	85 (80.2)	-0.58 (0.69)	(-1.94, 0.78)	-1.26 (1.03)	(-3.29, 0.77)	
		Irbesartan	103	69 (67.0)	0.68 (0.77)	(-0.83, 2.19)			
	Week 110	Sparsentan	106	80 (75.5)	-0.72 (0.71)	(-2.12, 0.68)	-0.51 (1.08)	(-2.63, 1.61)	
		Irbesartan	103	62 (60.2)	-0.21 (0.81)	(-1.79, 1.38)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
 Source Data: aqs, created on: 05MAR2024



Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.797
<= 18 years	Week 24	Sparsentan	9	5 (55.6)	5.12 (2.18)	(0.64, 9.61)	4.69 (3.18)	(-1.93, 11.30)	0.156
		Irbesartan	5	5 (100.0)	0.44 (2.23)	(-4.23, 5.10)			
	Week 48	Sparsentan	9	7 (77.8)	2.07 (1.87)	(-1.81, 5.95)	-2.48 (3.30)	(-9.28, 4.32)	0.459
		Irbesartan	5	3 (60.0)	4.55 (2.68)	(-0.95, 10.05)			
	Week 70	Sparsentan	9	7 (77.8)	2.68 (1.89)	(-1.26, 6.61)	2.19 (3.21)	(-4.49, 8.87)	0.503
		Irbesartan	5	4 (80.0)	0.49 (2.50)	(-4.70, 5.68)			
	Week 94	Sparsentan	9	5 (55.6)	-4.53 (2.16)	(-8.98, -0.08)	-5.03 (3.42)	(-12.13, 2.08)	0.157
		Irbesartan	5	4 (80.0)	0.50 (2.55)	(-4.81, 5.81)			
	Week 110	Sparsentan	9	4 (44.4)	-0.07 (2.45)	(-5.09, 4.96)	-0.36 (4.17)	(-8.92, 8.20)	0.932
		Irbesartan	5	2 (40.0)	0.29 (3.36)	(-6.59, 7.18)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
> 18 to 40 years	Week 24	Sparsentan	102	80 (78.4)	0.28 (0.68)	(-1.04, 1.61)	0.53 (0.98)	(-1.39, 2.45)	0.588	
		Irbesartan	109	73 (67.0)	-0.25 (0.71)	(-1.63, 1.14)				
	Week 48	Sparsentan	102	83 (81.4)	0.05 (0.66)	(-1.25, 1.36)	0.85 (0.98)	(-1.06, 2.77)	0.383	
		Irbesartan	109	69 (63.3)	-0.80 (0.72)	(-2.21, 0.61)				
	Week 70	Sparsentan	102	80 (78.4)	-1.01 (0.67)	(-2.33, 0.31)	0.05 (0.98)	(-1.88, 1.98)	0.957	
		Irbesartan	109	71 (65.1)	-1.06 (0.71)	(-2.46, 0.34)				
	Week 94	Sparsentan	102	76 (74.5)	-0.15 (0.69)	(-1.51, 1.20)	1.32 (1.00)	(-0.64, 3.29)	0.187	
		Irbesartan	109	69 (63.3)	-1.47 (0.72)	(-2.89, -0.05)				
	Week 110	Sparsentan	102	73 (71.6)	-0.77 (0.71)	(-2.16, 0.62)	-0.28 (1.03)	(-2.30, 1.74)	0.786	
		Irbesartan	109	65 (59.6)	-0.49 (0.75)	(-1.96, 0.98)				
> 40 years	Week 24	Sparsentan	91	74 (81.3)	0.20 (0.76)	(-1.29, 1.69)	0.10 (1.13)	(-2.13, 2.33)	0.929	
		Irbesartan	88	60 (68.2)	0.10 (0.84)	(-1.55, 1.75)				
	Week 48	Sparsentan	91	75 (82.4)	-0.13 (0.75)	(-1.60, 1.34)	-0.71 (1.14)	(-2.95, 1.53)	0.533	
		Irbesartan	88	55 (62.5)	0.58 (0.86)	(-1.11, 2.27)				
	Week 70	Sparsentan	91	76 (83.5)	-0.44 (0.75)	(-1.91, 1.03)	0.42 (1.13)	(-1.79, 2.64)	0.706	
		Irbesartan	88	60 (68.2)	-0.86 (0.84)	(-2.51, 0.79)				
	Week 94	Sparsentan	91	74 (81.3)	-0.51 (0.76)	(-2.00, 0.98)	-1.42 (1.13)	(-3.65, 0.81)	0.212	
		Irbesartan	88	60 (68.2)	0.91 (0.84)	(-0.75, 2.56)				
	Week 110	Sparsentan	91	73 (80.2)	-0.40 (0.77)	(-1.91, 1.10)	-0.08 (1.17)	(-2.38, 2.22)	0.945	
		Irbesartan	88	54 (61.4)	-0.32 (0.88)	(-2.06, 1.42)				

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.237
North America	Week 24	Sparsentan	35	23 (65.7)	0.05 (1.21)	(-2.34, 2.44)	0.24 (1.56)	(-2.84, 3.33)	0.878
		Irbesartan	46	37 (80.4)	-0.19 (0.97)	(-2.11, 1.72)			
	Week 48	Sparsentan	35	25 (71.4)	0.79 (1.16)	(-1.50, 3.08)	0.14 (1.54)	(-2.90, 3.19)	0.926
		Irbesartan	46	33 (71.7)	0.64 (1.01)	(-1.34, 2.63)			
	Week 70	Sparsentan	35	23 (65.7)	0.64 (1.19)	(-1.72, 2.99)	-0.24 (1.58)	(-3.36, 2.88)	0.877
		Irbesartan	46	31 (67.4)	0.88 (1.03)	(-1.16, 2.92)			
	Week 94	Sparsentan	35	24 (68.6)	1.63 (1.19)	(-0.72, 3.98)	1.46 (1.60)	(-1.69, 4.62)	0.362
		Irbesartan	46	30 (65.2)	0.16 (1.06)	(-1.92, 2.25)			
	Week 110	Sparsentan	35	21 (60.0)	-0.73 (1.27)	(-3.23, 1.78)	-1.73 (1.66)	(-5.01, 1.55)	0.299
		Irbesartan	46	31 (67.4)	1.01 (1.06)	(-1.09, 3.10)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Europe	Week 24	Sparsentan	98	75 (76.5)	0.95 (0.72)	(-0.45, 2.36)	1.61 (1.06)	(-0.46, 3.69)	0.127
		Irbesartan	115	65 (56.5)	-0.66 (0.77)	(-2.18, 0.86)			
	Week 48	Sparsentan	98	77 (78.6)	0.11 (0.71)	(-1.29, 1.50)	0.62 (1.06)	(-1.46, 2.69)	0.561
		Irbesartan	115	61 (53.0)	-0.51 (0.78)	(-2.05, 1.03)			
	Week 70	Sparsentan	98	76 (77.6)	-0.64 (0.72)	(-2.05, 0.76)	1.78 (1.03)	(-0.25, 3.80)	0.085
		Irbesartan	115	72 (62.6)	-2.42 (0.74)	(-3.87, -0.97)			
	Week 94	Sparsentan	98	71 (72.4)	-0.73 (0.74)	(-2.18, 0.72)	0.28 (1.04)	(-1.77, 2.33)	0.788
		Irbesartan	115	72 (62.6)	-1.01 (0.74)	(-2.45, 0.44)			
	Week 110	Sparsentan	98	68 (69.4)	0.29 (0.76)	(-1.19, 1.78)	1.56 (1.10)	(-0.59, 3.72)	0.155
		Irbesartan	115	61 (53.0)	-1.27 (0.79)	(-2.83, 0.29)			
Asia Pacific	Week 24	Sparsentan	69	61 (88.4)	-0.02 (0.83)	(-1.65, 1.60)	-0.67 (1.36)	(-3.34, 2.00)	0.620
		Irbesartan	41	36 (87.8)	0.65 (1.07)	(-1.46, 2.76)			
	Week 48	Sparsentan	69	63 (91.3)	-0.12 (0.82)	(-1.73, 1.48)	0.31 (1.38)	(-2.39, 3.02)	0.819
		Irbesartan	41	33 (80.5)	-0.44 (1.10)	(-2.61, 1.74)			
	Week 70	Sparsentan	69	64 (92.8)	-0.82 (0.81)	(-2.42, 0.78)	-0.85 (1.40)	(-3.60, 1.89)	0.541
		Irbesartan	41	32 (78.0)	0.03 (1.13)	(-2.19, 2.26)			
	Week 94	Sparsentan	69	60 (87.0)	-0.72 (0.83)	(-2.35, 0.91)	-0.65 (1.42)	(-3.44, 2.15)	0.649
		Irbesartan	41	31 (75.6)	-0.08 (1.15)	(-2.34, 2.19)			
	Week 110	Sparsentan	69	61 (88.4)	-1.35 (0.83)	(-2.98, 0.29)	-0.55 (1.45)	(-3.41, 2.31)	0.704
		Irbesartan	41	29 (70.7)	-0.79 (1.19)	(-3.14, 1.55)			

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
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Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Baseline BMI	Overall		Sparsentan							Interaction: 0.910
< 27 kg/m**2	Week 24		Sparsentan	83	66 (79.5)	0.35 (0.74)	(-1.10, 1.81)	0.64 (1.04)	(-1.40, 2.68)	0.540
			Irbesartan	94	69 (73.4)	-0.28 (0.73)	(-1.71, 1.14)			
	Week 48		Sparsentan	83	68 (81.9)	-0.33 (0.73)	(-1.76, 1.11)	0.83 (1.06)	(-1.25, 2.90)	0.435
			Irbesartan	94	60 (63.8)	-1.15 (0.76)	(-2.65, 0.34)			
	Week 70		Sparsentan	83	66 (79.5)	-1.47 (0.74)	(-2.93, -0.02)	0.03 (1.06)	(-2.06, 2.11)	0.980
			Irbesartan	94	62 (66.0)	-1.50 (0.76)	(-2.99, -0.01)			
	Week 94		Sparsentan	83	64 (77.1)	-1.06 (0.75)	(-2.53, 0.42)	-0.28 (1.06)	(-2.37, 1.81)	0.791
			Irbesartan	94	64 (68.1)	-0.78 (0.75)	(-2.25, 0.70)			
	Week 110		Sparsentan	83	63 (75.9)	-1.39 (0.76)	(-2.88, 0.10)	-0.98 (1.10)	(-3.13, 1.18)	0.373
			Irbesartan	94	57 (60.6)	-0.41 (0.79)	(-1.97, 1.15)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
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Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 27 kg/m**2	Week 24	Sparsentan	119	93 (78.2)	0.37 (0.66)	(-0.93, 1.67)	0.13 (1.01)	(-1.86, 2.12)	0.896
		Irbesartan	107	69 (64.5)	0.24 (0.77)	(-1.27, 1.75)			
	Week 48	Sparsentan	119	97 (81.5)	0.30 (0.65)	(-0.97, 1.57)	-0.63 (1.00)	(-2.60, 1.34)	0.531
		Irbesartan	107	67 (62.6)	0.93 (0.77)	(-0.57, 2.43)			
	Week 70	Sparsentan	119	97 (81.5)	0.01 (0.65)	(-1.26, 1.29)	0.31 (0.99)	(-1.63, 2.26)	0.751
		Irbesartan	107	73 (68.2)	-0.30 (0.75)	(-1.77, 1.17)			
	Week 94	Sparsentan	119	91 (76.5)	-0.04 (0.67)	(-1.35, 1.26)	-0.25 (1.02)	(-2.25, 1.75)	0.807
		Irbesartan	107	68 (63.6)	0.20 (0.77)	(-1.30, 1.71)			
	Week 110	Sparsentan	119	87 (73.1)	-0.06 (0.68)	(-1.40, 1.29)	0.05 (1.05)	(-2.02, 2.12)	0.964
		Irbesartan	107	63 (58.9)	-0.10 (0.80)	(-1.68, 1.47)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan						Interaction:	0.761
eGFR Low and UP High	Week 24	Sparsentan	71	54 (76.1)	-0.08 (0.90)	(-1.84, 1.69)	0.92 (1.34)	(-1.70, 3.55)	0.490
		Irbesartan	74	44 (59.5)	-1.00 (0.99)	(-2.94, 0.95)			
	Week 48	Sparsentan	71	53 (74.6)	0.11 (0.89)	(-1.65, 1.87)	0.53 (1.36)	(-2.15, 3.21)	0.697
		Irbesartan	74	38 (51.4)	-0.42 (1.03)	(-2.44, 1.59)			
	Week 70	Sparsentan	71	58 (81.7)	-1.05 (0.87)	(-2.77, 0.66)	0.11 (1.34)	(-2.52, 2.75)	0.932
		Irbesartan	74	41 (55.4)	-1.17 (1.01)	(-3.16, 0.83)			
	Week 94	Sparsentan	71	52 (73.2)	-0.55 (0.90)	(-2.33, 1.23)	0.07 (1.39)	(-2.66, 2.80)	0.962
		Irbesartan	74	38 (51.4)	-0.62 (1.05)	(-2.68, 1.45)			
	Week 110	Sparsentan	71	53 (74.6)	-0.76 (0.91)	(-2.55, 1.02)	-0.39 (1.42)	(-3.18, 2.40)	0.784
		Irbesartan	74	36 (48.6)	-0.37 (1.09)	(-2.52, 1.77)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
eGFR Low and UP Low	Week 24	Sparsentan	55	44 (80.0)	0.64 (0.92)	(-1.18, 2.46)	0.54 (1.36)	(-2.13, 3.22)	0.688
		Irbesartan	55	39 (70.9)	0.09 (0.98)	(-1.85, 2.03)			
	Week 48	Sparsentan	55	43 (78.2)	0.69 (0.93)	(-1.13, 2.52)	0.22 (1.37)	(-2.48, 2.91)	0.875
		Irbesartan	55	36 (65.5)	0.48 (1.00)	(-1.50, 2.45)			
	Week 70	Sparsentan	55	42 (76.4)	0.46 (0.94)	(-1.39, 2.31)	1.79 (1.35)	(-0.87, 4.46)	0.186
		Irbesartan	55	40 (72.7)	-1.33 (0.97)	(-3.23, 0.57)			
	Week 94	Sparsentan	55	42 (76.4)	0.56 (0.95)	(-1.30, 2.42)	0.56 (1.36)	(-2.12, 3.23)	0.683
		Irbesartan	55	40 (72.7)	0.00 (0.97)	(-1.91, 1.91)			
	Week 110	Sparsentan	55	38 (69.1)	-0.10 (0.99)	(-2.04, 1.85)	0.14 (1.44)	(-2.69, 2.98)	0.920
		Irbesartan	55	34 (61.8)	-0.24 (1.04)	(-2.29, 1.81)			
eGFR High and UP High	Week 24	Sparsentan	37	27 (73.0)	0.69 (1.11)	(-1.50, 2.89)	-0.30 (1.61)	(-3.48, 2.87)	0.851
		Irbesartan	36	26 (72.2)	1.00 (1.16)	(-1.30, 3.29)			
	Week 48	Sparsentan	37	34 (91.9)	-0.54 (1.02)	(-2.56, 1.48)	-0.17 (1.56)	(-3.24, 2.90)	0.912
		Irbesartan	36	25 (69.4)	-0.37 (1.17)	(-2.68, 1.94)			
	Week 70	Sparsentan	37	31 (83.8)	0.43 (1.06)	(-1.66, 2.52)	0.87 (1.58)	(-2.25, 4.00)	0.582
		Irbesartan	36	25 (69.4)	-0.44 (1.18)	(-2.76, 1.88)			
	Week 94	Sparsentan	37	27 (73.0)	-1.00 (1.12)	(-3.20, 1.21)	-0.86 (1.62)	(-4.06, 2.33)	0.594
		Irbesartan	36	25 (69.4)	-0.13 (1.17)	(-2.45, 2.19)			
	Week 110	Sparsentan	37	26 (70.3)	1.40 (1.15)	(-0.87, 3.67)	2.30 (1.69)	(-1.04, 5.64)	0.176
		Irbesartan	36	22 (61.1)	-0.90 (1.24)	(-3.34, 1.54)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024



Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
eGFR High and UP Low	Week 24	Sparsentan	39	34 (87.2)	0.76 (1.07)	(-1.36, 2.87)	0.58 (1.58)	(-2.53, 3.68)	0.715
		Irbesartan	37	29 (78.4)	0.18 (1.15)	(-2.09, 2.45)			
	Week 48	Sparsentan	39	35 (89.7)	-0.03 (1.06)	(-2.12, 2.06)	-0.21 (1.58)	(-3.32, 2.91)	
		Irbesartan	37	28 (75.7)	0.18 (1.17)	(-2.12, 2.48)			
	Week 70	Sparsentan	39	32 (82.1)	-1.98 (1.10)	(-4.15, 0.18)	-1.58 (1.60)	(-4.74, 1.58)	
		Irbesartan	37	29 (78.4)	-0.40 (1.16)	(-2.69, 1.88)			
	Week 94	Sparsentan	39	34 (87.2)	-0.97 (1.07)	(-3.09, 1.15)	-0.34 (1.57)	(-3.44, 2.76)	
		Irbesartan	37	30 (81.1)	-0.63 (1.14)	(-2.88, 1.62)			
	Week 110	Sparsentan	39	33 (84.6)	-2.54 (1.09)	(-4.69, -0.39)	-2.15 (1.60)	(-5.30, 0.99)	
		Irbesartan	37	29 (78.4)	-0.39 (1.16)	(-2.67, 1.90)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.107
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	99 (78.0)	0.21 (0.63)	(-1.03, 1.45)	0.59 (0.93)	(-1.23, 2.41)	0.524
		Irbesartan	129	85 (65.9)	-0.38 (0.68)	(-1.72, 0.95)			
	Week 48	Sparsentan	127	97 (76.4)	0.75 (0.63)	(-0.49, 1.98)	0.26 (0.94)	(-1.59, 2.12)	0.780
		Irbesartan	129	76 (58.9)	0.48 (0.70)	(-0.89, 1.86)			
	Week 70	Sparsentan	127	101 (79.5)	-0.31 (0.62)	(-1.53, 0.92)	0.95 (0.93)	(-0.88, 2.77)	0.308
		Irbesartan	129	83 (64.3)	-1.25 (0.68)	(-2.60, 0.09)			
	Week 94	Sparsentan	127	94 (74.0)	-0.17 (0.64)	(-1.43, 1.09)	-0.03 (0.95)	(-1.89, 1.84)	0.978
		Irbesartan	129	79 (61.2)	-0.14 (0.70)	(-1.51, 1.23)			
	Week 110	Sparsentan	127	93 (73.2)	-0.30 (0.65)	(-1.58, 0.98)	0.16 (0.98)	(-1.77, 2.09)	0.869
		Irbesartan	129	72 (55.8)	-0.46 (0.73)	(-1.91, 0.98)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	39 (79.6)	0.83 (1.01)	(-1.17, 2.82)	0.03 (1.50)	(-2.93, 2.99)	0.986
		Irbesartan	48	33 (68.8)	0.80 (1.11)	(-1.38, 2.98)			
	Week 48	Sparsentan	49	45 (91.8)	-0.94 (0.96)	(-2.83, 0.96)	0.69 (1.46)	(-2.19, 3.57)	0.637
		Irbesartan	48	33 (68.8)	-1.63 (1.10)	(-3.78, 0.53)			
	Week 70	Sparsentan	49	40 (81.6)	0.13 (1.00)	(-1.83, 2.10)	2.10 (1.49)	(-0.84, 5.04)	0.161
		Irbesartan	48	32 (66.7)	-1.97 (1.11)	(-4.15, 0.21)			
	Week 94	Sparsentan	49	41 (83.7)	-0.79 (1.00)	(-2.76, 1.18)	1.16 (1.47)	(-1.75, 4.06)	0.434
		Irbesartan	48	35 (72.9)	-1.95 (1.08)	(-4.07, 0.18)			
	Week 110	Sparsentan	49	36 (73.5)	-0.38 (1.06)	(-2.46, 1.70)	0.76 (1.54)	(-2.27, 3.79)	0.622
		Irbesartan	48	32 (66.7)	-1.14 (1.12)	(-3.34, 1.06)			
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	21 (80.8)	0.54 (1.23)	(-1.88, 2.96)	0.38 (1.76)	(-3.09, 3.84)	0.831
		Irbesartan	25	20 (80.0)	0.16 (1.26)	(-2.32, 2.64)			
	Week 48	Sparsentan	26	23 (88.5)	-0.77 (1.18)	(-3.10, 1.56)	-1.55 (1.78)	(-5.06, 1.96)	0.384
		Irbesartan	25	18 (72.0)	0.78 (1.32)	(-1.82, 3.39)			
	Week 70	Sparsentan	26	22 (84.6)	-2.90 (1.21)	(-5.28, -0.52)	-5.39 (1.75)	(-8.85, -1.93)	0.002 *
		Irbesartan	25	20 (80.0)	2.49 (1.26)	(-0.01, 4.98)			
	Week 94	Sparsentan	26	20 (76.9)	-1.18 (1.26)	(-3.66, 1.30)	-2.96 (1.81)	(-6.54, 0.61)	0.103
		Irbesartan	25	19 (76.0)	1.79 (1.30)	(-0.77, 4.34)			
	Week 110	Sparsentan	26	21 (80.8)	-2.55 (1.23)	(-4.97, -0.12)	-3.74 (1.84)	(-7.37, -0.12)	0.043 *
		Irbesartan	25	17 (68.0)	1.19 (1.36)	(-1.50, 3.89)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.053
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	64 (78.0)	0.46 (0.82)	(-1.16, 2.08)	0.96 (1.24)	(-1.47, 3.40)	0.436
		Irbesartan	80	51 (63.8)	-0.51 (0.92)	(-2.32, 1.31)			
	Week 48	Sparsentan	82	63 (76.8)	1.14 (0.83)	(-0.48, 2.77)	1.06 (1.26)	(-1.41, 3.53)	0.401
		Irbesartan	80	45 (56.3)	0.09 (0.95)	(-1.78, 1.95)			
	Week 70	Sparsentan	82	65 (79.3)	0.07 (0.82)	(-1.54, 1.69)	2.07 (1.23)	(-0.35, 4.49)	0.094
		Irbesartan	80	50 (62.5)	-2.00 (0.92)	(-3.80, -0.19)			
	Week 94	Sparsentan	82	61 (74.4)	-0.68 (0.84)	(-2.33, 0.97)	0.16 (1.26)	(-2.32, 2.63)	0.900
		Irbesartan	80	48 (60.0)	-0.83 (0.94)	(-2.68, 1.01)			
	Week 110	Sparsentan	82	57 (69.5)	0.40 (0.87)	(-1.31, 2.10)	1.74 (1.31)	(-0.84, 4.33)	0.185
		Irbesartan	80	44 (55.0)	-1.35 (0.98)	(-3.28, 0.59)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	35 (77.8)	-0.23 (0.96)	(-2.11, 1.66)	-0.02 (1.37)	(-2.72, 2.68)	0.989
		Irbesartan	49	34 (69.4)	-0.21 (0.97)	(-2.13, 1.71)			
	Week 48	Sparsentan	45	34 (75.6)	0.05 (0.96)	(-1.84, 1.93)	-1.16 (1.40)	(-3.91, 1.59)	0.407
		Irbesartan	49	31 (63.3)	1.21 (1.01)	(-0.78, 3.19)			
	Week 70	Sparsentan	45	36 (80.0)	-1.01 (0.94)	(-2.87, 0.85)	-0.94 (1.38)	(-3.65, 1.77)	0.495
		Irbesartan	49	33 (67.3)	-0.07 (0.99)	(-2.03, 1.89)			
	Week 94	Sparsentan	45	33 (73.3)	0.81 (0.98)	(-1.11, 2.73)	-0.21 (1.42)	(-3.00, 2.58)	0.884
		Irbesartan	49	31 (63.3)	1.02 (1.02)	(-0.99, 3.02)			
	Week 110	Sparsentan	45	36 (80.0)	-1.41 (0.95)	(-3.28, 0.46)	-2.45 (1.44)	(-5.28, 0.38)	0.089
		Irbesartan	49	28 (57.1)	1.04 (1.07)	(-1.07, 3.15)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	39 (79.6)	0.83 (1.01)	(-1.17, 2.82)	0.03 (1.50)	(-2.93, 2.99)	0.986
		Irbesartan	48	33 (68.8)	0.80 (1.11)	(-1.38, 2.98)			
	Week 48	Sparsentan	49	45 (91.8)	-0.94 (0.96)	(-2.83, 0.96)	0.69 (1.46)	(-2.19, 3.57)	0.637
		Irbesartan	48	33 (68.8)	-1.63 (1.10)	(-3.78, 0.53)			
	Week 70	Sparsentan	49	40 (81.6)	0.13 (1.00)	(-1.83, 2.10)	2.10 (1.49)	(-0.84, 5.04)	0.161
		Irbesartan	48	32 (66.7)	-1.97 (1.11)	(-4.15, 0.21)			
	Week 94	Sparsentan	49	41 (83.7)	-0.79 (1.00)	(-2.76, 1.18)	1.16 (1.47)	(-1.75, 4.06)	0.434
		Irbesartan	48	35 (72.9)	-1.95 (1.08)	(-4.07, 0.18)			
	Week 110	Sparsentan	49	36 (73.5)	-0.38 (1.06)	(-2.46, 1.70)	0.76 (1.54)	(-2.27, 3.79)	0.622
		Irbesartan	48	32 (66.7)	-1.14 (1.12)	(-3.34, 1.06)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	21 (80.8)	0.54 (1.23)	(-1.88, 2.96)	0.38 (1.76)	(-3.09, 3.84)	0.831
		Irbesartan	25	20 (80.0)	0.16 (1.26)	(-2.32, 2.64)			
	Week 48	Sparsentan	26	23 (88.5)	-0.77 (1.18)	(-3.10, 1.56)	-1.55 (1.78)	(-5.06, 1.96)	0.384
		Irbesartan	25	18 (72.0)	0.78 (1.32)	(-1.82, 3.39)			
	Week 70	Sparsentan	26	22 (84.6)	-2.90 (1.21)	(-5.28, -0.52)	-5.39 (1.75)	(-8.85, -1.93)	0.002 *
		Irbesartan	25	20 (80.0)	2.49 (1.26)	(-0.01, 4.98)			
	Week 94	Sparsentan	26	20 (76.9)	-1.18 (1.26)	(-3.66, 1.30)	-2.96 (1.81)	(-6.54, 0.61)	0.103
		Irbesartan	25	19 (76.0)	1.79 (1.30)	(-0.77, 4.34)			
	Week 110	Sparsentan	26	21 (80.8)	-2.55 (1.23)	(-4.97, -0.12)	-3.74 (1.84)	(-7.37, -0.12)	0.043 *
		Irbesartan	25	17 (68.0)	1.19 (1.36)	(-1.50, 3.89)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.182
<= 1.75 g/day	Week 24	Sparsentan	98	81 (82.7)	0.39 (0.71)	(-1.01, 1.78)	-0.02 (1.09)	(-2.16, 2.13)	0.988
		Irbesartan	93	59 (63.4)	0.40 (0.83)	(-1.22, 2.03)			
	Week 48	Sparsentan	98	83 (84.7)	0.33 (0.70)	(-1.04, 1.71)	0.09 (1.08)	(-2.04, 2.22)	0.932
		Irbesartan	93	57 (61.3)	0.24 (0.83)	(-1.38, 1.86)			
	Week 70	Sparsentan	98	77 (78.6)	-1.07 (0.72)	(-2.49, 0.34)	-0.89 (1.08)	(-3.02, 1.24)	0.412
		Irbesartan	93	62 (66.7)	-0.18 (0.81)	(-1.77, 1.41)			
	Week 94	Sparsentan	98	79 (80.6)	-0.55 (0.72)	(-1.96, 0.86)	-0.31 (1.08)	(-2.43, 1.81)	0.774
		Irbesartan	93	64 (68.8)	-0.24 (0.80)	(-1.81, 1.33)			
	Week 110	Sparsentan	98	76 (77.6)	-1.46 (0.73)	(-2.89, -0.02)	-0.88 (1.12)	(-3.07, 1.32)	0.433
		Irbesartan	93	57 (61.3)	-0.58 (0.84)	(-2.23, 1.08)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 1.75 g/day	Week 24	Sparsentan	104	78 (75.0)	0.41 (0.69)	(-0.93, 1.76)	0.89 (0.97)	(-1.02, 2.80)	0.359
		Irbesartan	109	79 (72.5)	-0.48 (0.69)	(-1.83, 0.87)			
	Week 48	Sparsentan	104	82 (78.8)	-0.20 (0.67)	(-1.51, 1.11)	0.16 (0.98)	(-1.76, 2.08)	
		Irbesartan	109	70 (64.2)	-0.36 (0.71)	(-1.76, 1.04)			
	Week 70	Sparsentan	104	86 (82.7)	-0.11 (0.66)	(-1.41, 1.18)	1.50 (0.97)	(-0.41, 3.40)	
		Irbesartan	109	73 (67.0)	-1.61 (0.71)	(-3.00, -0.22)			
	Week 94	Sparsentan	104	76 (73.1)	-0.29 (0.69)	(-1.65, 1.07)	0.16 (1.00)	(-1.81, 2.13)	
		Irbesartan	109	69 (63.3)	-0.45 (0.73)	(-1.88, 0.97)			
	Week 110	Sparsentan	104	74 (71.2)	0.33 (0.71)	(-1.06, 1.72)	0.69 (1.04)	(-1.34, 2.73)	
		Irbesartan	109	64 (58.7)	-0.36 (0.76)	(-1.84, 1.12)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024



Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan						Interaction:	0.807
Yes	Week 24	Sparsentan	90	68 (75.6)	0.52 (0.77)	(-1.00, 2.03)	0.19 (1.17)	(-2.11, 2.48)	0.874
		Irbesartan	88	52 (59.1)	0.33 (0.88)	(-1.40, 2.06)			
	Week 48	Sparsentan	90	68 (75.6)	0.34 (0.77)	(-1.17, 1.84)	-0.62 (1.18)	(-2.94, 1.70)	0.598
		Irbesartan	88	48 (54.5)	0.96 (0.90)	(-0.80, 2.72)			
	Week 70	Sparsentan	90	69 (76.7)	-0.96 (0.76)	(-2.46, 0.55)	0.08 (1.17)	(-2.21, 2.38)	0.942
		Irbesartan	88	51 (58.0)	-1.04 (0.88)	(-2.77, 0.69)			
	Week 94	Sparsentan	90	66 (73.3)	-0.37 (0.78)	(-1.90, 1.16)	-0.58 (1.15)	(-2.83, 1.68)	0.617
		Irbesartan	88	57 (64.8)	0.20 (0.84)	(-1.45, 1.86)			
	Week 110	Sparsentan	90	64 (71.1)	0.43 (0.80)	(-1.14, 1.99)	1.19 (1.19)	(-1.15, 3.53)	0.319
		Irbesartan	88	51 (58.0)	-0.76 (0.89)	(-2.50, 0.98)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
No	Week 24	Sparsentan	112	91 (81.3)	0.31 (0.64)	(-0.95, 1.57)	0.60 (0.92)	(-1.21, 2.41)	0.517
		Irbesartan	114	86 (75.4)	-0.29 (0.66)	(-1.59, 1.02)			
	Week 48	Sparsentan	112	97 (86.6)	-0.14 (0.62)	(-1.36, 1.09)	0.54 (0.92)	(-1.27, 2.36)	0.556
		Irbesartan	114	79 (69.3)	-0.68 (0.68)	(-2.02, 0.65)			
	Week 70	Sparsentan	112	94 (83.9)	-0.30 (0.63)	(-1.55, 0.94)	0.56 (0.92)	(-1.25, 2.37)	0.545
		Irbesartan	114	84 (73.7)	-0.86 (0.67)	(-2.18, 0.45)			
	Week 94	Sparsentan	112	89 (79.5)	-0.52 (0.65)	(-1.80, 0.75)	0.20 (0.96)	(-1.68, 2.08)	0.836
		Irbesartan	114	76 (66.7)	-0.72 (0.70)	(-2.10, 0.65)			
	Week 110	Sparsentan	112	86 (76.8)	-1.37 (0.66)	(-2.67, -0.07)	-1.27 (0.99)	(-3.21, 0.67)	0.198
		Irbesartan	114	70 (61.4)	-0.09 (0.73)	(-1.53, 1.34)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.567
<= 5 years	Week 24	Sparsentan	113	90 (79.6)	1.04 (0.67)	(-0.28, 2.37)	1.16 (0.95)	(-0.71, 3.03)	0.223
		Irbesartan	127	91 (71.7)	-0.12 (0.67)	(-1.43, 1.20)			
	Week 48	Sparsentan	113	93 (82.3)	0.03 (0.66)	(-1.27, 1.33)	-0.33 (0.95)	(-2.20, 1.53)	0.725
		Irbesartan	127	86 (67.7)	0.37 (0.68)	(-0.96, 1.70)			
	Week 70	Sparsentan	113	91 (80.5)	-0.27 (0.67)	(-1.58, 1.05)	0.03 (0.95)	(-1.84, 1.89)	0.979
		Irbesartan	127	88 (69.3)	-0.29 (0.68)	(-1.62, 1.04)			
	Week 94	Sparsentan	113	88 (77.9)	0.11 (0.68)	(-1.22, 1.44)	-0.12 (0.96)	(-2.01, 1.76)	0.898
		Irbesartan	127	88 (69.3)	0.24 (0.68)	(-1.10, 1.57)			
	Week 110	Sparsentan	113	88 (77.9)	0.49 (0.68)	(-0.85, 1.83)	0.34 (0.99)	(-1.60, 2.27)	0.733
		Irbesartan	127	80 (63.0)	0.15 (0.71)	(-1.24, 1.55)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 5 years	Week 24	Sparsentan	89	69 (77.5)	-0.58 (0.72)	(-1.99, 0.83)	-0.93 (1.14)	(-3.17, 1.31)	0.413
		Irbesartan	75	47 (62.7)	0.35 (0.88)	(-1.38, 2.08)			
	Week 48	Sparsentan	89	72 (80.9)	-0.04 (0.70)	(-1.42, 1.35)	0.59 (1.15)	(-1.67, 2.86)	
		Irbesartan	75	41 (54.7)	-0.63 (0.91)	(-2.42, 1.16)			
	Week 70	Sparsentan	89	72 (80.9)	-1.09 (0.71)	(-2.49, 0.30)	0.86 (1.13)	(-1.36, 3.08)	
		Irbesartan	75	47 (62.7)	-1.96 (0.87)	(-3.68, -0.24)			
	Week 94	Sparsentan	89	67 (75.3)	-1.30 (0.73)	(-2.73, 0.14)	-0.05 (1.16)	(-2.33, 2.22)	
		Irbesartan	75	45 (60.0)	-1.24 (0.89)	(-3.00, 0.51)			
	Week 110	Sparsentan	89	62 (69.7)	-2.22 (0.76)	(-3.71, -0.73)	-0.95 (1.20)	(-3.32, 1.42)	
		Irbesartan	75	41 (54.7)	-1.27 (0.93)	(-3.11, 0.56)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.390
Yes	Week 24	Sparsentan	155	121 (78.1)	0.52 (0.56)	(-0.59, 1.62)	0.58 (0.82)	(-1.04, 2.20)	0.483
		Irbesartan	161	106 (65.8)	-0.06 (0.60)	(-1.24, 1.12)			
	Week 48	Sparsentan	155	124 (80.0)	0.11 (0.55)	(-0.98, 1.20)	-0.23 (0.83)	(-1.86, 1.40)	0.782
		Irbesartan	161	97 (60.2)	0.34 (0.62)	(-0.87, 1.55)			
	Week 70	Sparsentan	155	125 (80.6)	-0.31 (0.56)	(-1.40, 0.78)	1.10 (0.82)	(-0.52, 2.71)	0.183
		Irbesartan	161	104 (64.6)	-1.41 (0.60)	(-2.60, -0.22)			
	Week 94	Sparsentan	155	118 (76.1)	-0.26 (0.57)	(-1.37, 0.85)	0.26 (0.83)	(-1.37, 1.90)	0.754
		Irbesartan	161	103 (64.0)	-0.52 (0.61)	(-1.71, 0.67)			
	Week 110	Sparsentan	155	113 (72.9)	-0.22 (0.58)	(-1.36, 0.92)	-0.04 (0.87)	(-1.74, 1.66)	0.965
		Irbesartan	161	92 (57.1)	-0.18 (0.64)	(-1.44, 1.08)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KPSC\_FSCM: KDQOL-SF12: Change from baseline in PCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in PCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
No	Week 24	Sparsentan	47	38 (80.9)	0.01 (1.04)	(-2.04, 2.06)	-0.17 (1.55)	(-3.22, 2.89)	0.914
		Irbesartan	41	32 (78.0)	0.18 (1.14)	(-2.08, 2.43)			
	Week 48	Sparsentan	47	41 (87.2)	-0.10 (1.01)	(-2.09, 1.89)	1.09 (1.54)	(-1.94, 4.13)	0.478
		Irbesartan	41	30 (73.2)	-1.20 (1.16)	(-3.48, 1.08)			
	Week 70	Sparsentan	47	38 (80.9)	-1.43 (1.04)	(-3.47, 0.62)	-2.25 (1.56)	(-5.32, 0.81)	0.149
		Irbesartan	41	31 (75.6)	0.83 (1.15)	(-1.45, 3.10)			
	Week 94	Sparsentan	47	37 (78.7)	-1.06 (1.06)	(-3.14, 1.02)	-1.48 (1.59)	(-4.60, 1.65)	0.352
		Irbesartan	41	30 (73.2)	0.42 (1.18)	(-1.90, 2.74)			
	Week 110	Sparsentan	47	37 (78.7)	-1.77 (1.06)	(-3.86, 0.33)	-0.77 (1.61)	(-3.95, 2.40)	0.631
		Irbesartan	41	29 (70.7)	-0.99 (1.21)	(-3.37, 1.38)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. PCS = physical component summary.  
 Source Data: aqs, created on: 05MAR2024

Figure PF2KPSC\_FSGM: KDQOL-SF12: Change from baseline in PCS by subgroup  
Full Analysis Set

Not done: No significant subgroup found.

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
PCS = physical component summary.  
Reference table: PT2KPSC\_FSCM

Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.942
Male	Sparsentan	139	22 (15.8)	NE		1.074	(0.569, 2.025)	0.826
	Irbesartan	143	20 (14.0)	117.0	(NE, NE)			
Female	Sparsentan	63	10 (15.9)	NE		1.274	(0.363, 4.476)	0.705
	Irbesartan	59	4 (6.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.167
<= 45 years	Sparsentan	96	10 (10.4)	NE		0.620	(0.239, 1.608)	0.325
	Irbesartan	99	10 (10.1)	NE				
> 45 years	Sparsentan	106	22 (20.8)	NE		1.367	(0.679, 2.754)	0.381
	Irbesartan	103	14 (13.6)	117.0	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: <0.001 #
<= 18 years	Sparsentan	9	1 (11.1)	NE		NE		NE
	Irbesartan	5	1 (20.0)	NE				
> 18 to 40 years	Sparsentan	102	11 (10.8)	NE		1.184	(0.500, 2.804)	0.701
	Irbesartan	109	11 (10.1)	NE				
> 40 years	Sparsentan	91	20 (22.0)	NE		1.318	(0.606, 2.867)	0.486
	Irbesartan	88	12 (13.6)	117.0	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan							Interaction test: <0.001 #
North America	Sparsentan	35	2 (5.7)	NE		0.109	(0.012, 1.024)	0.052
	Irbesartan	46	9 (19.6)	NE				
Europe	Sparsentan	98	12 (12.2)	NE		1.011	(0.382, 2.673)	0.983
	Irbesartan	115	9 (7.8)	117.0	(117.0, NE)			
Asia Pacific	Sparsentan	69	18 (26.1)	NE		1.708	(0.656, 4.444)	0.273
	Irbesartan	41	6 (14.6)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.674
< 27 kg/m**2	Sparsentan	83	13 (15.7)	NE		1.035	(0.385, 2.784)	0.945
	Irbesartan	94	8 (8.5)	NE				
≥ 27 kg/m**2	Sparsentan	119	19 (16.0)	NE		1.232	(0.604, 2.512)	0.566
	Irbesartan	107	16 (15.0)	117.0	(117.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan							Interaction test: <0.001 #
eGFR Low and UP High	Sparsentan	71	8 (11.3)	NE		0.794	(0.304, 2.070)	0.637
	Irbesartan	74	10 (13.5)	117.0	(NE, NE)			
eGFR Low and UP Low	Sparsentan	55	12 (21.8)	NE		0.947	(0.286, 3.142)	0.929
	Irbesartan	55	5 (9.1)	NE				
eGFR High and UP High	Sparsentan	37	3 (8.1)	NE		0.134	(0.016, 1.155)	0.067
	Irbesartan	36	5 (13.9)	NE				
eGFR High and UP Low	Sparsentan	39	9 (23.1)	NE		4.828	(1.153, 20.215)	0.031 *
	Irbesartan	37	4 (10.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.580
< 60 mL/min/1.73 m**2	Sparsentan	127	21 (16.5)	NE		1.098	(0.546, 2.206)	0.794
	Irbesartan	129	16 (12.4)	117.0	(NE, NE)			
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	7 (14.3)	116.1	(116.1, NE)	2.550	(0.603, 10.780)	0.203
	Irbesartan	48	6 (12.5)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	4 (15.4)	NE		0.916	(0.133, 6.290)	0.929
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.730
< 45 mL/min/1.73 m**2	Sparsentan	82	13 (15.9)	NE		0.891	(0.359, 2.207)	0.802
	Irbesartan	80	10 (12.5)	117.0	(NE, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	8 (17.8)	NE		1.321	(0.402, 4.336)	0.647
	Irbesartan	49	6 (12.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	7 (14.3)	116.1	(116.1, NE)	2.550	(0.603, 10.780)	0.203
	Irbesartan	48	6 (12.5)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	4 (15.4)	NE		0.916	(0.133, 6.290)	0.929
	Irbesartan	25	2 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.017 #
<= 1.75 g/day	Sparsentan	98	21 (21.4)	NE		1.976	(0.869, 4.493)	0.104
	Irbesartan	93	9 (9.7)	NE				
> 1.75 g/day	Sparsentan	104	11 (10.6)	NE		0.493	(0.209, 1.163)	0.106
	Irbesartan	109	15 (13.8)	117.0	(NE, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.393
Yes	Sparsentan	90	14 (15.6)	NE		0.635	(0.282, 1.427)	0.271
	Irbesartan	88	14 (15.9)	NE				
No	Sparsentan	112	18 (16.1)	NE		1.878	(0.818, 4.313)	0.137
	Irbesartan	114	10 (8.8)	117.0	(117.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.374
<= 5 years	Sparsentan	113	23 (20.4)	NE		1.284	(0.673, 2.449)	0.448
	Irbesartan	127	18 (14.2)	NE				
> 5 years	Sparsentan	89	9 (10.1)	NE		1.346	(0.397, 4.567)	0.634
	Irbesartan	75	6 (8.0)	117.0	(117.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

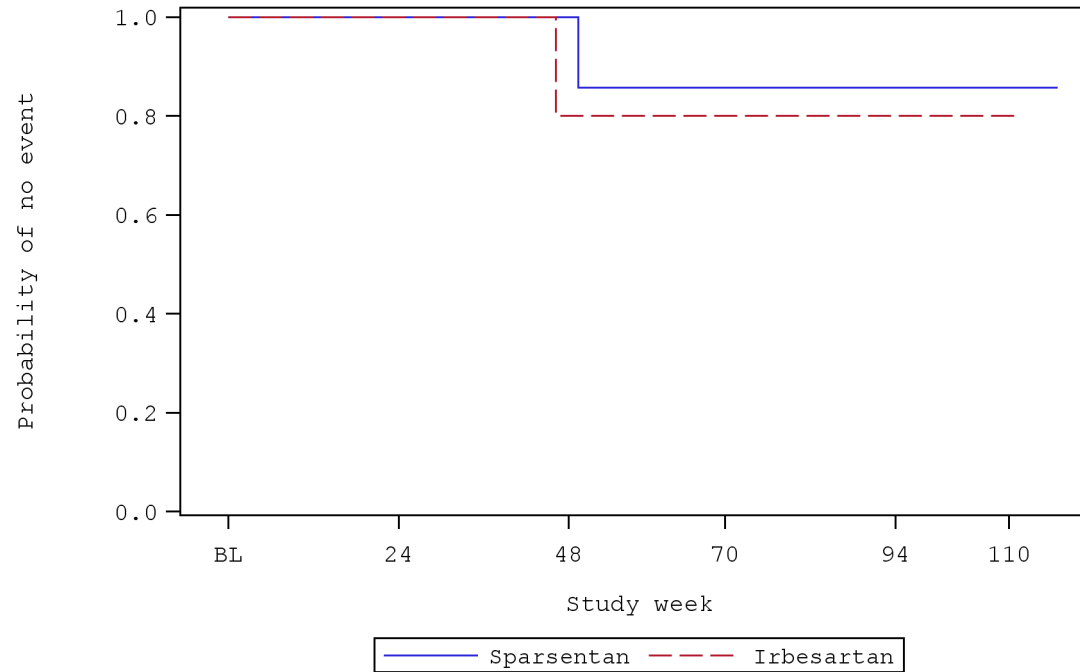
Table PT2KPSIT\_FSTM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.034 #
Yes	Sparsentan	155	24 (15.5)	NE		0.730	(0.383, 1.393)	0.340
	Irbesartan	161	20 (12.4)	117.0	(117.0, NE)			
No	Sparsentan	47	8 (17.0)	NE		4.536	(0.904, 22.756)	0.066
	Irbesartan	41	4 (9.8)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

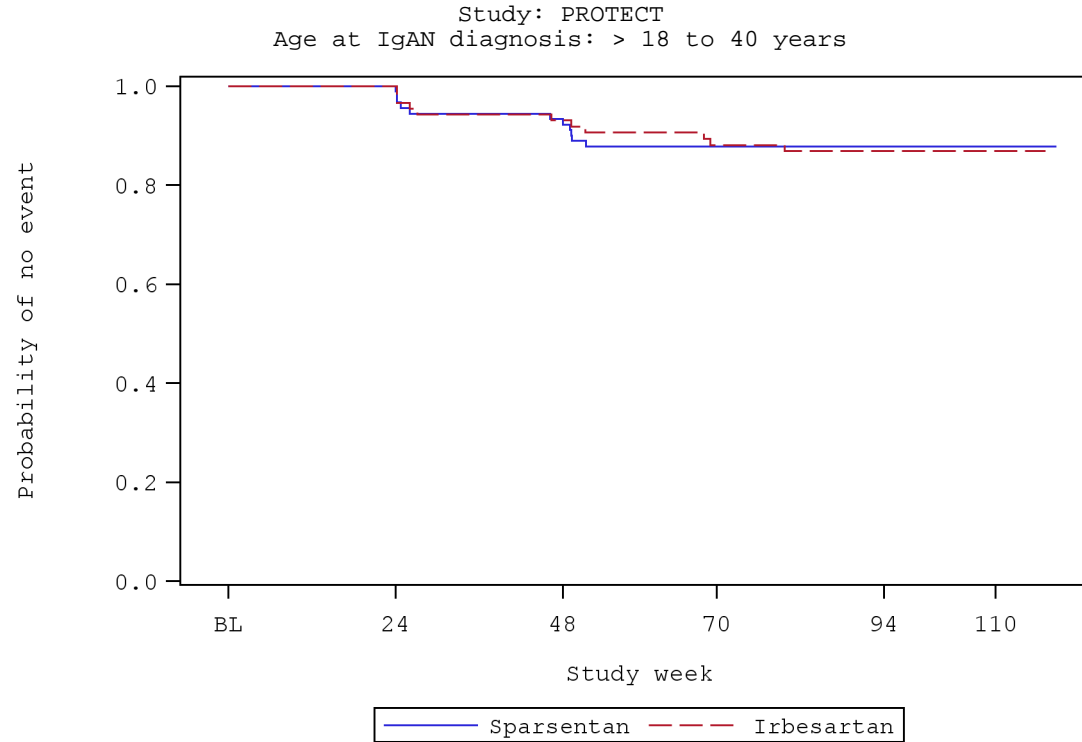
Study: PROTECT  
 Age at IgAN diagnosis: <= 18 years



Sparsentan	9	8	7	6	5	3
Irbesartan	5	5	4	4	3	2

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

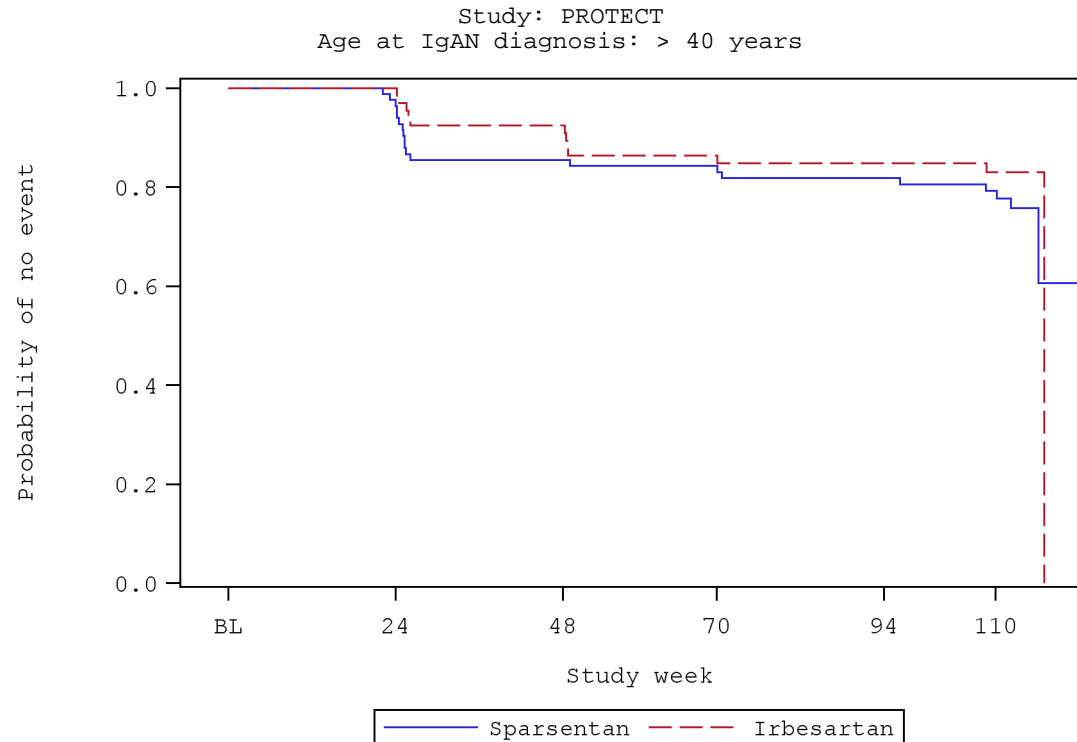
Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	102	91	84	77	71	62
Irbesartan	109	90	78	69	67	63

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

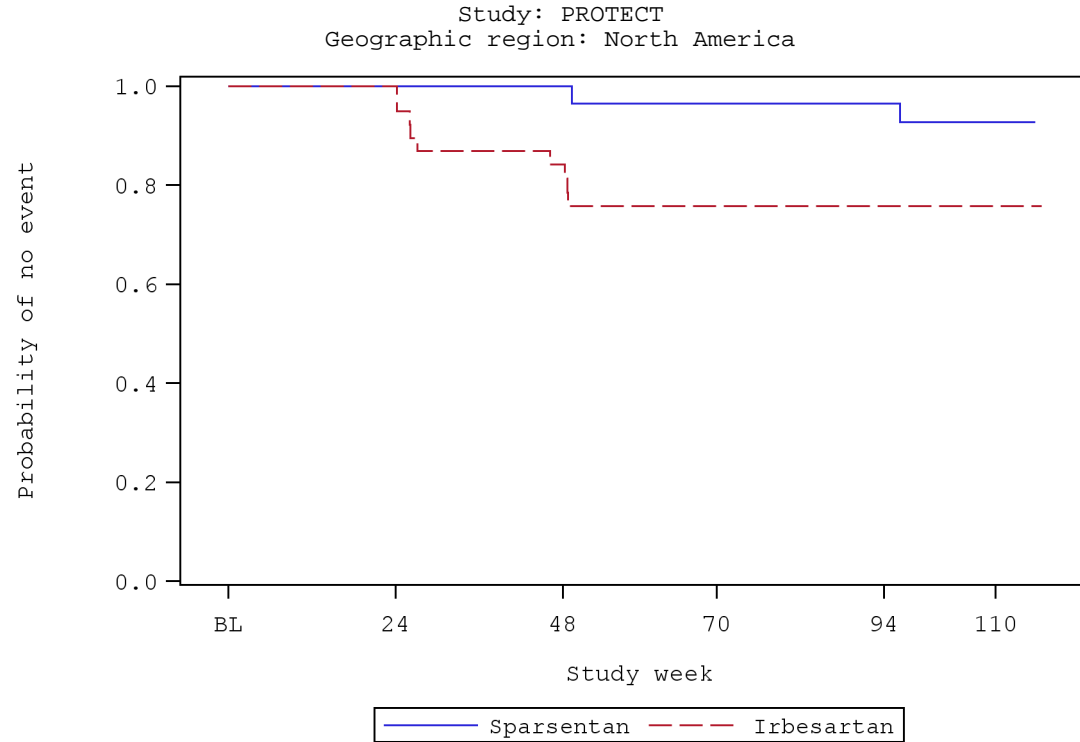
Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	91	81	70	69	64	54
Irbesartan	88	68	61	56	53	40

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

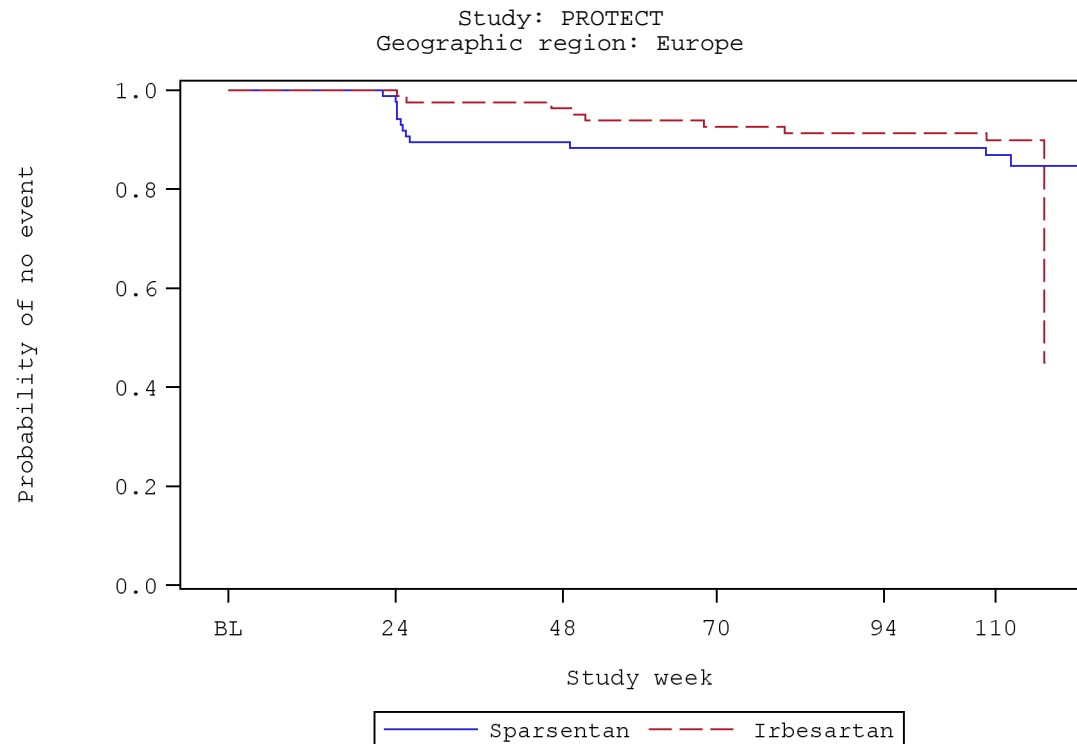
Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	35	30	29	28	26	20
Irbesartan	46	40	31	26	24	23

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

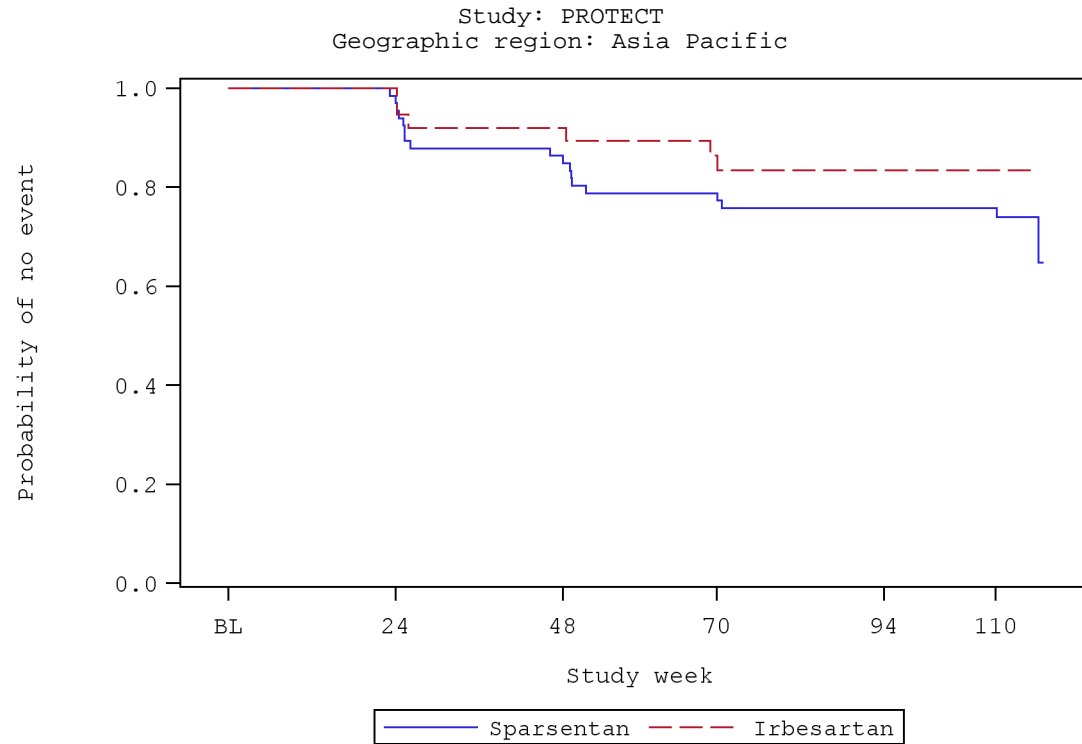


Sparsentan	98	85	75	72	67	56
Irbesartan	115	85	78	74	71	59

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM



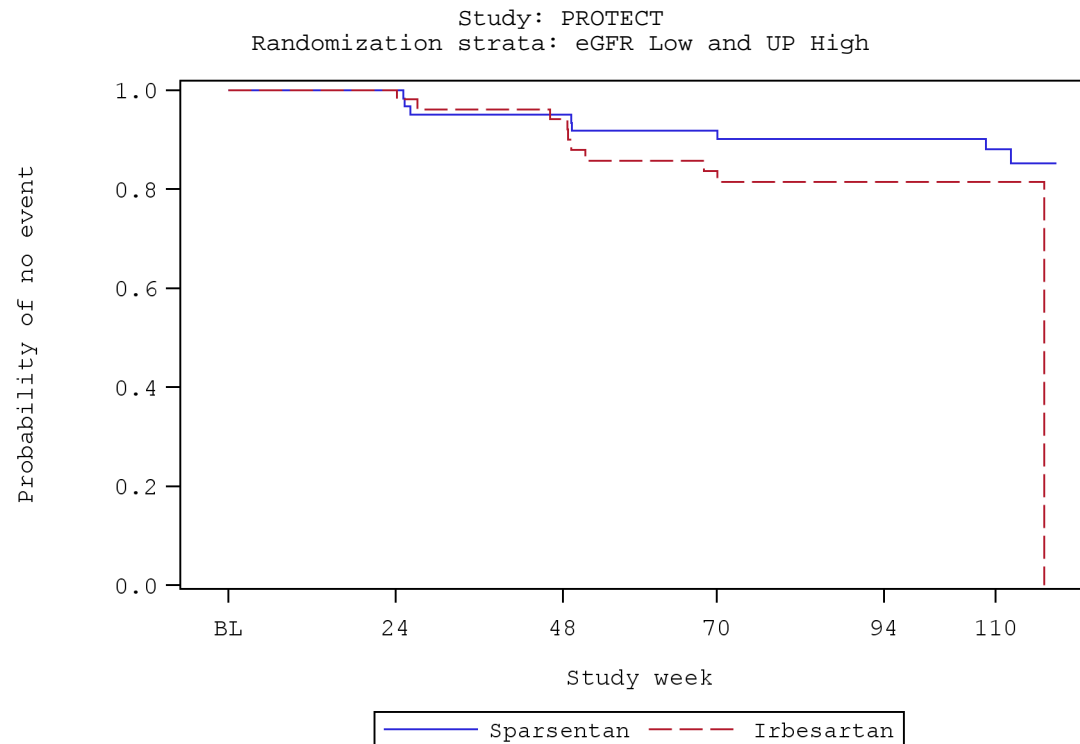
Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	69	65	57	52	47	43
Irbesartan	41	38	34	29	28	23

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

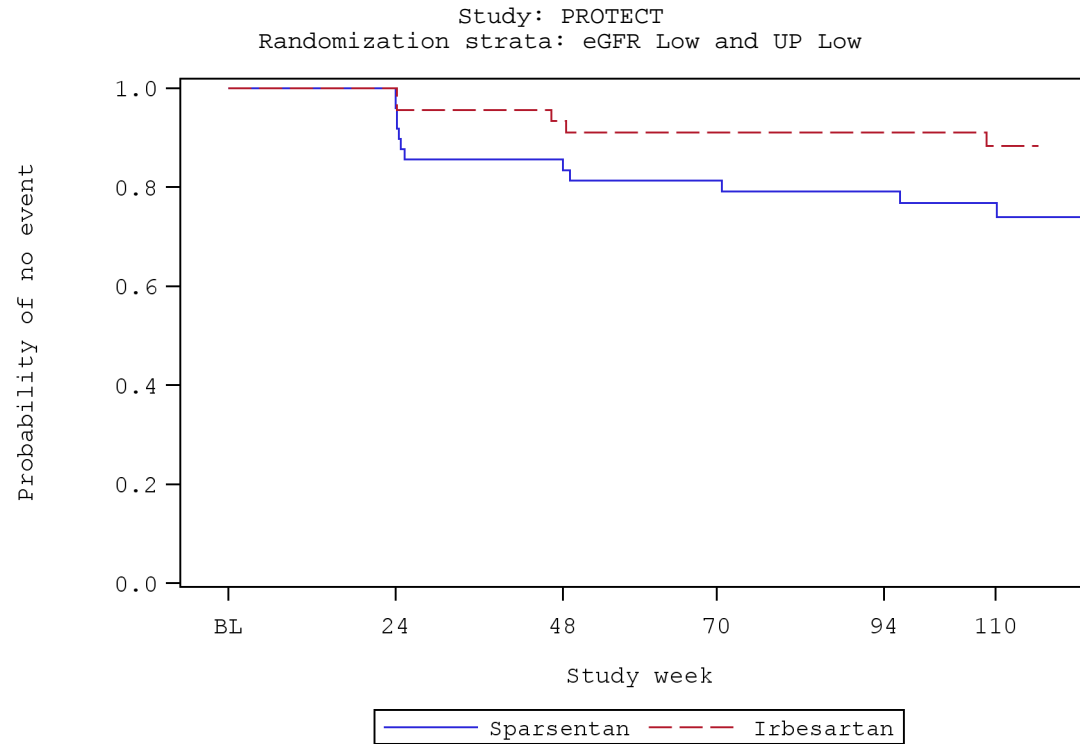
Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	71	61	58	55	50	41
Irbesartan	74	54	47	38	37	30

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

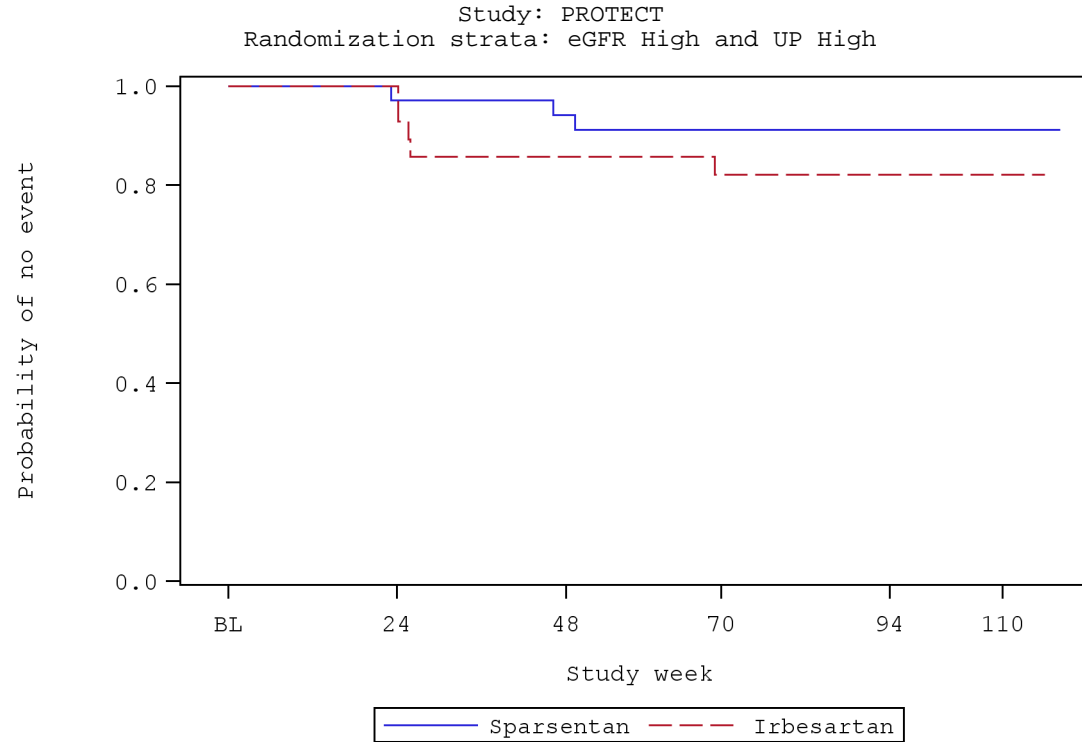
Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	55	49	40	38	34	28
Irbesartan	55	46	41	38	36	30

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

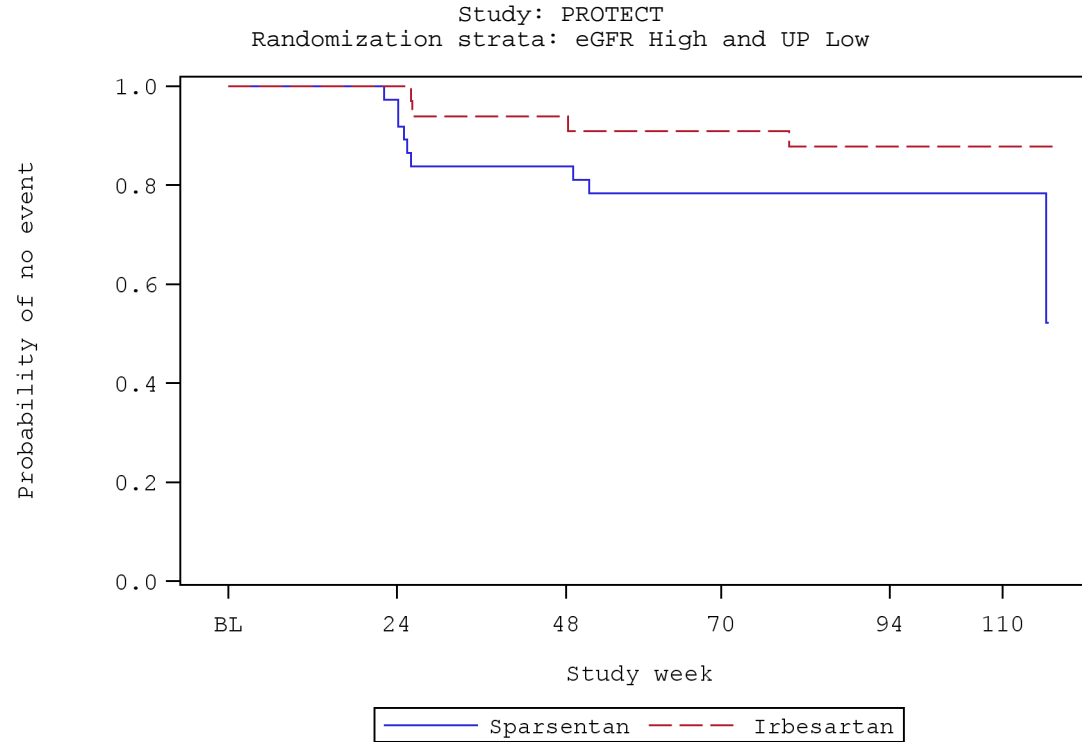
Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	37	34	32	30	27	23
Irbesartan	36	29	24	23	22	20

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

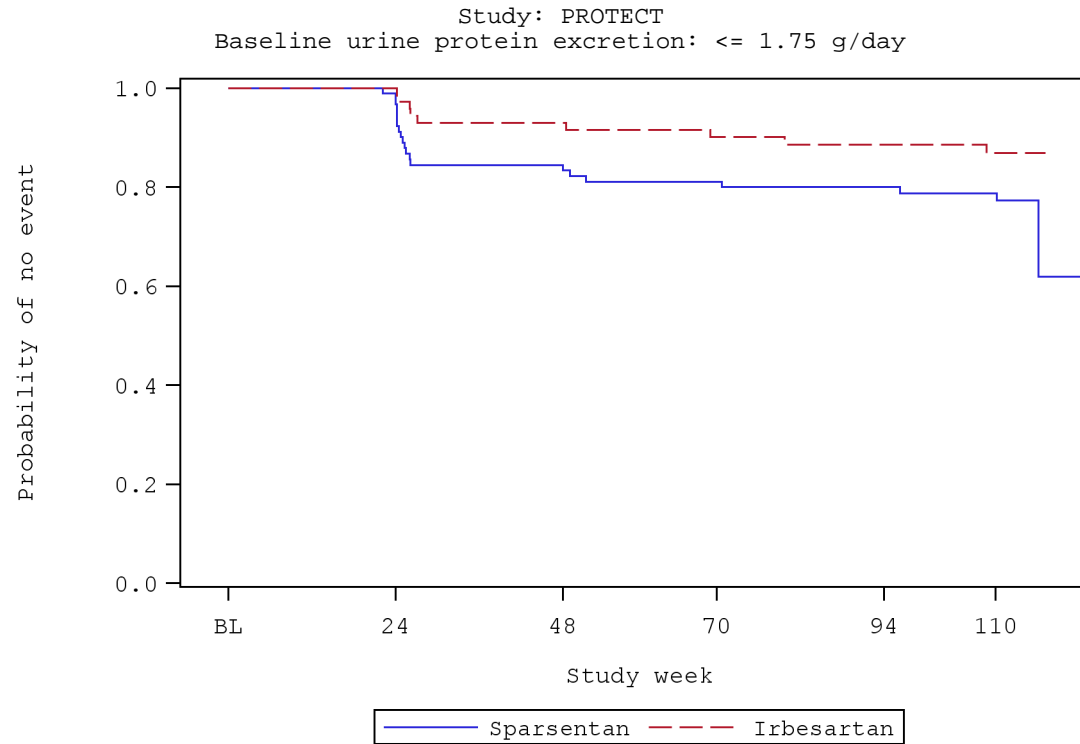
Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	39	36	31	29	29	27
Irbesartan	37	34	31	30	28	25

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

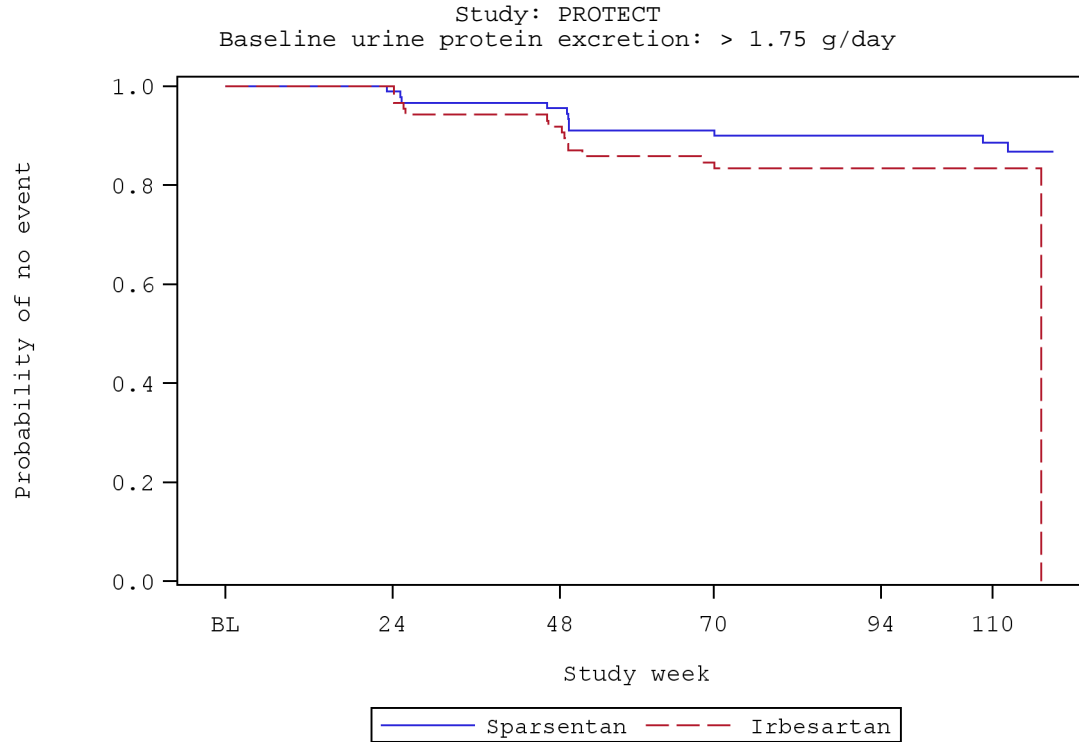
Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	98	90	75	72	69	57
Irbesartan	93	74	65	61	57	46

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

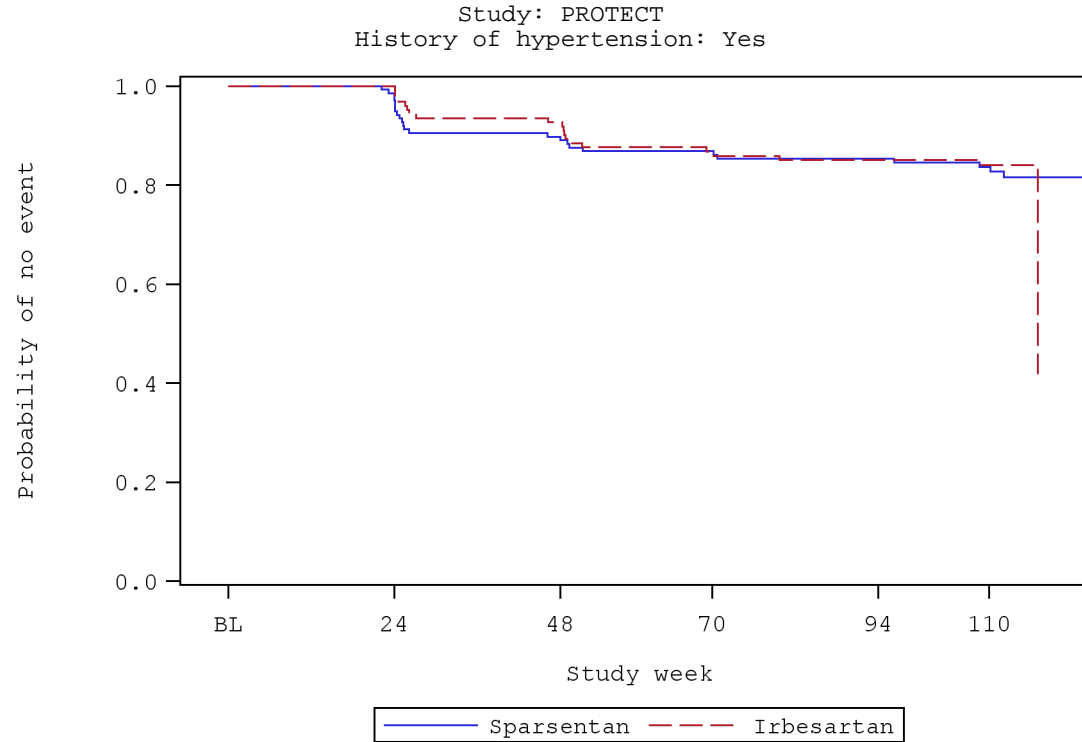
Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	104	90	86	80	71	62
Irbesartan	109	89	78	68	66	59

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

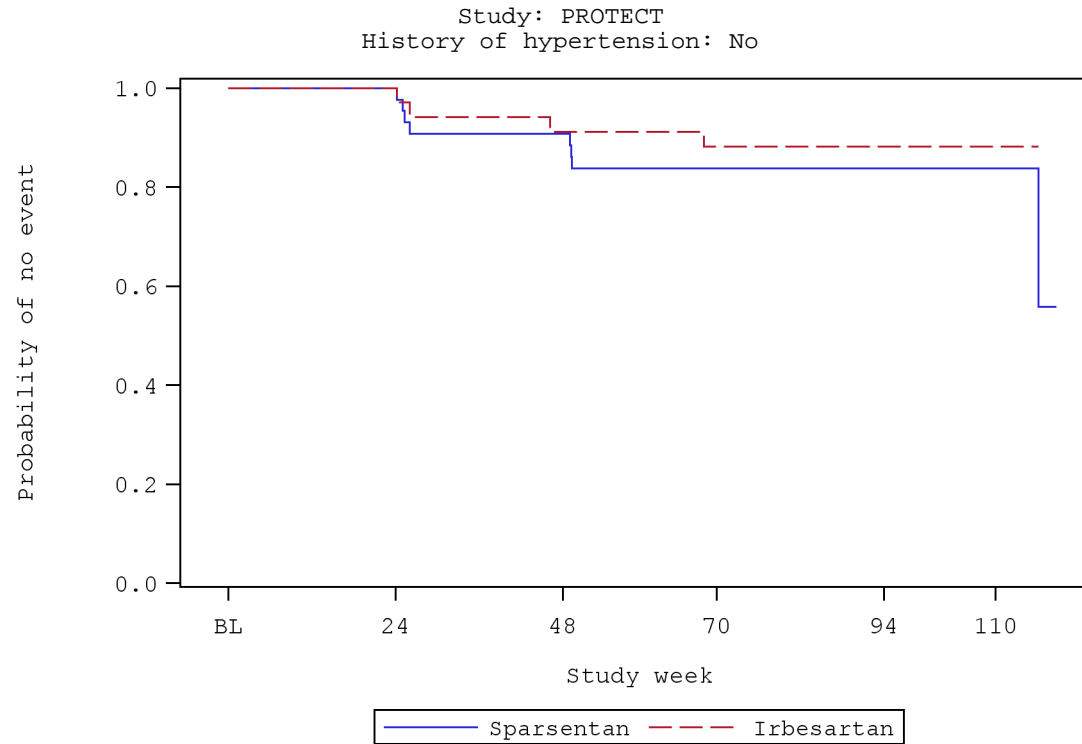


Sparsentan	155	136	122	117	108	91
Irbesartan	161	128	112	100	95	79

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM



Figure PF2KPSIT\_FSKM: KDQOL-SF12: Time to increase in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	47	44	39	35	32	28
Irbesartan	41	35	31	29	28	26

An increase reflects an improvement of the status of the patient.  
 Reference table: PT2KPSIT\_FSTM

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.797
Male	Sparsentan	139	37 (26.6)	NE		1.142	(0.701, 1.861)	0.594
	Irbesartan	143	29 (20.3)	NE				
Female	Sparsentan	63	21 (33.3)	NE		0.989	(0.512, 1.912)	0.974
	Irbesartan	59	17 (28.8)	116.0	(111.3, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.787
<= 45 years	Sparsentan	96	27 (28.1)	115.6	(114.3, NE)	1.055	(0.603, 1.848)	0.850
	Irbesartan	99	23 (23.2)	NE				
> 45 years	Sparsentan	106	31 (29.2)	NE		1.189	(0.691, 2.049)	0.532
	Irbesartan	103	23 (22.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: NE
<= 18 years	Sparsentan	9	2 (22.2) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	30 (29.4)	115.6	(114.3, NE)	1.082	(0.632, 1.854)	0.773
	Irbesartan	109	25 (22.9)	NE				
> 40 years	Sparsentan	91	26 (28.6)	NE		1.085	(0.606, 1.942)	0.784
	Irbesartan	88	21 (23.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.756
North America	Sparsentan	35	10 (28.6)	115.6	(111.6, NE)	0.916	(0.362, 2.317)	0.853
	Irbesartan	46	11 (23.9)	NE				
Europe	Sparsentan	98	26 (26.5)	NE		1.011	(0.585, 1.747)	0.969
	Irbesartan	115	27 (23.5)	NE				
Asia Pacific	Sparsentan	69	22 (31.9)	NE		1.610	(0.706, 3.670)	0.258
	Irbesartan	41	8 (19.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.510
< 27 kg/m**2	Sparsentan	83	24 (28.9)	NE		1.347	(0.735, 2.469)	0.335
	Irbesartan	94	19 (20.2)	NE				
≥ 27 kg/m**2	Sparsentan	119	34 (28.6)	NE		1.025	(0.613, 1.714)	0.925
	Irbesartan	107	26 (24.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.419
eGFR Low and UP High	Sparsentan	71	22 (31.0)	NE		1.457	(0.731, 2.903)	0.285
	Irbesartan	74	13 (17.6)	NE				
eGFR Low and UP Low	Sparsentan	55	9 (16.4)	NE		0.598	(0.257, 1.393)	0.233
	Irbesartan	55	14 (25.5)	NE				
eGFR High and UP High	Sparsentan	37	12 (32.4)	115.0	(113.6, NE)	1.226	(0.496, 3.033)	0.659
	Irbesartan	36	8 (22.2)	NE				
eGFR High and UP Low	Sparsentan	39	15 (38.5)	NE		1.262	(0.578, 2.756)	0.558
	Irbesartan	37	11 (29.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan							Interaction test: 0.040 #
< 60 mL/min/1.73 m**2	Sparsentan	127	31 (24.4)	NE		1.152	(0.676, 1.961)	0.603
	Irbesartan	129	26 (20.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	15 (30.6)	NE		0.628	(0.306, 1.289)	0.205
	Irbesartan	48	17 (35.4)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	12 (46.2)	114.3	(72.1, NE)	3.618	(1.017, 12.874)	0.047 *
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.001 #
< 45 mL/min/1.73 m**2	Sparsentan	82	17 (20.7)	NE		0.625	(0.331, 1.179)	0.146
	Irbesartan	80	22 (27.5)	116.0	(111.7, NE)			
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	14 (31.1)	NE		5.981	(1.744, 20.512)	0.004 *
	Irbesartan	49	4 (8.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	15 (30.6)	NE		0.628	(0.306, 1.289)	0.205
	Irbesartan	48	17 (35.4)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	12 (46.2)	114.3	(72.1, NE)	3.618	(1.017, 12.874)	0.047 *
	Irbesartan	25	3 (12.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.931
<= 1.75 g/day	Sparsentan	98	29 (29.6)	NE		1.099	(0.631, 1.916)	0.738
	Irbesartan	93	22 (23.7)	NE				
> 1.75 g/day	Sparsentan	104	29 (27.9)	NE		1.154	(0.669, 1.992)	0.606
	Irbesartan	109	24 (22.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.545
Yes	Sparsentan	90	24 (26.7)	NE		1.034	(0.573, 1.866)	0.913
	Irbesartan	88	21 (23.9)	NE				
No	Sparsentan	112	34 (30.4)	NE		1.259	(0.751, 2.111)	0.383
	Irbesartan	114	25 (21.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.809
<= 5 years	Sparsentan	113	30 (26.5)	NE		1.091	(0.658, 1.810)	0.736
	Irbesartan	127	31 (24.4)	NE				
> 5 years	Sparsentan	89	28 (31.5)	NE		1.203	(0.636, 2.278)	0.570
	Irbesartan	75	15 (20.0)	NE				

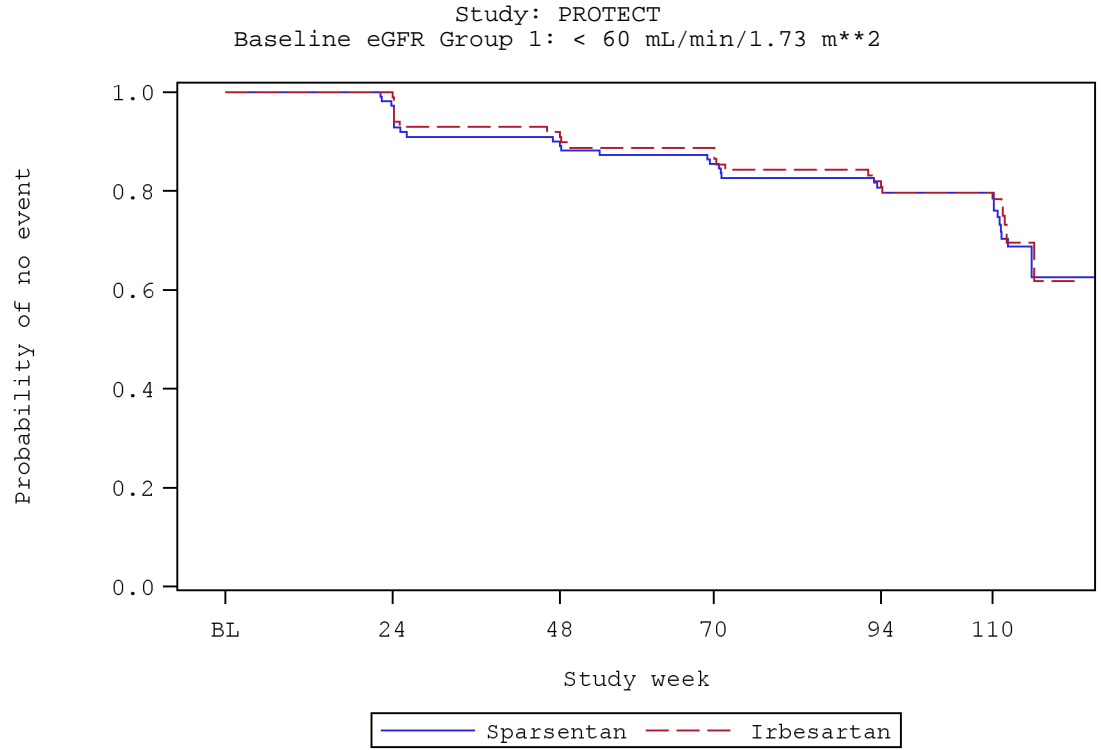
N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KPSDT\_FSTM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to decrease in PCS by at least 8.4				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.592
Yes	Sparsentan	155	41 (26.5)	NE		1.058	(0.678, 1.652)	0.804
	Irbesartan	161	37 (23.0)	NE				
No	Sparsentan	47	17 (36.2)	115.0	(111.6, NE)	1.244	(0.541, 2.858)	0.608
	Irbesartan	41	9 (22.0)	116.0	(116.0, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

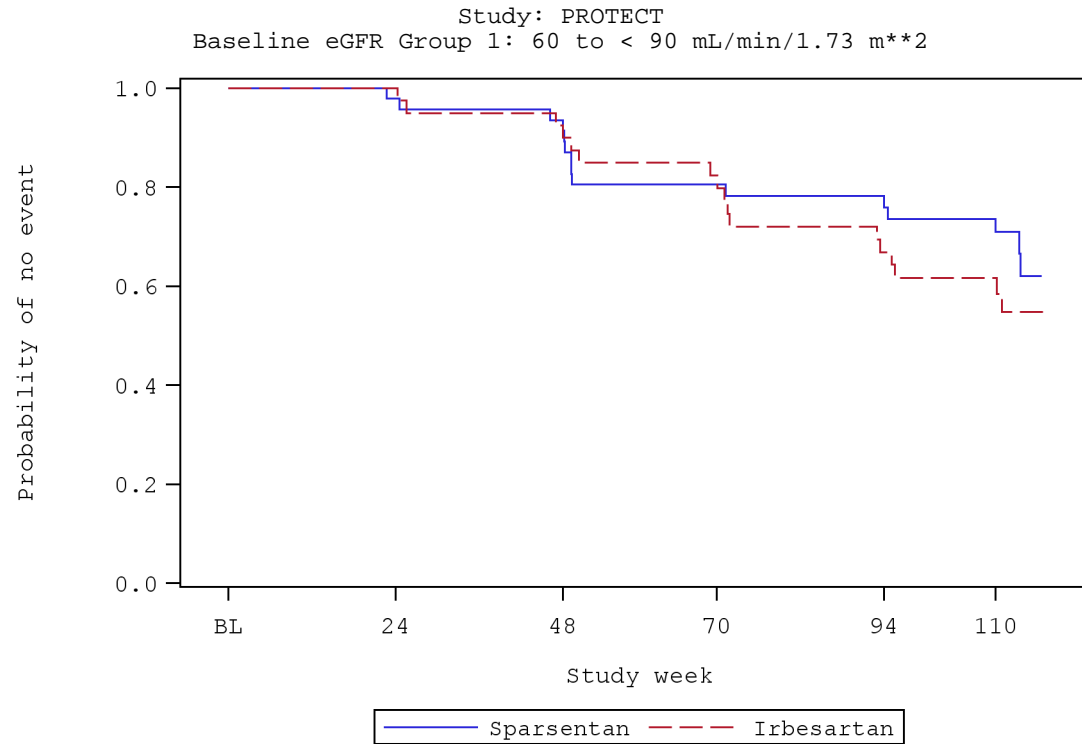
Figure PF2KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	127	108	98	92	81	68
Irbesartan	129	101	87	79	71	60

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KPSDT\_FSTM

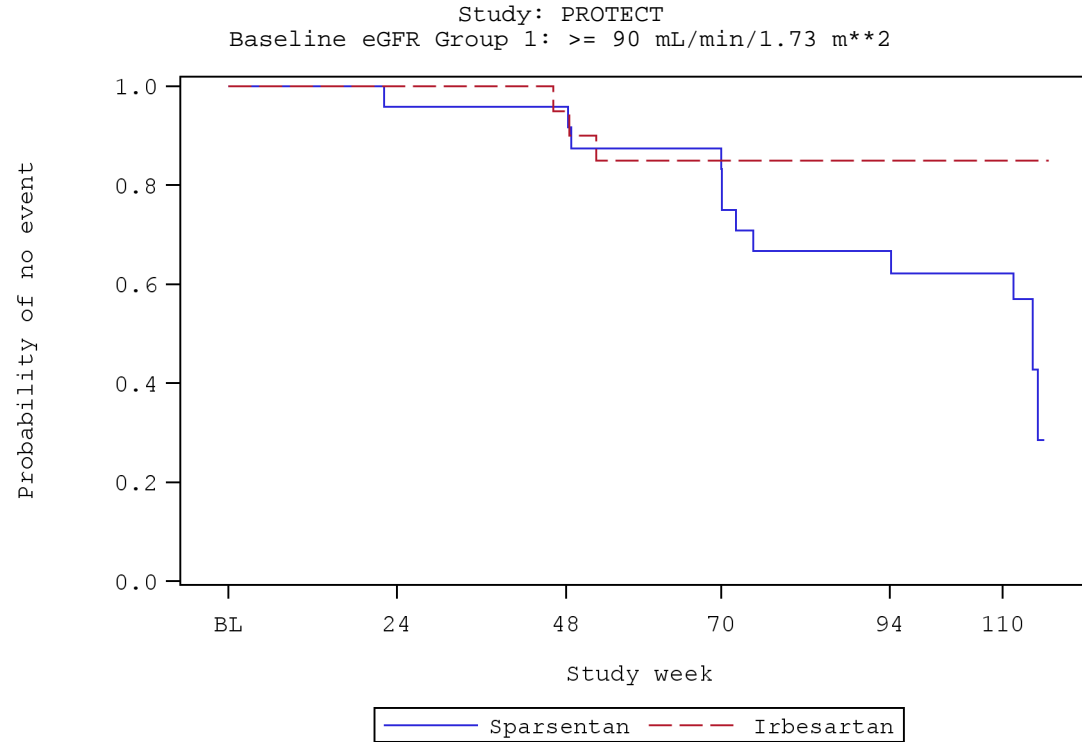
Figure PF2KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	49	46	43	36	34	28
Irbesartan	48	41	37	32	26	20

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KPSDT\_FSTM

Figure PF2KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set

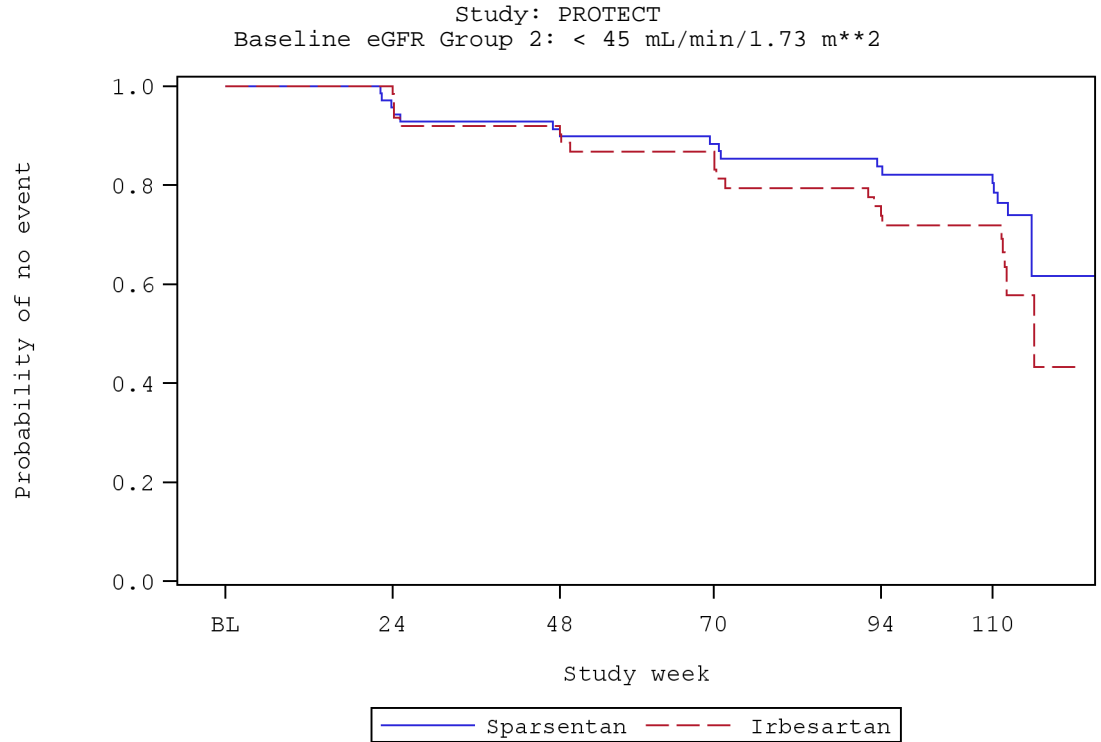


Sparsentan	26	23	23	21	15	13
Irbesartan	25	21	19	17	16	14

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KPSDT\_FSTM



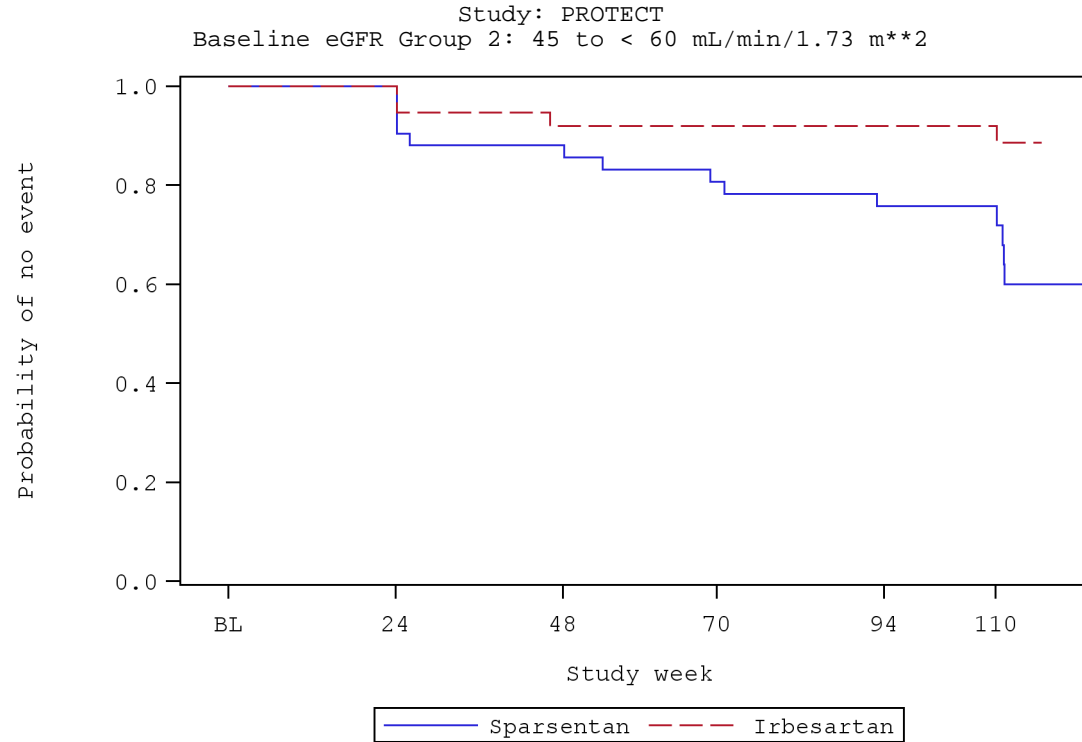
Figure PF2KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	82	66	62	59	52	46
Irbesartan	80	63	54	47	40	32

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KPSDT\_FSTM

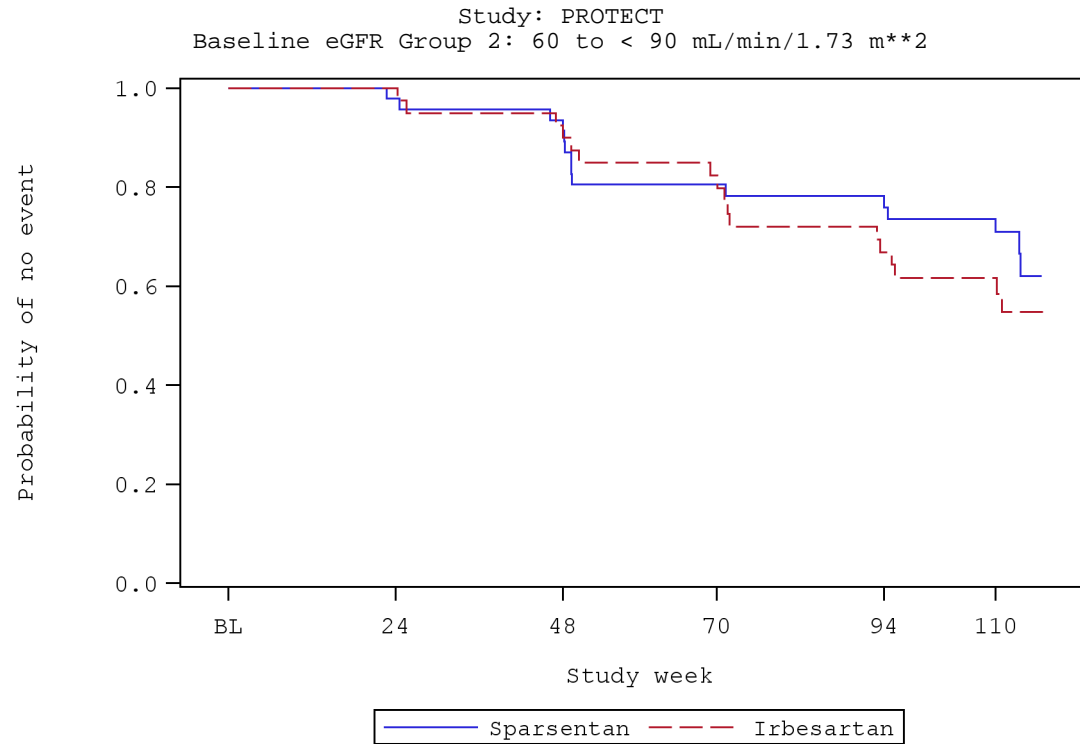
Figure PF2KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	45	42	36	33	29	22
Irbesartan	49	38	33	32	31	28

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KPSDT\_FSTM

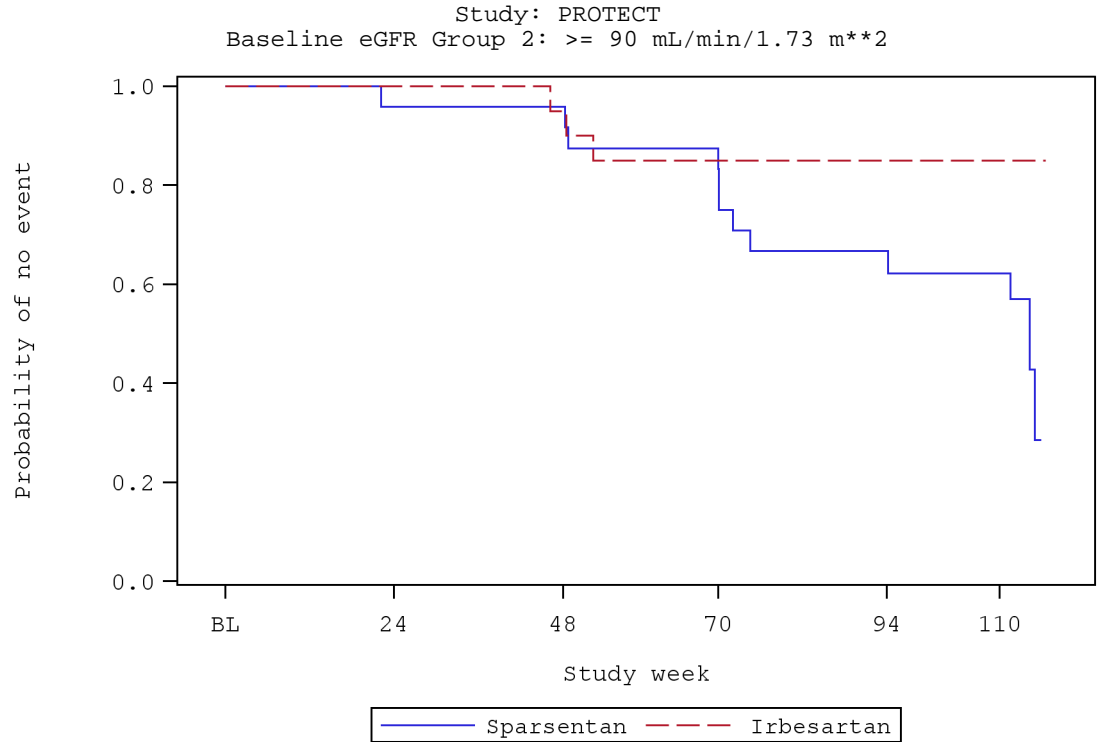
Figure PF2KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	49	46	43	36	34	28
Irbesartan	48	41	37	32	26	20

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KPSDT\_FSTM

Figure PF2KPSDT\_FSKM: KDQOL-SF12: Time to decrease in PCS by at least 8.4 points by subgroup  
 Full Analysis Set



Sparsentan	26	23	23	21	15	13
Irbesartan	25	21	19	17	16	14

A decrease reflects a worsening of the status of the patient.  
 Reference table: PT2KPSDT\_FSTM

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Sex														
Male	KDQOL-SF12: MCS	Baseline	Sparsentan	139	130 (93.5)	50.45 (7.97)	30.1	45.01	51.54	57.06	66.5			
			Irbesartan	143	132 (92.3)	51.61 (8.91)	24.3	48.28	53.36	57.16	66.6			
		Week 24	Sparsentan	139	110 (79.1)	51.41 (8.17)	20.8	47.15	52.23	57.16	68.7			
			Irbesartan	143	100 (69.9)	52.19 (8.39)	24.2	48.37	54.62	57.76	64.2			
		Week 48	Sparsentan	139	115 (82.7)	51.03 (8.39)	27.5	46.56	52.62	57.16	64.2			
			Irbesartan	143	90 (62.9)	51.71 (8.45)	28.5	47.59	54.20	57.16	64.6			
		Week 70	Sparsentan	139	117 (84.2)	50.84 (8.25)	25.4	45.40	52.87	57.16	65.5			
			Irbesartan	143	97 (67.8)	51.19 (8.51)	18.9	46.98	52.54	57.16	63.8			
		Week 94	Sparsentan	139	108 (77.7)	50.37 (8.83)	26.5	43.72	53.62	57.11	66.9			
			Irbesartan	143	99 (69.2)	50.05 (10.02)	21.9	43.32	54.20	57.53	63.4			
		Week 110	Sparsentan	139	104 (74.8)	50.79 (8.39)	20.8	45.43	51.31	57.16	68.0			
			Irbesartan	143	89 (62.2)	49.07 (10.05)	20.0	43.59	50.20	57.16	62.7			
			KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	139	110 (79.1)	0.52 (7.77)	-19.1	-4.29	0.00	4.84	25.3	-0.03 [-0.30, 0.24]
					Irbesartan	143	100 (69.9)	0.77 (6.94)	-15.4	-3.40	0.00	4.39	28.7	
	Week 48	Sparsentan		139	115 (82.7)	0.51 (7.75)	-20.8	-4.67	1.34	5.68	18.7	0.15 [-0.13, 0.43]		
		Irbesartan		143	90 (62.9)	-0.56 (6.49)	-17.3	-3.88	0.00	3.11	17.3			
	Week 70	Sparsentan		139	117 (84.2)	0.52 (8.06)	-21.8	-3.95	0.39	5.23	22.1	0.11 [-0.16, 0.38]		
		Irbesartan		143	97 (67.8)	-0.29 (6.89)	-27.2	-3.95	-0.10	3.64	17.4			
	Week 94	Sparsentan	139	108 (77.7)	-0.04 (7.60)	-22.4	-4.62	0.81	4.93	16.9	0.15 [-0.12, 0.43]			
		Irbesartan	143	99 (69.2)	-1.21 (7.69)	-21.0	-5.33	-0.84	2.72	23.3				
Week 110	Sparsentan	139	104 (74.8)	0.08 (7.71)	-17.6	-3.95	0.00	4.74	22.2	0.26 [-0.02, 0.55]				
	Irbesartan	143	89 (62.2)	-1.97 (8.02)	-29.3	-6.43	-2.20	2.07	19.1					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Female	KDQOL-SF12: MCS	Baseline	Sparsentan	63	58 (92.1)	50.37 (9.08)	16.2	44.79	52.28	56.44	65.2	
			Irbesartan	59	51 (86.4)	50.92 (7.92)	31.5	45.74	53.14	57.16	61.3	
		Week 24	Sparsentan	63	49 (77.8)	52.81 (7.44)	30.7	48.65	54.20	58.26	63.5	
			Irbesartan	59	38 (64.4)	52.76 (7.73)	32.5	48.87	54.29	57.66	62.6	
		Week 48	Sparsentan	63	50 (79.4)	50.97 (8.73)	27.5	44.18	51.75	59.31	64.7	
			Irbesartan	59	37 (62.7)	47.44 (11.50)	19.6	39.39	50.48	56.11	64.5	
		Week 70	Sparsentan	63	46 (73.0)	50.56 (8.64)	31.6	43.46	52.70	57.56	62.7	
			Irbesartan	59	38 (64.4)	51.06 (6.98)	29.2	48.81	52.32	55.51	62.1	
		Week 94	Sparsentan	63	47 (74.6)	50.54 (10.26)	16.0	45.18	51.83	57.94	64.7	
			Irbesartan	59	34 (57.6)	49.51 (9.42)	28.3	42.22	51.91	57.73	61.5	
	Week 110	Sparsentan	63	46 (73.0)	48.34 (9.73)	20.1	42.30	50.10	56.49	62.5		
		Irbesartan	59	32 (54.2)	50.04 (8.15)	32.4	43.58	52.18	56.91	60.9		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	63	49 (77.8)	2.01 (8.74)	-23.6	-1.40	0.61	7.30	26.4	0.13 [-0.30, 0.55]
			Irbesartan	59	38 (64.4)	1.02 (5.90)	-9.4	-2.96	1.32	3.95	20.2	
		Week 48	Sparsentan	63	50 (79.4)	-0.06 (10.26)	-23.7	-4.74	0.04	4.38	28.1	0.35 [-0.08, 0.77]
			Irbesartan	59	37 (62.7)	-3.51 (9.59)	-33.9	-6.62	-2.56	3.74	11.3	
		Week 70	Sparsentan	63	46 (73.0)	-0.43 (9.51)	-21.3	-5.39	0.04	3.17	26.5	0.07 [-0.36, 0.50]
			Irbesartan	59	38 (64.4)	-1.01 (6.24)	-18.6	-3.56	-0.45	3.80	9.4	
		Week 94	Sparsentan	63	47 (74.6)	-0.19 (9.89)	-28.8	-6.46	0.04	6.28	24.1	0.25 [-0.20, 0.69]
			Irbesartan	59	34 (57.6)	-2.40 (7.47)	-19.0	-6.22	-0.15	2.34	10.7	
Week 110		Sparsentan	63	46 (73.0)	-2.05 (11.45)	-31.2	-6.43	-2.47	3.42	33.3	-0.04 [-0.49, 0.41]	
		Irbesartan	59	32 (54.2)	-1.66 (6.75)	-16.1	-5.31	-0.77	3.79	10.5		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age														
<= 45 years	KDQOL-SF12: MCS	Baseline	Sparsentan	96	88 (91.7)	48.94 (8.49)	27.8	43.44	50.17	55.34	65.2			
			Irbesartan	99	92 (92.9)	50.34 (9.59)	24.3	44.08	52.45	57.16	64.2			
		Week 24	Sparsentan	96	72 (75.0)	50.21 (8.49)	20.8	45.18	50.62	57.16	63.5			
			Irbesartan	99	70 (70.7)	51.72 (8.88)	24.2	48.30	53.19	57.76	62.7			
		Week 48	Sparsentan	96	76 (79.2)	49.23 (8.82)	27.5	42.71	51.04	56.41	62.5			
			Irbesartan	99	62 (62.6)	49.88 (10.41)	21.5	44.16	51.51	57.16	64.6			
		Week 70	Sparsentan	96	75 (78.1)	49.92 (9.09)	25.4	42.45	52.82	57.16	65.5			
			Irbesartan	99	64 (64.6)	50.60 (8.73)	18.9	46.62	51.89	57.06	63.8			
		Week 94	Sparsentan	96	70 (72.9)	48.47 (9.76)	16.0	41.53	49.49	56.85	66.9			
			Irbesartan	99	64 (64.6)	48.37 (11.25)	21.9	41.83	51.48	57.66	63.4			
		Week 110	Sparsentan	96	70 (72.9)	48.87 (9.14)	20.1	42.67	49.92	56.27	67.0			
			Irbesartan	99	59 (59.6)	49.67 (9.74)	20.0	43.59	50.82	57.16	62.7			
			KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	96	72 (75.0)	0.59 (8.77)	-23.6	-4.30	0.44	5.35	26.4	-0.05 [-0.38, 0.28]
					Irbesartan	99	70 (70.7)	1.01 (7.00)	-14.1	-2.96	0.68	5.23	21.7	
		Week 48	Sparsentan	96	76 (79.2)	0.06 (8.89)	-23.7	-4.74	0.42	5.33	21.1	0.22 [-0.11, 0.56]		
			Irbesartan	99	62 (62.6)	-1.78 (7.25)	-23.5	-5.33	-0.35	2.43	13.4			
		Week 70	Sparsentan	96	75 (78.1)	0.77 (8.52)	-21.3	-4.37	0.97	5.22	26.4	0.23 [-0.10, 0.56]		
			Irbesartan	99	64 (64.6)	-0.96 (6.15)	-21.2	-5.33	-0.28	4.03	10.4			
		Week 94	Sparsentan	96	70 (72.9)	-0.60 (9.27)	-28.8	-5.53	-0.46	4.94	24.1	0.24 [-0.10, 0.58]		
			Irbesartan	99	64 (64.6)	-2.67 (8.05)	-21.0	-6.24	-2.96	2.31	16.1			
		Week 110	Sparsentan	96	70 (72.9)	-0.32 (9.30)	-24.6	-5.16	-0.11	4.18	26.4	0.15 [-0.20, 0.50]		
			Irbesartan	99	59 (59.6)	-1.61 (7.75)	-21.2	-7.53	-2.25	3.00	14.2			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 45 years	KDQOL-SF12: MCS	Baseline	Sparsentan	106	100 (94.3)	51.73 (7.95)	16.2	48.28	53.38	57.16	66.5		
			Irbesartan	103	91 (88.3)	52.51 (7.44)	26.4	49.06	53.45	57.16	66.6		
		Week 24	Sparsentan	106	87 (82.1)	53.19 (7.26)	34.3	48.65	54.37	57.47	68.7		
			Irbesartan	103	68 (66.0)	52.98 (7.43)	27.2	49.46	54.62	57.33	64.2		
		Week 48	Sparsentan	106	89 (84.0)	52.54 (7.89)	29.5	49.35	53.51	58.32	64.7		
			Irbesartan	103	65 (63.1)	51.01 (8.79)	19.6	47.58	53.63	57.06	64.5		
		Week 70	Sparsentan	106	88 (83.0)	51.47 (7.61)	27.6	46.05	52.75	57.16	65.4		
			Irbesartan	103	71 (68.9)	51.66 (7.49)	29.2	47.34	52.88	57.16	63.4		
		Week 94	Sparsentan	106	85 (80.2)	52.03 (8.54)	28.0	47.79	54.34	57.73	64.7		
			Irbesartan	103	69 (67.0)	51.34 (8.15)	30.5	45.79	54.37	57.33	61.5		
	Week 110	Sparsentan	106	80 (75.5)	51.06 (8.53)	20.8	46.61	51.76	57.24	68.0			
		Irbesartan	103	62 (60.2)	49.00 (9.46)	23.2	42.38	50.63	57.06	62.7			
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	106	87 (82.1)	1.31 (7.50)	-18.1	-2.79	0.37	4.80	25.9	0.09 [-0.22, 0.41]
	Irbesartan			103	68 (66.0)	0.65 (6.32)	-15.4	-3.40	0.00	3.83	28.7		
	Week 48		Sparsentan	106	89 (84.0)	0.58 (8.32)	-20.8	-4.58	1.48	5.23	28.1	0.20 [-0.12, 0.52]	
			Irbesartan	103	65 (63.1)	-1.08 (7.97)	-33.9	-3.76	-0.10	4.19	17.3		
	Week 70		Sparsentan	106	88 (83.0)	-0.19 (8.46)	-21.8	-4.78	-0.14	4.18	26.5	-0.02 [-0.33, 0.30]	
			Irbesartan	103	71 (68.9)	-0.07 (7.18)	-27.2	-3.03	0.00	3.22	17.4		
Week 94	Sparsentan		106	85 (80.2)	0.34 (7.49)	-22.4	-4.52	0.99	4.92	16.9	0.11 [-0.21, 0.42]		
	Irbesartan		103	69 (67.0)	-0.44 (7.11)	-19.0	-3.64	0.00	2.72	23.3			
Week 110	Sparsentan	106	80 (75.5)	-0.79 (8.85)	-31.2	-5.10	-1.13	4.33	33.3	0.16 [-0.17, 0.49]			
	Irbesartan	103	62 (60.2)	-2.15 (7.67)	-29.3	-4.70	-1.94	2.27	19.1				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024



Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Age at IgAN diagnosis														
<= 18 years	KDQOL-SF12: MCS	Baseline	Sparsentan	9	9 (100.0)	44.94 (11.76)	31.1	35.44	42.40	51.24	62.5			
			Irbesartan	5	5 (100.0)	50.09 (9.56)	34.8	46.63	54.94	56.91	57.2			
		Week 24	Sparsentan	9	5 (55.6)	42.94 (15.49)	20.8	39.14	41.07	51.21	62.5			
			Irbesartan	5	5 (100.0)	47.89 (10.99)	29.7	46.63	50.42	55.54	57.2			
		Week 48	Sparsentan	9	7 (77.8)	46.48 (6.88)	37.2	41.30	44.18	51.24	57.2			
			Irbesartan	5	3 (60.0)	52.01 (4.47)	49.3	49.29	49.58	57.16	57.2			
		Week 70	Sparsentan	9	7 (77.8)	46.88 (7.77)	37.3	39.56	46.14	57.06	57.2			
			Irbesartan	5	4 (80.0)	42.98 (11.96)	28.1	34.69	43.38	51.26	57.1			
		Week 94	Sparsentan	9	5 (55.6)	42.92 (10.92)	32.5	38.00	39.01	44.11	61.0			
			Irbesartan	5	4 (80.0)	39.90 (12.63)	29.6	30.49	36.41	49.32	57.2			
		Week 110	Sparsentan	9	4 (44.4)	48.77 (7.80)	38.4	43.46	49.75	54.08	57.2			
			Irbesartan	5	2 (40.0)	35.53 (12.08)	27.0	26.99	35.53	44.07	44.1			
			KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	9	5 (55.6)	-4.27 (15.99)	-19.1	-11.28	-10.30	-3.26	22.5	-0.18 [-1.42, 1.06]
		Irbesartan			5	5 (100.0)	-2.20 (2.46)	-5.1	-4.51	-1.37	0.00	0.0		
		Week 48		Sparsentan	9	7 (77.8)	1.77 (7.09)	-11.3	0.45	1.73	5.92	12.4	0.95 [-0.47, 2.36]	
				Irbesartan	5	3 (60.0)	-4.33 (3.91)	-7.6	-7.62	-5.36	0.00	0.0		
		Week 70		Sparsentan	9	7 (77.8)	2.17 (7.66)	-5.3	-5.11	4.12	5.82	15.6	1.14 [-0.18, 2.45]	
				Irbesartan	5	4 (80.0)	-5.41 (3.94)	-9.5	-8.10	-6.02	-2.72	-0.1		
Week 94	Sparsentan	9		5 (55.6)	0.65 (7.05)	-4.4	-2.95	-1.48	-0.94	13.0	1.29 [-0.16, 2.73]			
	Irbesartan	5		4 (80.0)	-8.48 (7.17)	-15.3	-14.38	-9.31	-2.58	0.0				
Week 110	Sparsentan	9		4 (44.4)	1.97 (12.53)	-11.5	-7.74	0.97	11.67	17.4	1.04 [-0.76, 2.83]			
	Irbesartan	5		2 (40.0)	-9.34 (2.16)	-10.9	-10.86	-9.34	-7.81	-7.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
> 18 to 40 years	KDQOL-SF12: MCS	Baseline	Sparsentan	102	94 (92.2)	50.15 (7.60)	27.8	44.79	51.32	55.70	65.2	
			Irbesartan	109	100 (91.7)	50.61 (9.27)	24.3	45.26	52.05	57.16	64.2	
		Week 24	Sparsentan	102	80 (78.4)	51.03 (7.40)	30.7	46.63	52.03	57.16	63.5	
			Irbesartan	109	73 (67.0)	52.24 (8.33)	24.2	48.87	54.45	57.76	62.7	
		Week 48	Sparsentan	102	83 (81.4)	49.95 (8.73)	27.5	43.09	51.77	57.06	62.5	
			Irbesartan	109	69 (63.3)	49.96 (10.03)	21.5	44.16	52.00	57.16	62.5	
		Week 70	Sparsentan	102	80 (78.4)	50.05 (8.84)	25.4	42.76	52.96	57.11	65.5	
			Irbesartan	109	71 (65.1)	50.93 (8.14)	18.9	46.63	52.09	57.06	62.4	
		Week 94	Sparsentan	102	76 (74.5)	49.28 (9.44)	16.0	43.23	51.75	56.95	66.9	
			Irbesartan	109	69 (63.3)	49.73 (10.62)	21.9	43.13	54.20	57.60	63.4	
	Week 110	Sparsentan	102	73 (71.6)	49.06 (9.09)	20.1	43.65	50.03	56.27	67.0		
		Irbesartan	109	65 (59.6)	50.15 (9.38)	20.0	44.45	51.05	57.16	62.7		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	102	80 (78.4)	0.58 (7.54)	-23.6	-3.61	0.00	4.96	26.4	-0.13 [-0.44, 0.19]
			Irbesartan	109	73 (67.0)	1.51 (7.22)	-14.1	-2.96	0.99	5.33	21.7	
		Week 48	Sparsentan	102	83 (81.4)	-0.19 (9.03)	-23.7	-5.08	0.00	5.23	21.1	0.17 [-0.15, 0.49]
			Irbesartan	109	69 (63.3)	-1.56 (7.07)	-23.5	-3.88	-0.10	2.43	13.4	
		Week 70	Sparsentan	102	80 (78.4)	-0.14 (8.87)	-21.8	-4.71	0.48	4.78	26.4	0.11 [-0.21, 0.43]
			Irbesartan	109	71 (65.1)	-1.02 (6.95)	-27.2	-4.16	-0.64	4.87	10.6	
		Week 94	Sparsentan	102	76 (74.5)	-0.72 (9.56)	-28.8	-6.19	0.52	5.25	24.1	0.09 [-0.23, 0.42]
			Irbesartan	109	69 (63.3)	-1.52 (7.63)	-21.0	-5.31	-0.41	2.96	16.1	
Week 110		Sparsentan	102	73 (71.6)	-0.64 (9.10)	-24.6	-5.33	-0.99	3.54	26.4	0.06 [-0.27, 0.40]	
		Irbesartan	109	65 (59.6)	-1.18 (7.38)	-21.2	-5.58	-1.14	2.75	14.2		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 40 years	KDQOL-SF12: MCS	Baseline	Sparsentan	91	85 (93.4)	51.32 (8.50)	16.2	46.65	53.37	57.16	66.5		
			Irbesartan	88	78 (88.6)	52.53 (7.66)	26.4	48.76	54.19	57.57	66.6		
		Week 24	Sparsentan	91	74 (81.3)	53.31 (7.49)	34.3	49.11	54.94	57.90	68.7		
			Irbesartan	88	60 (68.2)	52.84 (7.82)	27.2	48.57	54.29	57.46	64.2		
		Week 48	Sparsentan	91	75 (82.4)	52.61 (8.04)	29.5	48.77	54.20	58.72	64.7		
			Irbesartan	88	55 (62.5)	51.00 (9.30)	19.6	47.27	54.20	57.06	64.6		
		Week 70	Sparsentan	91	76 (83.5)	51.86 (7.74)	27.6	46.05	53.38	57.33	65.4		
			Irbesartan	88	60 (68.2)	51.96 (7.58)	29.2	47.81	52.75	57.16	63.8		
		Week 94	Sparsentan	91	74 (81.3)	52.10 (8.63)	28.0	47.79	54.27	57.90	64.7		
			Irbesartan	88	60 (68.2)	50.79 (8.43)	28.3	45.09	53.48	57.33	61.5		
	Week 110	Sparsentan	91	73 (80.2)	51.10 (8.67)	20.8	46.05	52.08	57.16	68.0			
		Irbesartan	88	54 (61.4)	48.85 (9.48)	23.2	42.38	50.63	55.94	62.7			
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	91	74 (81.3)	1.77 (7.95)	-18.1	-2.72	1.05	5.56	25.9	0.21 [-0.13, 0.55]
	Irbesartan			88	60 (68.2)	0.26 (6.09)	-15.4	-3.40	0.13	3.23	28.7		
	Week 48		Sparsentan	91	75 (82.4)	0.80 (8.21)	-17.5	-4.58	1.89	5.92	28.1	0.23 [-0.12, 0.58]	
			Irbesartan	88	55 (62.5)	-1.09 (8.41)	-33.9	-3.95	0.00	4.87	17.3		
	Week 70		Sparsentan	91	76 (83.5)	0.48 (8.19)	-17.1	-4.39	-0.14	5.13	26.5	0.00 [-0.34, 0.34]	
			Irbesartan	88	60 (68.2)	0.46 (6.40)	-18.6	-2.63	0.18	3.06	17.4		
	Week 94		Sparsentan	91	74 (81.3)	0.51 (6.99)	-15.5	-4.25	0.96	4.91	16.9	0.21 [-0.13, 0.56]	
			Irbesartan	88	60 (68.2)	-1.04 (7.54)	-19.0	-5.75	-0.09	2.46	23.3		
Week 110	Sparsentan		91	73 (80.2)	-0.65 (8.90)	-31.2	-4.29	-0.72	4.55	33.3	0.21 [-0.14, 0.56]		
	Irbesartan		88	54 (61.4)	-2.46 (8.04)	-29.3	-5.43	-2.06	0.87	19.1			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Geographic region													
North America	KDQOL-SF12: MCS	Baseline	Sparsentan	35	32 (91.4)	49.84 (8.24)	32.9	44.11	51.11	56.51	62.6		
			Irbesartan	46	43 (93.5)	53.85 (6.93)	35.7	48.11	56.91	58.15	64.2		
		Week 24	Sparsentan	35	23 (65.7)	52.94 (6.13)	40.6	49.25	54.20	57.16	68.7		
			Irbesartan	46	37 (80.4)	54.43 (7.66)	24.2	51.25	57.06	58.15	63.2		
		Week 48	Sparsentan	35	25 (71.4)	51.23 (9.62)	30.2	46.56	53.02	57.06	64.2		
			Irbesartan	46	33 (71.7)	50.23 (11.03)	19.6	46.91	54.69	57.16	62.4		
		Week 70	Sparsentan	35	23 (65.7)	50.66 (7.82)	35.5	45.99	52.62	57.16	63.5		
			Irbesartan	46	31 (67.4)	52.86 (6.47)	39.7	48.92	54.37	57.16	62.4		
		Week 94	Sparsentan	35	24 (68.6)	48.95 (10.33)	16.0	42.33	50.58	57.11	62.0		
			Irbesartan	46	30 (65.2)	51.08 (8.68)	31.8	45.91	53.45	57.34	63.4		
		Week 110	Sparsentan	35	21 (60.0)	51.15 (11.11)	20.1	46.13	54.37	57.16	68.0		
			Irbesartan	46	31 (67.4)	50.57 (8.91)	30.5	41.23	52.77	57.49	62.4		
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	35	23 (65.7)	1.40 (7.19)	-19.1	-0.99	0.00	5.33	14.0	0.21 [-0.32, 0.73]
			Irbesartan	46	37 (80.4)	0.12 (5.42)	-14.1	-2.54	0.27	2.45	13.0		
	Week 48		Sparsentan	35	25 (71.4)	1.22 (7.98)	-23.4	-1.59	1.14	6.04	16.3	0.51 [-0.02, 1.03]	
			Irbesartan	46	33 (71.7)	-2.97 (8.50)	-33.9	-5.33	-0.89	0.53	12.1		
	Week 70		Sparsentan	35	23 (65.7)	-0.11 (8.27)	-16.0	-5.33	-0.10	5.33	22.1	0.18 [-0.36, 0.72]	
			Irbesartan	46	31 (67.4)	-1.33 (5.60)	-21.2	-2.96	-0.10	0.69	10.4		
Week 94	Sparsentan	35	24 (68.6)	-1.97 (9.27)	-28.8	-6.45	0.41	2.80	13.6	0.08 [-0.46, 0.62]			
	Irbesartan	46	30 (65.2)	-2.60 (6.61)	-21.0	-5.50	-0.14	0.99	5.9				
Week 110	Sparsentan	35	21 (60.0)	0.15 (8.19)	-24.6	-2.14	0.42	2.96	15.0	0.41 [-0.15, 0.97]			
	Irbesartan	46	31 (67.4)	-3.03 (7.51)	-21.2	-7.59	-2.96	0.59	12.7				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Europe	KDQOL-SF12: MCS	Baseline	Sparsentan	98	89 (90.8)	50.64 (9.24)	16.2	45.43	51.83	57.16	66.5	
			Irbesartan	115	99 (86.1)	50.45 (9.15)	26.4	46.56	52.31	57.16	64.6	
		Week 24	Sparsentan	98	75 (76.5)	50.77 (8.57)	20.8	46.59	50.17	57.24	63.5	
			Irbesartan	115	65 (56.5)	51.60 (8.20)	27.2	47.77	52.71	57.73	62.7	
		Week 48	Sparsentan	98	77 (78.6)	50.28 (8.49)	29.5	43.09	51.65	57.16	64.7	
			Irbesartan	115	61 (53.0)	49.94 (9.29)	28.6	44.56	51.83	57.16	64.5	
		Week 70	Sparsentan	98	76 (77.6)	49.72 (8.14)	27.6	43.53	50.64	56.48	62.7	
			Irbesartan	115	72 (62.6)	49.66 (8.75)	18.9	45.42	51.02	56.31	63.4	
		Week 94	Sparsentan	98	71 (72.4)	50.10 (9.45)	30.9	43.32	52.92	57.73	64.7	
			Irbesartan	115	72 (62.6)	48.37 (10.78)	21.9	41.83	50.72	57.56	61.5	
	Week 110	Sparsentan	98	68 (69.4)	49.54 (8.40)	20.8	45.39	50.11	55.86	62.7		
		Irbesartan	115	61 (53.0)	47.92 (10.58)	20.0	42.26	49.61	57.06	62.7		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	98	75 (76.5)	-0.17 (7.44)	-17.0	-4.54	-0.17	3.27	25.9	-0.22 [-0.55, 0.12]
			Irbesartan	115	65 (56.5)	1.50 (7.91)	-15.4	-4.49	0.00	5.33	28.7	
		Week 48	Sparsentan	98	77 (78.6)	-0.87 (8.46)	-20.8	-5.18	0.00	4.34	28.1	0.04 [-0.30, 0.37]
			Irbesartan	115	61 (53.0)	-1.17 (7.34)	-23.5	-3.95	-0.74	3.45	17.3	
		Week 70	Sparsentan	98	76 (77.6)	-0.75 (7.87)	-21.3	-4.21	-0.16	3.37	26.5	-0.04 [-0.36, 0.28]
			Irbesartan	115	72 (62.6)	-0.46 (7.60)	-27.2	-4.60	-1.06	5.36	17.4	
		Week 94	Sparsentan	98	71 (72.4)	-0.37 (7.97)	-22.4	-5.33	0.00	4.92	15.7	0.12 [-0.21, 0.44]
			Irbesartan	115	72 (62.6)	-1.29 (8.02)	-17.4	-5.34	-1.75	2.72	23.3	
Week 110		Sparsentan	98	68 (69.4)	-1.01 (9.55)	-31.2	-6.66	-1.39	4.15	33.3	0.04 [-0.31, 0.38]	
		Irbesartan	115	61 (53.0)	-1.35 (8.23)	-29.3	-5.43	-0.37	3.48	19.1		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
Asia Pacific	KDQOL-SF12: MCS	Baseline	Sparsentan	69	67 (97.1)	50.42 (7.02)	27.8	45.01	51.25	55.70	64.2	
			Irbesartan	41	41 (100.0)	51.19 (8.65)	24.3	49.04	51.77	57.16	66.6	
		Week 24	Sparsentan	69	61 (88.4)	52.75 (7.72)	30.7	47.94	54.20	57.35	66.8	
			Irbesartan	41	36 (87.8)	51.53 (8.55)	30.8	47.62	52.00	57.24	64.2	
		Week 48	Sparsentan	69	63 (91.3)	51.81 (8.00)	27.5	48.28	52.60	57.33	62.4	
			Irbesartan	41	33 (80.5)	51.66 (8.76)	21.5	47.59	53.02	57.16	64.6	
		Week 70	Sparsentan	69	64 (92.8)	52.03 (8.69)	25.4	46.00	54.32	57.96	65.5	
			Irbesartan	41	32 (78.0)	52.87 (7.46)	33.9	50.83	54.32	58.47	63.8	
		Week 94	Sparsentan	69	60 (87.0)	51.40 (8.60)	26.5	47.75	54.20	57.16	66.9	
			Irbesartan	41	31 (75.6)	52.36 (8.03)	31.9	49.76	54.94	57.97	61.1	
	Week 110	Sparsentan	69	61 (88.4)	50.22 (8.62)	28.3	43.83	50.69	57.16	67.0		
		Irbesartan	41	29 (70.7)	50.96 (7.61)	32.5	47.72	51.14	57.06	62.5		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	69	61 (88.4)	2.23 (9.04)	-23.6	-2.29	1.28	7.00	26.4	0.24 [-0.17, 0.65]
			Irbesartan	41	36 (87.8)	0.36 (5.20)	-9.4	-2.77	0.18	3.36	15.5	
		Week 48	Sparsentan	69	63 (91.3)	1.46 (8.85)	-23.7	-4.61	1.97	7.21	21.1	0.22 [-0.20, 0.64]
			Irbesartan	41	33 (80.5)	-0.35 (7.12)	-18.5	-3.09	0.62	4.92	12.6	
		Week 70	Sparsentan	69	64 (92.8)	1.57 (9.16)	-21.8	-4.39	0.77	7.04	26.4	0.16 [-0.26, 0.59]
			Irbesartan	41	32 (78.0)	0.26 (5.47)	-10.7	-2.19	1.03	3.81	10.4	
		Week 94	Sparsentan	69	60 (87.0)	1.01 (8.32)	-16.8	-4.40	1.11	5.79	24.1	0.24 [-0.19, 0.68]
			Irbesartan	41	31 (75.6)	-0.97 (7.74)	-19.0	-6.26	0.00	4.86	14.5	
Week 110		Sparsentan	69	61 (88.4)	-0.33 (8.84)	-23.0	-4.08	-0.73	4.55	26.4	0.18 [-0.27, 0.62]	
		Irbesartan	41	29 (70.7)	-1.78 (6.72)	-13.3	-6.43	-1.97	0.87	12.9		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline BMI														
< 27 kg/m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	83	75 (90.4)	48.54 (9.00)	16.2	43.23	50.35	54.50	64.6			
			Irbesartan	94	86 (91.5)	51.32 (8.05)	27.9	48.11	52.55	57.16	62.7			
		Week 24	Sparsentan	83	66 (79.5)	50.04 (9.49)	20.8	42.93	50.09	57.16	66.8			
			Irbesartan	94	69 (73.4)	52.06 (8.79)	24.2	48.30	54.45	57.76	62.7			
		Week 48	Sparsentan	83	68 (81.9)	49.31 (9.39)	27.5	42.61	51.33	57.08	64.7			
			Irbesartan	94	60 (63.8)	49.52 (10.04)	19.6	45.36	51.02	57.11	64.5			
		Week 70	Sparsentan	83	66 (79.5)	49.22 (9.62)	25.4	41.23	52.38	57.16	65.4			
			Irbesartan	94	62 (66.0)	50.66 (8.16)	28.1	45.46	52.01	57.06	62.7			
		Week 94	Sparsentan	83	64 (77.1)	48.90 (10.07)	26.5	41.17	51.75	56.03	66.9			
			Irbesartan	94	64 (68.1)	50.22 (10.21)	24.5	43.32	54.55	57.73	63.4			
		Week 110	Sparsentan	83	63 (75.9)	48.51 (8.97)	20.8	41.87	48.69	57.06	65.1			
			Irbesartan	94	57 (60.6)	49.11 (9.46)	23.3	42.61	51.05	57.06	62.7			
			KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	83	66 (79.5)	1.25 (9.82)	-23.6	-4.54	-0.16	7.65	26.4	-0.01 [-0.34, 0.33]
					Irbesartan	94	69 (73.4)	1.29 (6.70)	-14.1	-2.96	0.39	5.23	28.7	
Week 48	Sparsentan	83		68 (81.9)	1.01 (9.34)	-23.4	-3.96	0.89	6.45	28.1	0.30 [-0.05, 0.65]			
	Irbesartan	94		60 (63.8)	-1.64 (8.08)	-33.9	-4.95	-1.67	2.79	17.3				
Week 70	Sparsentan	83		66 (79.5)	1.02 (10.13)	-21.8	-5.11	0.72	7.45	26.5	0.18 [-0.17, 0.53]			
	Irbesartan	94		62 (66.0)	-0.53 (6.64)	-27.2	-4.13	0.00	3.22	16.2				
Week 94	Sparsentan	83		64 (77.1)	1.17 (9.00)	-22.4	-3.73	1.30	6.97	24.1	0.23 [-0.12, 0.58]			
	Irbesartan	94		64 (68.1)	-0.79 (7.82)	-19.0	-4.59	-0.12	4.38	16.1				
Week 110	Sparsentan	83		63 (75.9)	0.33 (10.27)	-31.2	-5.37	-0.73	5.23	33.3	0.26 [-0.10, 0.62]			
	Irbesartan	94		57 (60.6)	-2.12 (8.18)	-29.3	-5.92	-2.14	2.12	19.1				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
>= 27 kg/m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	119	113 (95.0)	51.68 (7.59)	31.4	47.07	53.19	57.16	66.5	
			Irbesartan	107	96 (89.7)	51.76 (8.86)	24.3	47.57	54.14	57.45	66.6	
		Week 24	Sparsentan	119	93 (78.2)	53.12 (6.40)	37.4	48.65	54.20	57.35	68.7	
			Irbesartan	107	69 (64.5)	52.63 (7.61)	27.2	48.87	54.20	57.59	64.2	
		Week 48	Sparsentan	119	97 (81.5)	52.21 (7.58)	27.5	48.76	53.02	57.56	64.2	
			Irbesartan	107	67 (62.6)	51.30 (9.17)	28.4	47.01	54.20	57.16	64.6	
		Week 70	Sparsentan	119	97 (81.5)	51.81 (7.20)	35.5	46.11	52.82	57.16	65.5	
			Irbesartan	107	73 (68.2)	51.58 (8.05)	18.9	47.34	53.20	57.06	63.8	
		Week 94	Sparsentan	119	91 (76.5)	51.50 (8.53)	16.0	45.97	54.34	57.73	62.7	
			Irbesartan	107	68 (63.6)	50.03 (8.99)	23.6	42.60	51.28	57.33	61.1	
	Week 110	Sparsentan	119	87 (73.1)	51.15 (8.67)	20.1	45.57	51.74	57.16	68.0		
		Irbesartan	107	63 (58.9)	49.82 (9.51)	20.0	44.45	50.82	57.33	62.7		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	119	93 (78.2)	0.79 (6.63)	-18.1	-2.46	1.11	4.59	22.5	0.06 [-0.25, 0.37]
			Irbesartan	107	69 (64.5)	0.38 (6.62)	-15.4	-3.53	0.00	3.48	20.6	
		Week 48	Sparsentan	119	97 (81.5)	-0.13 (7.99)	-23.7	-4.74	0.99	4.59	18.7	0.14 [-0.17, 0.45]
			Irbesartan	107	67 (62.6)	-1.22 (7.20)	-23.5	-3.53	0.00	4.85	12.1	
		Week 70	Sparsentan	119	97 (81.5)	-0.27 (7.14)	-21.3	-3.95	0.01	3.68	22.1	0.03 [-0.28, 0.33]
			Irbesartan	107	73 (68.2)	-0.46 (6.80)	-21.2	-3.70	-0.17	3.87	17.4	
		Week 94	Sparsentan	119	91 (76.5)	-0.97 (7.74)	-28.8	-5.53	-0.17	3.82	14.4	0.15 [-0.16, 0.47]
			Irbesartan	107	68 (63.6)	-2.14 (7.48)	-21.0	-6.09	-1.16	2.31	23.3	
Week 110		Sparsentan	119	87 (73.1)	-1.23 (8.02)	-24.6	-4.83	-0.72	3.54	22.2	0.07 [-0.26, 0.39]	
		Irbesartan	107	63 (58.9)	-1.76 (7.29)	-21.3	-6.43	-1.91	2.75	12.9		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024



Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Randomization strata														
eGFR Low and UP High	KDQOL-SF12: MCS	Baseline	Sparsentan	71	64 (90.1)	51.84 (7.80)	31.4	45.00	54.20	57.25	64.6			
			Irbesartan	74	64 (86.5)	50.93 (8.84)	26.4	47.48	53.01	56.83	64.6			
		Week 24	Sparsentan	71	54 (76.1)	52.24 (7.56)	36.1	47.15	54.19	57.34	68.7			
			Irbesartan	74	44 (59.5)	52.68 (8.74)	27.2	49.07	56.15	58.23	62.6			
		Week 48	Sparsentan	71	53 (74.6)	51.84 (7.96)	33.7	46.92	54.20	57.33	64.7			
			Irbesartan	74	38 (51.4)	52.17 (10.94)	19.6	49.29	54.79	59.14	64.6			
		Week 70	Sparsentan	71	58 (81.7)	49.70 (7.77)	32.1	44.19	49.48	55.90	65.4			
			Irbesartan	74	41 (55.4)	52.10 (7.28)	28.1	49.02	54.02	56.94	63.8			
		Week 94	Sparsentan	71	52 (73.2)	52.10 (8.07)	32.1	45.80	54.45	57.92	64.7			
			Irbesartan	74	38 (51.4)	52.03 (9.60)	21.9	49.78	55.35	57.53	63.4			
		Week 110	Sparsentan	71	53 (74.6)	49.93 (7.84)	32.3	46.14	50.20	57.06	62.7			
			Irbesartan	74	36 (48.6)	49.16 (10.26)	23.2	45.33	50.51	57.11	62.7			
			KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	71	54 (76.1)	-0.46 (6.51)	-14.5	-4.09	-0.31	2.55	20.8	-0.16 [-0.56, 0.24]
					Irbesartan	74	44 (59.5)	0.61 (6.76)	-15.4	-2.96	0.34	3.27	28.7	
		Week 48	Sparsentan	71	53 (74.6)	-1.01 (7.52)	-20.8	-5.18	-0.37	4.45	16.3	-0.08 [-0.50, 0.33]		
			Irbesartan	74	38 (51.4)	-0.33 (9.00)	-33.9	-3.06	0.08	4.85	17.3			
		Week 70	Sparsentan	71	58 (81.7)	-2.73 (7.25)	-21.8	-7.40	-2.73	2.53	17.0	-0.32 [-0.72, 0.09]		
			Irbesartan	74	41 (55.4)	-0.40 (7.55)	-27.2	-3.13	0.00	2.00	17.4			
		Week 94	Sparsentan	71	52 (73.2)	-0.29 (7.81)	-22.4	-5.12	1.41	3.59	13.6	-0.17 [-0.59, 0.25]		
			Irbesartan	74	38 (51.4)	0.96 (7.10)	-17.6	-3.38	0.66	4.56	23.3			
		Week 110	Sparsentan	71	53 (74.6)	-2.22 (7.81)	-31.2	-5.37	-1.63	2.57	13.8	-0.14 [-0.57, 0.28]		
			Irbesartan	74	36 (48.6)	-1.19 (6.42)	-9.9	-5.31	-2.48	0.35	19.1			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR Low and UP Low	KDQOL-SF12: MCS	Baseline	Sparsentan	55	52 (94.5)	50.44 (9.63)	16.2	45.35	53.09	57.19	65.2	
			Irbesartan	55	50 (90.9)	52.45 (7.68)	31.9	49.36	52.36	57.73	66.6	
		Week 24	Sparsentan	55	44 (80.0)	52.68 (7.95)	34.3	47.52	54.58	58.08	63.9	
			Irbesartan	55	39 (70.9)	52.60 (8.16)	30.8	47.84	54.37	57.66	64.2	
		Week 48	Sparsentan	55	43 (78.2)	50.18 (8.77)	29.5	42.35	52.05	57.16	63.7	
			Irbesartan	55	36 (65.5)	49.38 (9.46)	29.2	42.71	52.89	57.06	62.5	
		Week 70	Sparsentan	55	42 (76.4)	50.98 (8.48)	27.6	44.95	53.34	57.33	64.2	
			Irbesartan	55	40 (72.7)	50.86 (8.16)	29.2	44.94	52.61	57.06	62.7	
		Week 94	Sparsentan	55	42 (76.4)	50.24 (10.63)	16.0	44.54	54.03	57.16	66.9	
			Irbesartan	55	40 (72.7)	49.10 (10.55)	23.6	40.85	52.13	57.66	61.3	
	Week 110	Sparsentan	55	38 (69.1)	50.54 (11.01)	20.1	43.84	51.04	58.77	68.0		
		Irbesartan	55	34 (61.8)	47.67 (9.60)	23.3	39.15	47.09	56.76	62.5		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	55	44 (80.0)	2.37 (8.05)	-17.0	-2.38	1.14	6.24	25.9	0.29 [-0.14, 0.73]
			Irbesartan	55	39 (70.9)	0.22 (6.50)	-9.4	-5.65	0.00	5.02	20.2	
		Week 48	Sparsentan	55	43 (78.2)	-0.41 (8.87)	-20.3	-5.08	-0.99	5.62	28.1	0.25 [-0.19, 0.69]
			Irbesartan	55	36 (65.5)	-2.53 (8.05)	-23.5	-7.17	-2.92	2.63	12.6	
		Week 70	Sparsentan	55	42 (76.4)	0.92 (8.97)	-21.3	-3.77	0.52	4.63	26.5	0.22 [-0.22, 0.65]
			Irbesartan	55	40 (72.7)	-0.85 (7.20)	-21.2	-3.54	-0.33	3.97	10.6	
		Week 94	Sparsentan	55	42 (76.4)	-0.08 (7.53)	-28.8	-3.72	-0.50	4.38	15.7	0.38 [-0.06, 0.81]
			Irbesartan	55	40 (72.7)	-3.17 (8.84)	-21.0	-7.98	-1.97	2.35	14.5	
Week 110		Sparsentan	55	38 (69.1)	-0.14 (10.09)	-24.6	-4.74	-1.20	5.43	33.3	0.42 [-0.05, 0.89]	
		Irbesartan	55	34 (61.8)	-4.03 (8.31)	-29.3	-6.82	-3.57	1.88	10.5		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP High	KDQOL-SF12: MCS	Baseline	Sparsentan	37	35 (94.6)	48.30 (7.97)	27.8	43.44	49.67	54.20	62.4	
			Irbesartan	36	33 (91.7)	52.02 (8.60)	27.9	46.56	54.45	57.33	64.2	
		Week 24	Sparsentan	37	27 (73.0)	49.15 (8.16)	20.8	46.59	51.83	54.37	57.3	
			Irbesartan	36	26 (72.2)	52.61 (7.05)	36.7	48.43	53.54	57.76	62.5	
		Week 48	Sparsentan	37	34 (91.9)	49.58 (8.82)	27.5	43.09	51.14	57.16	62.5	
			Irbesartan	36	25 (69.4)	49.95 (9.55)	28.4	47.01	51.01	57.06	62.4	
		Week 70	Sparsentan	37	31 (83.8)	49.86 (7.62)	35.9	43.60	51.83	56.60	65.5	
			Irbesartan	36	25 (69.4)	50.60 (8.63)	33.9	46.54	51.94	57.16	63.4	
		Week 94	Sparsentan	37	27 (73.0)	48.32 (7.67)	30.9	44.11	49.52	55.02	60.1	
			Irbesartan	36	25 (69.4)	50.11 (8.59)	24.5	45.91	52.85	57.16	59.5	
	Week 110	Sparsentan	37	26 (70.3)	48.72 (6.67)	28.3	46.08	50.45	53.78	57.3		
		Irbesartan	36	22 (61.1)	51.87 (7.70)	32.5	46.39	53.97	57.33	62.5		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	37	27 (73.0)	0.54 (9.21)	-19.1	-5.23	0.00	5.54	26.4	-0.09 [-0.63, 0.45]
			Irbesartan	36	26 (72.2)	1.26 (6.90)	-13.4	-2.54	0.51	3.77	21.7	
		Week 48	Sparsentan	37	34 (91.9)	1.63 (10.31)	-23.7	-2.43	2.70	7.36	21.1	0.28 [-0.24, 0.80]
			Irbesartan	36	25 (69.4)	-0.82 (6.18)	-15.3	-3.76	0.00	4.94	6.9	
		Week 70	Sparsentan	37	31 (83.8)	2.23 (8.30)	-12.5	-2.96	0.09	5.23	26.4	0.34 [-0.19, 0.88]
			Irbesartan	36	25 (69.4)	-0.31 (5.97)	-10.7	-3.70	-0.10	5.14	8.9	
		Week 94	Sparsentan	37	27 (73.0)	1.18 (10.07)	-14.5	-9.39	3.66	7.97	24.1	0.31 [-0.24, 0.85]
			Irbesartan	36	25 (69.4)	-1.36 (5.87)	-13.5	-5.86	-1.18	1.55	9.9	
Week 110		Sparsentan	37	26 (70.3)	0.84 (9.62)	-23.0	-5.33	0.95	4.11	26.4	0.12 [-0.45, 0.68]	
		Irbesartan	36	22 (61.1)	-0.20 (8.37)	-16.1	-5.58	0.03	5.68	12.9		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
eGFR High and UP Low	KDQOL-SF12: MCS	Baseline	Sparsentan	39	37 (94.9)	49.98 (7.21)	30.1	46.33	50.94	54.63	66.5	
			Irbesartan	37	36 (97.3)	50.30 (9.63)	24.3	45.44	51.94	57.16	62.5	
		Week 24	Sparsentan	39	34 (87.2)	52.25 (8.31)	30.7	48.65	52.32	58.72	66.8	
			Irbesartan	37	29 (78.4)	51.24 (8.65)	24.2	47.77	52.23	57.16	62.5	
		Week 48	Sparsentan	39	35 (89.7)	52.17 (8.55)	27.5	49.43	54.79	58.72	62.5	
			Irbesartan	37	28 (75.7)	50.00 (7.90)	28.6	44.36	51.55	55.51	62.5	
		Week 70	Sparsentan	39	32 (82.1)	53.25 (9.56)	25.4	50.37	57.07	60.07	62.7	
			Irbesartan	37	29 (78.4)	50.72 (8.86)	18.9	47.10	51.67	57.06	61.0	
		Week 94	Sparsentan	39	34 (87.2)	49.77 (10.17)	26.5	42.72	52.56	57.73	63.2	
			Irbesartan	37	30 (81.1)	48.14 (10.12)	24.5	41.17	48.77	57.92	60.1	
	Week 110	Sparsentan	39	33 (84.6)	50.70 (9.43)	30.0	43.83	53.39	58.47	64.0		
		Irbesartan	37	29 (78.4)	49.55 (9.93)	20.0	44.26	51.14	57.16	62.7		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	39	34 (87.2)	1.82 (9.32)	-23.6	-2.72	1.74	7.65	25.3	0.02 [-0.47, 0.52]
			Irbesartan	37	29 (78.4)	1.63 (6.72)	-13.9	-1.32	0.00	4.80	20.6	
		Week 48	Sparsentan	39	35 (89.7)	2.04 (7.66)	-18.6	-0.27	1.97	5.23	19.4	0.58 [0.07, 1.09]
			Irbesartan	37	28 (75.7)	-2.02 (6.08)	-17.3	-5.06	-1.01	0.62	13.4	
		Week 70	Sparsentan	39	32 (82.1)	2.87 (8.78)	-20.3	-3.20	4.13	8.25	17.0	0.43 [-0.08, 0.93]
			Irbesartan	37	29 (78.4)	-0.28 (5.51)	-10.6	-4.26	-0.51	3.87	10.4	
		Week 94	Sparsentan	39	34 (87.2)	-0.78 (8.77)	-16.8	-7.82	-0.46	5.23	16.9	0.22 [-0.27, 0.71]
			Irbesartan	37	30 (81.1)	-2.56 (7.35)	-16.7	-6.03	-3.01	1.58	16.1	
Week 110		Sparsentan	39	33 (84.6)	0.46 (9.14)	-15.6	-4.40	-1.41	6.06	22.2	0.23 [-0.27, 0.73]	
		Irbesartan	37	29 (78.4)	-1.51 (7.67)	-21.3	-6.43	-0.10	3.48	12.8		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 1													
< 60 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	127	117 (92.1)	51.41 (8.59)	16.2	46.65	53.71	57.22	65.2		
			Irbesartan	129	115 (89.1)	51.65 (8.12)	26.4	48.11	52.63	57.16	66.6		
		Sparsentan	127	99 (78.0)	52.44 (7.68)	34.3	47.15	54.20	57.65	68.7			
		Irbesartan	129	85 (65.9)	52.45 (8.22)	27.2	48.87	54.45	57.76	64.2			
	Week 24	Sparsentan	127	97 (76.4)	50.95 (8.29)	29.5	44.50	52.05	57.33	64.7			
		Irbesartan	129	76 (58.9)	50.21 (9.94)	19.6	43.74	53.32	57.16	64.5			
	Week 48	Sparsentan	127	101 (79.5)	50.33 (7.81)	27.6	44.73	51.83	57.06	65.4			
		Irbesartan	129	83 (64.3)	51.29 (7.70)	28.1	47.34	52.63	57.06	62.7			
	Week 70	Sparsentan	127	94 (74.0)	51.15 (8.72)	28.0	43.45	54.10	57.33	66.9			
		Irbesartan	129	79 (61.2)	50.84 (9.76)	21.9	42.46	54.94	57.73	63.4			
	Week 94	Sparsentan	127	93 (73.2)	50.61 (8.61)	20.8	45.57	50.99	57.16	68.0			
		Irbesartan	129	72 (55.8)	49.07 (9.81)	23.2	42.50	50.51	57.11	62.7			
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	127	99 (78.0)	0.66 (7.37)	-17.0	-3.26	0.00	3.97	25.9	0.03 [-0.26, 0.32]
				Irbesartan	129	85 (65.9)	0.48 (6.59)	-15.4	-4.36	0.27	4.26	28.7	
			Week 48	Sparsentan	127	97 (76.4)	-1.12 (8.18)	-20.8	-5.18	-0.99	4.34	28.1	0.09 [-0.21, 0.39]
				Irbesartan	129	76 (58.9)	-1.87 (8.69)	-33.9	-4.71	-0.94	3.28	17.3	
			Week 70	Sparsentan	127	101 (79.5)	-1.34 (7.90)	-21.8	-5.43	-1.80	2.92	26.5	-0.09 [-0.38, 0.20]
				Irbesartan	129	83 (64.3)	-0.61 (7.43)	-27.2	-3.70	-0.10	3.64	17.4	
		Week 94	Sparsentan	127	94 (74.0)	-0.51 (7.32)	-22.4	-5.02	0.03	3.53	15.7	0.05 [-0.25, 0.35]	
			Irbesartan	129	79 (61.2)	-0.89 (8.29)	-21.0	-5.31	0.00	4.07	23.3		
		Week 110	Sparsentan	127	93 (73.2)	-1.24 (8.51)	-31.2	-5.16	-1.40	2.96	33.3	0.10 [-0.21, 0.40]	
			Irbesartan	129	72 (55.8)	-2.03 (7.83)	-29.3	-5.39	-2.17	2.19	19.1		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	49	47 (95.9)	48.70 (7.66)	30.1	43.44	48.53	54.37	66.5		
			Irbesartan	48	44 (91.7)	51.20 (9.16)	27.9	44.31	54.27	57.45	64.2		
		Week 24	Sparsentan	49	39 (79.6)	50.15 (9.01)	20.8	47.83	51.83	57.16	66.8		
			Irbesartan	48	33 (68.8)	52.74 (8.14)	24.2	49.52	54.96	57.33	62.7		
		Week 48	Sparsentan	49	45 (91.8)	49.59 (9.53)	27.5	43.19	51.73	57.06	63.4		
			Irbesartan	48	33 (68.8)	51.32 (9.72)	28.6	49.58	51.36	57.16	64.6		
		Week 70	Sparsentan	49	40 (81.6)	50.63 (9.37)	25.4	43.53	54.41	57.19	62.7		
			Irbesartan	48	32 (66.7)	51.85 (7.95)	33.9	47.04	52.87	57.25	63.8		
		Week 94	Sparsentan	49	41 (83.7)	48.19 (11.16)	16.0	40.95	50.63	57.06	62.7		
			Irbesartan	48	35 (72.9)	48.50 (9.92)	24.5	42.73	51.83	57.16	60.1		
		Week 110	Sparsentan	49	36 (73.5)	47.61 (9.86)	20.1	41.91	49.71	54.92	62.5		
			Irbesartan	48	32 (66.7)	49.33 (8.52)	30.5	42.65	50.20	57.11	62.5		
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	49	39 (79.6)	0.67 (9.04)	-23.6	-3.11	0.37	5.54	25.3	-0.03 [-0.49, 0.44]
				Irbesartan	48	33 (68.8)	0.89 (6.37)	-13.9	-1.98	0.00	2.98	21.7	
	Week 48		Sparsentan	49	45 (91.8)	1.14 (8.19)	-23.7	-1.59	2.04	6.32	15.7	0.15 [-0.30, 0.60]	
			Irbesartan	48	33 (68.8)	0.07 (4.90)	-12.1	-3.03	0.59	3.95	6.7		
	Week 70		Sparsentan	49	40 (81.6)	2.33 (7.90)	-17.9	-2.84	2.41	6.54	17.0	0.29 [-0.18, 0.76]	
			Irbesartan	48	32 (66.7)	0.35 (5.27)	-9.5	-3.38	-0.05	5.06	8.9		
	Week 94		Sparsentan	49	41 (83.7)	-0.05 (9.62)	-28.8	-5.53	1.14	5.24	16.9	0.34 [-0.12, 0.79]	
			Irbesartan	48	35 (72.9)	-2.83 (6.20)	-16.2	-6.22	-3.06	0.59	9.4		
Week 110	Sparsentan	49	36 (73.5)	-0.70 (9.25)	-24.6	-4.92	-0.11	4.52	18.4	0.20 [-0.27, 0.68]			
	Irbesartan	48	32 (66.7)	-2.47 (8.10)	-21.3	-7.25	-2.75	2.31	12.9				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	26	24 (92.3)	49.00 (7.58)	27.8	47.06	50.17	53.71	61.5		
			Irbesartan	25	24 (96.0)	50.68 (10.24)	24.3	46.60	53.61	57.51	62.5		
		Week 24	Sparsentan	26	21 (80.8)	52.17 (7.01)	36.1	47.09	52.62	57.16	62.4		
			Irbesartan	25	20 (80.0)	51.21 (8.47)	36.1	46.40	50.33	59.86	62.5		
		Week 48	Sparsentan	26	23 (88.5)	54.06 (6.23)	39.9	50.34	55.02	59.46	62.5		
			Irbesartan	25	18 (72.0)	49.98 (8.15)	28.4	44.16	53.82	54.37	60.0		
		Week 70	Sparsentan	26	22 (84.6)	52.95 (8.73)	31.7	49.11	54.58	60.02	65.5		
			Irbesartan	25	20 (80.0)	49.48 (9.92)	18.9	45.86	51.81	56.81	60.1		
		Week 94	Sparsentan	26	20 (76.9)	51.59 (6.73)	38.5	46.18	51.75	57.16	63.2		
			Irbesartan	25	19 (76.0)	48.65 (10.10)	24.5	43.32	49.52	57.90	60.1		
		Week 110	Sparsentan	26	21 (80.8)	51.71 (7.66)	36.4	47.10	51.66	58.47	64.0		
			Irbesartan	25	17 (68.0)	50.44 (10.78)	20.0	47.77	52.48	57.16	62.7		
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	26	21 (80.8)	3.05 (9.49)	-13.0	-2.62	3.65	9.48	26.4	0.09 [-0.52, 0.71]
			Irbesartan	25	20 (80.0)	2.26 (7.48)	-13.4	-2.71	1.88	5.28	20.6		
	Week 48		Sparsentan	26	23 (88.5)	4.94 (9.36)	-14.6	-0.03	3.68	13.74	21.1	0.87 [0.22, 1.51]	
			Irbesartan	25	18 (72.0)	-2.25 (6.68)	-15.3	-6.63	-2.47	0.00	13.4		
	Week 70		Sparsentan	26	22 (84.6)	3.77 (10.35)	-20.3	-3.69	4.01	11.62	26.4	0.61 [-0.01, 1.22]	
			Irbesartan	25	20 (80.0)	-1.33 (5.58)	-10.6	-5.33	-0.72	1.95	9.2		
	Week 94		Sparsentan	26	20 (76.9)	1.82 (9.99)	-12.7	-6.21	1.87	9.56	24.1	0.40 [-0.23, 1.04]	
			Irbesartan	25	19 (76.0)	-1.68 (7.17)	-15.3	-7.82	-0.14	2.79	16.1		
Week 110	Sparsentan	26	21 (80.8)	2.58 (10.63)	-15.6	-3.98	1.71	7.90	26.4	0.31 [-0.33, 0.95]			
	Irbesartan	25	17 (68.0)	-0.19 (6.25)	-11.7	-2.86	0.00	5.16	12.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
Subgroup: Baseline eGFR Group 2													
< 45 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	82	73 (89.0)	50.19 (8.99)	16.2	44.41	52.00	57.16	64.2		
			Irbesartan	80	70 (87.5)	50.90 (8.98)	26.4	47.81	52.12	57.16	66.6		
		Week 24	Sparsentan	82	64 (78.0)	52.42 (8.03)	34.3	47.00	54.29	57.56	68.7		
			Irbesartan	80	51 (63.8)	52.07 (8.95)	27.2	48.87	54.45	58.17	64.2		
		Week 48	Sparsentan	82	63 (76.8)	50.54 (8.10)	29.5	44.50	51.65	57.16	64.2		
			Irbesartan	80	45 (56.3)	49.71 (10.72)	19.6	43.60	51.65	57.41	64.5		
		Week 70	Sparsentan	82	65 (79.3)	49.81 (8.06)	27.6	44.19	49.62	57.06	63.5		
			Irbesartan	80	50 (62.5)	50.06 (7.97)	28.1	44.95	51.92	55.41	62.4		
		Week 94	Sparsentan	82	61 (74.4)	51.27 (8.83)	28.0	46.14	54.20	57.33	66.9		
			Irbesartan	80	48 (60.0)	50.72 (9.57)	21.9	42.89	53.34	57.53	63.4		
		Week 110	Sparsentan	82	57 (69.5)	50.41 (9.02)	20.8	45.22	51.53	57.16	67.0		
			Irbesartan	80	44 (55.0)	48.47 (10.01)	23.2	42.44	50.51	56.91	62.7		
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	82	64 (78.0)	1.75 (7.44)	-13.0	-2.78	1.20	4.71	25.9	0.18 [-0.19, 0.55]
				Irbesartan	80	51 (63.8)	0.38 (7.56)	-15.4	-4.80	0.00	4.24	28.7	
		Week 48	Sparsentan	82	63 (76.8)	-0.18 (8.38)	-20.8	-5.16	0.00	4.49	28.1	0.19 [-0.19, 0.58]	
			Irbesartan	80	45 (56.3)	-1.96 (10.17)	-33.9	-6.62	-0.11	4.85	17.3		
		Week 70	Sparsentan	82	65 (79.3)	-1.07 (8.21)	-21.8	-5.33	0.01	3.36	26.5	0.03 [-0.34, 0.40]	
			Irbesartan	80	50 (62.5)	-1.33 (7.90)	-27.2	-4.16	-1.40	1.83	17.4		
		Week 94	Sparsentan	82	61 (74.4)	0.66 (6.60)	-22.4	-2.96	0.99	3.82	15.7	0.09 [-0.29, 0.47]	
			Irbesartan	80	48 (60.0)	0.00 (8.11)	-19.0	-5.01	-0.11	3.86	23.3		
		Week 110	Sparsentan	82	57 (69.5)	-0.65 (8.38)	-17.6	-4.74	-1.40	3.54	33.3	0.05 [-0.34, 0.45]	
			Irbesartan	80	44 (55.0)	-1.05 (6.77)	-13.3	-5.39	-2.17	2.87	19.1		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024



Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
45 to < 60 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	45	44 (97.8)	53.44 (7.56)	33.8	49.54	55.60	57.82	65.2	
		Week 24	Irbesartan	49	45 (91.8)	52.83 (6.48)	31.9	49.36	54.15	57.09	64.6	
			Sparsentan	45	35 (77.8)	52.46 (7.12)	40.6	47.77	53.04	57.90	63.9	
		Week 48	Irbesartan	49	34 (69.4)	53.03 (7.08)	33.5	48.43	54.62	57.16	63.2	
			Sparsentan	45	34 (75.6)	51.69 (8.70)	30.2	42.67	52.61	59.31	64.7	
		Week 70	Irbesartan	49	31 (63.3)	50.93 (8.81)	29.2	47.58	54.61	57.06	62.4	
			Sparsentan	45	36 (80.0)	51.28 (7.36)	35.9	45.95	52.28	56.40	65.4	
		Week 94	Irbesartan	49	33 (67.3)	53.16 (6.97)	36.7	50.51	55.36	57.16	62.7	
			Sparsentan	45	33 (73.3)	50.92 (8.62)	36.0	42.90	52.77	57.16	64.7	
		Week 110	Irbesartan	49	31 (63.3)	51.03 (10.20)	23.6	41.34	57.06	57.73	61.3	
	Sparsentan		45	36 (80.0)	50.92 (8.03)	33.4	46.10	49.91	57.16	68.0		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	45	35 (77.8)	-1.32 (6.89)	-17.0	-6.52	-0.77	2.37	14.8	-0.33 [-0.80, 0.15]
			Irbesartan	49	34 (69.4)	0.64 (4.88)	-8.6	-2.96	1.38	4.66	8.9	
		Week 48	Sparsentan	45	34 (75.6)	-2.86 (7.62)	-18.6	-5.50	-2.58	2.70	11.8	-0.16 [-0.65, 0.33]
			Irbesartan	49	31 (63.3)	-1.74 (6.08)	-17.3	-3.53	-1.14	2.45	8.2	
		Week 70	Sparsentan	45	36 (80.0)	-1.82 (7.40)	-17.1	-5.61	-2.27	1.45	15.2	-0.33 [-0.80, 0.15]
			Irbesartan	49	33 (67.3)	0.48 (6.62)	-21.2	-2.86	0.42	4.20	10.6	
		Week 94	Sparsentan	45	33 (73.3)	-2.67 (8.17)	-20.7	-8.51	-0.57	2.18	13.6	-0.05 [-0.54, 0.44]
			Irbesartan	49	31 (63.3)	-2.27 (8.50)	-21.0	-7.31	0.00	4.20	10.1	
		Week 110	Sparsentan	45	36 (80.0)	-2.16 (8.73)	-31.2	-6.70	-1.27	2.50	16.0	0.16 [-0.34, 0.65]
Irbesartan			49	28 (57.1)	-3.56 (9.19)	-29.3	-5.39	-2.37	1.84	10.9		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
60 to < 90 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	49	47 (95.9)	48.70 (7.66)	30.1	43.44	48.53	54.37	66.5		
			Irbesartan	48	44 (91.7)	51.20 (9.16)	27.9	44.31	54.27	57.45	64.2		
		Week 24	Sparsentan	49	39 (79.6)	50.15 (9.01)	20.8	47.83	51.83	57.16	66.8		
			Irbesartan	48	33 (68.8)	52.74 (8.14)	24.2	49.52	54.96	57.33	62.7		
		Week 48	Sparsentan	49	45 (91.8)	49.59 (9.53)	27.5	43.19	51.73	57.06	63.4		
			Irbesartan	48	33 (68.8)	51.32 (9.72)	28.6	49.58	51.36	57.16	64.6		
		Week 70	Sparsentan	49	40 (81.6)	50.63 (9.37)	25.4	43.53	54.41	57.19	62.7		
			Irbesartan	48	32 (66.7)	51.85 (7.95)	33.9	47.04	52.87	57.25	63.8		
		Week 94	Sparsentan	49	41 (83.7)	48.19 (11.16)	16.0	40.95	50.63	57.06	62.7		
			Irbesartan	48	35 (72.9)	48.50 (9.92)	24.5	42.73	51.83	57.16	60.1		
		Week 110	Sparsentan	49	36 (73.5)	47.61 (9.86)	20.1	41.91	49.71	54.92	62.5		
			Irbesartan	48	32 (66.7)	49.33 (8.52)	30.5	42.65	50.20	57.11	62.5		
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	49	39 (79.6)	0.67 (9.04)	-23.6	-3.11	0.37	5.54	25.3	-0.03 [-0.49, 0.44]
				Irbesartan	48	33 (68.8)	0.89 (6.37)	-13.9	-1.98	0.00	2.98	21.7	
	Week 48		Sparsentan	49	45 (91.8)	1.14 (8.19)	-23.7	-1.59	2.04	6.32	15.7	0.15 [-0.30, 0.60]	
			Irbesartan	48	33 (68.8)	0.07 (4.90)	-12.1	-3.03	0.59	3.95	6.7		
	Week 70		Sparsentan	49	40 (81.6)	2.33 (7.90)	-17.9	-2.84	2.41	6.54	17.0	0.29 [-0.18, 0.76]	
			Irbesartan	48	32 (66.7)	0.35 (5.27)	-9.5	-3.38	-0.05	5.06	8.9		
	Week 94		Sparsentan	49	41 (83.7)	-0.05 (9.62)	-28.8	-5.53	1.14	5.24	16.9	0.34 [-0.12, 0.79]	
			Irbesartan	48	35 (72.9)	-2.83 (6.20)	-16.2	-6.22	-3.06	0.59	9.4		
Week 110	Sparsentan	49	36 (73.5)	-0.70 (9.25)	-24.6	-4.92	-0.11	4.52	18.4	0.20 [-0.27, 0.68]			
	Irbesartan	48	32 (66.7)	-2.47 (8.10)	-21.3	-7.25	-2.75	2.31	12.9				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
>= 90 mL/min/1.73 m**2	KDQOL-SF12: MCS	Baseline	Sparsentan	26	24 (92.3)	49.00 (7.58)	27.8	47.06	50.17	53.71	61.5		
			Irbesartan	25	24 (96.0)	50.68 (10.24)	24.3	46.60	53.61	57.51	62.5		
	Week 24	Sparsentan	26	21 (80.8)	52.17 (7.01)	36.1	47.09	52.62	57.16	62.4			
		Irbesartan	25	20 (80.0)	51.21 (8.47)	36.1	46.40	50.33	59.86	62.5			
	Week 48	Sparsentan	26	23 (88.5)	54.06 (6.23)	39.9	50.34	55.02	59.46	62.5			
		Irbesartan	25	18 (72.0)	49.98 (8.15)	28.4	44.16	53.82	54.37	60.0			
	Week 70	Sparsentan	26	22 (84.6)	52.95 (8.73)	31.7	49.11	54.58	60.02	65.5			
		Irbesartan	25	20 (80.0)	49.48 (9.92)	18.9	45.86	51.81	56.81	60.1			
	Week 94	Sparsentan	26	20 (76.9)	51.59 (6.73)	38.5	46.18	51.75	57.16	63.2			
		Irbesartan	25	19 (76.0)	48.65 (10.10)	24.5	43.32	49.52	57.90	60.1			
	Week 110	Sparsentan	26	21 (80.8)	51.71 (7.66)	36.4	47.10	51.66	58.47	64.0			
		Irbesartan	25	17 (68.0)	50.44 (10.78)	20.0	47.77	52.48	57.16	62.7			
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	26	21 (80.8)	3.05 (9.49)	-13.0	-2.62	3.65	9.48	26.4	0.09 [-0.52, 0.71]
			Irbesartan	25	20 (80.0)	2.26 (7.48)	-13.4	-2.71	1.88	5.28	20.6		
	Week 48		Sparsentan	26	23 (88.5)	4.94 (9.36)	-14.6	-0.03	3.68	13.74	21.1	0.87 [0.22, 1.51]	
			Irbesartan	25	18 (72.0)	-2.25 (6.68)	-15.3	-6.63	-2.47	0.00	13.4		
	Week 70		Sparsentan	26	22 (84.6)	3.77 (10.35)	-20.3	-3.69	4.01	11.62	26.4	0.61 [-0.01, 1.22]	
			Irbesartan	25	20 (80.0)	-1.33 (5.58)	-10.6	-5.33	-0.72	1.95	9.2		
	Week 94		Sparsentan	26	20 (76.9)	1.82 (9.99)	-12.7	-6.21	1.87	9.56	24.1	0.40 [-0.23, 1.04]	
			Irbesartan	25	19 (76.0)	-1.68 (7.17)	-15.3	-7.82	-0.14	2.79	16.1		
Week 110	Sparsentan		26	21 (80.8)	2.58 (10.63)	-15.6	-3.98	1.71	7.90	26.4	0.31 [-0.33, 0.95]		
	Irbesartan		25	17 (68.0)	-0.19 (6.25)	-11.7	-2.86	0.00	5.16	12.8			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline urine protein excretion														
<= 1.75 g/day	KDQOL-SF12: MCS	Baseline	Sparsentan	98	93 (94.9)	50.20 (8.75)	16.2	45.91	51.55	56.44	66.5			
			Irbesartan	93	83 (89.2)	50.95 (8.77)	24.3	47.81	52.09	57.16	66.6			
		Week 24	Sparsentan	98	81 (82.7)	51.60 (8.82)	20.8	46.86	52.62	57.90	68.7			
			Irbesartan	93	59 (63.4)	52.10 (8.36)	24.2	47.77	54.20	57.33	64.2			
		Week 48	Sparsentan	98	83 (84.7)	50.58 (9.28)	27.5	44.32	52.05	58.32	64.2			
			Irbesartan	93	57 (61.3)	50.11 (9.11)	19.6	44.56	52.77	57.06	62.5			
		Week 70	Sparsentan	98	77 (78.6)	51.51 (8.58)	25.4	47.07	53.24	57.33	63.5			
			Irbesartan	93	62 (66.7)	50.66 (8.61)	18.9	46.98	52.31	57.06	62.7			
		Week 94	Sparsentan	98	79 (80.6)	49.87 (9.73)	26.5	42.72	51.67	57.73	66.9			
			Irbesartan	93	64 (68.8)	48.76 (10.45)	21.9	41.68	51.80	57.66	61.3			
		Week 110	Sparsentan	98	76 (77.6)	49.82 (9.68)	20.8	43.25	50.09	57.63	68.0			
			Irbesartan	93	57 (61.3)	48.64 (10.10)	20.0	42.62	50.66	57.16	62.7			
			KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	98	81 (82.7)	1.33 (8.38)	-23.6	-2.79	0.87	5.84	25.9	0.08 [-0.25, 0.42]
		Irbesartan			93	59 (63.4)	0.71 (6.13)	-13.9	-3.53	0.00	4.66	20.6		
Week 48	Sparsentan	98		83 (84.7)	0.37 (9.03)	-23.7	-4.58	0.70	5.23	28.1	0.22 [-0.12, 0.55]			
	Irbesartan	93		57 (61.3)	-1.46 (7.61)	-33.9	-3.53	-0.10	2.43	13.4				
Week 70	Sparsentan	98		77 (78.6)	1.37 (8.93)	-21.3	-3.80	1.81	5.39	26.5	0.29 [-0.04, 0.63]			
	Irbesartan	93		62 (66.7)	-0.93 (6.15)	-18.6	-4.26	-0.54	3.87	10.4				
Week 94	Sparsentan	98		79 (80.6)	-0.35 (8.08)	-20.7	-5.56	-0.17	4.94	16.9	0.24 [-0.09, 0.57]			
	Irbesartan	93		64 (68.8)	-2.22 (7.44)	-19.0	-5.49	-2.36	1.94	16.1				
Week 110	Sparsentan	98		76 (77.6)	-0.36 (9.53)	-23.0	-5.59	-1.81	5.33	33.3	0.20 [-0.14, 0.54]			
	Irbesartan	93		57 (61.3)	-2.12 (7.74)	-29.3	-4.85	-0.68	2.75	12.8				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 1.75 g/day	KDQOL-SF12: MCS	Baseline	Sparsentan	104	95 (91.3)	50.65 (7.88)	27.8	44.79	51.83	57.16	64.6		
			Irbesartan	109	100 (91.7)	51.80 (8.54)	26.4	47.17	54.16	57.33	64.6		
	Week 24	Sparsentan	104	78 (75.0)	52.09 (6.99)	36.1	47.15	54.20	57.16	63.9			
		Irbesartan	109	79 (72.5)	52.53 (8.11)	27.2	48.87	54.45	58.15	62.6			
	Week 48	Sparsentan	104	82 (78.8)	51.45 (7.59)	33.7	46.61	52.61	57.33	64.7			
		Irbesartan	109	70 (64.2)	50.75 (10.02)	21.5	46.91	53.54	57.41	64.6			
	Week 70	Sparsentan	104	86 (82.7)	50.09 (8.11)	32.1	44.07	52.23	57.06	65.5			
		Irbesartan	109	73 (67.0)	51.57 (7.65)	28.1	47.45	52.54	57.06	63.8			
	Week 94	Sparsentan	104	76 (73.1)	51.00 (8.75)	16.0	45.84	54.15	57.16	64.7			
		Irbesartan	109	69 (63.3)	50.98 (9.18)	23.6	46.53	54.79	57.34	63.4			
	Week 110	Sparsentan	104	74 (71.2)	50.27 (7.99)	20.1	46.98	51.46	57.06	65.1			
		Irbesartan	109	64 (58.7)	49.94 (9.10)	23.2	44.26	50.93	57.16	62.7			
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	104	78 (75.0)	0.62 (7.80)	-19.1	-3.26	0.00	4.20	26.4	-0.04 [-0.36, 0.27]
	Irbesartan			109	79 (72.5)	0.93 (7.06)	-15.4	-2.96	0.72	4.24	28.7		
	Week 48		Sparsentan	104	82 (78.8)	0.31 (8.12)	-20.8	-4.74	1.06	5.68	21.1	0.21 [-0.11, 0.53]	
			Irbesartan	109	70 (64.2)	-1.39 (7.65)	-23.5	-4.01	-0.42	4.19	17.3		
	Week 70		Sparsentan	104	86 (82.7)	-0.75 (7.96)	-21.8	-5.11	-1.13	3.36	26.4	-0.08 [-0.39, 0.23]	
			Irbesartan	109	73 (67.0)	-0.12 (7.16)	-27.2	-3.56	0.00	2.96	17.4		
	Week 94		Sparsentan	104	76 (73.1)	0.19 (8.62)	-28.8	-4.77	1.41	5.00	24.1	0.13 [-0.20, 0.45]	
			Irbesartan	109	69 (63.3)	-0.85 (7.80)	-21.0	-5.16	0.00	3.66	23.3		
Week 110	Sparsentan		104	74 (71.2)	-0.80 (8.55)	-31.2	-4.83	0.00	2.96	26.4	0.11 [-0.23, 0.44]		
	Irbesartan		109	64 (58.7)	-1.67 (7.68)	-21.2	-6.83	-2.48	2.00	19.1			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Baseline use of antihypertensives														
Yes	KDQOL-SF12: MCS	Baseline	Sparsentan	90	80 (88.9)	49.67 (9.27)	16.2	43.84	51.69	57.16	65.2			
			Irbesartan	88	77 (87.5)	52.31 (7.80)	27.2	48.59	54.15	57.16	66.6			
		Week 24	Sparsentan	90	68 (75.6)	51.15 (8.01)	20.8	47.00	52.03	57.16	63.9			
			Irbesartan	88	52 (59.1)	53.14 (7.55)	30.8	50.30	55.37	58.16	64.2			
		Week 48	Sparsentan	90	68 (75.6)	51.13 (8.02)	29.5	44.25	52.52	57.25	63.7			
			Irbesartan	88	48 (54.5)	51.39 (8.57)	28.5	49.52	53.32	57.11	64.5			
		Week 70	Sparsentan	90	69 (76.7)	50.28 (8.06)	27.6	45.81	51.83	57.06	64.2			
			Irbesartan	88	51 (58.0)	51.18 (7.52)	29.2	48.30	52.63	56.18	62.1			
		Week 94	Sparsentan	90	66 (73.3)	50.64 (9.29)	16.0	44.60	53.62	57.16	63.2			
			Irbesartan	88	57 (64.8)	51.33 (9.57)	21.9	44.47	56.10	57.73	61.5			
		Week 110	Sparsentan	90	64 (71.1)	49.53 (8.76)	20.1	45.51	50.51	56.26	68.0			
			Irbesartan	88	51 (58.0)	49.59 (9.40)	23.3	42.38	51.05	57.16	62.7			
			KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	90	68 (75.6)	0.98 (7.49)	-18.1	-2.91	0.73	4.82	25.9	0.11 [-0.25, 0.47]
		Irbesartan			88	52 (59.1)	0.21 (6.21)	-15.4	-4.23	0.08	3.32	28.7		
Week 48	Sparsentan	90		68 (75.6)	0.79 (8.40)	-17.5	-4.16	1.63	5.33	28.1	0.20 [-0.17, 0.57]			
	Irbesartan	88		48 (54.5)	-0.87 (7.85)	-23.5	-3.74	0.08	3.82	17.3				
Week 70	Sparsentan	90		69 (76.7)	0.08 (8.64)	-21.8	-5.11	-0.14	4.63	26.5	0.16 [-0.20, 0.53]			
	Irbesartan	88		51 (58.0)	-1.16 (5.77)	-18.6	-4.26	-1.54	2.43	16.2				
Week 94	Sparsentan	90		66 (73.3)	0.74 (8.16)	-28.8	-4.25	1.21	5.09	15.7	0.23 [-0.13, 0.58]			
	Irbesartan	88		57 (64.8)	-0.96 (6.67)	-16.7	-5.31	-0.84	2.72	14.5				
Week 110	Sparsentan	90		64 (71.1)	-0.39 (8.98)	-24.6	-5.59	0.00	4.37	33.3	0.12 [-0.25, 0.49]			
	Irbesartan	88		51 (58.0)	-1.46 (8.50)	-29.3	-5.43	-1.91	3.72	19.1				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
No	KDQOL-SF12: MCS	Baseline	Sparsentan	112	108 (96.4)	50.99 (7.51)	27.8	45.98	51.65	56.51	66.5			
			Irbesartan	114	106 (93.0)	50.76 (9.17)	24.3	45.98	52.16	57.16	64.6			
		Week 24	Sparsentan	112	91 (81.3)	52.36 (7.91)	30.7	47.15	54.20	57.65	68.7			
			Irbesartan	114	86 (75.4)	51.86 (8.56)	24.2	47.84	52.47	57.66	63.2			
		Week 48	Sparsentan	112	97 (86.6)	50.93 (8.80)	27.5	46.56	51.73	57.33	64.7			
			Irbesartan	114	79 (69.3)	49.90 (10.18)	19.6	43.89	52.00	57.16	64.6			
		Week 70	Sparsentan	112	94 (83.9)	51.11 (8.56)	25.4	44.26	52.99	57.33	65.5			
			Irbesartan	114	84 (73.7)	51.14 (8.45)	18.9	46.58	52.01	57.16	63.8			
		Week 94	Sparsentan	112	89 (79.5)	50.26 (9.27)	26.5	43.32	51.83	57.16	66.9			
			Irbesartan	114	76 (66.7)	48.84 (9.96)	23.6	41.83	51.20	57.16	63.4			
		Week 110	Sparsentan	112	86 (76.8)	50.42 (8.97)	28.3	43.84	51.59	57.33	67.0			
			Irbesartan	114	70 (61.4)	49.14 (9.74)	20.0	43.59	50.51	57.16	62.7			
		KDQOL-SF12: change from baseline in MCS	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	112	91 (81.3)	0.98 (8.54)	-23.6	-4.29	0.00	5.32	26.4	-0.03 [-0.32, 0.26]
					Irbesartan	114	86 (75.4)	1.22 (6.91)	-14.1	-2.96	0.18	5.23	21.7	
				Week 48	Sparsentan	112	97 (86.6)	0.02 (8.70)	-23.7	-5.14	0.39	5.23	21.1	0.22 [-0.08, 0.52]
					Irbesartan	114	79 (69.3)	-1.76 (7.48)	-33.9	-5.33	-0.60	2.79	13.4	
				Week 70	Sparsentan	112	94 (83.9)	0.37 (8.40)	-21.3	-4.06	0.90	4.97	26.4	0.06 [-0.24, 0.35]
					Irbesartan	114	84 (73.7)	-0.09 (7.21)	-27.2	-3.51	0.00	4.48	17.4	
				Week 94	Sparsentan	112	89 (79.5)	-0.70 (8.44)	-22.4	-5.60	-0.11	4.56	24.1	0.15 [-0.16, 0.45]
					Irbesartan	114	76 (66.7)	-1.93 (8.30)	-21.0	-6.19	-0.29	2.54	23.3	
Week 110	Sparsentan			112	86 (76.8)	-0.71 (9.13)	-31.2	-5.16	-1.11	3.95	26.4	0.18 [-0.14, 0.50]		
	Irbesartan			114	70 (61.4)	-2.19 (7.07)	-21.9	-6.84	-2.06	1.63	12.8			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: Time since renal biopsy														
<= 5 years	KDQOL-SF12: MCS	Baseline	Sparsentan	113	105 (92.9)	50.34 (8.81)	16.2	45.01	52.02	57.16	66.5			
			Irbesartan	127	118 (92.9)	51.30 (9.07)	24.3	46.98	53.77	57.32	66.6			
		Week 24	Sparsentan	113	90 (79.6)	52.35 (8.46)	20.8	47.61	54.20	57.90	68.7			
			Irbesartan	127	91 (71.7)	52.57 (8.44)	24.2	48.30	54.45	58.16	64.2			
		Week 48	Sparsentan	113	93 (82.3)	52.25 (7.78)	30.2	48.76	52.64	58.72	64.7			
			Irbesartan	127	86 (67.7)	50.09 (10.19)	19.6	44.56	52.74	57.16	64.6			
		Week 70	Sparsentan	113	91 (80.5)	51.46 (8.27)	31.6	45.81	53.44	57.16	65.5			
			Irbesartan	127	88 (69.3)	51.67 (7.93)	18.9	48.23	52.31	57.16	63.8			
		Week 94	Sparsentan	113	88 (77.9)	50.43 (9.64)	16.0	43.58	53.39	57.16	66.9			
			Irbesartan	127	88 (69.3)	49.44 (10.27)	21.9	42.04	53.48	57.73	63.4			
		Week 110	Sparsentan	113	88 (77.9)	51.07 (8.41)	20.1	46.85	51.81	57.11	68.0			
			Irbesartan	127	80 (63.0)	50.17 (9.91)	20.0	43.10	52.71	57.25	62.7			
			KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	113	90 (79.6)	1.71 (8.68)	-23.6	-2.96	1.86	6.42	26.4	0.05 [-0.25, 0.34]
				Irbesartan	127	91 (71.7)	1.35 (7.06)	-14.1	-2.88	0.72	5.23	28.7		
Week 48	Sparsentan	113		93 (82.3)	1.79 (8.07)	-18.6	-3.97	2.39	6.48	28.1	0.42 [0.12, 0.71]			
	Irbesartan	127		86 (67.7)	-1.56 (8.01)	-33.9	-4.01	-0.35	3.11	17.3				
Week 70	Sparsentan	113		91 (80.5)	1.05 (9.13)	-20.3	-5.04	0.13	5.66	26.5	0.16 [-0.13, 0.45]			
	Irbesartan	127		88 (69.3)	-0.20 (6.09)	-21.2	-3.51	0.00	3.81	16.2				
Week 94	Sparsentan	113	88 (77.9)	-0.04 (8.74)	-28.8	-5.37	1.02	5.08	24.1	0.26 [-0.04, 0.56]				
	Irbesartan	127	88 (69.3)	-2.20 (7.92)	-21.0	-6.19	-2.52	2.35	23.3					
Week 110	Sparsentan	113	88 (77.9)	0.63 (9.78)	-31.2	-4.02	0.00	5.23	33.3	0.22 [-0.08, 0.53]				
	Irbesartan	127	80 (63.0)	-1.40 (8.29)	-29.3	-5.75	-0.91	3.60	19.1					

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024



Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]	
> 5 years	KDQOL-SF12: MCS	Baseline	Sparsentan	89	83 (93.3)	50.53 (7.66)	30.1	44.79	51.40	56.59	64.2		
			Irbesartan	75	65 (86.7)	51.63 (7.83)	26.4	47.72	52.31	57.16	63.0		
	Week 24	Sparsentan	89	69 (77.5)	51.18 (7.24)	34.3	47.15	52.01	57.16	62.7			
		Irbesartan	75	47 (62.7)	51.90 (7.75)	29.7	48.43	53.00	57.16	62.7			
	Week 48	Sparsentan	89	72 (80.9)	49.42 (9.09)	27.5	42.54	51.68	56.53	63.4			
		Irbesartan	75	41 (54.7)	51.24 (8.28)	29.2	47.59	53.45	57.33	61.4			
	Week 70	Sparsentan	89	72 (80.9)	49.87 (8.39)	25.4	43.53	51.74	57.16	61.3			
		Irbesartan	75	47 (62.7)	50.20 (8.37)	28.1	45.46	52.88	57.06	62.7			
	Week 94	Sparsentan	89	67 (75.3)	50.41 (8.79)	26.5	46.14	52.92	57.06	62.7			
		Irbesartan	75	45 (60.0)	50.83 (8.99)	23.6	47.78	52.88	57.33	61.3			
	Week 110	Sparsentan	89	62 (69.7)	48.58 (9.34)	20.8	41.87	49.41	57.16	67.0			
		Irbesartan	75	41 (54.7)	47.70 (8.74)	23.2	44.07	47.72	53.16	62.4			
		KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	89	69 (77.5)	0.03 (7.19)	-19.1	-3.11	0.00	3.27	22.5	0.03 [-0.34, 0.40]
	Irbesartan			75	47 (62.7)	-0.16 (5.71)	-15.4	-4.36	0.00	3.55	20.2		
	Week 48		Sparsentan	89	72 (80.9)	-1.53 (8.87)	-23.7	-5.38	-0.58	3.60	19.4	-0.05 [-0.43, 0.33]	
			Irbesartan	75	41 (54.7)	-1.13 (6.75)	-18.5	-3.95	-0.10	3.95	12.6		
	Week 70		Sparsentan	89	72 (80.9)	-0.76 (7.51)	-21.8	-4.17	0.05	3.26	17.0	0.04 [-0.33, 0.40]	
			Irbesartan	75	47 (62.7)	-1.03 (7.75)	-27.2	-4.26	-0.57	3.22	17.4		
	Week 94		Sparsentan	89	67 (75.3)	-0.14 (7.82)	-22.4	-4.85	0.00	4.94	13.6	0.00 [-0.37, 0.38]	
			Irbesartan	75	45 (60.0)	-0.16 (6.91)	-19.0	-2.85	0.00	4.56	14.5		
Week 110	Sparsentan		89	62 (69.7)	-2.28 (7.62)	-23.0	-6.34	-1.60	2.71	13.8	0.08 [-0.32, 0.47]		
	Irbesartan		75	41 (54.7)	-2.83 (6.32)	-21.9	-6.97	-2.20	1.80	10.9			

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]		
Subgroup: History of hypertension														
Yes	KDQOL-SF12: MCS	Baseline	Sparsentan	155	142 (91.6)	50.36 (8.29)	16.2	44.79	51.80	57.06	65.2			
			Irbesartan	161	143 (88.8)	51.74 (8.63)	26.4	48.02	54.10	57.33	66.6			
		Week 24	Sparsentan	155	121 (78.1)	52.17 (8.07)	20.8	47.83	53.96	57.34	68.7			
			Irbesartan	161	106 (65.8)	52.79 (7.74)	27.2	49.35	54.41	57.76	64.2			
		Week 48	Sparsentan	155	124 (80.0)	51.02 (8.46)	29.5	44.41	52.24	57.33	64.7			
			Irbesartan	161	97 (60.2)	50.67 (9.32)	19.6	47.01	53.45	57.16	64.6			
		Week 70	Sparsentan	155	125 (80.6)	51.10 (7.88)	27.6	45.40	52.87	57.16	65.5			
			Irbesartan	161	104 (64.6)	51.13 (8.18)	18.9	47.81	52.75	57.06	63.8			
		Week 94	Sparsentan	155	118 (76.1)	50.70 (9.17)	16.0	44.47	53.39	57.16	66.9			
			Irbesartan	161	103 (64.0)	50.33 (9.72)	21.9	44.47	54.20	57.60	61.3			
		Week 110	Sparsentan	155	113 (72.9)	50.25 (8.95)	20.1	43.65	51.53	57.16	68.0			
			Irbesartan	161	92 (57.1)	49.54 (9.74)	20.0	44.17	51.43	57.16	62.7			
			KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	155	121 (78.1)	1.46 (7.73)	-17.0	-2.96	0.99	5.28	25.9	0.08 [-0.18, 0.34]
		Irbesartan			161	106 (65.8)	0.87 (6.71)	-15.4	-3.28	0.13	4.24	28.7		
Week 48	Sparsentan	155		124 (80.0)	0.53 (8.15)	-23.4	-4.62	1.17	5.77	28.1	0.23 [-0.04, 0.50]			
	Irbesartan	161		97 (60.2)	-1.31 (7.77)	-33.9	-3.88	-0.10	2.79	17.3				
Week 70	Sparsentan	155		125 (80.6)	0.71 (7.85)	-17.1	-4.24	0.39	4.97	26.5	0.15 [-0.11, 0.41]			
	Irbesartan	161		104 (64.6)	-0.34 (6.51)	-21.2	-3.63	-0.17	3.82	17.4				
Week 94	Sparsentan	155		118 (76.1)	0.24 (8.19)	-28.8	-5.02	0.96	5.09	16.9	0.17 [-0.09, 0.44]			
	Irbesartan	161		103 (64.0)	-1.13 (7.57)	-21.0	-5.33	-0.11	2.79	23.3				
Week 110	Sparsentan	155		113 (72.9)	-0.42 (8.86)	-31.2	-4.40	0.00	4.55	33.3	0.14 [-0.13, 0.42]			
	Irbesartan	161		92 (57.1)	-1.64 (8.20)	-29.3	-6.18	-1.30	3.50	19.1				

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSHM: KDQOL-SF12: Course of MCS by subgroup  
 Full Analysis Set

Subgroup	Parameter	Time	Treatment	N	n (%)	Mean (SD)	Min	Q25	Q50	Q75	Max	Hedge's G [95% CI]
No	KDQOL-SF12: MCS	Baseline	Sparsentan	47	46 (97.9)	50.65 (8.43)	27.8	46.33	51.51	56.59	66.5	
			Irbesartan	41	40 (97.6)	50.28 (8.64)	24.3	45.85	51.25	57.16	62.5	
		Week 24	Sparsentan	47	38 (80.9)	50.79 (7.57)	30.7	46.52	52.54	57.16	62.7	
			Irbesartan	41	32 (78.0)	50.85 (9.52)	24.2	46.40	53.30	57.24	62.6	
		Week 48	Sparsentan	47	41 (87.2)	50.99 (8.59)	27.5	46.89	52.05	57.06	62.5	
			Irbesartan	41	30 (73.2)	49.79 (10.57)	21.5	43.60	50.99	57.56	64.5	
		Week 70	Sparsentan	47	38 (80.9)	49.62 (9.71)	25.4	41.34	52.37	57.16	62.7	
			Irbesartan	41	31 (75.6)	51.23 (7.90)	34.7	45.09	51.94	57.16	63.4	
		Week 94	Sparsentan	47	37 (78.7)	49.54 (9.57)	26.5	42.90	51.83	56.85	62.5	
			Irbesartan	41	30 (73.2)	48.47 (10.26)	30.5	41.07	51.69	57.16	63.4	
	Week 110	Sparsentan	47	37 (78.7)	49.40 (8.68)	28.3	45.81	50.20	54.96	64.0		
		Irbesartan	41	29 (70.7)	48.66 (9.12)	30.5	42.61	48.93	57.06	62.5		
	KDQOL-SF12: change from baseline in MCS	Week 24	Sparsentan	47	38 (80.9)	-0.55 (9.05)	-23.6	-4.31	-0.05	3.86	26.4	-0.16 [-0.63, 0.31]
			Irbesartan	41	32 (78.0)	0.71 (6.55)	-14.1	-2.92	0.08	4.74	15.5	
		Week 48	Sparsentan	47	41 (87.2)	-0.25 (9.78)	-23.7	-4.74	-0.24	5.04	21.1	0.17 [-0.30, 0.65]
			Irbesartan	41	30 (73.2)	-1.78 (7.15)	-17.6	-7.62	-0.30	3.95	12.6	
		Week 70	Sparsentan	47	38 (80.9)	-1.27 (10.24)	-21.8	-5.78	-0.42	3.36	26.4	-0.03 [-0.51, 0.44]
			Irbesartan	41	31 (75.6)	-0.99 (7.41)	-27.2	-5.33	0.00	3.22	10.6	
		Week 94	Sparsentan	47	37 (78.7)	-1.12 (8.78)	-16.8	-7.87	-0.42	4.10	24.1	0.20 [-0.28, 0.68]
			Irbesartan	41	30 (73.2)	-2.81 (7.81)	-19.0	-7.82	-2.85	1.23	14.5	
Week 110		Sparsentan	47	37 (78.7)	-1.04 (9.67)	-23.0	-5.33	-1.41	2.96	26.4	0.20 [-0.29, 0.68]	
		Irbesartan	41	29 (70.7)	-2.65 (5.78)	-16.1	-5.34	-2.25	0.00	10.4		

N = total number of patients in analysis set. n = number of evaluable values. SD = standard deviation. Q = quantile.  
 95% CI = 95% confidence interval for Hedges G. A decrease reflects a worsening of the status of the patient.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Sex	Overall	Sparsentan						Interaction:	0.896
Male	Week 24	Sparsentan	139	110 (79.1)	0.40 (0.66)	(-0.89, 1.69)	-0.44 (0.96)	(-2.32, 1.43)	0.645
		Irbesartan	143	100 (69.9)	0.84 (0.69)	(-0.52, 2.20)			
	Week 48	Sparsentan	139	115 (82.7)	0.20 (0.64)	(-1.06, 1.46)	0.51 (0.96)	(-1.38, 2.40)	0.594
		Irbesartan	143	90 (62.9)	-0.31 (0.71)	(-1.72, 1.09)			
	Week 70	Sparsentan	139	117 (84.2)	0.17 (0.64)	(-1.10, 1.43)	0.49 (0.95)	(-1.37, 2.36)	0.605
		Irbesartan	143	97 (67.8)	-0.33 (0.70)	(-1.70, 1.05)			
	Week 94	Sparsentan	139	108 (77.7)	-0.27 (0.66)	(-1.57, 1.03)	0.84 (0.96)	(-1.05, 2.73)	0.384
		Irbesartan	143	99 (69.2)	-1.11 (0.70)	(-2.47, 0.26)			
	Week 110	Sparsentan	139	104 (74.8)	-0.22 (0.68)	(-1.56, 1.11)	1.78 (1.00)	(-0.17, 3.74)	0.074
		Irbesartan	143	89 (62.2)	-2.01 (0.73)	(-3.44, -0.57)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Female	Week 24	Sparsentan	63	49 (77.8)	1.66 (1.10)	(-0.51, 3.83)	0.32 (1.67)	(-2.96, 3.61)	0.847
		Irbesartan	59	38 (64.4)	1.34 (1.25)	(-1.13, 3.80)			
	Week 48	Sparsentan	63	50 (79.4)	-0.26 (1.09)	(-2.40, 1.88)	3.33 (1.66)	(0.05, 6.60)	0.046 *
		Irbesartan	59	37 (62.7)	-3.58 (1.26)	(-6.06, -1.11)			
	Week 70	Sparsentan	63	46 (73.0)	-0.68 (1.13)	(-2.90, 1.54)	-0.06 (1.69)	(-3.38, 3.25)	0.971
		Irbesartan	59	38 (64.4)	-0.62 (1.25)	(-3.08, 1.85)			
	Week 94	Sparsentan	63	47 (74.6)	-0.57 (1.12)	(-2.78, 1.64)	1.54 (1.73)	(-1.86, 4.94)	0.375
		Irbesartan	59	34 (57.6)	-2.11 (1.31)	(-4.69, 0.47)			
	Week 110	Sparsentan	63	46 (73.0)	-2.42 (1.14)	(-4.66, -0.18)	-1.20 (1.78)	(-4.70, 2.29)	0.499
		Irbesartan	59	32 (54.2)	-1.22 (1.36)	(-3.89, 1.46)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age	Overall	Sparsentan						Interaction:	0.693
<= 45 years	Week 24	Sparsentan	96	72 (75.0)	0.40 (0.88)	(-1.34, 2.14)	-0.73 (1.26)	(-3.21, 1.75)	0.564
		Irbesartan	99	70 (70.7)	1.13 (0.90)	(-0.64, 2.89)			
	Week 48	Sparsentan	96	76 (79.2)	-0.46 (0.86)	(-2.15, 1.23)	0.95 (1.27)	(-1.55, 3.45)	0.454
		Irbesartan	99	62 (62.6)	-1.41 (0.93)	(-3.25, 0.42)			
	Week 70	Sparsentan	96	75 (78.1)	0.53 (0.87)	(-1.18, 2.24)	1.19 (1.28)	(-1.32, 3.69)	0.353
		Irbesartan	99	64 (64.6)	-0.66 (0.93)	(-2.49, 1.18)			
	Week 94	Sparsentan	96	70 (72.9)	-0.74 (0.89)	(-2.50, 1.01)	1.55 (1.29)	(-0.99, 4.10)	0.231
		Irbesartan	99	64 (64.6)	-2.30 (0.94)	(-4.13, -0.46)			
	Week 110	Sparsentan	96	70 (72.9)	-0.93 (0.90)	(-2.70, 0.85)	0.47 (1.33)	(-2.15, 3.08)	0.727
		Irbesartan	99	59 (59.6)	-1.39 (0.97)	(-3.31, 0.52)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 45 years	Week 24	Sparsentan	106	87 (82.1)	1.25 (0.73)	(-0.19, 2.68)	0.35 (1.11)	(-1.82, 2.52)	0.752
		Irbesartan	103	68 (66.0)	0.90 (0.83)	(-0.73, 2.53)			
	Week 48	Sparsentan	106	89 (84.0)	0.60 (0.72)	(-0.81, 2.02)	1.73 (1.11)	(-0.45, 3.90)	
		Irbesartan	103	65 (63.1)	-1.12 (0.84)	(-2.76, 0.52)			
	Week 70	Sparsentan	106	88 (83.0)	-0.49 (0.73)	(-1.92, 0.94)	-0.27 (1.09)	(-2.41, 1.88)	
		Irbesartan	103	71 (68.9)	-0.22 (0.81)	(-1.81, 1.37)			
	Week 94	Sparsentan	106	85 (80.2)	0.06 (0.74)	(-1.40, 1.51)	0.62 (1.11)	(-1.56, 2.79)	
		Irbesartan	103	69 (67.0)	-0.56 (0.82)	(-2.17, 1.05)			
	Week 110	Sparsentan	106	80 (75.5)	-0.76 (0.76)	(-2.26, 0.73)	1.57 (1.15)	(-0.70, 3.84)	
		Irbesartan	103	62 (60.2)	-2.34 (0.87)	(-4.04, -0.64)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Age at IgAN diagnosis	Overall	Sparsentan						Interaction:	0.250
<= 18 years	Week 24	Sparsentan	9	5 (55.6)	-4.46 (3.73)	(-12.23, 3.32)	-4.20 (5.50)	(-15.91, 7.52)	0.457
		Irbesartan	5	5 (100.0)	-0.26 (3.91)	(-8.69, 8.17)			
	Week 48	Sparsentan	9	7 (77.8)	-0.58 (3.20)	(-7.38, 6.23)	1.72 (5.69)	(-10.23, 13.66)	0.766
		Irbesartan	5	3 (60.0)	-2.29 (4.58)	(-11.81, 7.22)			
	Week 70	Sparsentan	9	7 (77.8)	0.21 (3.25)	(-6.74, 7.15)	4.40 (5.56)	(-7.45, 16.26)	0.441
		Irbesartan	5	4 (80.0)	-4.19 (4.32)	(-13.35, 4.96)			
	Week 94	Sparsentan	9	5 (55.6)	-1.46 (3.72)	(-9.21, 6.30)	5.74 (5.97)	(-6.89, 18.37)	0.350
		Irbesartan	5	4 (80.0)	-7.20 (4.43)	(-16.65, 2.25)			
	Week 110	Sparsentan	9	4 (44.4)	0.09 (4.18)	(-8.59, 8.78)	8.33 (7.11)	(-6.44, 23.10)	0.254
		Irbesartan	5	2 (40.0)	-8.24 (5.71)	(-20.03, 3.56)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024



Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 18 to 40 years	Week 24	Sparsentan	102	80 (78.4)	0.50 (0.83)	(-1.13, 2.12)	-0.94 (1.20)	(-3.29, 1.41)	0.434
		Irbesartan	109	73 (67.0)	1.44 (0.86)	(-0.26, 3.13)			
	Week 48	Sparsentan	102	83 (81.4)	-0.51 (0.81)	(-2.11, 1.08)	0.86 (1.20)	(-1.49, 3.21)	0.475
		Irbesartan	109	69 (63.3)	-1.37 (0.88)	(-3.09, 0.35)			
	Week 70	Sparsentan	102	80 (78.4)	-0.28 (0.83)	(-1.90, 1.34)	0.46 (1.20)	(-1.91, 2.82)	0.705
		Irbesartan	109	71 (65.1)	-0.73 (0.87)	(-2.45, 0.98)			
	Week 94	Sparsentan	102	76 (74.5)	-0.95 (0.85)	(-2.61, 0.71)	0.45 (1.23)	(-1.96, 2.85)	0.715
		Irbesartan	109	69 (63.3)	-1.40 (0.89)	(-3.14, 0.34)			
	Week 110	Sparsentan	102	73 (71.6)	-1.17 (0.87)	(-2.87, 0.53)	-0.03 (1.26)	(-2.51, 2.45)	0.983
		Irbesartan	109	65 (59.6)	-1.14 (0.92)	(-2.94, 0.66)			
> 40 years	Week 24	Sparsentan	91	74 (81.3)	1.53 (0.80)	(-0.03, 3.10)	0.91 (1.19)	(-1.43, 3.25)	0.445
		Irbesartan	88	60 (68.2)	0.62 (0.88)	(-1.11, 2.36)			
	Week 48	Sparsentan	91	75 (82.4)	0.81 (0.79)	(-0.73, 2.36)	1.85 (1.20)	(-0.51, 4.22)	0.125
		Irbesartan	88	55 (62.5)	-1.04 (0.91)	(-2.83, 0.75)			
	Week 70	Sparsentan	91	76 (83.5)	0.15 (0.79)	(-1.40, 1.69)	-0.04 (1.18)	(-2.36, 2.28)	0.973
		Irbesartan	88	60 (68.2)	0.19 (0.88)	(-1.55, 1.92)			
	Week 94	Sparsentan	91	74 (81.3)	0.35 (0.80)	(-1.21, 1.91)	1.37 (1.19)	(-0.96, 3.71)	0.249
		Irbesartan	88	60 (68.2)	-1.02 (0.88)	(-2.76, 0.71)			
	Week 110	Sparsentan	91	73 (80.2)	-0.62 (0.80)	(-2.19, 0.96)	1.88 (1.23)	(-0.53, 4.30)	0.127
		Irbesartan	88	54 (61.4)	-2.50 (0.93)	(-4.33, -0.67)			

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Geographic region	Overall	Sparsentan						Interaction:	0.811
North America	Week 24	Sparsentan	35	23 (65.7)	1.40 (1.48)	(-1.52, 4.31)	1.02 (1.90)	(-2.72, 4.76)	0.591
		Irbesartan	46	37 (80.4)	0.37 (1.18)	(-1.95, 2.70)			
	Week 48	Sparsentan	35	25 (71.4)	0.11 (1.43)	(-2.70, 2.92)	2.67 (1.89)	(-1.06, 6.39)	0.160
		Irbesartan	46	33 (71.7)	-2.55 (1.23)	(-4.97, -0.14)			
	Week 70	Sparsentan	35	23 (65.7)	-0.25 (1.47)	(-3.15, 2.66)	1.21 (1.95)	(-2.63, 5.06)	0.535
		Irbesartan	46	31 (67.4)	-1.46 (1.26)	(-3.95, 1.04)			
	Week 94	Sparsentan	35	24 (68.6)	-2.26 (1.46)	(-5.13, 0.62)	-0.18 (1.96)	(-4.04, 3.68)	0.928
		Irbesartan	46	30 (65.2)	-2.08 (1.29)	(-4.62, 0.46)			
	Week 110	Sparsentan	35	21 (60.0)	-0.26 (1.55)	(-3.32, 2.79)	2.40 (2.02)	(-1.58, 6.38)	0.236
		Irbesartan	46	31 (67.4)	-2.66 (1.28)	(-5.19, -0.13)			

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A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
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Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
Europe	Week 24	Sparsentan	98	75 (76.5)	-0.07 (0.83)	(-1.69, 1.55)	-1.37 (1.21)	(-3.76, 1.01)	0.258	
		Irbesartan	115	65 (56.5)	1.30 (0.89)	(-0.44, 3.05)				
	Week 48	Sparsentan	98	77 (78.6)	-0.46 (0.81)	(-2.07, 1.14)	1.10 (1.21)	(-1.29, 3.49)	0.366	
		Irbesartan	115	61 (53.0)	-1.56 (0.90)	(-3.33, 0.20)				
	Week 70	Sparsentan	98	76 (77.6)	-0.87 (0.82)	(-2.49, 0.74)	-0.19 (1.18)	(-2.51, 2.13)	0.872	
		Irbesartan	115	72 (62.6)	-0.68 (0.85)	(-2.35, 0.98)				
	Week 94	Sparsentan	98	71 (72.4)	-0.32 (0.85)	(-1.98, 1.35)	1.48 (1.20)	(-0.88, 3.84)	0.218	
		Irbesartan	115	72 (62.6)	-1.80 (0.85)	(-3.46, -0.14)				
	Week 110	Sparsentan	98	68 (69.4)	-0.91 (0.87)	(-2.62, 0.79)	1.06 (1.26)	(-1.42, 3.54)	0.400	
		Irbesartan	115	61 (53.0)	-1.98 (0.91)	(-3.77, -0.18)				
Asia Pacific	Week 24	Sparsentan	69	61 (88.4)	1.96 (0.94)	(0.12, 3.81)	1.42 (1.54)	(-1.60, 4.45)	0.355	
		Irbesartan	41	36 (87.8)	0.54 (1.22)	(-1.85, 2.93)				
	Week 48	Sparsentan	69	63 (91.3)	1.11 (0.92)	(-0.71, 2.92)	0.93 (1.56)	(-2.14, 4.00)	0.553	
		Irbesartan	41	33 (80.5)	0.18 (1.25)	(-2.28, 2.64)				
	Week 70	Sparsentan	69	64 (92.8)	1.25 (0.92)	(-0.55, 3.06)	0.38 (1.58)	(-2.74, 3.50)	0.810	
		Irbesartan	41	32 (78.0)	0.87 (1.28)	(-1.65, 3.40)				
	Week 94	Sparsentan	69	60 (87.0)	0.61 (0.94)	(-1.24, 2.45)	0.73 (1.62)	(-2.45, 3.91)	0.653	
		Irbesartan	41	31 (75.6)	-0.12 (1.31)	(-2.69, 2.45)				
	Week 110	Sparsentan	69	61 (88.4)	-0.74 (0.94)	(-2.59, 1.11)	0.32 (1.65)	(-2.93, 3.57)	0.848	
		Irbesartan	41	29 (70.7)	-1.06 (1.35)	(-3.71, 1.60)				

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A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline BMI	Overall	Sparsentan						Interaction:	0.829
< 27 kg/m**2	Week 24	Sparsentan	83	66 (79.5)	0.81 (0.97)	(-1.11, 2.72)	-0.88 (1.37)	(-3.58, 1.81)	0.521
		Irbesartan	94	69 (73.4)	1.69 (0.96)	(-0.20, 3.57)			
	Week 48	Sparsentan	83	68 (81.9)	0.46 (0.96)	(-1.43, 2.35)	1.77 (1.39)	(-0.97, 4.51)	0.204
		Irbesartan	94	60 (63.8)	-1.31 (1.00)	(-3.28, 0.66)			
	Week 70	Sparsentan	83	66 (79.5)	0.25 (0.98)	(-1.66, 2.17)	0.27 (1.40)	(-2.49, 3.02)	0.849
		Irbesartan	94	62 (66.0)	-0.01 (1.00)	(-1.98, 1.95)			
	Week 94	Sparsentan	83	64 (77.1)	0.09 (0.99)	(-1.85, 2.04)	0.48 (1.41)	(-2.28, 3.25)	0.731
		Irbesartan	94	64 (68.1)	-0.39 (0.99)	(-2.35, 1.56)			
	Week 110	Sparsentan	83	63 (75.9)	-0.31 (1.00)	(-2.28, 1.67)	1.43 (1.45)	(-1.43, 4.29)	0.326
		Irbesartan	94	57 (60.6)	-1.74 (1.05)	(-3.79, 0.32)			

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Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 27 kg/m**2	Week 24	Sparsentan	119	93 (78.2)	0.81 (0.68)	(-0.52, 2.15)	0.48 (1.04)	(-1.57, 2.53)	0.647
		Irbesartan	107	69 (64.5)	0.33 (0.79)	(-1.22, 1.89)			
	Week 48	Sparsentan	119	97 (81.5)	-0.20 (0.66)	(-1.51, 1.10)	1.01 (1.03)	(-1.03, 3.04)	0.331
		Irbesartan	107	67 (62.6)	-1.21 (0.79)	(-2.77, 0.35)			
	Week 70	Sparsentan	119	97 (81.5)	-0.34 (0.67)	(-1.65, 0.97)	0.33 (1.02)	(-1.67, 2.33)	0.746
		Irbesartan	107	73 (68.2)	-0.67 (0.77)	(-2.18, 0.84)			
	Week 94	Sparsentan	119	91 (76.5)	-0.74 (0.69)	(-2.08, 0.61)	1.40 (1.05)	(-0.66, 3.46)	0.183
		Irbesartan	107	68 (63.6)	-2.13 (0.79)	(-3.69, -0.58)			
	Week 110	Sparsentan	119	87 (73.1)	-1.31 (0.70)	(-2.69, 0.07)	0.66 (1.09)	(-1.47, 2.79)	0.542
		Irbesartan	107	63 (58.9)	-1.97 (0.83)	(-3.60, -0.35)			

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Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Randomization strata	Overall	Sparsentan						Interaction:	0.163
eGFR Low and UP High	Week 24	Sparsentan	71	54 (76.1)	-0.26 (0.93)	(-2.09, 1.56)	-0.75 (1.38)	(-3.47, 1.97)	0.586
		Irbesartan	74	44 (59.5)	0.49 (1.03)	(-1.52, 2.51)			
	Week 48	Sparsentan	71	53 (74.6)	-0.79 (0.93)	(-2.62, 1.03)	-0.06 (1.43)	(-2.86, 2.75)	0.968
		Irbesartan	74	38 (51.4)	-0.74 (1.08)	(-2.86, 1.39)			
	Week 70	Sparsentan	71	58 (81.7)	-2.60 (0.90)	(-4.37, -0.83)	-1.69 (1.39)	(-4.42, 1.04)	0.223
		Irbesartan	74	41 (55.4)	-0.91 (1.06)	(-2.98, 1.17)			
	Week 94	Sparsentan	71	52 (73.2)	-0.26 (0.94)	(-2.10, 1.59)	-0.71 (1.44)	(-3.55, 2.13)	0.623
		Irbesartan	74	38 (51.4)	0.46 (1.10)	(-1.70, 2.61)			
	Week 110	Sparsentan	71	53 (74.6)	-2.41 (0.94)	(-4.26, -0.56)	-0.44 (1.47)	(-3.33, 2.46)	0.768
		Irbesartan	74	36 (48.6)	-1.97 (1.13)	(-4.20, 0.26)			

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 Full Analysis Set

KDQOL-SF12: change from baseline in MCS	Subgroup	Time	Treatment	N	n (%)	Repeated measures analysis				
						Change from Baseline		Treatment Difference		p-value
						LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	
eGFR Low and UP Low	Week 24	Sparsentan	55	44 (80.0)	2.02 (1.12)	(-0.19, 4.22)	1.42 (1.64)	(-1.81, 4.64)	0.388	
		Irbesartan	55	39 (70.9)	0.60 (1.20)	(-1.75, 2.95)				
	Week 48	Sparsentan	55	43 (78.2)	-0.62 (1.13)	(-2.84, 1.59)	1.60 (1.66)	(-1.67, 4.87)	0.337	
		Irbesartan	55	36 (65.5)	-2.22 (1.22)	(-4.63, 0.18)				
	Week 70	Sparsentan	55	42 (76.4)	0.27 (1.14)	(-1.98, 2.52)	0.61 (1.64)	(-2.61, 3.84)	0.708	
		Irbesartan	55	40 (72.7)	-0.34 (1.18)	(-2.66, 1.97)				
	Week 94	Sparsentan	55	42 (76.4)	-0.49 (1.15)	(-2.75, 1.76)	2.22 (1.65)	(-1.02, 5.46)	0.179	
		Irbesartan	55	40 (72.7)	-2.71 (1.18)	(-5.03, -0.39)				
	Week 110	Sparsentan	55	38 (69.1)	-0.23 (1.20)	(-2.59, 2.14)	3.50 (1.75)	(0.06, 6.94)	0.046 *	
		Irbesartan	55	34 (61.8)	-3.73 (1.27)	(-6.23, -1.24)				
eGFR High and UP High	Week 24	Sparsentan	37	27 (73.0)	0.31 (1.34)	(-2.33, 2.94)	-2.12 (1.93)	(-5.93, 1.70)	0.275	
		Irbesartan	36	26 (72.2)	2.42 (1.39)	(-0.32, 5.17)				
	Week 48	Sparsentan	37	34 (91.9)	0.64 (1.22)	(-1.78, 3.05)	0.75 (1.87)	(-2.94, 4.44)	0.688	
		Irbesartan	36	25 (69.4)	-0.11 (1.41)	(-2.89, 2.66)				
	Week 70	Sparsentan	37	31 (83.8)	1.18 (1.27)	(-1.32, 3.69)	0.43 (1.91)	(-3.33, 4.19)	0.821	
		Irbesartan	36	25 (69.4)	0.75 (1.41)	(-2.03, 3.53)				
	Week 94	Sparsentan	37	27 (73.0)	-0.06 (1.35)	(-2.72, 2.59)	0.19 (1.96)	(-3.68, 4.05)	0.924	
		Irbesartan	36	25 (69.4)	-0.25 (1.41)	(-3.03, 2.53)				
	Week 110	Sparsentan	37	26 (70.3)	-0.18 (1.38)	(-2.91, 2.55)	-1.42 (2.05)	(-5.46, 2.61)	0.488	
		Irbesartan	36	22 (61.1)	1.24 (1.49)	(-1.70, 4.19)				

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Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
eGFR High and UP Low	Week 24	Sparsentan	39	34 (87.2)	1.38 (1.28)	(-1.14, 3.91)	0.18 (1.88)	(-3.51, 3.88)	0.922
		Irbesartan	37	29 (78.4)	1.20 (1.37)	(-1.51, 3.91)			
	Week 48	Sparsentan	39	35 (89.7)	1.85 (1.27)	(-0.65, 4.35)	3.30 (1.87)	(-0.39, 6.99)	0.079
		Irbesartan	37	28 (75.7)	-1.45 (1.38)	(-4.17, 1.26)			
	Week 70	Sparsentan	39	32 (82.1)	2.70 (1.30)	(0.13, 5.27)	3.41 (1.89)	(-0.32, 7.14)	0.073
		Irbesartan	37	29 (78.4)	-0.71 (1.37)	(-3.41, 1.99)			
	Week 94	Sparsentan	39	34 (87.2)	-0.52 (1.28)	(-3.04, 2.01)	2.11 (1.87)	(-1.57, 5.79)	0.259
		Irbesartan	37	30 (81.1)	-2.63 (1.36)	(-5.30, 0.05)			
	Week 110	Sparsentan	39	33 (84.6)	0.27 (1.30)	(-2.29, 2.83)	1.64 (1.89)	(-2.10, 5.37)	0.388
		Irbesartan	37	29 (78.4)	-1.36 (1.38)	(-4.08, 1.35)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024



Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 1	Overall	Sparsentan						Interaction:	0.181
< 60 mL/min/1.73 m**2	Week 24	Sparsentan	127	99 (78.0)	0.62 (0.71)	(-0.76, 2.01)	0.10 (1.04)	(-1.94, 2.14)	0.925
		Irbesartan	129	85 (65.9)	0.53 (0.76)	(-0.97, 2.02)			
	Week 48	Sparsentan	127	97 (76.4)	-1.06 (0.71)	(-2.45, 0.33)	0.92 (1.07)	(-1.17, 3.01)	0.389
		Irbesartan	129	76 (58.9)	-1.98 (0.79)	(-3.54, -0.42)			
	Week 70	Sparsentan	127	101 (79.5)	-1.48 (0.70)	(-2.85, -0.10)	-0.74 (1.04)	(-2.79, 1.30)	0.474
		Irbesartan	129	83 (64.3)	-0.73 (0.77)	(-2.24, 0.78)			
	Week 94	Sparsentan	127	94 (74.0)	-0.63 (0.72)	(-2.05, 0.78)	0.23 (1.07)	(-1.86, 2.33)	0.827
		Irbesartan	129	79 (61.2)	-0.87 (0.79)	(-2.41, 0.68)			
	Week 110	Sparsentan	127	93 (73.2)	-1.33 (0.73)	(-2.76, 0.10)	0.96 (1.10)	(-1.20, 3.13)	0.383
		Irbesartan	129	72 (55.8)	-2.30 (0.83)	(-3.92, -0.67)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	39 (79.6)	0.21 (1.16)	(-2.09, 2.50)	-1.30 (1.73)	(-4.72, 2.12)	0.455
		Irbesartan	48	33 (68.8)	1.51 (1.27)	(-1.00, 4.02)			
	Week 48	Sparsentan	49	45 (91.8)	0.41 (1.11)	(-1.77, 2.59)	-0.25 (1.69)	(-3.57, 3.08)	0.884
		Irbesartan	48	33 (68.8)	0.66 (1.26)	(-1.83, 3.14)			
	Week 70	Sparsentan	49	40 (81.6)	1.47 (1.15)	(-0.80, 3.74)	0.80 (1.73)	(-2.60, 4.20)	0.644
		Irbesartan	48	32 (66.7)	0.67 (1.27)	(-1.84, 3.18)			
	Week 94	Sparsentan	49	41 (83.7)	-0.74 (1.15)	(-3.02, 1.53)	1.70 (1.70)	(-1.66, 5.06)	0.320
		Irbesartan	48	35 (72.9)	-2.45 (1.24)	(-4.89, 0.00)			
	Week 110	Sparsentan	49	36 (73.5)	-1.27 (1.22)	(-3.67, 1.13)	0.25 (1.78)	(-3.26, 3.76)	0.889
		Irbesartan	48	32 (66.7)	-1.52 (1.29)	(-4.06, 1.02)			
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	21 (80.8)	2.82 (1.58)	(-0.30, 5.95)	1.03 (2.26)	(-3.45, 5.50)	0.651
		Irbesartan	25	20 (80.0)	1.80 (1.62)	(-1.40, 5.00)			
	Week 48	Sparsentan	26	23 (88.5)	4.66 (1.53)	(1.64, 7.68)	6.14 (2.27)	(1.65, 10.62)	0.008 *
		Irbesartan	25	18 (72.0)	-1.48 (1.67)	(-4.78, 1.83)			
	Week 70	Sparsentan	26	22 (84.6)	3.70 (1.55)	(0.63, 6.77)	4.59 (2.26)	(0.13, 9.06)	0.044 *
		Irbesartan	25	20 (80.0)	-0.90 (1.63)	(-4.13, 2.33)			
	Week 94	Sparsentan	26	20 (76.9)	2.05 (1.61)	(-1.13, 5.23)	3.57 (2.31)	(-1.01, 8.15)	0.125
		Irbesartan	25	19 (76.0)	-1.52 (1.66)	(-4.80, 1.77)			
	Week 110	Sparsentan	26	21 (80.8)	2.14 (1.59)	(-1.01, 5.30)	2.10 (2.36)	(-2.57, 6.77)	0.375
		Irbesartan	25	17 (68.0)	0.04 (1.74)	(-3.40, 3.48)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline eGFR Group 2	Overall	Sparsentan						Interaction:	0.179
< 45 mL/min/1.73 m**2	Week 24	Sparsentan	82	64 (78.0)	1.61 (0.88)	(-0.13, 3.34)	1.05 (1.33)	(-1.55, 3.66)	0.427
		Irbesartan	80	51 (63.8)	0.55 (0.99)	(-1.39, 2.50)			
	Week 48	Sparsentan	82	63 (76.8)	-0.28 (0.89)	(-2.02, 1.46)	2.03 (1.36)	(-0.64, 4.70)	0.136
		Irbesartan	80	45 (56.3)	-2.31 (1.03)	(-4.34, -0.28)			
	Week 70	Sparsentan	82	65 (79.3)	-1.15 (0.88)	(-2.87, 0.58)	0.23 (1.32)	(-2.38, 2.83)	0.864
		Irbesartan	80	50 (62.5)	-1.37 (0.99)	(-3.32, 0.57)			
	Week 94	Sparsentan	82	61 (74.4)	0.46 (0.90)	(-1.31, 2.23)	0.76 (1.35)	(-1.90, 3.42)	0.576
		Irbesartan	80	48 (60.0)	-0.30 (1.01)	(-2.29, 1.69)			
	Week 110	Sparsentan	82	57 (69.5)	-0.59 (0.93)	(-2.43, 1.24)	1.14 (1.41)	(-1.63, 3.92)	0.419
		Irbesartan	80	44 (55.0)	-1.74 (1.06)	(-3.82, 0.35)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
45 to < 60 mL/min/1.73 m**2	Week 24	Sparsentan	45	35 (77.8)	-1.17 (1.17)	(-3.47, 1.13)	-1.64 (1.67)	(-4.92, 1.65)	0.328
		Irbesartan	49	34 (69.4)	0.47 (1.19)	(-1.87, 2.81)			
	Week 48	Sparsentan	45	34 (75.6)	-2.41 (1.18)	(-4.73, -0.10)	-0.93 (1.71)	(-4.30, 2.44)	
		Irbesartan	49	31 (63.3)	-1.49 (1.24)	(-3.92, 0.95)			
	Week 70	Sparsentan	45	36 (80.0)	-2.09 (1.15)	(-4.36, 0.17)	-2.36 (1.67)	(-5.66, 0.93)	
		Irbesartan	49	33 (67.3)	0.27 (1.21)	(-2.11, 2.65)			
	Week 94	Sparsentan	45	33 (73.3)	-2.62 (1.20)	(-4.98, -0.27)	-0.78 (1.73)	(-4.18, 2.62)	
		Irbesartan	49	31 (63.3)	-1.84 (1.24)	(-4.29, 0.61)			
	Week 110	Sparsentan	45	36 (80.0)	-2.52 (1.16)	(-4.80, -0.24)	0.78 (1.75)	(-2.67, 4.23)	
		Irbesartan	49	28 (57.1)	-3.30 (1.31)	(-5.88, -0.72)			
60 to < 90 mL/min/1.73 m**2	Week 24	Sparsentan	49	39 (79.6)	0.21 (1.16)	(-2.09, 2.50)	-1.30 (1.73)	(-4.72, 2.12)	0.455
		Irbesartan	48	33 (68.8)	1.51 (1.27)	(-1.00, 4.02)			
	Week 48	Sparsentan	49	45 (91.8)	0.41 (1.11)	(-1.77, 2.59)	-0.25 (1.69)	(-3.57, 3.08)	
		Irbesartan	48	33 (68.8)	0.66 (1.26)	(-1.83, 3.14)			
	Week 70	Sparsentan	49	40 (81.6)	1.47 (1.15)	(-0.80, 3.74)	0.80 (1.73)	(-2.60, 4.20)	
		Irbesartan	48	32 (66.7)	0.67 (1.27)	(-1.84, 3.18)			
	Week 94	Sparsentan	49	41 (83.7)	-0.74 (1.15)	(-3.02, 1.53)	1.70 (1.70)	(-1.66, 5.06)	
		Irbesartan	48	35 (72.9)	-2.45 (1.24)	(-4.89, 0.00)			
	Week 110	Sparsentan	49	36 (73.5)	-1.27 (1.22)	(-3.67, 1.13)	0.25 (1.78)	(-3.26, 3.76)	
		Irbesartan	48	32 (66.7)	-1.52 (1.29)	(-4.06, 1.02)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
>= 90 mL/min/1.73 m**2	Week 24	Sparsentan	26	21 (80.8)	2.82 (1.58)	(-0.30, 5.95)	1.03 (2.26)	(-3.45, 5.50)	0.651
		Irbesartan	25	20 (80.0)	1.80 (1.62)	(-1.40, 5.00)			
	Week 48	Sparsentan	26	23 (88.5)	4.66 (1.53)	(1.64, 7.68)	6.14 (2.27)	(1.65, 10.62)	0.008 *
		Irbesartan	25	18 (72.0)	-1.48 (1.67)	(-4.78, 1.83)			
	Week 70	Sparsentan	26	22 (84.6)	3.70 (1.55)	(0.63, 6.77)	4.59 (2.26)	(0.13, 9.06)	0.044 *
		Irbesartan	25	20 (80.0)	-0.90 (1.63)	(-4.13, 2.33)			
	Week 94	Sparsentan	26	20 (76.9)	2.05 (1.61)	(-1.13, 5.23)	3.57 (2.31)	(-1.01, 8.15)	0.125
		Irbesartan	25	19 (76.0)	-1.52 (1.66)	(-4.80, 1.77)			
	Week 110	Sparsentan	26	21 (80.8)	2.14 (1.59)	(-1.01, 5.30)	2.10 (2.36)	(-2.57, 6.77)	0.375
		Irbesartan	25	17 (68.0)	0.04 (1.74)	(-3.40, 3.48)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
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Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline urine protein excretion	Overall	Sparsentan						Interaction:	0.440
<= 1.75 g/day	Week 24	Sparsentan	98	81 (82.7)	1.22 (0.82)	(-0.38, 2.83)	0.36 (1.26)	(-2.12, 2.83)	0.776
		Irbesartan	93	59 (63.4)	0.86 (0.96)	(-1.02, 2.74)			
	Week 48	Sparsentan	98	83 (84.7)	0.16 (0.81)	(-1.43, 1.74)	1.27 (1.26)	(-1.20, 3.73)	0.313
		Irbesartan	93	57 (61.3)	-1.11 (0.96)	(-3.00, 0.78)			
	Week 70	Sparsentan	98	77 (78.6)	0.97 (0.83)	(-0.67, 2.61)	1.69 (1.25)	(-0.77, 4.15)	0.178
		Irbesartan	93	62 (66.7)	-0.72 (0.93)	(-2.55, 1.12)			
	Week 94	Sparsentan	98	79 (80.6)	-0.49 (0.83)	(-2.12, 1.13)	1.65 (1.24)	(-0.79, 4.09)	0.185
		Irbesartan	93	64 (68.8)	-2.14 (0.92)	(-3.96, -0.32)			
	Week 110	Sparsentan	98	76 (77.6)	-0.49 (0.84)	(-2.15, 1.17)	1.56 (1.29)	(-0.97, 4.10)	0.227
		Irbesartan	93	57 (61.3)	-2.05 (0.97)	(-3.96, -0.13)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 1.75 g/day	Week 24	Sparsentan	104	78 (75.0)	0.44 (0.79)	(-1.11, 1.99)	-0.56 (1.12)	(-2.75, 1.64)	0.618
		Irbesartan	109	79 (72.5)	1.00 (0.79)	(-0.56, 2.55)			
	Week 48	Sparsentan	104	82 (78.8)	0.07 (0.77)	(-1.44, 1.58)	1.51 (1.13)	(-0.70, 3.72)	
		Irbesartan	109	70 (64.2)	-1.44 (0.82)	(-3.06, 0.17)			
	Week 70	Sparsentan	104	86 (82.7)	-0.97 (0.76)	(-2.46, 0.52)	-0.74 (1.11)	(-2.93, 1.45)	
		Irbesartan	109	73 (67.0)	-0.23 (0.81)	(-1.83, 1.37)			
	Week 94	Sparsentan	104	76 (73.1)	-0.14 (0.80)	(-1.71, 1.42)	0.57 (1.16)	(-1.70, 2.84)	
		Irbesartan	109	69 (63.3)	-0.71 (0.84)	(-2.35, 0.93)			
	Week 110	Sparsentan	104	74 (71.2)	-1.19 (0.81)	(-2.79, 0.41)	0.51 (1.19)	(-1.84, 2.85)	
		Irbesartan	109	64 (58.7)	-1.69 (0.87)	(-3.40, 0.01)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
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Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
 Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Baseline use of antihypertensives	Overall	Sparsentan							Interaction: 0.536
Yes	Week 24	Sparsentan	90	68 (75.6)	0.44 (0.84)	(-1.21, 2.08)	-0.60 (1.27)	(-3.10, 1.90)	0.637
		Irbesartan	88	52 (59.1)	1.04 (0.95)	(-0.84, 2.91)			
	Week 48	Sparsentan	90	68 (75.6)	0.40 (0.83)	(-1.24, 2.03)	0.92 (1.28)	(-1.60, 3.45)	0.473
		Irbesartan	88	48 (54.5)	-0.53 (0.98)	(-2.44, 1.39)			
	Week 70	Sparsentan	90	69 (76.7)	-0.37 (0.83)	(-2.00, 1.26)	0.69 (1.27)	(-1.81, 3.18)	0.589
		Irbesartan	88	51 (58.0)	-1.05 (0.96)	(-2.94, 0.83)			
	Week 94	Sparsentan	90	66 (73.3)	0.13 (0.85)	(-1.53, 1.79)	0.43 (1.25)	(-2.02, 2.89)	0.730
		Irbesartan	88	57 (64.8)	-0.30 (0.92)	(-2.10, 1.50)			
	Week 110	Sparsentan	90	64 (71.1)	-1.12 (0.86)	(-2.82, 0.57)	0.15 (1.29)	(-2.40, 2.69)	0.910
		Irbesartan	88	51 (58.0)	-1.27 (0.96)	(-3.16, 0.62)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
 A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
 A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
 Source Data: aqs, created on: 05MAR2024



Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
No	Week 24	Sparsentan	112	91 (81.3)	1.12 (0.77)	(-0.40, 2.64)	0.15 (1.11)	(-2.04, 2.34)	0.893
		Irbesartan	114	86 (75.4)	0.97 (0.80)	(-0.60, 2.54)			
	Week 48	Sparsentan	112	97 (86.6)	-0.13 (0.75)	(-1.61, 1.35)	1.59 (1.11)	(-0.60, 3.77)	0.155
		Irbesartan	114	79 (69.3)	-1.72 (0.82)	(-3.33, -0.11)			
	Week 70	Sparsentan	112	94 (83.9)	0.14 (0.76)	(-1.36, 1.64)	0.20 (1.11)	(-1.99, 2.38)	0.858
		Irbesartan	114	84 (73.7)	-0.06 (0.81)	(-1.65, 1.53)			
	Week 94	Sparsentan	112	89 (79.5)	-0.73 (0.78)	(-2.27, 0.81)	1.47 (1.15)	(-0.79, 3.73)	0.202
		Irbesartan	114	76 (66.7)	-2.20 (0.85)	(-3.86, -0.54)			
	Week 110	Sparsentan	112	86 (76.8)	-0.71 (0.80)	(-2.28, 0.87)	1.54 (1.19)	(-0.80, 3.88)	0.198
		Irbesartan	114	70 (61.4)	-2.24 (0.88)	(-3.98, -0.51)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
Time since renal biopsy	Overall	Sparsentan						Interaction:	0.223
<= 5 years	Week 24	Sparsentan	113	90 (79.6)	1.53 (0.77)	(0.02, 3.05)	0.15 (1.09)	(-1.99, 2.28)	0.894
		Irbesartan	127	91 (71.7)	1.39 (0.77)	(-0.12, 2.89)			
	Week 48	Sparsentan	113	93 (82.3)	1.52 (0.76)	(0.04, 3.01)	2.89 (1.08)	(0.76, 5.02)	0.008 *
		Irbesartan	127	86 (67.7)	-1.37 (0.78)	(-2.89, 0.16)			
	Week 70	Sparsentan	113	91 (80.5)	0.73 (0.76)	(-0.77, 2.23)	0.94 (1.09)	(-1.19, 3.08)	0.386
		Irbesartan	127	88 (69.3)	-0.22 (0.77)	(-1.74, 1.30)			
	Week 94	Sparsentan	113	88 (77.9)	-0.27 (0.78)	(-1.79, 1.25)	1.64 (1.10)	(-0.51, 3.80)	0.135
		Irbesartan	127	88 (69.3)	-1.91 (0.78)	(-3.44, -0.39)			
	Week 110	Sparsentan	113	88 (77.9)	0.27 (0.78)	(-1.27, 1.81)	1.34 (1.13)	(-0.88, 3.55)	0.236
		Irbesartan	127	80 (63.0)	-1.07 (0.81)	(-2.66, 0.53)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
> 5 years	Week 24	Sparsentan	89	69 (77.5)	-0.02 (0.83)	(-1.64, 1.61)	-0.26 (1.31)	(-2.84, 2.32)	0.843
		Irbesartan	75	47 (62.7)	0.24 (1.01)	(-1.75, 2.24)			
	Week 48	Sparsentan	89	72 (80.9)	-1.72 (0.81)	(-3.32, -0.13)	-0.72 (1.34)	(-3.34, 1.91)	0.593
		Irbesartan	75	41 (54.7)	-1.00 (1.06)	(-3.09, 1.08)			
	Week 70	Sparsentan	89	72 (80.9)	-1.05 (0.82)	(-2.65, 0.56)	-0.22 (1.30)	(-2.77, 2.34)	0.868
		Irbesartan	75	47 (62.7)	-0.83 (1.01)	(-2.81, 1.15)			
	Week 94	Sparsentan	89	67 (75.3)	-0.41 (0.84)	(-2.06, 1.24)	-0.11 (1.33)	(-2.72, 2.51)	0.936
		Irbesartan	75	45 (60.0)	-0.30 (1.03)	(-2.33, 1.72)			
	Week 110	Sparsentan	89	62 (69.7)	-2.43 (0.87)	(-4.15, -0.71)	0.71 (1.39)	(-2.02, 3.44)	0.608
		Irbesartan	75	41 (54.7)	-3.14 (1.08)	(-5.26, -1.03)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
					Change from Baseline		Treatment Difference		
Subgroup	Time	Treatment	N	n (%)	LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
History of hypertension	Overall	Sparsentan						Interaction:	0.997
Yes	Week 24	Sparsentan	155	121 (78.1)	1.25 (0.63)	(0.01, 2.49)	-0.01 (0.93)	(-1.83, 1.80)	0.989
		Irbesartan	161	106 (65.8)	1.26 (0.68)	(-0.07, 2.58)			
	Week 48	Sparsentan	155	124 (80.0)	0.28 (0.62)	(-0.94, 1.50)	1.42 (0.93)	(-0.40, 3.25)	0.127
		Irbesartan	161	97 (60.2)	-1.14 (0.69)	(-2.50, 0.22)			
	Week 70	Sparsentan	155	125 (80.6)	0.35 (0.62)	(-0.87, 1.57)	0.67 (0.92)	(-1.14, 2.48)	0.467
		Irbesartan	161	104 (64.6)	-0.32 (0.68)	(-1.65, 1.01)			
	Week 94	Sparsentan	155	118 (76.1)	0.03 (0.64)	(-1.22, 1.28)	0.93 (0.93)	(-0.91, 2.76)	0.322
		Irbesartan	161	103 (64.0)	-0.89 (0.68)	(-2.23, 0.44)			
	Week 110	Sparsentan	155	113 (72.9)	-0.72 (0.65)	(-2.00, 0.57)	0.85 (0.97)	(-1.05, 2.76)	0.380
		Irbesartan	161	92 (57.1)	-1.57 (0.72)	(-2.99, -0.16)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Table PT2KMSC\_FSCM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

KDQOL-SF12: change from baseline in MCS					Repeated measures analysis				
Subgroup	Time	Treatment	N	n (%)	Change from Baseline		Treatment Difference		
					LS-Mean (STE)	95% CI	LS-Mean (STE)	95% CI	p-value
No	Week 24	Sparsentan	47	38 (80.9)	-0.60 (1.28)	(-3.12, 1.91)	-0.68 (1.90)	(-4.42, 3.06)	0.722
		Irbesartan	41	32 (78.0)	0.07 (1.40)	(-2.68, 2.83)			
	Week 48	Sparsentan	47	41 (87.2)	-0.62 (1.24)	(-3.05, 1.82)	1.14 (1.89)	(-2.58, 4.86)	0.547
		Irbesartan	41	30 (73.2)	-1.76 (1.42)	(-4.56, 1.05)			
	Week 70	Sparsentan	47	38 (80.9)	-1.42 (1.28)	(-3.93, 1.09)	-0.57 (1.91)	(-4.33, 3.19)	0.766
		Irbesartan	41	31 (75.6)	-0.85 (1.42)	(-3.65, 1.94)			
	Week 94	Sparsentan	47	37 (78.7)	-1.60 (1.30)	(-4.15, 0.96)	1.54 (1.95)	(-2.30, 5.38)	0.429
		Irbesartan	41	30 (73.2)	-3.14 (1.45)	(-5.99, -0.29)			
	Week 110	Sparsentan	47	37 (78.7)	-1.42 (1.30)	(-3.99, 1.14)	1.40 (1.98)	(-2.50, 5.30)	0.480
		Irbesartan	41	29 (70.7)	-2.82 (1.48)	(-5.74, 0.09)			

N = number of patients in analysis set. n (%) = number of non-missing values included in the model. MMRM = Mixed Model Repeated Measures. LS-Mean = Least square mean. STE = Standard error. 95% CI = 95% confidence interval. \* = significant treatment effect. # = significant interaction.  
A MMRM was applied with fixed factors for treatment, time, interaction treatment-by-time, randomization strata and baseline as covariate. Baseline covariates/factors, which are part of considered subgroup, were excluded from the model. For interaction test, subgroup and interaction subgroup\*treatment was added to the model. All observed values are included. A first order regressive covariance structure was used.  
A decrease reflects a worsening of the status of the patient. eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day. eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index. MCS = mental component summary.  
Source Data: aqs, created on: 05MAR2024

Figure PF2KMSC\_FSGM: KDQOL-SF12: Change from baseline in MCS by subgroup  
Full Analysis Set

Not done: No significant subgroup found.

Number of available records are presented for key visits up to Week 110 only. LS-Mean = Least square mean. 95% CI = 95% confidence interval. A decrease reflects a worsening of the status of the patient.

eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.

eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.

MCS = mental component summary.

Reference table: PT2KMSC\_FSCM

Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.284
Male	Sparsentan	139	35 (25.2)	NE		1.622	(0.923, 2.848)	0.093
	Irbesartan	143	20 (14.0)	NE				
Female	Sparsentan	63	16 (25.4)	NE		3.300	(1.152, 9.451)	0.026 *
	Irbesartan	59	5 (8.5)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.850
<= 45 years	Sparsentan	96	28 (29.2)	NE		1.754	(0.940, 3.272)	0.078
	Irbesartan	99	16 (16.2)	NE				
> 45 years	Sparsentan	106	23 (21.7)	NE		2.516	(1.126, 5.625)	0.025 *
	Irbesartan	103	9 (8.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan						Interaction test:	NE
<= 18 years	Sparsentan	9	2 (22.2) No events in 1 group	NE		NE		NE
	Irbesartan	5	0 (0.0)	NE				
> 18 to 40 years	Sparsentan	102	28 (27.5)	NE		1.993	(1.069, 3.716)	0.030 *
	Irbesartan	109	18 (16.5)	NE				
> 40 years	Sparsentan	91	21 (23.1)	NE		3.061	(1.252, 7.484)	0.014 *
	Irbesartan	88	7 (8.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.261
North America	Sparsentan	35	11 (31.4)	NE		2.323	(0.783, 6.888)	0.129
	Irbesartan	46	5 (10.9)	NE				
Europe	Sparsentan	98	17 (17.3)	NE		1.347	(0.621, 2.924)	0.451
	Irbesartan	115	13 (11.3)	NE				
Asia Pacific	Sparsentan	69	23 (33.3)	NE		3.617	(1.446, 9.046)	0.006 *
	Irbesartan	41	7 (17.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.777
< 27 kg/m**2	Sparsentan	83	26 (31.3)	NE		2.147	(0.981, 4.696)	0.056
	Irbesartan	94	9 (9.6)	NE				
≥ 27 kg/m**2	Sparsentan	119	25 (21.0)	NE		1.959	(0.995, 3.858)	0.052
	Irbesartan	107	16 (15.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.627
eGFR Low and UP High	Sparsentan	71	13 (18.3)	NE		2.675	(0.935, 7.650)	0.066
	Irbesartan	74	6 (8.1)	NE				
eGFR Low and UP Low	Sparsentan	55	13 (23.6)	NE		1.331	(0.513, 3.458)	0.557
	Irbesartan	55	7 (12.7)	NE				
eGFR High and UP High	Sparsentan	37	12 (32.4)	NE		1.680	(0.615, 4.588)	0.311
	Irbesartan	36	6 (16.7)	NE				
eGFR High and UP Low	Sparsentan	39	13 (33.3)	NE		2.498	(0.926, 6.737)	0.071
	Irbesartan	37	6 (16.2)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.221
< 60 mL/min/1.73 m**2	Sparsentan	127	25 (19.7)	NE		1.512	(0.789, 2.896)	0.213
	Irbesartan	129	15 (11.6)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	14 (28.6)	NE		1.893	(0.705, 5.082)	0.206
	Irbesartan	48	6 (12.5)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	12 (46.2)	114.3	(46.1, NE)	13.813	(2.534, 75.308)	0.002 *
	Irbesartan	25	4 (16.0)	115.3	(115.3, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.388
< 45 mL/min/1.73 m**2	Sparsentan	82	17 (20.7)	NE		1.581	(0.709, 3.526)	0.263
	Irbesartan	80	10 (12.5)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	8 (17.8)	NE		1.244	(0.373, 4.147)	0.722
	Irbesartan	49	5 (10.2)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	14 (28.6)	NE		1.893	(0.705, 5.082)	0.206
	Irbesartan	48	6 (12.5)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	12 (46.2)	114.3	(46.1, NE)	13.813	(2.534, 75.308)	0.002 *
	Irbesartan	25	4 (16.0)	115.3	(115.3, NE)			

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.223
<= 1.75 g/day	Sparsentan	98	27 (27.6)	NE		2.549	(1.186, 5.479)	0.017 *
	Irbesartan	93	9 (9.7)	NE				
> 1.75 g/day	Sparsentan	104	24 (23.1)	NE		1.620	(0.838, 3.133)	0.151
	Irbesartan	109	16 (14.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.508
Yes	Sparsentan	90	21 (23.3)	NE		2.975	(1.168, 7.578)	0.022 *
	Irbesartan	88	6 (6.8)	NE				
No	Sparsentan	112	30 (26.8)	NE		1.903	(1.045, 3.467)	0.035 *
	Irbesartan	114	19 (16.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan							Interaction test: 0.537
<= 5 years	Sparsentan	113	35 (31.0)	NE		2.210	(1.234, 3.959)	0.008 *
	Irbesartan	127	18 (14.2)	NE				
> 5 years	Sparsentan	89	16 (18.0)	NE		1.788	(0.714, 4.479)	0.215
	Irbesartan	75	7 (9.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSIT\_FSTM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to increase in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan							Interaction test: 0.508
Yes	Sparsentan	155	39 (25.2)	NE		2.171	(1.230, 3.832)	0.007 *
	Irbesartan	161	18 (11.2)	NE				
No	Sparsentan	47	12 (25.5)	NE		1.721	(0.556, 5.322)	0.346
	Irbesartan	41	7 (17.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
An increase reflects an improvement of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Figure PF2KMSIT\_FSKM: KDQOL-SF12: Time to increase in MCS by at least 9.0 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

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An increase reflects an improvement of the status of the patient.  
Reference table: PT2KMSIT\_FSTM

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Sex	Sparsentan						Interaction test:	0.441
Male	Sparsentan	139	34 (24.5)	NE		1.018	(0.626, 1.655)	0.943
	Irbesartan	143	33 (23.1)	NE				
Female	Sparsentan	63	23 (36.5)	NE		1.499	(0.776, 2.896)	0.228
	Irbesartan	59	15 (25.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age	Sparsentan						Interaction test:	0.440
<= 45 years	Sparsentan	96	26 (27.1)	NE		0.964	(0.554, 1.676)	0.896
	Irbesartan	99	28 (28.3)	116.0	(111.1, NE)			
> 45 years	Sparsentan	106	31 (29.2)	NE		1.372	(0.777, 2.421)	0.275
	Irbesartan	103	20 (19.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Age at IgAN diagnosis	Sparsentan							Interaction test: 0.993
<= 18 years	Sparsentan	9	3 (33.3)	NE		2.696	(0.254, 28.581)	0.410
	Irbesartan	5	2 (40.0)	NE				
> 18 to 40 years	Sparsentan	102	30 (29.4)	NE		1.176	(0.697, 1.984)	0.544
	Irbesartan	109	29 (26.6)	116.0	(114.3, NE)			
> 40 years	Sparsentan	91	24 (26.4)	NE		1.130	(0.604, 2.114)	0.702
	Irbesartan	88	17 (19.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Geographic region	Sparsentan						Interaction test:	0.731
North America	Sparsentan	35	9 (25.7)	NE		0.876	(0.347, 2.207)	0.778
	Irbesartan	46	12 (26.1)	NE				
Europe	Sparsentan	98	32 (32.7)	NE		1.441	(0.850, 2.441)	0.175
	Irbesartan	115	26 (22.6)	NE				
Asia Pacific	Sparsentan	69	16 (23.2)	NE		1.010	(0.448, 2.276)	0.981
	Irbesartan	41	10 (24.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

Subgroup	Treatment	N	nev (%)	Kaplan-Meier analysis		Cox regression		
				Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline BMI	Sparsentan						Interaction test:	0.491
< 27 kg/m**2	Sparsentan	83	26 (31.3)	NE		1.292	(0.731, 2.286)	0.378
	Irbesartan	94	24 (25.5)	NE				
≥ 27 kg/m**2	Sparsentan	119	31 (26.1)	NE		1.026	(0.598, 1.760)	0.926
	Irbesartan	107	24 (22.4)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024



Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Randomization strata	Sparsentan						Interaction test:	0.543
eGFR Low and UP High	Sparsentan	71	19 (26.8)	NE		1.296	(0.639, 2.629)	0.472
	Irbesartan	74	13 (17.6)	NE				
eGFR Low and UP Low	Sparsentan	55	15 (27.3)	NE		0.886	(0.435, 1.806)	0.739
	Irbesartan	55	16 (29.1)	NE				
eGFR High and UP High	Sparsentan	37	15 (40.5)	114.6	(94.1, NE)	1.726	(0.725, 4.110)	0.217
	Irbesartan	36	9 (25.0)	NE				
eGFR High and UP Low	Sparsentan	39	8 (20.5)	NE		0.739	(0.290, 1.888)	0.528
	Irbesartan	37	10 (27.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 1	Sparsentan						Interaction test:	0.794
< 60 mL/min/1.73 m**2	Sparsentan	127	36 (28.3)	NE		1.189	(0.725, 1.948)	0.493
	Irbesartan	129	30 (23.3)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	14 (28.6)	NE		1.386	(0.601, 3.196)	0.443
	Irbesartan	48	11 (22.9)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	7 (26.9)	NE		0.756	(0.252, 2.266)	0.617
	Irbesartan	25	7 (28.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline eGFR Group 2	Sparsentan						Interaction test:	0.520
< 45 mL/min/1.73 m**2	Sparsentan	82	18 (22.0)	NE		0.797	(0.413, 1.539)	0.500
	Irbesartan	80	18 (22.5)	NE				
45 to < 60 mL/min/1.73 m**2	Sparsentan	45	18 (40.0)	NE		1.828	(0.856, 3.904)	0.119
	Irbesartan	49	12 (24.5)	NE				
60 to < 90 mL/min/1.73 m**2	Sparsentan	49	14 (28.6)	NE		1.386	(0.601, 3.196)	0.443
	Irbesartan	48	11 (22.9)	NE				
≥ 90 mL/min/1.73 m**2	Sparsentan	26	7 (26.9)	NE		0.756	(0.252, 2.266)	0.617
	Irbesartan	25	7 (28.0)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
\* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline urine protein excretion	Sparsentan							Interaction test: 0.663
<= 1.75 g/day	Sparsentan	98	27 (27.6)	NE		0.979	(0.554, 1.730)	0.942
	Irbesartan	93	22 (23.7)	NE				
> 1.75 g/day	Sparsentan	104	30 (28.8)	NE		1.341	(0.781, 2.303)	0.288
	Irbesartan	109	26 (23.9)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Baseline use of antihypertensives	Sparsentan							Interaction test: 0.738
Yes	Sparsentan	90	26 (28.9)	NE		1.255	(0.694, 2.271)	0.452
	Irbesartan	88	21 (23.9)	NE				
No	Sparsentan	112	31 (27.7)	NE		1.123	(0.664, 1.901)	0.665
	Irbesartan	114	27 (23.7)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
Time since renal biopsy	Sparsentan						Interaction test:	0.386
<= 5 years	Sparsentan	113	28 (24.8)	NE		0.948	(0.567, 1.586)	0.839
	Irbesartan	127	32 (25.2)	NE				
> 5 years	Sparsentan	89	29 (32.6)	NE		1.433	(0.774, 2.655)	0.252
	Irbesartan	75	16 (21.3)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Table PT2KMSDT\_FSTM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
 Full Analysis Set

KDQOL-SF12: time to decrease in MCS by at least 9.0				Kaplan-Meier analysis		Cox regression		
Subgroup	Treatment	N	nev (%)	Median time (weeks)	95% CI	Hazard Ratio	95% CI	p-value
History of hypertension	Sparsentan						Interaction test:	0.967
Yes	Sparsentan	155	39 (25.2)	NE		1.133	(0.710, 1.809)	0.600
	Irbesartan	161	34 (21.1)	NE				
No	Sparsentan	47	18 (38.3)	NE		1.055	(0.521, 2.135)	0.882
	Irbesartan	41	14 (34.1)	NE				

N = number of patients in analysis set. nev (%) = number of patients with at least one event. NE = not evaluable.  
 95% CI = 95% confidence interval (median: Brookmeyer/Crowley hazard ratio: coming from Cox model).  
 A Cox proportional model was applied with factors treatment, randomization strata and baseline as covariate.  
 \* = significant treatment effect. # = significant interaction. For interaction test, subgroup and interaction subgroup\*treatment was added to the model.  
 A decrease reflects a worsening of the status of the patient. Time is presented in weeks.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aqs\_tte, created on: 26MAR2024

Figure PF2KMSDT\_FSKM: KDQOL-SF12: Time to decrease in MCS by at least 9.0 points by subgroup  
Full Analysis Set

Not done. No significant subgroups.

\_\_\_\_\_ A decrease reflects a worsening of the status of the patient.  
Reference table: PT2KMSDT\_FSTM



Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.396
Male	Double-blind period	Sparsentan	139	128 (92.1)	1.037 [0.961, 1.119]	1.466 [0.655, 3.282]	3.3 [-4.3, 10.8]	0.420
		Irbesartan	143	127 (88.8)				
Female	Double-blind period	Sparsentan	63	59 (93.7)	1.105 [0.974, 1.253]	2.655 [0.771, 9.143]	8.9 [-3.7, 21.5]	0.145
		Irbesartan	59	50 (84.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.950
<= 45 years	Double-blind period	Sparsentan	96	90 (93.8)	1.055 [0.967, 1.150]	1.875 [0.665, 5.290]	4.9 [-4.0, 13.7]	0.311
		Irbesartan	99	88 (88.9)				
> 45 years	Double-blind period	Sparsentan	106	97 (91.5)	1.059 [0.962, 1.166]	1.695 [0.699, 4.110]	5.1 [-4.3, 14.5]	0.274
		Irbesartan	103	89 (86.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.943
<= 18 years	Double-blind period	Sparsentan	9	9 (100.0)	1.036 + [0.783, 1.371]	1.727 + [0.030, 99.953]	0.0 [NE, NE]	NE
		Irbesartan	5	5 (100.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	95 (93.1)	1.047 [0.962, 1.139]	1.679 [0.634, 4.447]	4.1 [-4.5, 12.8]	0.342
		Irbesartan	109	97 (89.0)				
> 40 years	Double-blind period	Sparsentan	91	83 (91.2)	1.070 [0.961, 1.192]	1.798 [0.706, 4.578]	6.0 [-4.6, 16.5]	0.250
		Irbesartan	88	75 (85.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.896
North America	Double-blind period	Sparsentan	35	32 (91.4)	1.026 [0.889, 1.184]	1.301 [0.289, 5.854]	2.3 [-13.1, 17.7]	1.000
		Irbesartan	46	41 (89.1)				
Europe	Double-blind period	Sparsentan	98	90 (91.8)	1.067 [0.971, 1.172]	1.818 [0.743, 4.452]	5.7 [-3.5, 15.0]	0.201
		Irbesartan	115	99 (86.1)				
Asia Pacific	Double-blind period	Sparsentan	69	65 (94.2)	1.044 [0.929, 1.173]	1.757 [0.415, 7.441]	4.0 [-8.6, 16.5]	0.468
		Irbesartan	41	37 (90.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.865
< 27 kg/m**2	Double-blind period	Sparsentan	83	76 (91.6)	1.050 [0.949, 1.161]	1.589 [0.595, 4.246]	4.3 [-5.8, 14.5]	0.467
		Irbesartan	94	82 (87.2)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	111 (93.3)	1.062 [0.975, 1.156]	1.919 [0.763, 4.827]	5.4 [-3.1, 14.0]	0.176
		Irbesartan	107	94 (87.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.663
eGFR Low and UP High	Double-blind period	Sparsentan	71	67 (94.4)	1.074 [0.970, 1.190]	2.319 [0.680, 7.905]	6.5 [-4.0, 17.1]	0.246
		Irbesartan	74	65 (87.8)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	49 (89.1)	1.000 [0.877, 1.140]	1.000 [0.302, 3.316]	0.0 [-13.5, 13.5]	1.000
		Irbesartan	55	49 (89.1)				
eGFR High and UP High	Double-blind period	Sparsentan	37	35 (94.6)	1.135 [0.962, 1.339]	3.500 [0.657, 18.648]	11.3 [-5.7, 28.2]	0.152
		Irbesartan	36	30 (83.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	36 (92.3)	1.035 [0.896, 1.196]	1.455 [0.303, 6.989]	3.1 [-12.6, 18.8]	0.708
		Irbesartan	37	33 (89.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.239
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	118 (92.9)	1.061 [0.978, 1.150]	1.856 [0.788, 4.371]	5.3 [-2.7, 13.3]	0.206
		Irbesartan	129	113 (87.6)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	45 (91.8)	0.980 [0.877, 1.095]	0.750 [0.159, 3.544]	-1.9 [-14.3, 10.4]	1.000
		Irbesartan	48	45 (93.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	24 (92.3)	1.215 [0.949, 1.554]	3.789 [0.686, 20.946]	16.3 [-7.2, 39.9]	0.140
		Irbesartan	25	19 (76.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.366
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	75 (91.5)	1.045 [0.940, 1.162]	1.531 [0.552, 4.242]	4.0 [-6.7, 14.6]	0.452
		Irbesartan	80	70 (87.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	43 (95.6)	1.089 [0.964, 1.230]	3.000 [0.573, 15.702]	7.8 [-5.3, 20.9]	0.271
		Irbesartan	49	43 (87.8)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	45 (91.8)	0.980 [0.877, 1.095]	0.750 [0.159, 3.544]	-1.9 [-14.3, 10.4]	1.000
		Irbesartan	48	45 (93.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	24 (92.3)	1.215 [0.949, 1.554]	3.789 [0.686, 20.946]	16.3 [-7.2, 39.9]	0.140
		Irbesartan	25	19 (76.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.481
<= 1.75 g/day	Double-blind period	Sparsentan	98	88 (89.8)	1.031 [0.930, 1.143]	1.304 [0.534, 3.180]	2.7 [-7.4, 12.8]	0.652
		Irbesartan	93	81 (87.1)				
> 1.75 g/day	Double-blind period	Sparsentan	104	99 (95.2)	1.081 [0.996, 1.173]	2.681 [0.921, 7.808]	7.1 [-1.2, 15.4]	0.084
		Irbesartan	109	96 (88.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
Safety Set

TEAEs					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.481
Yes	Double-blind period	Sparsentan	90	81 (90.0)	1.029	1.286	2.5	0.641
		Irbesartan	88	77 (87.5)	[0.926, 1.142]	[0.505, 3.273]	[-7.9, 12.9]	
No	Double-blind period	Sparsentan	112	106 (94.6)	1.079	2.473	6.9	0.099
		Irbesartan	114	100 (87.7)	[0.994, 1.171]	[0.915, 6.687]	[-1.3, 15.1]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.634
<= 5 years	Double-blind period	Sparsentan	113	105 (92.9)	1.044 [0.964, 1.131]	1.626 [0.656, 4.033]	3.9 [-4.1, 12.0]	0.372
		Irbesartan	127	113 (89.0)				
> 5 years	Double-blind period	Sparsentan	89	82 (92.1)	1.080 [0.966, 1.207]	2.013 [0.739, 5.486]	6.8 [-4.2, 17.8]	0.212
		Irbesartan	75	64 (85.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2A\_SSIM: Incidence of TEAEs during double-blind period by subgroup  
 Safety Set

TEAEs									
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
History of hypertension	Double-blind period	Sparsentan							Interaction test: 0.039 #
Yes	Double-blind period	Sparsentan	155	145 (93.5)	1.091 [1.012, 1.177]	2.417 [1.110, 5.262]	7.8 [0.6, 15.1]	0.027	*
		Irbesartan	161	138 (85.7)					
No	Double-blind period	Sparsentan	47	42 (89.4)	0.939 [0.833, 1.060]	0.431 [0.079, 2.350]	-5.8 [-19.1, 7.5]	0.442	
		Irbesartan	41	39 (95.1)					

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.488
Male	Double-blind period	Sparsentan	139	51 (36.7)	0.937 [0.695, 1.263]	0.900 [0.556, 1.457]	-2.5 [-14.5, 9.6]	0.713
		Irbesartan	143	56 (39.2)				
Female	Double-blind period	Sparsentan	63	20 (31.7)	1.171 [0.673, 2.036]	1.250 [0.572, 2.731]	4.6 [-13.2, 22.4]	0.692
		Irbesartan	59	16 (27.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.246
<= 45 years	Double-blind period	Sparsentan	96	39 (40.6)	1.149 [0.802, 1.647]	1.251 [0.701, 2.233]	5.3 [-9.4, 19.9]	0.464
		Irbesartan	99	35 (35.4)				
> 45 years	Double-blind period	Sparsentan	106	32 (30.2)	0.840 [0.570, 1.238]	0.771 [0.433, 1.375]	-5.7 [-19.4, 8.0]	0.462
		Irbesartan	103	37 (35.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.571
<= 18 years	Double-blind period	Sparsentan	9	7 (77.8)	1.296 [0.585, 2.874]	2.333 [0.216, 25.245]	17.8 [-48.6, 84.1]	0.580
		Irbesartan	5	3 (60.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	33 (32.4)	0.860 [0.594, 1.246]	0.793 [0.450, 1.399]	-5.3 [-19.1, 8.5]	0.472
		Irbesartan	109	41 (37.6)				
> 40 years	Double-blind period	Sparsentan	91	31 (34.1)	1.071 [0.704, 1.627]	1.107 [0.593, 2.066]	2.2 [-12.6, 17.1]	0.754
		Irbesartan	88	28 (31.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.825
North America	Double-blind period	Sparsentan	35	12 (34.3)	1.127 [0.598, 2.122]	1.193 [0.466, 3.050]	3.9 [-19.3, 27.0]	0.811
		Irbesartan	46	14 (30.4)				
Europe	Double-blind period	Sparsentan	98	38 (38.8)	0.949 [0.681, 1.322]	0.916 [0.528, 1.589]	-2.1 [-16.2, 12.0]	0.780
		Irbesartan	115	47 (40.9)				
Asia Pacific	Double-blind period	Sparsentan	69	21 (30.4)	1.134 [0.611, 2.106]	1.193 [0.505, 2.821]	3.6 [-15.7, 22.9]	0.829
		Irbesartan	41	11 (26.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.048 #
< 27 kg/m**2	Double-blind period	Sparsentan	83	24 (28.9)	0.715 [0.471, 1.086]	0.599 [0.320, 1.124]	-11.5 [-26.6, 3.5]	0.117
		Irbesartan	94	38 (40.4)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	47 (39.5)	1.243 [0.871, 1.774]	1.402 [0.810, 2.425]	7.7 [-5.6, 21.1]	0.267
		Irbesartan	107	34 (31.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.989
eGFR Low and UP High	Double-blind period	Sparsentan	71	33 (46.5)	0.955 [0.679, 1.345]	0.917 [0.478, 1.760]	-2.2 [-19.8, 15.5]	0.868
		Irbesartan	74	36 (48.6)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	18 (32.7)	1.000 [0.585, 1.709]	1.000 [0.451, 2.218]	0.0 [-19.4, 19.4]	1.000
		Irbesartan	55	18 (32.7)				
eGFR High and UP High	Double-blind period	Sparsentan	37	12 (32.4)	1.061 [0.539, 2.090]	1.091 [0.406, 2.931]	1.9 [-22.2, 25.9]	1.000
		Irbesartan	36	11 (30.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	8 (20.5)	1.084 [0.437, 2.691]	1.106 [0.357, 3.430]	1.6 [-18.9, 22.1]	1.000
		Irbesartan	37	7 (18.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.505
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	50 (39.4)	0.996 [0.735, 1.349]	0.993 [0.602, 1.639]	-0.2 [-12.9, 12.6]	1.000
		Irbesartan	129	51 (39.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	14 (28.6)	0.807 [0.450, 1.448]	0.729 [0.310, 1.718]	-6.8 [-27.4, 13.7]	0.518
		Irbesartan	48	17 (35.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	7 (26.9)	1.683 [0.561, 5.050]	1.934 [0.488, 7.660]	10.9 [-15.3, 37.1]	0.499
		Irbesartan	25	4 (16.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.618
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	35 (42.7)	1.067 [0.739, 1.541]	1.117 [0.597, 2.088]	2.7 [-13.7, 19.1]	0.752
		Irbesartan	80	32 (40.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	15 (33.3)	0.860 [0.500, 1.479]	0.789 [0.339, 1.838]	-5.4 [-27.0, 16.1]	0.669
		Irbesartan	49	19 (38.8)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	14 (28.6)	0.807 [0.450, 1.448]	0.729 [0.310, 1.718]	-6.8 [-27.4, 13.7]	0.518
		Irbesartan	48	17 (35.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	7 (26.9)	1.683 [0.561, 5.050]	1.934 [0.488, 7.660]	10.9 [-15.3, 37.1]	0.499
		Irbesartan	25	4 (16.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.315
<= 1.75 g/day	Double-blind period	Sparsentan	98	28 (28.6)	1.208 [0.747, 1.954]	1.291 [0.675, 2.470]	4.9 [-8.6, 18.4]	0.511
		Irbesartan	93	22 (23.7)				
> 1.75 g/day	Double-blind period	Sparsentan	104	43 (41.3)	0.901 [0.663, 1.225]	0.832 [0.484, 1.431]	-4.5 [-18.8, 9.7]	0.581
		Irbesartan	109	50 (45.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.043 #
Yes	Double-blind period	Sparsentan	90	40 (44.4)	1.304 [0.899, 1.890]	1.547 [0.844, 2.835]	10.4 [-5.0, 25.7]	0.170
		Irbesartan	88	30 (34.1)				
No	Double-blind period	Sparsentan	112	31 (27.7)	0.751 [0.512, 1.103]	0.656 [0.374, 1.151]	-9.2 [-22.2, 3.8]	0.156
		Irbesartan	114	42 (36.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.242
<= 5 years	Double-blind period	Sparsentan	113	44 (38.9)	1.124 [0.806, 1.567]	1.203 [0.711, 2.035]	4.3 [-8.8, 17.3]	0.505
		Irbesartan	127	44 (34.6)				
> 5 years	Double-blind period	Sparsentan	89	27 (30.3)	0.813 [0.528, 1.249]	0.731 [0.381, 1.401]	-7.0 [-22.8, 8.8]	0.407
		Irbesartan	75	28 (37.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AN\_SSIM: Incidence of non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.002 #
Yes	Double-blind period	Sparsentan	155	65 (41.9)	1.228 [0.925, 1.630]	1.392 [0.882, 2.196]	7.8 [-3.5, 19.1]	0.166
		Irbesartan	161	55 (34.2)				
No	Double-blind period	Sparsentan	47	6 (12.8)	0.308 [0.134, 0.707]	0.207 [0.072, 0.595]	-28.7 [-48.8, -8.6]	0.003 *
		Irbesartan	41	17 (41.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
 Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.298
Male	Double-blind period	Sparsentan	139	19 (13.7)	0.977 [0.546, 1.751]	0.974 [0.495, 1.915]	-0.3 [-9.1, 8.5]	1.000
		Irbesartan	143	20 (14.0)				
Female	Double-blind period	Sparsentan	63	5 (7.9)	0.520 [0.185, 1.463]	0.479 [0.151, 1.523]	-7.3 [-20.3, 5.7]	0.260
		Irbesartan	59	9 (15.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.453
<= 45 years	Double-blind period	Sparsentan	96	16 (16.7)	0.971 [0.521, 1.808]	0.965 [0.456, 2.040]	-0.5 [-12.1, 11.0]	1.000
		Irbesartan	99	17 (17.2)				
> 45 years	Double-blind period	Sparsentan	106	8 (7.5)	0.648 [0.276, 1.519]	0.619 [0.242, 1.583]	-4.1 [-13.0, 4.8]	0.354
		Irbesartan	103	12 (11.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.958
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	1.111 [0.131, 9.416]	1.143 [0.077, 16.947]	2.2 [-57.7, 62.1]	1.000
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	13 (12.7)	0.817 [0.418, 1.597]	0.790 [0.363, 1.722]	-2.9 [-13.2, 7.5]	0.694
		Irbesartan	109	17 (15.6)				
> 40 years	Double-blind period	Sparsentan	91	9 (9.9)	0.791 [0.345, 1.816]	0.768 [0.302, 1.956]	-2.6 [-13.0, 7.7]	0.640
		Irbesartan	88	11 (12.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.650
North America	Double-blind period	Sparsentan	35	7 (20.0)	1.314 [0.508, 3.402]	1.393 [0.439, 4.419]	4.8 [-14.6, 24.1]	0.768
		Irbesartan	46	7 (15.2)				
Europe	Double-blind period	Sparsentan	98	12 (12.2)	0.782 [0.397, 1.543]	0.752 [0.343, 1.650]	-3.4 [-13.6, 6.8]	0.555
		Irbesartan	115	18 (15.7)				
Asia Pacific	Double-blind period	Sparsentan	69	5 (7.2)	0.743 [0.211, 2.610]	0.723 [0.183, 2.860]	-2.5 [-15.4, 10.4]	0.725
		Irbesartan	41	4 (9.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.688
< 27 kg/m**2	Double-blind period	Sparsentan	83	9 (10.8)	0.728 [0.333, 1.594]	0.695 [0.284, 1.701]	-4.1 [-15.0, 6.9]	0.505
		Irbesartan	94	14 (14.9)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	15 (12.6)	0.899 [0.462, 1.751]	0.885 [0.410, 1.908]	-1.4 [-11.2, 8.4]	0.845
		Irbesartan	107	15 (14.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.646
eGFR Low and UP High	Double-blind period	Sparsentan	71	12 (16.9)	1.042	1.051	0.7	1.000
		Irbesartan	74	12 (16.2)	[0.502, 2.165]	[0.438, 2.523]	[-12.8, 14.2]	
eGFR Low and UP Low	Double-blind period	Sparsentan	55	4 (7.3)	0.444	0.401	-9.1	0.237
		Irbesartan	55	9 (16.4)	[0.145, 1.358]	[0.116, 1.390]	[-22.9, 4.7]	
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	0.973	0.970	-0.3	1.000
		Irbesartan	36	4 (11.1)	[0.263, 3.598]	[0.223, 4.212]	[-17.4, 16.8]	
eGFR High and UP Low	Double-blind period	Sparsentan	39	4 (10.3)	0.949	0.943	-0.6	1.000
		Irbesartan	37	4 (10.8)	[0.256, 3.519]	[0.218, 4.081]	[-17.0, 15.9]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.762
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	16 (12.6)	0.739 [0.407, 1.340]	0.701 [0.349, 1.407]	-4.5 [-13.9, 5.0]	0.380
		Irbesartan	129	22 (17.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	1.176 [0.384, 3.596]	1.200 [0.340, 4.230]	1.8 [-12.8, 16.5]	1.000
		Irbesartan	48	5 (10.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)	0.962 [0.147, 6.311]	0.958 [0.124, 7.383]	-0.3 [-19.0, 18.4]	1.000
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.882
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	12 (14.6)	0.780 [0.390, 1.562]	0.743 [0.324, 1.705]	-4.1 [-16.8, 8.6]	0.532
		Irbesartan	80	15 (18.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	0.622 [0.195, 1.985]	0.585 [0.159, 2.151]	-5.4 [-20.4, 9.6]	0.528
		Irbesartan	49	7 (14.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	1.176 [0.384, 3.596]	1.200 [0.340, 4.230]	1.8 [-12.8, 16.5]	1.000
		Irbesartan	48	5 (10.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)	0.962 [0.147, 6.311]	0.958 [0.124, 7.383]	-0.3 [-19.0, 18.4]	1.000
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.731
<= 1.75 g/day	Double-blind period	Sparsentan	98	9 (9.2)	0.949 [0.394, 2.286]	0.944 [0.357, 2.492]	-0.5 [-9.8, 8.8]	1.000
		Irbesartan	93	9 (9.7)				
> 1.75 g/day	Double-blind period	Sparsentan	104	15 (14.4)	0.786 [0.426, 1.451]	0.750 [0.361, 1.558]	-3.9 [-14.8, 6.9]	0.465
		Irbesartan	109	20 (18.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
 Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.911
Yes	Double-blind period	Sparsentan	90	13 (14.4)	0.847 [0.428, 1.676]	0.822 [0.366, 1.845]	-2.6 [-14.4, 9.2]	0.684
		Irbesartan	88	15 (17.0)				
No	Double-blind period	Sparsentan	112	11 (9.8)	0.800 [0.380, 1.685]	0.778 [0.337, 1.796]	-2.5 [-11.5, 6.6]	0.673
		Irbesartan	114	14 (12.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.779
<= 5 years	Double-blind period	Sparsentan	113	18 (15.9)	0.843 [0.483, 1.470]	0.813 [0.415, 1.592]	-3.0 [-13.4, 7.5]	0.611
		Irbesartan	127	24 (18.9)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	1.011 [0.321, 3.182]	1.012 [0.296, 3.458]	0.1 [-8.8, 9.0]	1.000
		Irbesartan	75	5 (6.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSIM: Incidence of severe TEAEs during double-blind period by subgroup  
 Safety Set

Severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.757
Yes	Double-blind period	Sparsentan	155	20 (12.9)	0.866 [0.499, 1.502]	0.846 [0.446, 1.603]	-2.0 [-10.3, 6.3]	0.629
		Irbesartan	161	24 (14.9)				
No	Double-blind period	Sparsentan	47	4 (8.5)	0.698 [0.201, 2.427]	0.670 [0.167, 2.682]	-3.7 [-18.8, 11.4]	0.728
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.635
Male	Double-blind period	Sparsentan	139	48 (34.5)	1.008 [0.730, 1.391]	1.012 [0.619, 1.654]	0.3 [-11.5, 12.1]	1.000
		Irbesartan	143	49 (34.3)				
Female	Double-blind period	Sparsentan	63	27 (42.9)	1.149 [0.743, 1.779]	1.261 [0.610, 2.608]	5.6 [-13.4, 24.6]	0.582
		Irbesartan	59	22 (37.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.965
<= 45 years	Double-blind period	Sparsentan	96	42 (43.8)	1.056 [0.762, 1.464]	1.100 [0.624, 1.942]	2.3 [-12.6, 17.2]	0.773
		Irbesartan	99	41 (41.4)				
> 45 years	Double-blind period	Sparsentan	106	33 (31.1)	1.069 [0.707, 1.616]	1.100 [0.609, 1.987]	2.0 [-11.4, 15.4]	0.765
		Irbesartan	103	30 (29.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.763
<= 18 years	Double-blind period	Sparsentan	9	6 (66.7)	0.833 [0.441, 1.575]	0.500 [0.037, 6.683]	-13.3 [-75.6, 48.9]	1.000
		Irbesartan	5	4 (80.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	35 (34.3)	1.100 [0.747, 1.620]	1.152 [0.648, 2.049]	3.1 [-10.5, 16.7]	0.661
		Irbesartan	109	34 (31.2)				
> 40 years	Double-blind period	Sparsentan	91	34 (37.4)	0.996 [0.682, 1.455]	0.994 [0.543, 1.821]	-0.1 [-15.4, 15.2]	1.000
		Irbesartan	88	33 (37.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.591
North America	Double-blind period	Sparsentan	35	15 (42.9)	1.232 [0.711, 2.136]	1.406 [0.570, 3.471]	8.1 [-15.8, 32.0]	0.496
		Irbesartan	46	16 (34.8)				
Europe	Double-blind period	Sparsentan	98	37 (37.8)	1.113 [0.777, 1.596]	1.182 [0.674, 2.073]	3.8 [-10.0, 17.7]	0.569
		Irbesartan	115	39 (33.9)				
Asia Pacific	Double-blind period	Sparsentan	69	23 (33.3)	0.854 [0.514, 1.419]	0.781 [0.350, 1.743]	-5.7 [-26.3, 14.9]	0.681
		Irbesartan	41	16 (39.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.732
< 27 kg/m**2	Double-blind period	Sparsentan	83	31 (37.3)	1.003 [0.684, 1.471]	1.005 [0.546, 1.850]	0.1 [-15.3, 15.5]	1.000
		Irbesartan	94	35 (37.2)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	44 (37.0)	1.099 [0.771, 1.567]	1.157 [0.669, 2.000]	3.3 [-10.0, 16.7]	0.676
		Irbesartan	107	36 (33.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.916
eGFR Low and UP High	Double-blind period	Sparsentan	71	29 (40.8)	1.119 [0.742, 1.688]	1.202 [0.615, 2.347]	4.4 [-12.9, 21.6]	0.613
		Irbesartan	74	27 (36.5)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	17 (30.9)	1.063 [0.600, 1.881]	1.090 [0.482, 2.466]	1.8 [-17.1, 20.8]	1.000
		Irbesartan	55	16 (29.1)				
eGFR High and UP High	Double-blind period	Sparsentan	37	14 (37.8)	1.135 [0.611, 2.109]	1.217 [0.466, 3.179]	4.5 [-20.2, 29.2]	0.808
		Irbesartan	36	12 (33.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	15 (38.5)	0.889 [0.517, 1.529]	0.820 [0.328, 2.050]	-4.8 [-29.5, 19.9]	0.816
		Irbesartan	37	16 (43.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.746
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	47 (37.0)	1.038 [0.750, 1.435]	1.060 [0.637, 1.764]	1.3 [-11.2, 13.9]	0.897
		Irbesartan	129	46 (35.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	19 (38.8)	0.980 [0.597, 1.608]	0.967 [0.428, 2.185]	-0.8 [-22.3, 20.7]	1.000
		Irbesartan	48	19 (39.6)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	9 (34.6)	1.442 [0.601, 3.460]	1.676 [0.494, 5.694]	10.6 [-18.1, 39.3]	0.541
		Irbesartan	25	6 (24.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.608
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	32 (39.0)	1.201 [0.792, 1.820]	1.329 [0.698, 2.533]	6.5 [-9.4, 22.5]	0.416
		Irbesartan	80	26 (32.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	15 (33.3)	0.817 [0.479, 1.392]	0.725 [0.312, 1.682]	-7.5 [-29.1, 14.1]	0.524
		Irbesartan	49	20 (40.8)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	19 (38.8)	0.980 [0.597, 1.608]	0.967 [0.428, 2.185]	-0.8 [-22.3, 20.7]	1.000
		Irbesartan	48	19 (39.6)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	9 (34.6)	1.442 [0.601, 3.460]	1.676 [0.494, 5.694]	10.6 [-18.1, 39.3]	0.541
		Irbesartan	25	6 (24.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.945
<= 1.75 g/day	Double-blind period	Sparsentan	98	36 (36.7)	1.068 [0.728, 1.565]	1.107 [0.612, 2.003]	2.3 [-12.3, 17.0]	0.764
		Irbesartan	93	32 (34.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	39 (37.5)	1.048 [0.736, 1.492]	1.077 [0.617, 1.881]	1.7 [-12.2, 15.6]	0.887
		Irbesartan	109	39 (35.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.178
Yes	Double-blind period	Sparsentan	90	30 (33.3)	0.863 [0.582, 1.278]	0.794 [0.430, 1.466]	-5.3 [-20.5, 9.9]	0.533
		Irbesartan	88	34 (38.6)				
No	Double-blind period	Sparsentan	112	45 (40.2)	1.238 [0.874, 1.753]	1.398 [0.811, 2.409]	7.7 [-5.7, 21.1]	0.269
		Irbesartan	114	37 (32.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.021 #
<= 5 years	Double-blind period	Sparsentan	113	42 (37.2)	0.858 [0.629, 1.172]	0.774 [0.461, 1.301]	-6.1 [-19.4, 7.1]	0.358
		Irbesartan	127	55 (43.3)				
> 5 years	Double-blind period	Sparsentan	89	33 (37.1)	1.738 [1.042, 2.900]	2.173 [1.079, 4.377]	15.7 [0.9, 30.6]	0.039 *
		Irbesartan	75	16 (21.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSIM: Incidence of serious TEAEs during double-blind period by subgroup  
 Safety Set

Serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.405
Yes	Double-blind period	Sparsentan	155	58 (37.4)	1.004 [0.755, 1.336]	1.007 [0.638, 1.588]	0.2 [-11.1, 11.5]	1.000
		Irbesartan	161	60 (37.3)				
No	Double-blind period	Sparsentan	47	17 (36.2)	1.348 [0.716, 2.537]	1.545 [0.621, 3.846]	9.3 [-12.2, 30.9]	0.370
		Irbesartan	41	11 (26.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.936
Male	Double-blind period	Sparsentan	139	15 (10.8)	1.187 [0.587, 2.403]	1.210 [0.553, 2.645]	1.7 [-6.0, 9.4]	0.693
		Irbesartan	143	13 (9.1)				
Female	Double-blind period	Sparsentan	63	6 (9.5)	1.124 [0.362, 3.487]	1.137 [0.328, 3.943]	1.0 [-10.7, 12.8]	1.000
		Irbesartan	59	5 (8.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.250
<= 45 years	Double-blind period	Sparsentan	96	10 (10.4)	0.859 [0.390, 1.895]	0.843 [0.346, 2.054]	-1.7 [-11.6, 8.2]	0.822
		Irbesartan	99	12 (12.1)				
> 45 years	Double-blind period	Sparsentan	106	11 (10.4)	1.781 [0.684, 4.639]	1.872 [0.666, 5.265]	4.6 [-3.8, 12.9]	0.313
		Irbesartan	103	6 (5.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.587
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	0.556 [0.109, 2.826]	0.429 [0.040, 4.637]	-17.8 [-84.1, 48.6]	0.580
		Irbesartan	5	2 (40.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	11 (10.8)	1.069 [0.485, 2.357]	1.077 [0.445, 2.604]	0.7 [-8.5, 9.9]	1.000
		Irbesartan	109	11 (10.1)				
> 40 years	Double-blind period	Sparsentan	91	8 (8.8)	1.547 [0.526, 4.548]	1.600 [0.503, 5.094]	3.1 [-5.6, 11.8]	0.567
		Irbesartan	88	5 (5.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
 Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.691
North America	Double-blind period	Sparsentan	35	4 (11.4)	0.876 [0.268, 2.870]	0.860 [0.223, 3.316]	-1.6 [-18.5, 15.2]	1.000
		Irbesartan	46	6 (13.0)				
Europe	Double-blind period	Sparsentan	98	12 (12.2)	1.565 [0.688, 3.557]	1.643 [0.662, 4.082]	4.4 [-4.7, 13.5]	0.358
		Irbesartan	115	9 (7.8)				
Asia Pacific	Double-blind period	Sparsentan	69	5 (7.2)	0.990 [0.250, 3.929]	0.990 [0.224, 4.376]	-0.1 [-12.1, 11.9]	1.000
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
 Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.754
< 27 kg/m**2	Double-blind period	Sparsentan	83	8 (9.6)	1.294 [0.490, 3.416]	1.326 [0.459, 3.828]	2.2 [-7.2, 11.6]	0.788
		Irbesartan	94	7 (7.4)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	13 (10.9)	1.063 [0.497, 2.271]	1.070 [0.458, 2.502]	0.6 [-8.3, 9.6]	1.000
		Irbesartan	107	11 (10.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.216
eGFR Low and UP High	Double-blind period	Sparsentan	71	9 (12.7)	0.853 [0.376, 1.933]	0.831 [0.322, 2.146]	-2.2 [-14.8, 10.4]	0.811
		Irbesartan	74	11 (14.9)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	7 (12.7)	7.000 [0.891, 55.014]	7.875 [0.935, 66.337]	10.9 [-0.4, 22.2]	0.060
		Irbesartan	55	1 (1.8)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)	0.584 [0.150, 2.265]	0.547 [0.121, 2.481]	-5.8 [-22.8, 11.3]	0.479
		Irbesartan	36	5 (13.9)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)	1.897 [0.180, 20.054]	1.946 [0.169, 22.413]	2.4 [-8.9, 13.7]	1.000
		Irbesartan	37	1 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.749
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	16 (12.6)	1.354 [0.668, 2.747]	1.405 [0.636, 3.104]	3.3 [-5.1, 11.7]	0.429
		Irbesartan	129	12 (9.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.784 [0.224, 2.744]	0.764 [0.192, 3.037]	-2.3 [-15.9, 11.4]	0.740
		Irbesartan	48	5 (10.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.774
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	9 (11.0)	1.098 [0.446, 2.703]	1.110 [0.406, 3.036]	1.0 [-9.7, 11.6]	1.000
		Irbesartan	80	8 (10.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	7 (15.6)	1.906 [0.597, 6.078]	2.072 [0.564, 7.621]	7.4 [-7.8, 22.6]	0.342
		Irbesartan	49	4 (8.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.784 [0.224, 2.744]	0.764 [0.192, 3.037]	-2.3 [-15.9, 11.4]	0.740
		Irbesartan	48	5 (10.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.576
<= 1.75 g/day	Double-blind period	Sparsentan	98	8 (8.2)	1.518 [0.515, 4.474]	1.564 [0.493, 4.967]	2.8 [-5.4, 10.9]	0.569
		Irbesartan	93	5 (5.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	13 (12.5)	1.048 [0.510, 2.154]	1.055 [0.464, 2.396]	0.6 [-9.2, 10.3]	1.000
		Irbesartan	109	13 (11.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.824
Yes	Double-blind period	Sparsentan	90	10 (11.1)	1.086 [0.464, 2.545]	1.097 [0.423, 2.845]	0.9 [-9.3, 11.1]	1.000
		Irbesartan	88	9 (10.2)				
No	Double-blind period	Sparsentan	112	11 (9.8)	1.244 [0.536, 2.886]	1.271 [0.505, 3.196]	1.9 [-6.4, 10.2]	0.646
		Irbesartan	114	9 (7.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.347
<= 5 years	Double-blind period	Sparsentan	113	12 (10.6)	1.499 [0.656, 3.424]	1.558 [0.631, 3.848]	3.5 [-4.5, 11.6]	0.367
		Irbesartan	127	9 (7.1)				
> 5 years	Double-blind period	Sparsentan	89	9 (10.1)	0.843 [0.353, 2.014]	0.825 [0.310, 2.198]	-1.9 [-12.8, 9.0]	0.804
		Irbesartan	75	9 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AT\_SSIM: Incidence of TEAEs leading to study drug discontinuation during double-blind period by subgroup  
 Safety Set

TEAEs leading to study drug discontinuation								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.582
Yes	Double-blind period	Sparsentan	155	16 (10.3)	1.278 [0.636, 2.569]	1.310 [0.608, 2.823]	2.2 [-4.8, 9.3]	0.561
		Irbesartan	161	13 (8.1)				
No	Double-blind period	Sparsentan	47	5 (10.6)	0.872 [0.272, 2.801]	0.857 [0.230, 3.199]	-1.6 [-17.2, 14.1]	1.000
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AD\_SSIM: Incidence of fatal TEAEs during double-blind period by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.853
Male	Double-blind period	Sparsentan	139	128 (92.1)	1.071 [0.986, 1.162]	1.892 [0.871, 4.112]	6.1 [-1.9, 14.0]	0.128
		Irbesartan	143	123 (86.0)				
Female	Double-blind period	Sparsentan	63	58 (92.1)	1.086 [0.954, 1.238]	2.088 [0.657, 6.639]	7.3 [-5.7, 20.3]	0.260
		Irbesartan	59	50 (84.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.814
<= 45 years	Double-blind period	Sparsentan	96	90 (93.8)	1.067 [0.975, 1.167]	2.069 [0.744, 5.757]	5.9 [-3.2, 14.9]	0.216
		Irbesartan	99	87 (87.9)				
> 45 years	Double-blind period	Sparsentan	106	96 (90.6)	1.085 [0.976, 1.205]	1.898 [0.825, 4.367]	7.1 [-3.0, 17.1]	0.151
		Irbesartan	103	86 (83.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.842
<= 18 years	Double-blind period	Sparsentan	9	9 (100.0)	1.036 + [0.783, 1.371]	1.727 + [0.030, 99.953]	0.0 [NE, NE]	NE
		Irbesartan	5	5 (100.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	95 (93.1)	1.057 [0.970, 1.153]	1.838 [0.703, 4.808]	5.1 [-3.7, 13.8]	0.245
		Irbesartan	109	96 (88.1)				
> 40 years	Double-blind period	Sparsentan	91	82 (90.1)	1.101 [0.977, 1.241]	2.025 [0.843, 4.861]	8.3 [-3.0, 19.5]	0.133
		Irbesartan	88	72 (81.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.718
North America	Double-blind period	Sparsentan	35	32 (91.4)	1.026 [0.889, 1.184]	1.301 [0.289, 5.854]	2.3 [-13.1, 17.7]	1.000
		Irbesartan	46	41 (89.1)				
Europe	Double-blind period	Sparsentan	98	90 (91.8)	1.100 [0.995, 1.216]	2.227 [0.928, 5.340]	8.4 [-1.3, 18.0]	0.097
		Irbesartan	115	96 (83.5)				
Asia Pacific	Double-blind period	Sparsentan	69	64 (92.8)	1.056 [0.926, 1.205]	1.778 [0.482, 6.557]	4.9 [-8.7, 18.6]	0.496
		Irbesartan	41	36 (87.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.535
< 27 kg/m**2	Double-blind period	Sparsentan	83	75 (90.4)	1.049 [0.942, 1.167]	1.505 [0.591, 3.833]	4.2 [-6.4, 14.8]	0.487
		Irbesartan	94	81 (86.2)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	111 (93.3)	1.097 [0.999, 1.204]	2.440 [0.999, 5.957]	8.2 [-0.8, 17.2]	0.053
		Irbesartan	107	91 (85.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.389
eGFR Low and UP High	Double-blind period	Sparsentan	71	67 (94.4)	1.108 [0.992, 1.238]	2.925 [0.885, 9.661]	9.2 [-1.9, 20.3]	0.100
		Irbesartan	74	63 (85.1)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	48 (87.3)	0.980 [0.854, 1.123]	0.840 [0.263, 2.681]	-1.8 [-15.7, 12.1]	1.000
		Irbesartan	55	49 (89.1)				
eGFR High and UP High	Double-blind period	Sparsentan	37	35 (94.6)	1.174 [0.983, 1.403]	4.224 [0.814, 21.923]	14.0 [-3.5, 31.6]	0.085
		Irbesartan	36	29 (80.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	36 (92.3)	1.067 [0.913, 1.248]	1.875 [0.415, 8.475]	5.8 [-10.6, 22.3]	0.475
		Irbesartan	37	32 (86.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.486
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	117 (92.1)	1.071 [0.982, 1.167]	1.897 [0.839, 4.288]	6.1 [-2.3, 14.5]	0.160
		Irbesartan	129	111 (86.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	45 (91.8)	1.025 [0.902, 1.165]	1.308 [0.329, 5.198]	2.3 [-11.4, 15.9]	0.740
		Irbesartan	48	43 (89.6)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	24 (92.3)	1.215 [0.949, 1.554]	3.789 [0.686, 20.946]	16.3 [-7.2, 39.9]	0.140
		Irbesartan	25	19 (76.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.577
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	74 (90.2)	1.046 [0.935, 1.171]	1.475 [0.560, 3.882]	4.0 [-7.2, 15.1]	0.472
		Irbesartan	80	69 (86.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	43 (95.6)	1.115 [0.978, 1.270]	3.583 [0.704, 18.252]	9.8 [-3.8, 23.5]	0.162
		Irbesartan	49	42 (85.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	45 (91.8)	1.025 [0.902, 1.165]	1.308 [0.329, 5.198]	2.3 [-11.4, 15.9]	0.740
		Irbesartan	48	43 (89.6)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	24 (92.3)	1.215 [0.949, 1.554]	3.789 [0.686, 20.946]	16.3 [-7.2, 39.9]	0.140
		Irbesartan	25	19 (76.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.644
<= 1.75 g/day	Double-blind period	Sparsentan	98	88 (89.8)	1.057 [0.948, 1.178]	1.559 [0.656, 3.709]	4.8 [-5.6, 15.3]	0.384
		Irbesartan	93	79 (84.9)				
> 1.75 g/day	Double-blind period	Sparsentan	104	98 (94.2)	1.093 [1.000, 1.194]	2.606 [0.970, 7.001]	8.0 [-0.8, 16.8]	0.066
		Irbesartan	109	94 (86.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.457
Yes	Double-blind period	Sparsentan	90	80 (88.9)	1.043 [0.931, 1.168]	1.387 [0.574, 3.352]	3.7 [-7.3, 14.6]	0.509
		Irbesartan	88	75 (85.2)				
No	Double-blind period	Sparsentan	112	106 (94.6)	1.101 [1.010, 1.200]	2.884 [1.085, 7.667]	8.7 [0.2, 17.2]	0.042 *
		Irbesartan	114	98 (86.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.894
<= 5 years	Double-blind period	Sparsentan	113	105 (92.9)	1.073 [0.985, 1.168]	2.028 [0.840, 4.899]	6.3 [-2.1, 14.7]	0.139
		Irbesartan	127	110 (86.6)				
> 5 years	Double-blind period	Sparsentan	89	81 (91.0)	1.083 [0.962, 1.220]	1.929 [0.743, 5.003]	7.0 [-4.4, 18.4]	0.231
		Irbesartan	75	63 (84.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AA\_SSIM: Incidence of non-disease related TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.076
Yes	Double-blind period	Sparsentan	155	144 (92.9)	1.108 [1.022, 1.201]	2.521 [1.199, 5.300]	9.1 [1.4, 16.7]	0.014 *
		Irbesartan	161	135 (83.9)				
No	Double-blind period	Sparsentan	47	42 (89.4)	0.964 [0.846, 1.099]	0.663 [0.148, 2.963]	-3.3 [-17.5, 10.8]	0.719
		Irbesartan	41	38 (92.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.982
Male	Double-blind period	Sparsentan	139	128 (92.1)	1.088 [0.999, 1.185]	2.116 [0.984, 4.548]	7.5 [-0.7, 15.6]	0.064
		Irbesartan	143	121 (84.6)				
Female	Double-blind period	Sparsentan	63	58 (92.1)	1.086 [0.954, 1.238]	2.088 [0.657, 6.639]	7.3 [-5.7, 20.3]	0.260
		Irbesartan	59	50 (84.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.817
<= 45 years	Double-blind period	Sparsentan	96	90 (93.8)	1.079 [0.984, 1.184]	2.267 [0.825, 6.235]	6.9 [-2.4, 16.1]	0.147
		Irbesartan	99	86 (86.9)				
> 45 years	Double-blind period	Sparsentan	106	96 (90.6)	1.097 [0.985, 1.223]	2.033 [0.890, 4.645]	8.0 [-2.1, 18.2]	0.106
		Irbesartan	103	85 (82.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.810
<= 18 years	Double-blind period	Sparsentan	9	9 (100.0)	1.036 + [0.783, 1.371]	1.727 + [0.030, 99.953]	0.0 [NE, NE]	NE
		Irbesartan	5	5 (100.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	95 (93.1)	1.069 [0.977, 1.168]	2.000 [0.773, 5.176]	6.0 [-2.9, 14.9]	0.172
		Irbesartan	109	95 (87.2)				
> 40 years	Double-blind period	Sparsentan	91	82 (90.1)	1.117 [0.988, 1.263]	2.182 [0.916, 5.198]	9.4 [-2.0, 20.8]	0.091
		Irbesartan	88	71 (80.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.562
North America	Double-blind period	Sparsentan	35	32 (91.4)	1.026 [0.889, 1.184]	1.301 [0.289, 5.854]	2.3 [-13.1, 17.7]	1.000
		Irbesartan	46	41 (89.1)				
Europe	Double-blind period	Sparsentan	98	90 (91.8)	1.124 [1.012, 1.247]	2.513 [1.059, 5.964]	10.1 [0.3, 19.9]	0.044 *
		Irbesartan	115	94 (81.7)				
Asia Pacific	Double-blind period	Sparsentan	69	64 (92.8)	1.056 [0.926, 1.205]	1.778 [0.482, 6.557]	4.9 [-8.7, 18.6]	0.496
		Irbesartan	41	36 (87.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.558
< 27 kg/m**2	Double-blind period	Sparsentan	83	75 (90.4)	1.062 [0.951, 1.185]	1.641 [0.651, 4.133]	5.3 [-5.5, 16.0]	0.364
		Irbesartan	94	80 (85.1)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	111 (93.3)	1.109 [1.008, 1.220]	2.621 [1.082, 6.351]	9.2 [0.0, 18.3]	0.034 *
		Irbesartan	107	90 (84.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.537
eGFR Low and UP High	Double-blind period	Sparsentan	71	67 (94.4)	1.108 [0.992, 1.238]	2.925 [0.885, 9.661]	9.2 [-1.9, 20.3]	0.100
		Irbesartan	74	63 (85.1)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	48 (87.3)	1.000 [0.867, 1.153]	1.000 [0.326, 3.069]	0.0 [-14.3, 14.3]	1.000
		Irbesartan	55	48 (87.3)				
eGFR High and UP High	Double-blind period	Sparsentan	37	35 (94.6)	1.174 [0.983, 1.403]	4.224 [0.814, 21.923]	14.0 [-3.5, 31.6]	0.085
		Irbesartan	36	29 (80.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	36 (92.3)	1.102 [0.931, 1.304]	2.323 [0.536, 10.069]	8.5 [-8.6, 25.7]	0.303
		Irbesartan	37	31 (83.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.595
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	117 (92.1)	1.080 [0.989, 1.180]	2.021 [0.900, 4.537]	6.9 [-1.6, 15.3]	0.114
		Irbesartan	129	110 (85.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	45 (91.8)	1.050 [0.916, 1.202]	1.607 [0.424, 6.096]	4.3 [-9.8, 18.5]	0.524
		Irbesartan	48	42 (87.5)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	24 (92.3)	1.215 [0.949, 1.554]	3.789 [0.686, 20.946]	16.3 [-7.2, 39.9]	0.140
		Irbesartan	25	19 (76.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.582
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	74 (90.2)	1.046 [0.935, 1.171]	1.475 [0.560, 3.882]	4.0 [-7.2, 15.1]	0.472
		Irbesartan	80	69 (86.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	43 (95.6)	1.142 [0.994, 1.312]	4.195 [0.841, 20.934]	11.9 [-2.2, 26.0]	0.094
		Irbesartan	49	41 (83.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	45 (91.8)	1.050 [0.916, 1.202]	1.607 [0.424, 6.096]	4.3 [-9.8, 18.5]	0.524
		Irbesartan	48	42 (87.5)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	24 (92.3)	1.215 [0.949, 1.554]	3.789 [0.686, 20.946]	16.3 [-7.2, 39.9]	0.140
		Irbesartan	25	19 (76.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.919
<= 1.75 g/day	Double-blind period	Sparsentan	98	88 (89.8)	1.085 [0.968, 1.216]	1.829 [0.784, 4.266]	7.0 [-3.8, 17.8]	0.206
		Irbesartan	93	77 (82.8)				
> 1.75 g/day	Double-blind period	Sparsentan	104	98 (94.2)	1.093 [1.000, 1.194]	2.606 [0.970, 7.001]	8.0 [-0.8, 16.8]	0.066
		Irbesartan	109	94 (86.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.495
Yes	Double-blind period	Sparsentan	90	80 (88.9)	1.057 [0.941, 1.188]	1.514 [0.634, 3.616]	4.8 [-6.4, 15.9]	0.386
		Irbesartan	88	74 (84.1)				
No	Double-blind period	Sparsentan	112	106 (94.6)	1.112 [1.018, 1.215]	3.096 [1.173, 8.173]	9.6 [0.9, 18.2]	0.026 *
		Irbesartan	114	97 (85.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.827
<= 5 years	Double-blind period	Sparsentan	113	105 (92.9)	1.083 [0.992, 1.181]	2.167 [0.904, 5.199]	7.1 [-1.4, 15.6]	0.097
		Irbesartan	127	109 (85.8)				
> 5 years	Double-blind period	Sparsentan	89	81 (91.0)	1.101 [0.974, 1.244]	2.123 [0.829, 5.439]	8.3 [-3.3, 20.0]	0.159
		Irbesartan	75	62 (82.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAN\_SSIM: Incidence of non-disease related non-severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related non-severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.052
Yes	Double-blind period	Sparsentan	155	144 (92.9)	1.125 [1.035, 1.222]	2.756 [1.320, 5.754]	10.3 [2.5, 18.0]	0.006 *
		Irbesartan	161	133 (82.6)				
No	Double-blind period	Sparsentan	47	42 (89.4)	0.964 [0.846, 1.099]	0.663 [0.148, 2.963]	-3.3 [-17.5, 10.8]	0.719
		Irbesartan	41	38 (92.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.178
Male	Double-blind period	Sparsentan	139	16 (11.5)	1.097 [0.565, 2.133]	1.110 [0.526, 2.342]	1.0 [-7.0, 9.0]	0.850
		Irbesartan	143	15 (10.5)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	0.401 [0.109, 1.480]	0.371 [0.091, 1.510]	-7.1 [-18.5, 4.3]	0.195
		Irbesartan	59	7 (11.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.744
<= 45 years	Double-blind period	Sparsentan	96	11 (11.5)	0.945 [0.438, 2.039]	0.938 [0.393, 2.242]	-0.7 [-10.7, 9.4]	1.000
		Irbesartan	99	12 (12.1)				
> 45 years	Double-blind period	Sparsentan	106	8 (7.5)	0.777 [0.319, 1.892]	0.759 [0.287, 2.007]	-2.2 [-10.7, 6.4]	0.628
		Irbesartan	103	10 (9.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.615
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.200 + [0.010, 4.166]	0.158 + [0.005, 4.691]	-20.0 [-70.6, 30.6]	0.357
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	10 (9.8)	0.971 [0.431, 2.190]	0.968 [0.393, 2.388]	-0.3 [-9.3, 8.7]	1.000
		Irbesartan	109	11 (10.1)				
> 40 years	Double-blind period	Sparsentan	91	9 (9.9)	0.870 [0.371, 2.039]	0.856 [0.330, 2.219]	-1.5 [-11.6, 8.7]	0.811
		Irbesartan	88	10 (11.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.663
North America	Double-blind period	Sparsentan	35	6 (17.1)	1.127 [0.415, 3.056]	1.153 [0.350, 3.795]	1.9 [-16.8, 20.7]	1.000
		Irbesartan	46	7 (15.2)				
Europe	Double-blind period	Sparsentan	98	8 (8.2)	0.722 [0.312, 1.670]	0.697 [0.276, 1.759]	-3.1 [-12.0, 5.7]	0.496
		Irbesartan	115	13 (11.3)				
Asia Pacific	Double-blind period	Sparsentan	69	5 (7.2)	1.486 [0.302, 7.311]	1.523 [0.282, 8.235]	2.4 [-8.6, 13.3]	1.000
		Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.836
< 27 kg/m**2	Double-blind period	Sparsentan	83	7 (8.4)	0.793 [0.316, 1.989]	0.774 [0.281, 2.134]	-2.2 [-12.0, 7.6]	0.799
		Irbesartan	94	10 (10.6)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	12 (10.1)	0.899 [0.422, 1.916]	0.888 [0.381, 2.070]	-1.1 [-10.1, 7.8]	0.831
		Irbesartan	107	12 (11.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.497
eGFR Low and UP High	Double-blind period	Sparsentan	71	9 (12.7)	1.173 [0.479, 2.870]	1.198 [0.435, 3.300]	1.9 [-10.0, 13.7]	0.800
		Irbesartan	74	8 (10.8)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5)	0.375 [0.105, 1.340]	0.339 [0.085, 1.353]	-9.1 [-22.0, 3.8]	0.202
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)	0.973 [0.210, 4.507]	0.971 [0.183, 5.158]	-0.2 [-15.6, 15.1]	1.000
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	4 (10.3)	1.265 [0.303, 5.274]	1.295 [0.270, 6.223]	2.1 [-13.4, 17.7]	1.000
		Irbesartan	37	3 (8.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.574
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	12 (9.4)	0.717 [0.357, 1.440]	0.687 [0.314, 1.505]	-3.7 [-12.3, 4.8]	0.431
		Irbesartan	129	17 (13.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	1.633 [0.413, 6.457]	1.705 [0.384, 7.567]	4.0 [-9.0, 16.9]	0.715
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)	0.962 [0.147, 6.311]	0.958 [0.124, 7.383]	-0.3 [-19.0, 18.4]	1.000
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
 Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.754
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	8 (9.8)	0.780 [0.325, 1.876]	0.757 [0.282, 2.027]	-2.7 [-13.7, 8.2]	0.624
		Irbesartan	80	10 (12.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	0.622 [0.195, 1.985]	0.585 [0.159, 2.151]	-5.4 [-20.4, 9.6]	0.528
		Irbesartan	49	7 (14.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	1.633 [0.413, 6.457]	1.705 [0.384, 7.567]	4.0 [-9.0, 16.9]	0.715
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	2 (7.7)	0.962 [0.147, 6.311]	0.958 [0.124, 7.383]	-0.3 [-19.0, 18.4]	1.000
		Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.558
<= 1.75 g/day	Double-blind period	Sparsentan	98	9 (9.2)	1.068 [0.430, 2.650]	1.074 [0.396, 2.914]	0.6 [-8.5, 9.7]	1.000
		Irbesartan	93	8 (8.6)				
> 1.75 g/day	Double-blind period	Sparsentan	104	10 (9.6)	0.749 [0.348, 1.610]	0.722 [0.305, 1.706]	-3.2 [-12.6, 6.2]	0.520
		Irbesartan	109	14 (12.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.806
Yes	Double-blind period	Sparsentan	90	9 (10.0)	0.800 [0.349, 1.836]	0.778 [0.305, 1.980]	-2.5 [-12.9, 7.9]	0.641
		Irbesartan	88	11 (12.5)				
No	Double-blind period	Sparsentan	112	10 (8.9)	0.925 [0.409, 2.092]	0.918 [0.374, 2.256]	-0.7 [-9.2, 7.7]	1.000
		Irbesartan	114	11 (9.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.772
<= 5 years	Double-blind period	Sparsentan	113	15 (13.3)	0.887 [0.474, 1.662]	0.870 [0.419, 1.806]	-1.7 [-11.3, 8.0]	0.853
		Irbesartan	127	19 (15.0)				
> 5 years	Double-blind period	Sparsentan	89	4 (4.5)	1.124 [0.260, 4.863]	1.129 [0.245, 5.213]	0.5 [-6.9, 7.9]	1.000
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AAC\_SSIM: Incidence of non-disease related severe TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related severe TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.619
Yes	Double-blind period	Sparsentan	155	17 (11.0)	0.929 [0.502, 1.721]	0.921 [0.459, 1.845]	-0.8 [-8.5, 6.8]	0.861
		Irbesartan	161	19 (11.8)				
No	Double-blind period	Sparsentan	47	2 (4.3)	0.582 [0.102, 3.312]	0.563 [0.089, 3.547]	-3.1 [-15.2, 9.1]	0.661
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.848
Male	Double-blind period	Sparsentan	139	43 (30.9)	1.053 [0.738, 1.503]	1.077 [0.648, 1.792]	1.6 [-9.9, 13.0]	0.796
		Irbesartan	143	42 (29.4)				
Female	Double-blind period	Sparsentan	63	25 (39.7)	1.115 [0.705, 1.764]	1.190 [0.571, 2.480]	4.1 [-14.7, 22.9]	0.710
		Irbesartan	59	21 (35.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.559
<= 45 years	Double-blind period	Sparsentan	96	35 (36.5)	1.003 [0.692, 1.453]	1.004 [0.560, 1.800]	0.1 [-14.4, 14.6]	1.000
		Irbesartan	99	36 (36.4)				
> 45 years	Double-blind period	Sparsentan	106	33 (31.1)	1.188 [0.772, 1.826]	1.272 [0.697, 2.322]	4.9 [-8.3, 18.1]	0.449
		Irbesartan	103	27 (26.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.190
<= 18 years	Double-blind period	Sparsentan	9	3 (33.3)	0.417 [0.150, 1.159]	0.125 [0.009, 1.671]	-46.7 [-100.0, 15.6]	0.266
		Irbesartan	5	4 (80.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	31 (30.4)	1.142 [0.744, 1.753]	1.204 [0.662, 2.192]	3.8 [-9.3, 16.9]	0.546
		Irbesartan	109	29 (26.6)				
> 40 years	Double-blind period	Sparsentan	91	34 (37.4)	1.096 [0.739, 1.625]	1.153 [0.625, 2.127]	3.3 [-11.9, 18.4]	0.755
		Irbesartan	88	30 (34.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.715
North America	Double-blind period	Sparsentan	35	14 (40.0)	1.150 [0.652, 2.027]	1.250 [0.504, 3.101]	5.2 [-18.6, 29.0]	0.650
		Irbesartan	46	16 (34.8)				
Europe	Double-blind period	Sparsentan	98	33 (33.7)	1.173 [0.786, 1.751]	1.262 [0.705, 2.258]	5.0 [-8.5, 18.4]	0.460
		Irbesartan	115	33 (28.7)				
Asia Pacific	Double-blind period	Sparsentan	69	21 (30.4)	0.891 [0.512, 1.552]	0.844 [0.370, 1.924]	-3.7 [-23.8, 16.4]	0.833
		Irbesartan	41	14 (34.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.586
< 27 kg/m**2	Double-blind period	Sparsentan	83	28 (33.7)	0.991 [0.656, 1.497]	0.986 [0.529, 1.840]	-0.3 [-15.4, 14.8]	1.000
		Irbesartan	94	32 (34.0)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	40 (33.6)	1.160 [0.786, 1.713]	1.241 [0.706, 2.184]	4.6 [-8.3, 17.6]	0.476
		Irbesartan	107	31 (29.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.825
eGFR Low and UP High	Double-blind period	Sparsentan	71	26 (36.6)	1.290 [0.803, 2.073]	1.458 [0.725, 2.933]	8.2 [-8.3, 24.8]	0.375
		Irbesartan	74	21 (28.4)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	16 (29.1)	1.000 [0.558, 1.792]	1.000 [0.439, 2.277]	0.0 [-18.8, 18.8]	1.000
		Irbesartan	55	16 (29.1)				
eGFR High and UP High	Double-blind period	Sparsentan	37	11 (29.7)	0.973 [0.484, 1.956]	0.962 [0.354, 2.614]	-0.8 [-24.6, 23.0]	1.000
		Irbesartan	36	11 (30.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	15 (38.5)	0.949 [0.544, 1.655]	0.917 [0.365, 2.301]	-2.1 [-26.7, 22.5]	1.000
		Irbesartan	37	15 (40.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.876
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	43 (33.9)	1.092 [0.766, 1.556]	1.139 [0.675, 1.923]	2.9 [-9.4, 15.1]	0.689
		Irbesartan	129	40 (31.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	17 (34.7)	0.980 [0.570, 1.684]	0.969 [0.421, 2.231]	-0.7 [-21.8, 20.3]	1.000
		Irbesartan	48	17 (35.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	8 (30.8)	1.282 [0.519, 3.169]	1.407 [0.408, 4.860]	6.8 [-21.5, 35.1]	0.755
		Irbesartan	25	6 (24.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.675
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	28 (34.1)	1.301 [0.809, 2.091]	1.457 [0.741, 2.863]	7.9 [-7.4, 23.2]	0.307
		Irbesartan	80	21 (26.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	15 (33.3)	0.860 [0.500, 1.479]	0.789 [0.339, 1.838]	-5.4 [-27.0, 16.1]	0.669
		Irbesartan	49	19 (38.8)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	17 (34.7)	0.980 [0.570, 1.684]	0.969 [0.421, 2.231]	-0.7 [-21.8, 20.3]	1.000
		Irbesartan	48	17 (35.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	8 (30.8)	1.282 [0.519, 3.169]	1.407 [0.408, 4.860]	6.8 [-21.5, 35.1]	0.755
		Irbesartan	25	6 (24.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.976
<= 1.75 g/day	Double-blind period	Sparsentan	98	35 (35.7)	1.071 [0.724, 1.585]	1.111 [0.612, 2.019]	2.4 [-12.1, 16.9]	0.762
		Irbesartan	93	31 (33.3)				
> 1.75 g/day	Double-blind period	Sparsentan	104	33 (31.7)	1.081 [0.721, 1.621]	1.118 [0.624, 2.004]	2.4 [-10.9, 15.7]	0.767
		Irbesartan	109	32 (29.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.320
Yes	Double-blind period	Sparsentan	90	25 (27.8)	0.905 [0.573, 1.431]	0.869 [0.455, 1.659]	-2.9 [-17.4, 11.6]	0.742
		Irbesartan	88	27 (30.7)				
No	Double-blind period	Sparsentan	112	43 (38.4)	1.216 [0.850, 1.739]	1.350 [0.780, 2.337]	6.8 [-6.5, 20.1]	0.329
		Irbesartan	114	36 (31.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.032 #
<= 5 years	Double-blind period	Sparsentan	113	37 (32.7)	0.866 [0.613, 1.224]	0.801 [0.471, 1.364]	-5.1 [-18.0, 7.9]	0.421
		Irbesartan	127	48 (37.8)				
> 5 years	Double-blind period	Sparsentan	89	31 (34.8)	1.742 [1.021, 2.972]	2.138 [1.047, 4.367]	14.8 [0.2, 29.5]	0.038 *
		Irbesartan	75	15 (20.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AAS\_SSIM: Incidence of non-disease related serious TEAEs during double-blind period by subgroup  
Safety Set

Non-disease related serious TEAEs								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.546
Yes	Double-blind period	Sparsentan	155	53 (34.2)	1.039 [0.762, 1.417]	1.059 [0.664, 1.689]	1.3 [-9.8, 12.3]	0.813
		Irbesartan	161	53 (32.9)				
No	Double-blind period	Sparsentan	47	15 (31.9)	1.309 [0.662, 2.587]	1.453 [0.567, 3.721]	7.5 [-13.5, 28.5]	0.484
		Irbesartan	41	10 (24.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Sex	Double-blind period	Sparsentan					Interaction test:	0.558
	Male	Double-blind period	Sparsentan	139	15 (10.8)	1.929 [0.845, 4.405]	2.041 [0.837, 4.981]	5.2 [-1.9, 12.3]	0.130
			Irbesartan	143	8 (5.6)				
	Female	Double-blind period	Sparsentan	63	15 (23.8)	2.810 [1.089, 7.249]	3.375 [1.141, 9.980]	15.3 [1.0, 29.7]	0.028 *
			Irbesartan	59	5 (8.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Age	Double-blind period	Sparsentan					Interaction test:	0.720
	<= 45 years	Double-blind period	Sparsentan	96	14 (14.6)	2.063 [0.870, 4.887]	2.244 [0.864, 5.830]	7.5 [-2.2, 17.2]	0.108
			Irbesartan	99	7 (7.1)				
	> 45 years	Double-blind period	Sparsentan	106	16 (15.1)	2.591 [1.055, 6.362]	2.874 [1.077, 7.667]	9.3 [0.1, 18.4]	0.041 *
			Irbesartan	103	6 (5.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Age at IgAN diagnosis	Double-blind period	Sparsentan					Interaction test:	0.959
	<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
			Irbesartan	5	0 (0.0)				
	> 18 to 40 years	Double-blind period	Sparsentan	102	16 (15.7)	2.137 [0.956, 4.778]	2.349 [0.959, 5.755]	8.3 [-1.2, 17.9]	0.081
			Irbesartan	109	8 (7.3)				
	> 40 years	Double-blind period	Sparsentan	91	13 (14.3)	2.514 [0.935, 6.759]	2.767 [0.943, 8.120]	8.6 [-1.2, 18.4]	0.080
			Irbesartan	88	5 (5.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Geographic region	Double-blind period	Sparsentan					Interaction test:	0.326
	North America	Double-blind period	Sparsentan	35	4 (11.4)	2.629 [0.510, 13.543]	2.839 [0.489, 16.475]	7.1 [-7.5, 21.7]	0.395
			Irbesartan	46	2 (4.3)				
	Europe	Double-blind period	Sparsentan	98	7 (7.1)	4.107 [0.873, 19.316]	4.346 [0.881, 21.431]	5.4 [-1.2, 12.0]	0.084
			Irbesartan	115	2 (1.7)				
	Asia Pacific	Double-blind period	Sparsentan	69	19 (27.5)	1.254 [0.628, 2.507]	1.351 [0.545, 3.353]	5.6 [-12.8, 24.0]	0.652
			Irbesartan	41	9 (22.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.964
	< 27 kg/m**2	Double-blind period	Sparsentan	83	17 (20.5)	2.407 [1.096, 5.286]	2.769 [1.126, 6.807]	12.0 [0.5, 23.5]	0.030 *
			Irbesartan	94	8 (8.5)				
	>= 27 kg/m**2	Double-blind period	Sparsentan	119	13 (10.9)	2.338 [0.862, 6.341]	2.502 [0.861, 7.269]	6.3 [-1.5, 14.0]	0.091
			Irbesartan	107	5 (4.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Randomization strata	Double-blind period	Sparsentan					Interaction test:	0.650
	eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	1.390 [0.508, 3.805]	1.439 [0.473, 4.378]	3.2 [-7.9, 14.2]	0.582
			Irbesartan	74	6 (8.1)				
	eGFR Low and UP Low	Double-blind period	Sparsentan	55	10 (18.2)	2.500 [0.834, 7.493]	2.833 [0.831, 9.663]	10.9 [-3.2, 25.0]	0.151
			Irbesartan	55	4 (7.3)				
	eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	3.892 [0.457, 33.169]	4.242 [0.451, 39.943]	8.0 [-6.1, 22.1]	0.358
			Irbesartan	36	1 (2.8)				
	eGFR High and UP Low	Double-blind period	Sparsentan	39	8 (20.5)	3.795 [0.861, 16.717]	4.516 [0.891, 22.892]	15.1 [-2.1, 32.4]	0.087
			Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Baseline eGFR Group 1	Double-blind period	Sparsentan					Interaction test:	0.608
	< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	19 (15.0)	1.930 [0.934, 3.987]	2.094 [0.932, 4.700]	7.2 [-1.3, 15.7]	0.078
			Irbesartan	129	10 (7.8)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	5.878 [0.735, 47.011]	6.558 [0.759, 56.698]	10.2 [-1.9, 22.3]	0.111
			Irbesartan	48	1 (2.1)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	5 (19.2)	2.404 [0.513, 11.271]	2.738 [0.479, 15.651]	11.2 [-11.2, 33.7]	0.419
			Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Baseline eGFR Group 2	Double-blind period	Sparsentan					Interaction test:	0.733
	< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	10 (12.2)	1.626 [0.620, 4.264]	1.713 [0.592, 4.958]	4.7 [-5.7, 15.1]	0.431
			Irbesartan	80	6 (7.5)				
	45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	9 (20.0)	2.450 [0.811, 7.405]	2.813 [0.800, 9.882]	11.8 [-4.3, 27.9]	0.136
			Irbesartan	49	4 (8.2)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	5.878 [0.735, 47.011]	6.558 [0.759, 56.698]	10.2 [-1.9, 22.3]	0.111
			Irbesartan	48	1 (2.1)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	5 (19.2)	2.404 [0.513, 11.271]	2.738 [0.479, 15.651]	11.2 [-11.2, 33.7]	0.419
			Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Baseline urine protein excretion	Double-blind period	Sparsentan					Interaction test:	0.161
	<= 1.75 g/day	Double-blind period	Sparsentan	98	19 (19.4)	3.606 [1.404, 9.263]	4.233 [1.510, 11.867]	14.0 [3.9, 24.1]	0.004 *
			Irbesartan	93	5 (5.4)				
	> 1.75 g/day	Double-blind period	Sparsentan	104	11 (10.6)	1.441 [0.604, 3.440]	1.493 [0.576, 3.874]	3.2 [-5.4, 11.9]	0.475
			Irbesartan	109	8 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.912
	Yes	Double-blind period	Sparsentan	90	9 (10.0)	2.200 [0.703, 6.883]	2.333 [0.691, 7.877]	5.5 [-3.2, 14.2]	0.249
			Irbesartan	88	4 (4.5)				
	No	Double-blind period	Sparsentan	112	21 (18.8)	2.375 [1.138, 4.958]	2.692 [1.174, 6.173]	10.9 [1.2, 20.5]	0.019 *
			Irbesartan	114	9 (7.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Dizziness	Time since renal biopsy	Double-blind period	Sparsentan					Interaction test:	0.972
	<= 5 years	Double-blind period	Sparsentan	113	19 (16.8)	2.373 [1.119, 5.031]	2.650 [1.146, 6.127]	9.7 [0.7, 18.8]	0.026 *
			Irbesartan	127	9 (7.1)				
	> 5 years	Double-blind period	Sparsentan	89	11 (12.4)	2.317 [0.770, 6.978]	2.503 [0.763, 8.217]	7.0 [-2.7, 16.8]	0.174
			Irbesartan	75	4 (5.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs										
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
PT: Dizziness	History of hypertension	Double-blind period	Interaction test:							0.397
			Yes	Double-blind period	Sparsentan	155	21 (13.5)	2.727 [1.245, 5.971]	2.997 [1.285, 6.989]	8.6 [1.6, 15.6]
				Irbesartan	161	8 (5.0)				
	No	Double-blind period	Sparsentan	47	9 (19.1)	1.570 [0.572, 4.310]	1.705 [0.522, 5.574]	7.0 [-10.4, 24.3]	0.401	
			Irbesartan	41	5 (12.2)					

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Sex	Double-blind period	Sparsentan					Interaction test:	0.904
	Male	Double-blind period	Sparsentan	139	13 (9.4)	3.344 [1.117, 10.005]	3.585 [1.139, 11.281]	6.6 [0.3, 12.8]	0.024 *
			Irbesartan	143	4 (2.8)				
	Female	Double-blind period	Sparsentan	63	13 (20.6)	3.044 [1.051, 8.811]	3.575 [1.094, 11.684]	13.9 [0.3, 27.4]	0.036 *
			Irbesartan	59	4 (6.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Age	Double-blind period	Sparsentan					Interaction test:	0.255
	<= 45 years	Double-blind period	Sparsentan	96	10 (10.4)	2.063 [0.732, 5.813]	2.186 [0.719, 6.651]	5.4 [-3.1, 13.9]	0.187
			Irbesartan	99	5 (5.1)				
	> 45 years	Double-blind period	Sparsentan	106	16 (15.1)	5.182 [1.556, 17.257]	5.926 [1.672, 21.008]	12.2 [3.7, 20.7]	0.003 *
			Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Age at IgAN diagnosis	Double-blind period	Sparsentan					Interaction test:	0.656
	<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	3.000 + [0.171, 52.527]	3.667 + [0.145, 92.653]	22.2 [-20.5, 64.9]	0.505
			Irbesartan	5	0 (0.0)				
	> 18 to 40 years	Double-blind period	Sparsentan	102	10 (9.8)	2.137 [0.756, 6.041]	2.261 [0.745, 6.857]	5.2 [-2.7, 13.1]	0.182
			Irbesartan	109	5 (4.6)				
	> 40 years	Double-blind period	Sparsentan	91	14 (15.4)	4.513 [1.343, 15.164]	5.152 [1.426, 18.612]	12.0 [2.5, 21.4]	0.009 *
			Irbesartan	88	3 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Geographic region	Double-blind period	Sparsentan					Interaction test:	0.814
	North America	Double-blind period	Sparsentan	35	4 (11.4)	2.629 [0.510, 13.543]	2.839 [0.489, 16.475]	7.1 [-7.5, 21.7]	0.395
			Irbesartan	46	2 (4.3)				
	Europe	Double-blind period	Sparsentan	98	14 (14.3)	4.107 [1.398, 12.070]	4.625 [1.469, 14.560]	10.8 [2.2, 19.4]	0.006 *
			Irbesartan	115	4 (3.5)				
	Asia Pacific	Double-blind period	Sparsentan	69	8 (11.6)	2.377 [0.530, 10.658]	2.557 [0.516, 12.676]	6.7 [-5.3, 18.7]	0.316
			Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Baseline BMI	Double-blind period	Sparsentan					Interaction test:	0.728
	< 27 kg/m**2	Double-blind period	Sparsentan	83	13 (15.7)	2.945 [1.096, 7.911]	3.306 [1.125, 9.714]	10.3 [0.2, 20.5]	0.027 *
			Irbesartan	94	5 (5.3)				
	>= 27 kg/m**2	Double-blind period	Sparsentan	119	13 (10.9)	3.896 [1.141, 13.303]	4.252 [1.177, 15.356]	8.1 [0.8, 15.4]	0.020 *
			Irbesartan	107	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Randomization strata	Double-blind period	Sparsentan					Interaction test:	0.801
	eGFR Low and UP High	Double-blind period	Sparsentan	71	9 (12.7)	2.345 [0.756, 7.273]	2.540 [0.745, 8.660]	7.3 [-3.4, 17.9]	0.153
			Irbesartan	74	4 (5.4)				
	eGFR Low and UP Low	Double-blind period	Sparsentan	55	9 (16.4)	4.500 [1.018, 19.885]	5.185 [1.066, 25.229]	12.7 [-0.0, 25.5]	0.052
			Irbesartan	55	2 (3.6)				
	eGFR High and UP High	Double-blind period	Sparsentan	37	6 (16.2)	5.838 [0.739, 46.111]	6.774 [0.772, 59.419]	13.4 [-2.3, 29.2]	0.107
			Irbesartan	36	1 (2.8)				
	eGFR High and UP Low	Double-blind period	Sparsentan	39	2 (5.1)	1.897 [0.180, 20.054]	1.946 [0.169, 22.413]	2.4 [-8.9, 13.7]	1.000
			Irbesartan	37	1 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Baseline eGFR Group 1	Double-blind period	Sparsentan					Interaction test:	0.531
	< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	19 (15.0)	3.217 [1.328, 7.790]	3.606 [1.390, 9.358]	10.3 [2.3, 18.3]	0.006 *
			Irbesartan	129	6 (4.7)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	8.820 + [0.488, 159.505]	9.593 + [0.502, 183.220]	8.2 [-1.6, 17.9]	0.117
			Irbesartan	48	0 (0.0)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	3 (11.5)	1.442 [0.263, 7.918]	1.500 [0.229, 9.832]	3.5 [-16.6, 23.7]	1.000
			Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.541
	< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	9 (11.0)	2.195 [0.704, 6.842]	2.342 [0.691, 7.941]	6.0 [-3.5, 15.5]	0.247
			Irbesartan	80	4 (5.0)				
	45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	10 (22.2)	5.444 [1.260, 23.520]	6.714 [1.383, 32.597]	18.1 [2.7, 33.6]	0.012 *
			Irbesartan	49	2 (4.1)				
	60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	8.820 + [0.488, 159.505]	9.593 + [0.502, 183.220]	8.2 [-1.6, 17.9]	0.117
			Irbesartan	48	0 (0.0)				
	>= 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	3 (11.5)	1.442 [0.263, 7.918]	1.500 [0.229, 9.832]	3.5 [-16.6, 23.7]	1.000
			Irbesartan	25	2 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Baseline urine protein excretion	Double-blind period	Sparsentan					Interaction test:	0.900
	<= 1.75 g/day	Double-blind period	Sparsentan	98	11 (11.2)	3.480 [1.002, 12.080]	3.793 [1.023, 14.060]	8.0 [-0.3, 16.3]	0.050 *
			Irbesartan	93	3 (3.2)				
	> 1.75 g/day	Double-blind period	Sparsentan	104	15 (14.4)	3.144 [1.185, 8.343]	3.506 [1.226, 10.027]	9.8 [1.1, 18.6]	0.018 *
			Irbesartan	109	5 (4.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.525
	Yes	Double-blind period	Sparsentan	90	7 (7.8)	2.281 [0.609, 8.542]	2.390 [0.598, 9.555]	4.4 [-3.5, 12.2]	0.330
			Irbesartan	88	3 (3.4)				
	No	Double-blind period	Sparsentan	112	19 (17.0)	3.868 [1.496, 10.001]	4.454 [1.601, 12.391]	12.6 [3.8, 21.4]	0.002 *
			Irbesartan	114	5 (4.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

TEAEs									
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
PT: Hypotension	Time since renal biopsy	Double-blind period	Sparsentan					Interaction test:	0.572
	<= 5 years	Double-blind period	Sparsentan	113	14 (12.4)	3.934 [1.333, 11.605]	4.348 [1.388, 13.628]	9.2 [1.6, 16.9]	0.012 *
			Irbesartan	127	4 (3.1)				
	> 5 years	Double-blind period	Sparsentan	89	12 (13.5)	2.528 [0.851, 7.512]	2.766 [0.853, 8.972]	8.1 [-1.8, 18.1]	0.113
			Irbesartan	75	4 (5.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2A\_SSSM: Incidence of TEAEs during double-blind period by SOC/PT by subgroups  
 Safety Set

TEAEs										
SOC/PT	Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value	
PT: Hypotension	History of hypertension	Double-blind period	Interaction test:							0.053
			Yes	Double-blind period	Sparsentan	155	21 (13.5)	5.453 [1.915, 15.525]	6.151 [2.060, 18.365]	11.1 [4.5, 17.6]
				Irbesartan	161	4 (2.5)				
	No	Double-blind period	Interaction test:							1.000
			Sparsentan	47	5 (10.6)	1.090 [0.314, 3.792]	1.101 [0.275, 4.408]	0.9 [-14.1, 15.8]		
			Irbesartan	41	4 (9.8)					

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
 Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AC\_SSSM: Incidence of severe TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

Not done: No significant soc/pts found.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AS\_SSSM: Incidence of serious TEAEs during double-blind period by SOC/PT by subgroups  
Safety Set

Not done: No significant soc/pts found.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. TEAE = treatment emergent adverse event. SOC = system organ class. PT = preferred term.  
Only events are reported which demonstrate with significant treatment effects in main analysis. 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.559
Male	Double-blind period	Sparsentan	139	5 (3.6)	0.857 [0.268, 2.745]	0.852 [0.254, 2.858]	-0.6 [-5.8, 4.6]	1.000
		Irbesartan	143	6 (4.2)				
Female	Double-blind period	Sparsentan	63	0 (0.0)	0.313 + [0.013, 7.523]	0.307 + [0.012, 7.688]	-1.7 [-6.6, 3.2]	0.484
		Irbesartan	59	1 (1.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
 Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: NE
<= 45 years	Double-blind period	Sparsentan	96	1 (1.0) all n<10				NE
		Irbesartan	99	6 (6.1)				
> 45 years	Double-blind period	Sparsentan	106	4 (3.8) all n<10				NE
		Irbesartan	103	1 (1.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
 Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: NE
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0) all n<10				NE
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	2 (2.0) all n<10				NE
		Irbesartan	109	5 (4.6)				
> 40 years	Double-blind period	Sparsentan	91	3 (3.3) all n<10				NE
		Irbesartan	88	1 (1.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: NE
North America	Double-blind period	Sparsentan	35	2 (5.7) all n<10				NE
		Irbesartan	46	3 (6.5)				
Europe	Double-blind period	Sparsentan	98	3 (3.1) all n<10				NE
		Irbesartan	115	3 (2.6)				
Asia Pacific	Double-blind period	Sparsentan	69	0 (0.0) all n<10				NE
		Irbesartan	41	1 (2.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
 Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: NE
< 27 kg/m**2	Double-blind period	Sparsentan	83	2 (2.4)				NE
		Irbesartan	94	4 (4.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	3 (2.5)				NE
		Irbesartan	107	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	1 (1.4) all n<10				NE
		Irbesartan	74	4 (5.4)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	1 (1.8) all n<10				NE
		Irbesartan	55	0 (0.0)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4) all n<10				NE
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6) all n<10				NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
 Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: NE
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	2 (1.6)				NE
		Irbesartan	129	4 (3.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)				NE
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
 Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	2 (2.4)				NE
		Irbesartan	80	3 (3.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	0 (0.0)				NE
		Irbesartan	49	1 (2.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)				NE
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: NE
<= 1.75 g/day	Double-blind period	Sparsentan	98	2 (2.0) all n<10				NE
		Irbesartan	93	2 (2.2)				
> 1.75 g/day	Double-blind period	Sparsentan	104	3 (2.9) all n<10				NE
		Irbesartan	109	5 (4.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: NE
Yes	Double-blind period	Sparsentan	90	1 (1.1) all n<10				NE
		Irbesartan	88	2 (2.3)				
No	Double-blind period	Sparsentan	112	4 (3.6) all n<10				NE
		Irbesartan	114	5 (4.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
 Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: NE
<= 5 years	Double-blind period	Sparsentan	113	3 (2.7) all n<10				NE
		Irbesartan	127	4 (3.1)				
> 5 years	Double-blind period	Sparsentan	89	2 (2.2) all n<10				NE
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEF\_SSIM: Incidence of AESI abnormal liver function during double-blind period by subgroup  
Safety Set

AESI abnormal liver function								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.880
Yes	Double-blind period	Sparsentan	155	4 (2.6)	0.692 [0.199, 2.407]	0.684 [0.189, 2.473]	-1.1 [-5.6, 3.3]	0.750
		Irbesartan	161	6 (3.7)				
No	Double-blind period	Sparsentan	47	1 (2.1)	0.872 [0.056, 13.510]	0.870 [0.053, 14.357]	-0.3 [-8.9, 8.2]	1.000
		Irbesartan	41	1 (2.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.559
Male	Double-blind period	Sparsentan	139	5 (3.6)	0.857 [0.268, 2.745]	0.852 [0.254, 2.858]	-0.6 [-5.8, 4.6]	1.000
		Irbesartan	143	6 (4.2)				
Female	Double-blind period	Sparsentan	63	0 (0.0)	0.313 + [0.013, 7.523]	0.307 + [0.012, 7.688]	-1.7 [-6.6, 3.2]	0.484
		Irbesartan	59	1 (1.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
 Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: NE
<= 45 years	Double-blind period	Sparsentan	96	1 (1.0) all n<10				NE
		Irbesartan	99	6 (6.1)				
> 45 years	Double-blind period	Sparsentan	106	4 (3.8) all n<10				NE
		Irbesartan	103	1 (1.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: NE
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0) all n<10				NE
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	2 (2.0) all n<10				NE
		Irbesartan	109	5 (4.6)				
> 40 years	Double-blind period	Sparsentan	91	3 (3.3) all n<10				NE
		Irbesartan	88	1 (1.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: NE
North America	Double-blind period	Sparsentan	35	2 (5.7) all n<10				NE
		Irbesartan	46	3 (6.5)				
Europe	Double-blind period	Sparsentan	98	3 (3.1) all n<10				NE
		Irbesartan	115	3 (2.6)				
Asia Pacific	Double-blind period	Sparsentan	69	0 (0.0) all n<10				NE
		Irbesartan	41	1 (2.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
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p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
 Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: NE
< 27 kg/m**2	Double-blind period	Sparsentan	83	2 (2.4)				NE
		Irbesartan	94	4 (4.3)				
>= 27 kg/m**2	Double-blind period	Sparsentan	119	3 (2.5)				NE
		Irbesartan	107	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	1 (1.4)				NE
		Irbesartan	74	4 (5.4)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	1 (1.8)				NE
		Irbesartan	55	0 (0.0)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4)				NE
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)				NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
 Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: NE
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	2 (1.6)				NE
		Irbesartan	129	4 (3.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)				NE
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	2 (2.4)				NE
		Irbesartan	80	3 (3.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	0 (0.0)				NE
		Irbesartan	49	1 (2.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)				NE
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)				NE
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: NE
<= 1.75 g/day	Double-blind period	Sparsentan	98	2 (2.0) all n<10				NE
		Irbesartan	93	2 (2.2)				
> 1.75 g/day	Double-blind period	Sparsentan	104	3 (2.9) all n<10				NE
		Irbesartan	109	5 (4.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: NE
Yes	Double-blind period	Sparsentan	90	1 (1.1) all n<10				NE
		Irbesartan	88	2 (2.3)				
No	Double-blind period	Sparsentan	112	4 (3.6) all n<10				NE
		Irbesartan	114	5 (4.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
 Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: NE
<= 5 years	Double-blind period	Sparsentan	113	3 (2.7) all n<10				NE
		Irbesartan	127	4 (3.1)				
> 5 years	Double-blind period	Sparsentan	89	2 (2.2) all n<10				NE
		Irbesartan	75	3 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEFN\_SSIM: Incidence of AESI abnormal liver function during double-blind period - non-severe by subgroup  
 Safety Set

AESI abnormal liver function - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.880
Yes	Double-blind period	Sparsentan	155	4 (2.6)	0.692 [0.199, 2.407]	0.684 [0.189, 2.473]	-1.1 [-5.6, 3.3]	0.750
		Irbesartan	161	6 (3.7)				
No	Double-blind period	Sparsentan	47	1 (2.1)	0.872 [0.056, 13.510]	0.870 [0.053, 14.357]	-0.3 [-8.9, 8.2]	1.000
		Irbesartan	41	1 (2.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEFC\_SSIM: Incidence of AESI abnormal liver function during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEFS\_SSIM: Incidence of AESI abnormal liver function during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.865
Male	Double-blind period	Sparsentan	139	33 (23.7)	1.171 [0.753, 1.819]	1.224 [0.696, 2.152]	3.5 [-6.9, 13.8]	0.565
		Irbesartan	143	29 (20.3)				
Female	Double-blind period	Sparsentan	63	20 (31.7)	1.102 [0.642, 1.891]	1.149 [0.530, 2.491]	2.9 [-15.0, 20.9]	0.844
		Irbesartan	59	17 (28.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.499
<= 45 years	Double-blind period	Sparsentan	96	30 (31.3)	1.289 [0.816, 2.037]	1.420 [0.756, 2.668]	7.0 [-6.6, 20.6]	0.337
		Irbesartan	99	24 (24.2)				
> 45 years	Double-blind period	Sparsentan	106	23 (21.7)	1.016 [0.605, 1.705]	1.020 [0.527, 1.974]	0.3 [-11.8, 12.4]	1.000
		Irbesartan	103	22 (21.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.079
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	0.185 [0.026, 1.343]	0.083 [0.005, 1.294]	-48.9 [-100.0, 14.3]	0.095
		Irbesartan	5	3 (60.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	29 (28.4)	1.550 [0.938, 2.559]	1.768 [0.924, 3.381]	10.1 [-2.2, 22.4]	0.103
		Irbesartan	109	20 (18.3)				
> 40 years	Double-blind period	Sparsentan	91	23 (25.3)	0.967 [0.588, 1.591]	0.956 [0.489, 1.869]	-0.9 [-14.8, 13.1]	1.000
		Irbesartan	88	23 (26.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.238
North America	Double-blind period	Sparsentan	35	10 (28.6)	1.643 [0.724, 3.727]	1.900 [0.660, 5.472]	11.2 [-9.9, 32.2]	0.285
		Irbesartan	46	8 (17.4)				
Europe	Double-blind period	Sparsentan	98	33 (33.7)	1.335 [0.878, 2.032]	1.506 [0.831, 2.726]	8.5 [-4.8, 21.7]	0.226
		Irbesartan	115	29 (25.2)				
Asia Pacific	Double-blind period	Sparsentan	69	10 (14.5)	0.660 [0.293, 1.489]	0.603 [0.222, 1.635]	-7.5 [-24.6, 9.6]	0.434
		Irbesartan	41	9 (22.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.327
< 27 kg/m**2	Double-blind period	Sparsentan	83	19 (22.9)	0.936 [0.550, 1.591]	0.916 [0.457, 1.837]	-1.6 [-15.2, 12.1]	0.861
		Irbesartan	94	23 (24.5)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	34 (28.6)	1.329 [0.839, 2.106]	1.461 [0.795, 2.686]	7.1 [-5.1, 19.2]	0.283
		Irbesartan	107	23 (21.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.067
eGFR Low and UP High	Double-blind period	Sparsentan	71	19 (26.8)	2.200 [1.068, 4.535]	2.639 [1.102, 6.317]	14.6 [0.5, 28.7]	0.035 *
		Irbesartan	74	9 (12.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	14 (25.5)	1.077 [0.559, 2.075]	1.103 [0.463, 2.630]	1.8 [-16.1, 19.7]	1.000
		Irbesartan	55	13 (23.6)				
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	0.389 [0.134, 1.129]	0.315 [0.089, 1.120]	-17.0 [-37.4, 3.5]	0.081
		Irbesartan	36	10 (27.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	16 (41.0)	1.084 [0.620, 1.896]	1.143 [0.455, 2.871]	3.2 [-21.4, 27.8]	0.818
		Irbesartan	37	14 (37.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.104
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	33 (26.0)	1.397 [0.877, 2.224]	1.536 [0.847, 2.784]	7.4 [-3.6, 18.3]	0.178
		Irbesartan	129	24 (18.6)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	10 (20.4)	0.612 [0.309, 1.212]	0.513 [0.205, 1.284]	-12.9 [-32.5, 6.6]	0.174
		Irbesartan	48	16 (33.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	10 (38.5)	1.603 [0.685, 3.750]	1.979 [0.590, 6.644]	14.5 [-14.6, 43.5]	0.368
		Irbesartan	25	6 (24.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.080
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	22 (26.8)	1.951 [1.014, 3.756]	2.300 [1.031, 5.130]	13.1 [-0.4, 26.5]	0.051
		Irbesartan	80	11 (13.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	11 (24.4)	0.921 [0.461, 1.843]	0.896 [0.354, 2.270]	-2.1 [-21.8, 17.7]	1.000
		Irbesartan	49	13 (26.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	10 (20.4)	0.612 [0.309, 1.212]	0.513 [0.205, 1.284]	-12.9 [-32.5, 6.6]	0.174
		Irbesartan	48	16 (33.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	10 (38.5)	1.603 [0.685, 3.750]	1.979 [0.590, 6.644]	14.5 [-14.6, 43.5]	0.368
		Irbesartan	25	6 (24.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.581
<= 1.75 g/day	Double-blind period	Sparsentan	98	31 (31.6)	1.051 [0.687, 1.607]	1.074 [0.581, 1.986]	1.5 [-12.6, 15.7]	0.876
		Irbesartan	93	28 (30.1)				
> 1.75 g/day	Double-blind period	Sparsentan	104	22 (21.2)	1.281 [0.730, 2.247]	1.356 [0.680, 2.706]	4.6 [-6.8, 16.1]	0.483
		Irbesartan	109	18 (16.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.709
Yes	Double-blind period	Sparsentan	90	23 (25.6)	1.071 [0.641, 1.789]	1.095 [0.554, 2.165]	1.7 [-12.1, 15.5]	0.863
		Irbesartan	88	21 (23.9)				
No	Double-blind period	Sparsentan	112	30 (26.8)	1.221 [0.769, 1.939]	1.302 [0.708, 2.396]	4.9 [-7.2, 16.9]	0.440
		Irbesartan	114	25 (21.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
 Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.559
<= 5 years	Double-blind period	Sparsentan	113	32 (28.3)	1.090 [0.720, 1.650]	1.125 [0.636, 1.990]	2.3 [-9.8, 14.4]	0.771
		Irbesartan	127	33 (26.0)				
> 5 years	Double-blind period	Sparsentan	89	21 (23.6)	1.361 [0.733, 2.530]	1.473 [0.680, 3.189]	6.3 [-7.3, 19.8]	0.342
		Irbesartan	75	13 (17.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEV\_SSIM: Incidence of AESI COVID-19 during double-blind period by subgroup  
Safety Set

AESI COVID-19								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.410
Yes	Double-blind period	Sparsentan	155	39 (25.2)	1.066 [0.723, 1.572]	1.088 [0.651, 1.819]	1.6 [-8.5, 11.7]	0.794
		Irbesartan	161	38 (23.6)				
No	Double-blind period	Sparsentan	47	14 (29.8)	1.527 [0.713, 3.268]	1.750 [0.648, 4.727]	10.3 [-9.8, 30.4]	0.328
		Irbesartan	41	8 (19.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
 Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.887
Male	Double-blind period	Sparsentan	139	30 (21.6)	1.102 [0.696, 1.744]	1.130 [0.634, 2.015]	2.0 [-8.1, 12.1]	0.768
		Irbesartan	143	28 (19.6)				
Female	Double-blind period	Sparsentan	63	19 (30.2)	1.047 [0.604, 1.813]	1.067 [0.490, 2.325]	1.3 [-16.5, 19.2]	1.000
		Irbesartan	59	17 (28.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.894
<= 45 years	Double-blind period	Sparsentan	96	26 (27.1)	1.117 [0.692, 1.803]	1.161 [0.610, 2.209]	2.8 [-10.4, 16.1]	0.743
		Irbesartan	99	24 (24.2)				
> 45 years	Double-blind period	Sparsentan	106	23 (21.7)	1.064 [0.629, 1.800]	1.082 [0.556, 2.105]	1.3 [-10.7, 13.3]	0.866
		Irbesartan	103	21 (20.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
 Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.154
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	0.185 [0.026, 1.343]	0.083 [0.005, 1.294]	-48.9 [-100.0, 14.3]	0.095
		Irbesartan	5	3 (60.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	25 (24.5)	1.336 [0.792, 2.252]	1.445 [0.745, 2.802]	6.2 [-5.9, 18.2]	0.315
		Irbesartan	109	20 (18.3)				
> 40 years	Double-blind period	Sparsentan	91	23 (25.3)	1.011 [0.610, 1.676]	1.015 [0.516, 1.994]	0.3 [-13.6, 14.1]	1.000
		Irbesartan	88	22 (25.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.221
North America	Double-blind period	Sparsentan	35	9 (25.7)	1.479 [0.635, 3.442]	1.644 [0.561, 4.818]	8.3 [-12.3, 29.0]	0.416
		Irbesartan	46	8 (17.4)				
Europe	Double-blind period	Sparsentan	98	31 (31.6)	1.299 [0.842, 2.006]	1.438 [0.787, 2.625]	7.3 [-5.8, 20.3]	0.283
		Irbesartan	115	28 (24.3)				
Asia Pacific	Double-blind period	Sparsentan	69	9 (13.0)	0.594 [0.257, 1.375]	0.533 [0.193, 1.477]	-8.9 [-25.8, 8.0]	0.288
		Irbesartan	41	9 (22.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.333
< 27 kg/m**2	Double-blind period	Sparsentan	83	18 (21.7)	0.886 [0.516, 1.523]	0.855 [0.423, 1.726]	-2.8 [-16.3, 10.8]	0.723
		Irbesartan	94	23 (24.5)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	31 (26.1)	1.267 [0.784, 2.047]	1.361 [0.730, 2.536]	5.5 [-6.4, 17.4]	0.350
		Irbesartan	107	22 (20.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.106
eGFR Low and UP High	Double-blind period	Sparsentan	71	17 (23.9)	1.969 [0.940, 4.124]	2.274 [0.938, 5.508]	11.8 [-2.0, 25.6]	0.083
		Irbesartan	74	9 (12.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	13 (23.6)	1.000 [0.511, 1.958]	1.000 [0.415, 2.410]	0.0 [-17.7, 17.7]	1.000
		Irbesartan	55	13 (23.6)				
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	0.389 [0.134, 1.129]	0.315 [0.089, 1.120]	-17.0 [-37.4, 3.5]	0.081
		Irbesartan	36	10 (27.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	15 (38.5)	1.095 [0.606, 1.977]	1.154 [0.454, 2.935]	3.3 [-21.0, 27.6]	0.815
		Irbesartan	37	13 (35.1)				

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N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
 Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.085
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	30 (23.6)	1.325 [0.816, 2.151]	1.425 [0.775, 2.621]	5.8 [-4.9, 16.5]	0.282
		Irbesartan	129	23 (17.8)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	9 (18.4)	0.551 [0.270, 1.124]	0.450 [0.176, 1.152]	-15.0 [-34.2, 4.3]	0.108
		Irbesartan	48	16 (33.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	10 (38.5)	1.603 [0.685, 3.750]	1.979 [0.590, 6.644]	14.5 [-14.6, 43.5]	0.368
		Irbesartan	25	6 (24.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
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 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.086
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	20 (24.4)	1.774 [0.909, 3.460]	2.023 [0.899, 4.556]	10.6 [-2.6, 23.8]	0.110
		Irbesartan	80	11 (13.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	10 (22.2)	0.907 [0.435, 1.893]	0.881 [0.338, 2.296]	-2.3 [-21.5, 17.0]	0.813
		Irbesartan	49	12 (24.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	9 (18.4)	0.551 [0.270, 1.124]	0.450 [0.176, 1.152]	-15.0 [-34.2, 4.3]	0.108
		Irbesartan	48	16 (33.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	10 (38.5)	1.603 [0.685, 3.750]	1.979 [0.590, 6.644]	14.5 [-14.6, 43.5]	0.368
		Irbesartan	25	6 (24.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.719
<= 1.75 g/day	Double-blind period	Sparsentan	98	29 (29.6)	1.019 [0.656, 1.584]	1.027 [0.551, 1.916]	0.6 [-13.4, 14.5]	1.000
		Irbesartan	93	27 (29.0)				
> 1.75 g/day	Double-blind period	Sparsentan	104	20 (19.2)	1.165 [0.654, 2.074]	1.204 [0.596, 2.430]	2.7 [-8.5, 14.0]	0.721
		Irbesartan	109	18 (16.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.952
Yes	Double-blind period	Sparsentan	90	22 (24.4)	1.076 [0.633, 1.826]	1.100 [0.550, 2.199]	1.7 [-11.9, 15.3]	0.861
		Irbesartan	88	20 (22.7)				
No	Double-blind period	Sparsentan	112	27 (24.1)	1.099 [0.682, 1.772]	1.131 [0.608, 2.102]	2.2 [-9.7, 14.0]	0.753
		Irbesartan	114	25 (21.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.374
<= 5 years	Double-blind period	Sparsentan	113	29 (25.7)	0.988 [0.643, 1.518]	0.983 [0.551, 1.755]	-0.3 [-12.2, 11.6]	1.000
		Irbesartan	127	33 (26.0)				
> 5 years	Double-blind period	Sparsentan	89	20 (22.5)	1.404 [0.736, 2.681]	1.522 [0.689, 3.363]	6.5 [-6.8, 19.7]	0.328
		Irbesartan	75	12 (16.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEVN\_SSIM: Incidence of AESI COVID-19 during double-blind period - non-severe by subgroup  
Safety Set

AESI COVID-19 - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.447
Yes	Double-blind period	Sparsentan	155	36 (23.2)	1.011 [0.676, 1.511]	1.014 [0.601, 1.711]	0.2 [-9.7, 10.2]	1.000
		Irbesartan	161	37 (23.0)				
No	Double-blind period	Sparsentan	47	13 (27.7)	1.418 [0.653, 3.076]	1.577 [0.579, 4.298]	8.1 [-11.8, 28.1]	0.456
		Irbesartan	41	8 (19.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVC\_SSIM: Incidence of AESI COVID-19 during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.884
Male	Double-blind period	Sparsentan	139	25 (18.0)	1.072 [0.644, 1.783]	1.087 [0.587, 2.014]	1.2 [-8.4, 10.8]	0.875
		Irbesartan	143	24 (16.8)				
Female	Double-blind period	Sparsentan	63	17 (27.0)	1.137 [0.617, 2.097]	1.188 [0.524, 2.692]	3.3 [-13.8, 20.3]	0.835
		Irbesartan	59	14 (23.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.549
<= 45 years	Double-blind period	Sparsentan	96	24 (25.0)	1.238 [0.734, 2.087]	1.317 [0.671, 2.583]	4.8 [-8.0, 17.6]	0.494
		Irbesartan	99	20 (20.2)				
> 45 years	Double-blind period	Sparsentan	106	18 (17.0)	0.972 [0.536, 1.761]	0.966 [0.471, 1.981]	-0.5 [-11.7, 10.7]	1.000
		Irbesartan	103	18 (17.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.120
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	0.185 [0.026, 1.343]	0.083 [0.005, 1.294]	-48.9 [-100.0, 14.3]	0.095
		Irbesartan	5	3 (60.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	22 (21.6)	1.469 [0.819, 2.637]	1.598 [0.786, 3.251]	6.9 [-4.4, 18.2]	0.213
		Irbesartan	109	16 (14.7)				
> 40 years	Double-blind period	Sparsentan	91	19 (20.9)	0.967 [0.550, 1.700]	0.958 [0.468, 1.962]	-0.7 [-13.8, 12.4]	1.000
		Irbesartan	88	19 (21.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.106
North America	Double-blind period	Sparsentan	35	10 (28.6)	1.878 [0.794, 4.437]	2.229 [0.750, 6.619]	13.4 [-7.4, 34.1]	0.174
		Irbesartan	46	7 (15.2)				
Europe	Double-blind period	Sparsentan	98	24 (24.5)	1.280 [0.767, 2.136]	1.371 [0.713, 2.637]	5.4 [-6.7, 17.4]	0.404
		Irbesartan	115	22 (19.1)				
Asia Pacific	Double-blind period	Sparsentan	69	8 (11.6)	0.528 [0.221, 1.261]	0.466 [0.164, 1.325]	-10.4 [-27.1, 6.3]	0.177
		Irbesartan	41	9 (22.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
 Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.168
< 27 kg/m**2	Double-blind period	Sparsentan	83	15 (18.1)	0.809 [0.447, 1.464]	0.767 [0.366, 1.608]	-4.3 [-17.2, 8.7]	0.576
		Irbesartan	94	21 (22.3)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	27 (22.7)	1.428 [0.826, 2.470]	1.554 [0.793, 3.045]	6.8 [-4.3, 17.9]	0.240
		Irbesartan	107	17 (15.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.118
eGFR Low and UP High	Double-blind period	Sparsentan	71	16 (22.5)	2.085 [0.952, 4.564]	2.400 [0.955, 6.029]	11.7 [-1.7, 25.1]	0.074
		Irbesartan	74	8 (10.8)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	11 (20.0)	1.000 [0.474, 2.112]	1.000 [0.393, 2.546]	0.0 [-16.8, 16.8]	1.000
		Irbesartan	55	11 (20.0)				
eGFR High and UP High	Double-blind period	Sparsentan	37	2 (5.4)	0.278 [0.062, 1.250]	0.237 [0.046, 1.229]	-14.0 [-31.6, 3.5]	0.085
		Irbesartan	36	7 (19.4)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	13 (33.3)	1.028 [0.540, 1.955]	1.042 [0.400, 2.714]	0.9 [-22.9, 24.7]	1.000
		Irbesartan	37	12 (32.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.327
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	27 (21.3)	1.306 [0.780, 2.186]	1.389 [0.738, 2.612]	5.0 [-5.4, 15.3]	0.339
		Irbesartan	129	21 (16.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	9 (18.4)	0.678 [0.320, 1.437]	0.606 [0.231, 1.587]	-8.7 [-27.4, 9.9]	0.341
		Irbesartan	48	13 (27.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	6 (23.1)	1.442 [0.461, 4.509]	1.575 [0.386, 6.423]	7.1 [-18.5, 32.7]	0.726
		Irbesartan	25	4 (16.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.195
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	18 (22.0)	1.951 [0.932, 4.083]	2.219 [0.931, 5.288]	10.7 [-1.9, 23.3]	0.091
		Irbesartan	80	9 (11.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	9 (20.0)	0.817 [0.381, 1.753]	0.771 [0.290, 2.051]	-4.5 [-23.4, 14.4]	0.630
		Irbesartan	49	12 (24.5)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	9 (18.4)	0.678 [0.320, 1.437]	0.606 [0.231, 1.587]	-8.7 [-27.4, 9.9]	0.341
		Irbesartan	48	13 (27.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	6 (23.1)	1.442 [0.461, 4.509]	1.575 [0.386, 6.423]	7.1 [-18.5, 32.7]	0.726
		Irbesartan	25	4 (16.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.543
<= 1.75 g/day	Double-blind period	Sparsentan	98	25 (25.5)	0.989 [0.610, 1.602]	0.985 [0.514, 1.885]	-0.3 [-13.7, 13.1]	1.000
		Irbesartan	93	24 (25.8)				
> 1.75 g/day	Double-blind period	Sparsentan	104	17 (16.3)	1.273 [0.662, 2.448]	1.326 [0.617, 2.849]	3.5 [-6.9, 13.9]	0.561
		Irbesartan	109	14 (12.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.429
Yes	Double-blind period	Sparsentan	90	17 (18.9)	0.923 [0.510, 1.673]	0.906 [0.432, 1.897]	-1.6 [-14.4, 11.2]	0.852
		Irbesartan	88	18 (20.5)				
No	Double-blind period	Sparsentan	112	25 (22.3)	1.272 [0.751, 2.155]	1.351 [0.701, 2.603]	4.8 [-6.5, 16.1]	0.407
		Irbesartan	114	20 (17.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.452
<= 5 years	Double-blind period	Sparsentan	113	24 (21.2)	0.999 [0.613, 1.627]	0.999 [0.537, 1.856]	-0.0 [-11.2, 11.2]	1.000
		Irbesartan	127	27 (21.3)				
> 5 years	Double-blind period	Sparsentan	89	18 (20.2)	1.379 [0.696, 2.734]	1.475 [0.648, 3.358]	5.6 [-7.2, 18.4]	0.414
		Irbesartan	75	11 (14.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEVS\_SSIM: Incidence of AESI COVID-19 during double-blind period - serious by subgroup  
Safety Set

AESI COVID-19 - serious								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.706
Yes	Double-blind period	Sparsentan	155	33 (21.3)	1.071 [0.694, 1.652]	1.090 [0.632, 1.882]	1.4 [-8.1, 11.0]	0.782
		Irbesartan	161	32 (19.9)				
No	Double-blind period	Sparsentan	47	9 (19.1)	1.309 [0.509, 3.364]	1.382 [0.446, 4.279]	4.5 [-13.4, 22.4]	0.777
		Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.040 #
Male	Double-blind period	Sparsentan	139	39 (28.1)	0.854 [0.599, 1.217]	0.797 [0.479, 1.325]	-4.8 [-16.2, 6.6]	0.438
		Irbesartan	143	47 (32.9)				
Female	Double-blind period	Sparsentan	63	24 (38.1)	1.729 [0.974, 3.069]	2.178 [0.980, 4.839]	16.1 [-1.6, 33.7]	0.076
		Irbesartan	59	13 (22.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.165
<= 45 years	Double-blind period	Sparsentan	96	24 (25.0)	0.825 [0.522, 1.304]	0.767 [0.408, 1.440]	-5.3 [-18.9, 8.3]	0.428
		Irbesartan	99	30 (30.3)				
> 45 years	Double-blind period	Sparsentan	106	39 (36.8)	1.263 [0.854, 1.868]	1.416 [0.793, 2.530]	7.7 [-6.0, 21.3]	0.244
		Irbesartan	103	30 (29.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
 Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.153
<= 18 years	Double-blind period	Sparsentan	9	5 (55.6)	1.389 [0.409, 4.715]	1.875 [0.204, 17.269]	15.6 [-53.8, 84.9]	1.000
		Irbesartan	5	2 (40.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	23 (22.5)	0.745 [0.471, 1.178]	0.671 [0.361, 1.245]	-7.7 [-20.5, 5.1]	0.216
		Irbesartan	109	33 (30.3)				
> 40 years	Double-blind period	Sparsentan	91	35 (38.5)	1.354 [0.888, 2.063]	1.575 [0.841, 2.948]	10.1 [-4.8, 24.9]	0.205
		Irbesartan	88	25 (28.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.357
North America	Double-blind period	Sparsentan	35	6 (17.1)	0.607 [0.256, 1.436]	0.525 [0.177, 1.560]	-11.1 [-31.7, 9.4]	0.296
		Irbesartan	46	13 (28.3)				
Europe	Double-blind period	Sparsentan	98	37 (37.8)	1.206 [0.832, 1.748]	1.331 [0.755, 2.348]	6.5 [-7.3, 20.2]	0.385
		Irbesartan	115	36 (31.3)				
Asia Pacific	Double-blind period	Sparsentan	69	20 (29.0)	1.080 [0.578, 2.020]	1.113 [0.469, 2.643]	2.2 [-17.1, 21.4]	1.000
		Irbesartan	41	11 (26.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
 Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.294
< 27 kg/m**2	Double-blind period	Sparsentan	83	22 (26.5)	0.859 [0.538, 1.373]	0.808 [0.420, 1.556]	-4.3 [-18.8, 10.1]	0.618
		Irbesartan	94	29 (30.9)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	41 (34.5)	1.189 [0.808, 1.750]	1.289 [0.734, 2.264]	5.5 [-7.5, 18.5]	0.394
		Irbesartan	107	31 (29.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
 Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.980
eGFR Low and UP High	Double-blind period	Sparsentan	71	24 (33.8)	1.137 [0.705, 1.834]	1.207 [0.599, 2.431]	4.1 [-12.5, 20.6]	0.721
		Irbesartan	74	22 (29.7)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	17 (30.9)	1.000 [0.572, 1.749]	1.000 [0.445, 2.245]	0.0 [-19.1, 19.1]	1.000
		Irbesartan	55	17 (30.9)				
eGFR High and UP High	Double-blind period	Sparsentan	37	12 (32.4)	0.973 [0.505, 1.874]	0.960 [0.361, 2.549]	-0.9 [-25.2, 23.4]	1.000
		Irbesartan	36	12 (33.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	10 (25.6)	1.054 [0.483, 2.300]	1.073 [0.379, 3.034]	1.3 [-20.8, 23.4]	1.000
		Irbesartan	37	9 (24.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.953
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	41 (32.3)	1.068 [0.742, 1.536]	1.100 [0.648, 1.867]	2.1 [-10.1, 14.2]	0.788
		Irbesartan	129	39 (30.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	16 (32.7)	0.980 [0.556, 1.727]	0.970 [0.416, 2.261]	-0.7 [-21.5, 20.1]	1.000
		Irbesartan	48	16 (33.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	6 (23.1)	1.154 [0.403, 3.305]	1.200 [0.315, 4.578]	3.1 [-23.4, 29.5]	1.000
		Irbesartan	25	5 (20.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.879
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	26 (31.7)	1.208 [0.743, 1.963]	1.304 [0.660, 2.579]	5.5 [-9.7, 20.6]	0.491
		Irbesartan	80	21 (26.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	15 (33.3)	0.907 [0.522, 1.577]	0.861 [0.368, 2.013]	-3.4 [-24.8, 18.0]	0.830
		Irbesartan	49	18 (36.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	16 (32.7)	0.980 [0.556, 1.727]	0.970 [0.416, 2.261]	-0.7 [-21.5, 20.1]	1.000
		Irbesartan	48	16 (33.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	6 (23.1)	1.154 [0.403, 3.305]	1.200 [0.315, 4.578]	3.1 [-23.4, 29.5]	1.000
		Irbesartan	25	5 (20.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.117
<= 1.75 g/day	Double-blind period	Sparsentan	98	31 (31.6)	1.401 [0.871, 2.254]	1.586 [0.831, 3.027]	9.1 [-4.5, 22.6]	0.194
		Irbesartan	93	21 (22.6)				
> 1.75 g/day	Double-blind period	Sparsentan	104	32 (30.8)	0.860 [0.587, 1.261]	0.798 [0.450, 1.413]	-5.0 [-18.6, 8.6]	0.470
		Irbesartan	109	39 (35.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.987
Yes	Double-blind period	Sparsentan	90	28 (31.1)	1.053 [0.674, 1.644]	1.077 [0.568, 2.041]	1.6 [-13.1, 16.2]	0.871
		Irbesartan	88	26 (29.5)				
No	Double-blind period	Sparsentan	112	35 (31.3)	1.048 [0.707, 1.553]	1.070 [0.607, 1.884]	1.4 [-11.5, 14.3]	0.885
		Irbesartan	114	34 (29.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.728
<= 5 years	Double-blind period	Sparsentan	113	35 (31.0)	1.009 [0.690, 1.474]	1.012 [0.585, 1.753]	0.3 [-12.3, 12.8]	1.000
		Irbesartan	127	39 (30.7)				
> 5 years	Double-blind period	Sparsentan	89	28 (31.5)	1.124 [0.699, 1.807]	1.180 [0.602, 2.316]	3.5 [-11.8, 18.7]	0.732
		Irbesartan	75	21 (28.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEC\_SSIM: Incidence of AESI cardiovascular system during double-blind period by subgroup  
Safety Set

AESI cardiovascular system								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.276
Yes	Double-blind period	Sparsentan	155	55 (35.5)	1.143 [0.835, 1.563]	1.221 [0.764, 1.951]	4.4 [-6.6, 15.4]	0.474
		Irbesartan	161	50 (31.1)				
No	Double-blind period	Sparsentan	47	8 (17.0)	0.698 [0.304, 1.600]	0.636 [0.224, 1.804]	-7.4 [-26.6, 11.9]	0.436
		Irbesartan	41	10 (24.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.075
Male	Double-blind period	Sparsentan	139	39 (28.1)	0.892 [0.622, 1.278]	0.849 [0.509, 1.416]	-3.4 [-14.8, 8.0]	0.603
		Irbesartan	143	45 (31.5)				
Female	Double-blind period	Sparsentan	63	23 (36.5)	1.657 [0.928, 2.959]	2.035 [0.913, 4.534]	14.5 [-3.1, 32.0]	0.112
		Irbesartan	59	13 (22.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.200
<= 45 years	Double-blind period	Sparsentan	96	24 (25.0)	0.853 [0.538, 1.355]	0.805 [0.427, 1.515]	-4.3 [-17.8, 9.2]	0.523
		Irbesartan	99	29 (29.3)				
> 45 years	Double-blind period	Sparsentan	106	38 (35.8)	1.273 [0.853, 1.900]	1.426 [0.795, 2.559]	7.7 [-5.9, 21.3]	0.240
		Irbesartan	103	29 (28.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.183
<= 18 years	Double-blind period	Sparsentan	9	5 (55.6)	1.389 [0.409, 4.715]	1.875 [0.204, 17.269]	15.6 [-53.8, 84.9]	1.000
		Irbesartan	5	2 (40.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	23 (22.5)	0.768 [0.484, 1.220]	0.701 [0.377, 1.304]	-6.8 [-19.5, 5.9]	0.276
		Irbesartan	109	32 (29.4)				
> 40 years	Double-blind period	Sparsentan	91	34 (37.4)	1.370 [0.889, 2.112]	1.591 [0.845, 2.995]	10.1 [-4.6, 24.8]	0.155
		Irbesartan	88	24 (27.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.458
North America	Double-blind period	Sparsentan	35	6 (17.1)	0.657 [0.274, 1.578]	0.586 [0.196, 1.758]	-8.9 [-29.3, 11.4]	0.423
		Irbesartan	46	12 (26.1)				
Europe	Double-blind period	Sparsentan	98	36 (36.7)	1.207 [0.826, 1.764]	1.327 [0.750, 2.350]	6.3 [-7.4, 20.0]	0.382
		Irbesartan	115	35 (30.4)				
Asia Pacific	Double-blind period	Sparsentan	69	20 (29.0)	1.080 [0.578, 2.020]	1.113 [0.469, 2.643]	2.2 [-17.1, 21.4]	1.000
		Irbesartan	41	11 (26.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.167
< 27 kg/m**2	Double-blind period	Sparsentan	83	21 (25.3)	0.820 [0.509, 1.322]	0.759 [0.392, 1.470]	-5.5 [-19.9, 8.8]	0.504
		Irbesartan	94	29 (30.9)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	41 (34.5)	1.271 [0.854, 1.892]	1.414 [0.800, 2.499]	7.4 [-5.5, 20.2]	0.252
		Irbesartan	107	29 (27.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.989
eGFR Low and UP High	Double-blind period	Sparsentan	71	23 (32.4)	1.142 [0.697, 1.871]	1.209 [0.595, 2.457]	4.0 [-12.3, 20.4]	0.718
		Irbesartan	74	21 (28.4)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	17 (30.9)	1.000 [0.572, 1.749]	1.000 [0.445, 2.245]	0.0 [-19.1, 19.1]	1.000
		Irbesartan	55	17 (30.9)				
eGFR High and UP High	Double-blind period	Sparsentan	37	12 (32.4)	1.061 [0.539, 2.090]	1.091 [0.406, 2.931]	1.9 [-22.2, 25.9]	1.000
		Irbesartan	36	11 (30.6)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	10 (25.6)	1.054 [0.483, 2.300]	1.073 [0.379, 3.034]	1.3 [-20.8, 23.4]	1.000
		Irbesartan	37	9 (24.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.937
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	40 (31.5)	1.098 [0.756, 1.596]	1.143 [0.670, 1.951]	2.8 [-9.2, 14.8]	0.683
		Irbesartan	129	37 (28.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	16 (32.7)	0.980 [0.556, 1.727]	0.970 [0.416, 2.261]	-0.7 [-21.5, 20.1]	1.000
		Irbesartan	48	16 (33.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	6 (23.1)	1.154 [0.403, 3.305]	1.200 [0.315, 4.578]	3.1 [-23.4, 29.5]	1.000
		Irbesartan	25	5 (20.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.820
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	26 (31.7)	1.268 [0.773, 2.080]	1.393 [0.700, 2.770]	6.7 [-8.4, 21.8]	0.386
		Irbesartan	80	20 (25.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	14 (31.1)	0.897 [0.502, 1.602]	0.850 [0.359, 2.014]	-3.6 [-24.7, 17.5]	0.827
		Irbesartan	49	17 (34.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	16 (32.7)	0.980 [0.556, 1.727]	0.970 [0.416, 2.261]	-0.7 [-21.5, 20.1]	1.000
		Irbesartan	48	16 (33.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	6 (23.1)	1.154 [0.403, 3.305]	1.200 [0.315, 4.578]	3.1 [-23.4, 29.5]	1.000
		Irbesartan	25	5 (20.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.088
<= 1.75 g/day	Double-blind period	Sparsentan	98	31 (31.6)	1.471 [0.905, 2.390]	1.689 [0.879, 3.244]	10.1 [-3.4, 23.6]	0.141
		Irbesartan	93	20 (21.5)				
> 1.75 g/day	Double-blind period	Sparsentan	104	31 (29.8)	0.855 [0.578, 1.264]	0.793 [0.446, 1.411]	-5.1 [-18.5, 8.4]	0.466
		Irbesartan	109	38 (34.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.943
Yes	Double-blind period	Sparsentan	90	27 (30.0)	1.056 [0.668, 1.669]	1.080 [0.566, 2.061]	1.6 [-12.9, 16.1]	0.870
		Irbesartan	88	25 (28.4)				
No	Double-blind period	Sparsentan	112	35 (31.3)	1.080 [0.725, 1.607]	1.116 [0.632, 1.970]	2.3 [-10.5, 15.1]	0.772
		Irbesartan	114	33 (28.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.952
<= 5 years	Double-blind period	Sparsentan	113	35 (31.0)	1.063 [0.722, 1.565]	1.091 [0.628, 1.897]	1.8 [-10.6, 14.3]	0.779
		Irbesartan	127	37 (29.1)				
> 5 years	Double-blind period	Sparsentan	89	27 (30.3)	1.083 [0.670, 1.752]	1.120 [0.569, 2.204]	2.3 [-12.8, 17.5]	0.863
		Irbesartan	75	21 (28.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AECN\_SSIM: Incidence of AESI cardiovascular system during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiovascular system - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.256
Yes	Double-blind period	Sparsentan	155	54 (34.8)	1.169 [0.848, 1.610]	1.259 [0.785, 2.019]	5.0 [-5.9, 16.0]	0.400
		Irbesartan	161	48 (29.8)				
No	Double-blind period	Sparsentan	47	8 (17.0)	0.698 [0.304, 1.600]	0.636 [0.224, 1.804]	-7.4 [-26.6, 11.9]	0.436
		Irbesartan	41	10 (24.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AECC\_SSIM: Incidence of AESI cardiovascular system during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AECS\_SSIM: Incidence of AESI cardiovascular system during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
 Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.494
Male	Double-blind period	Sparsentan	139	34 (24.5)	2.058 [1.207, 3.507]	2.400 [1.269, 4.539]	12.6 [3.0, 22.2]	0.008 *
		Irbesartan	143	17 (11.9)				
Female	Double-blind period	Sparsentan	63	24 (38.1)	2.810 [1.372, 5.755]	3.923 [1.591, 9.671]	24.5 [8.1, 41.0]	0.004 *
		Irbesartan	59	8 (13.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
 Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.465
<= 45 years	Double-blind period	Sparsentan	96	27 (28.1)	1.989 [1.112, 3.557]	2.376 [1.157, 4.878]	14.0 [1.6, 26.3]	0.022 *
		Irbesartan	99	14 (14.1)				
> 45 years	Double-blind period	Sparsentan	106	31 (29.2)	2.738 [1.455, 5.153]	3.457 [1.629, 7.336]	18.6 [7.1, 30.0]	<0.001 *
		Irbesartan	103	11 (10.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
 Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.541
<= 18 years	Double-blind period	Sparsentan	9	3 (33.3)	4.200 + [0.259, 68.038]	5.923 + [0.248, 141.482]	33.3 [-13.0, 79.7]	0.258
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	30 (29.4)	1.886 [1.110, 3.205]	2.255 [1.154, 4.407]	13.8 [1.7, 25.9]	0.020 *
		Irbesartan	109	17 (15.6)				
> 40 years	Double-blind period	Sparsentan	91	25 (27.5)	3.022 [1.441, 6.335]	3.788 [1.602, 8.954]	18.4 [6.3, 30.5]	0.002 *
		Irbesartan	88	8 (9.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.135
North America	Double-blind period	Sparsentan	35	8 (22.9)	2.629 [0.860, 8.031]	3.111 [0.853, 11.347]	14.2 [-4.5, 32.8]	0.114
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	24 (24.5)	3.129 [1.528, 6.410]	3.820 [1.680, 8.687]	16.7 [5.9, 27.4]	0.001 *
		Irbesartan	115	9 (7.8)				
Asia Pacific	Double-blind period	Sparsentan	69	26 (37.7)	1.287 [0.732, 2.264]	1.461 [0.637, 3.353]	8.4 [-11.6, 28.4]	0.412
		Irbesartan	41	12 (29.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.856
< 27 kg/m**2	Double-blind period	Sparsentan	83	28 (33.7)	2.265 [1.281, 4.004]	2.909 [1.405, 6.022]	18.8 [5.2, 32.4]	0.004 *
		Irbesartan	94	14 (14.9)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	30 (25.2)	2.452 [1.294, 4.649]	2.942 [1.391, 6.219]	14.9 [4.3, 25.5]	0.005 *
		Irbesartan	107	11 (10.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.626
eGFR Low and UP High	Double-blind period	Sparsentan	71	18 (25.4)	1.706 [0.868, 3.352]	1.945 [0.845, 4.480]	10.5 [-3.9, 24.8]	0.147
		Irbesartan	74	11 (14.9)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	18 (32.7)	2.250 [1.069, 4.736]	2.858 [1.119, 7.299]	18.2 [0.9, 35.5]	0.042 *
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	10 (27.0)	3.243 [0.971, 10.831]	4.074 [1.018, 16.305]	18.7 [-1.0, 38.4]	0.064
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	12 (30.8)	3.795 [1.163, 12.381]	5.037 [1.290, 19.671]	22.7 [3.1, 42.2]	0.020 *
		Irbesartan	37	3 (8.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.448
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	38 (29.9)	2.031 [1.241, 3.327]	2.472 [1.333, 4.584]	15.2 [4.4, 26.0]	0.004 *
		Irbesartan	129	19 (14.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	14 (28.6)	4.571 [1.402, 14.903]	6.000 [1.598, 22.525]	22.3 [5.9, 38.8]	0.006 *
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	6 (23.1)	1.923 [0.539, 6.865]	2.200 [0.485, 9.983]	11.1 [-13.5, 35.6]	0.465
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.577
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	20 (24.4)	1.774 [0.909, 3.460]	2.023 [0.899, 4.556]	10.6 [-2.6, 23.8]	0.110
		Irbesartan	80	11 (13.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	18 (40.0)	2.450 [1.183, 5.073]	3.417 [1.303, 8.960]	23.7 [3.9, 43.5]	0.012 *
		Irbesartan	49	8 (16.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	14 (28.6)	4.571 [1.402, 14.903]	6.000 [1.598, 22.525]	22.3 [5.9, 38.8]	0.006 *
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	6 (23.1)	1.923 [0.539, 6.865]	2.200 [0.485, 9.983]	11.1 [-13.5, 35.6]	0.465
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.117
<= 1.75 g/day	Double-blind period	Sparsentan	98	30 (30.6)	3.559 [1.721, 7.358]	4.688 [2.018, 10.886]	22.0 [10.2, 33.8]	<0.001 *
		Irbesartan	93	8 (8.6)				
> 1.75 g/day	Double-blind period	Sparsentan	104	28 (26.9)	1.726 [1.007, 2.961]	1.994 [1.015, 3.915]	11.3 [-0.5, 23.2]	0.046 *
		Irbesartan	109	17 (15.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
 Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.458
Yes	Double-blind period	Sparsentan	90	17 (18.9)	1.847 [0.870, 3.920]	2.044 [0.858, 4.871]	8.7 [-2.7, 20.1]	0.137
		Irbesartan	88	9 (10.2)				
No	Double-blind period	Sparsentan	112	41 (36.6)	2.608 [1.558, 4.368]	3.537 [1.840, 6.799]	22.6 [10.7, 34.4]	<0.001 *
		Irbesartan	114	16 (14.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.963
<= 5 years	Double-blind period	Sparsentan	113	31 (27.4)	2.323 [1.324, 4.074]	2.823 [1.431, 5.567]	15.6 [4.8, 26.4]	0.003 *
		Irbesartan	127	15 (11.8)				
> 5 years	Double-blind period	Sparsentan	89	27 (30.3)	2.275 [1.179, 4.390]	2.831 [1.266, 6.329]	17.0 [3.5, 30.5]	0.014 *
		Irbesartan	75	10 (13.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEH\_SSIM: Incidence of AESI hypotension during double-blind period by subgroup  
Safety Set

AESI hypotension								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.209
Yes	Double-blind period	Sparsentan	155	42 (27.1)	2.727 [1.602, 4.641]	3.368 [1.801, 6.300]	17.2 [8.1, 26.2]	<0.001 *
		Irbesartan	161	16 (9.9)				
No	Double-blind period	Sparsentan	47	16 (34.0)	1.551 [0.769, 3.126]	1.835 [0.707, 4.766]	12.1 [-8.7, 32.9]	0.243
		Irbesartan	41	9 (22.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
 Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.557
Male	Double-blind period	Sparsentan	139	34 (24.5)	2.058 [1.207, 3.507]	2.400 [1.269, 4.539]	12.6 [3.0, 22.2]	0.008 *
		Irbesartan	143	17 (11.9)				
Female	Double-blind period	Sparsentan	63	23 (36.5)	2.692 [1.308, 5.542]	3.666 [1.483, 9.059]	22.9 [6.6, 39.3]	0.004 *
		Irbesartan	59	8 (13.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.514
<= 45 years	Double-blind period	Sparsentan	96	27 (28.1)	1.989 [1.112, 3.557]	2.376 [1.157, 4.878]	14.0 [1.6, 26.3]	0.022 *
		Irbesartan	99	14 (14.1)				
> 45 years	Double-blind period	Sparsentan	106	30 (28.3)	2.650 [1.404, 5.003]	3.301 [1.552, 7.022]	17.6 [6.2, 29.0]	0.002 *
		Irbesartan	103	11 (10.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.588
<= 18 years	Double-blind period	Sparsentan	9	3 (33.3)	4.200 + [0.259, 68.038]	5.923 + [0.248, 141.482]	33.3 [-13.0, 79.7]	0.258
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	30 (29.4)	1.886 [1.110, 3.205]	2.255 [1.154, 4.407]	13.8 [1.7, 25.9]	0.020 *
		Irbesartan	109	17 (15.6)				
> 40 years	Double-blind period	Sparsentan	91	24 (26.4)	2.901 [1.378, 6.108]	3.582 [1.511, 8.495]	17.3 [5.3, 29.3]	0.003 *
		Irbesartan	88	8 (9.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.159
North America	Double-blind period	Sparsentan	35	8 (22.9)	2.629 [0.860, 8.031]	3.111 [0.853, 11.347]	14.2 [-4.5, 32.8]	0.114
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	23 (23.5)	2.999 [1.457, 6.173]	3.612 [1.582, 8.245]	15.6 [5.0, 26.3]	0.002 *
		Irbesartan	115	9 (7.8)				
Asia Pacific	Double-blind period	Sparsentan	69	26 (37.7)	1.287 [0.732, 2.264]	1.461 [0.637, 3.353]	8.4 [-11.6, 28.4]	0.412
		Irbesartan	41	12 (29.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.792
< 27 kg/m**2	Double-blind period	Sparsentan	83	27 (32.5)	2.184 [1.230, 3.878]	2.755 [1.327, 5.719]	17.6 [4.1, 31.2]	0.007 *
		Irbesartan	94	14 (14.9)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	30 (25.2)	2.452 [1.294, 4.649]	2.942 [1.391, 6.219]	14.9 [4.3, 25.5]	0.005 *
		Irbesartan	107	11 (10.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.570
eGFR Low and UP High	Double-blind period	Sparsentan	71	17 (23.9)	1.611 [0.812, 3.195]	1.803 [0.778, 4.181]	9.1 [-5.1, 23.3]	0.208
		Irbesartan	74	11 (14.9)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	18 (32.7)	2.250 [1.069, 4.736]	2.858 [1.119, 7.299]	18.2 [0.9, 35.5]	0.042 *
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	10 (27.0)	3.243 [0.971, 10.831]	4.074 [1.018, 16.305]	18.7 [-1.0, 38.4]	0.064
		Irbesartan	36	3 (8.3)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	12 (30.8)	3.795 [1.163, 12.381]	5.037 [1.290, 19.671]	22.7 [3.1, 42.2]	0.020 *
		Irbesartan	37	3 (8.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
 Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.429
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	37 (29.1)	1.978 [1.205, 3.248]	2.380 [1.281, 4.422]	14.4 [3.6, 25.2]	0.006 *
		Irbesartan	129	19 (14.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	14 (28.6)	4.571 [1.402, 14.903]	6.000 [1.598, 22.525]	22.3 [5.9, 38.8]	0.006 *
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	6 (23.1)	1.923 [0.539, 6.865]	2.200 [0.485, 9.983]	11.1 [-13.5, 35.6]	0.465
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.587
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	20 (24.4)	1.774 [0.909, 3.460]	2.023 [0.899, 4.556]	10.6 [-2.6, 23.8]	0.110
		Irbesartan	80	11 (13.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	17 (37.8)	2.314 [1.108, 4.833]	3.112 [1.182, 8.192]	21.5 [1.8, 41.1]	0.021 *
		Irbesartan	49	8 (16.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	14 (28.6)	4.571 [1.402, 14.903]	6.000 [1.598, 22.525]	22.3 [5.9, 38.8]	0.006 *
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	6 (23.1)	1.923 [0.539, 6.865]	2.200 [0.485, 9.983]	11.1 [-13.5, 35.6]	0.465
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.101
<= 1.75 g/day	Double-blind period	Sparsentan	98	30 (30.6)	3.559 [1.721, 7.358]	4.688 [2.018, 10.886]	22.0 [10.2, 33.8]	<0.001 *
		Irbesartan	93	8 (8.6)				
> 1.75 g/day	Double-blind period	Sparsentan	104	27 (26.0)	1.665 [0.966, 2.868]	1.898 [0.963, 3.739]	10.4 [-1.4, 22.1]	0.065
		Irbesartan	109	17 (15.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.387
Yes	Double-blind period	Sparsentan	90	16 (17.8)	1.738 [0.811, 3.724]	1.898 [0.790, 4.558]	7.6 [-3.7, 18.8]	0.196
		Irbesartan	88	9 (10.2)				
No	Double-blind period	Sparsentan	112	41 (36.6)	2.608 [1.558, 4.368]	3.537 [1.840, 6.799]	22.6 [10.7, 34.4]	<0.001 *
		Irbesartan	114	16 (14.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.895
<= 5 years	Double-blind period	Sparsentan	113	31 (27.4)	2.323 [1.324, 4.074]	2.823 [1.431, 5.567]	15.6 [4.8, 26.4]	0.003 *
		Irbesartan	127	15 (11.8)				
> 5 years	Double-blind period	Sparsentan	89	26 (29.2)	2.191 [1.131, 4.245]	2.683 [1.196, 6.015]	15.9 [2.5, 29.3]	0.022 *
		Irbesartan	75	10 (13.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEHN\_SSIM: Incidence of AESI hypotension during double-blind period - non-severe by subgroup  
 Safety Set

AESI hypotension - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.230
Yes	Double-blind period	Sparsentan	155	41 (26.5)	2.662 [1.560, 4.540]	3.259 [1.740, 6.106]	16.5 [7.5, 25.5]	<0.001 *
		Irbesartan	161	16 (9.9)				
No	Double-blind period	Sparsentan	47	16 (34.0)	1.551 [0.769, 3.126]	1.835 [0.707, 4.766]	12.1 [-8.7, 32.9]	0.243
		Irbesartan	41	9 (22.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEHC\_SSIM: Incidence of AESI hypotension during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEHS\_SSIM: Incidence of AESI hypotension during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
 Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.130
Male	Double-blind period	Sparsentan	139	17 (12.2)	1.943 [0.897, 4.212]	2.075 [0.892, 4.827]	5.9 [-1.5, 13.4]	0.101
		Irbesartan	143	9 (6.3)				
Female	Double-blind period	Sparsentan	63	1 (1.6)	0.312 [0.033, 2.918]	0.301 [0.030, 2.979]	-3.5 [-11.5, 4.5]	0.353
		Irbesartan	59	3 (5.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.106
<= 45 years	Double-blind period	Sparsentan	96	5 (5.2)	0.737 [0.242, 2.241]	0.722 [0.221, 2.359]	-1.9 [-9.6, 5.9]	0.768
		Irbesartan	99	7 (7.1)				
> 45 years	Double-blind period	Sparsentan	106	13 (12.3)	2.526 [0.934, 6.834]	2.740 [0.940, 7.985]	7.4 [-1.0, 15.9]	0.083
		Irbesartan	103	5 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.404
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.200 + [0.010, 4.166]	0.158 + [0.005, 4.691]	-20.0 [-70.6, 30.6]	0.357
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	10 (9.8)	1.781 [0.672, 4.724]	1.866 [0.653, 5.335]	4.3 [-3.8, 12.4]	0.301
		Irbesartan	109	6 (5.5)				
> 40 years	Double-blind period	Sparsentan	91	8 (8.8)	1.547 [0.526, 4.548]	1.600 [0.503, 5.094]	3.1 [-5.6, 11.8]	0.567
		Irbesartan	88	5 (5.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.353
North America	Double-blind period	Sparsentan	35	5 (14.3)	1.643 [0.476, 5.672]	1.750 [0.433, 7.067]	5.6 [-11.1, 22.3]	0.490
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	10 (10.2)	2.347 [0.830, 6.634]	2.500 [0.824, 7.582]	5.9 [-2.1, 13.9]	0.112
		Irbesartan	115	5 (4.3)				
Asia Pacific	Double-blind period	Sparsentan	69	3 (4.3)	0.594 [0.126, 2.808]	0.576 [0.111, 2.996]	-3.0 [-14.2, 8.3]	0.669
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.282
< 27 kg/m**2	Double-blind period	Sparsentan	83	6 (7.2)	0.971 [0.340, 2.773]	0.968 [0.312, 3.006]	-0.2 [-9.0, 8.6]	1.000
		Irbesartan	94	7 (7.4)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	12 (10.1)	2.158 [0.786, 5.925]	2.288 [0.779, 6.723]	5.4 [-2.2, 13.0]	0.138
		Irbesartan	107	5 (4.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.720
eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	1.668 [0.573, 4.856]	1.752 [0.545, 5.637]	4.5 [-6.2, 15.2]	0.394
		Irbesartan	74	5 (6.8)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)	1.000 [0.146, 6.848]	1.000 [0.136, 7.364]	0.0 [-8.8, 8.8]	1.000
		Irbesartan	55	2 (3.6)				
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	0.973 [0.263, 3.598]	0.970 [0.223, 4.212]	-0.3 [-17.4, 16.8]	1.000
		Irbesartan	36	4 (11.1)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	4 (10.3)	3.795 [0.444, 32.404]	4.114 [0.438, 38.653]	7.6 [-5.9, 21.0]	0.359
		Irbesartan	37	1 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
 Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.219
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	10 (7.9)	1.451 [0.570, 3.694]	1.490 [0.549, 4.044]	2.4 [-4.4, 9.3]	0.463
		Irbesartan	129	7 (5.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	7 (14.3)	3.429 [0.750, 15.680]	3.833 [0.754, 19.490]	10.1 [-3.3, 23.5]	0.159
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.321 [0.036, 2.880]	0.293 [0.028, 3.029]	-8.2 [-26.8, 10.5]	0.350
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.370
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	8 (9.8)	1.561 [0.533, 4.569]	1.622 [0.507, 5.186]	3.5 [-6.1, 13.1]	0.565
		Irbesartan	80	5 (6.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	2 (4.4)	1.089 [0.160, 7.410]	1.093 [0.147, 8.102]	0.4 [-10.0, 10.7]	1.000
		Irbesartan	49	2 (4.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	7 (14.3)	3.429 [0.750, 15.680]	3.833 [0.754, 19.490]	10.1 [-3.3, 23.5]	0.159
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.321 [0.036, 2.880]	0.293 [0.028, 3.029]	-8.2 [-26.8, 10.5]	0.350
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.540
<= 1.75 g/day	Double-blind period	Sparsentan	98	6 (6.1)	1.139 [0.360, 3.605]	1.148 [0.338, 3.897]	0.7 [-6.9, 8.4]	1.000
		Irbesartan	93	5 (5.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	12 (11.5)	1.797 [0.736, 4.387]	1.901 [0.718, 5.033]	5.1 [-3.5, 13.7]	0.233
		Irbesartan	109	7 (6.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
 Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.181
Yes	Double-blind period	Sparsentan	90	9 (10.0)	2.933 [0.821, 10.478]	3.148 [0.823, 12.042]	6.6 [-1.8, 15.0]	0.133
		Irbesartan	88	3 (3.4)				
No	Double-blind period	Sparsentan	112	9 (8.0)	1.018 [0.420, 2.470]	1.019 [0.389, 2.671]	0.1 [-7.8, 8.1]	1.000
		Irbesartan	114	9 (7.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.135
<= 5 years	Double-blind period	Sparsentan	113	9 (8.0)	1.012 [0.426, 2.400]	1.013 [0.396, 2.588]	0.1 [-7.6, 7.8]	1.000
		Irbesartan	127	10 (7.9)				
> 5 years	Double-blind period	Sparsentan	89	9 (10.1)	3.792 [0.845, 17.013]	4.106 [0.859, 19.632]	7.4 [-1.0, 15.9]	0.067
		Irbesartan	75	2 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEL\_SSIM: Incidence of AESI hepatic disorders during double-blind period by subgroup  
Safety Set

AESI hepatic disorders								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.358
Yes	Double-blind period	Sparsentan	155	14 (9.0)	1.818 [0.785, 4.211]	1.899 [0.773, 4.663]	4.1 [-2.2, 10.3]	0.187
		Irbesartan	161	8 (5.0)				
No	Double-blind period	Sparsentan	47	4 (8.5)	0.872 [0.233, 3.269]	0.860 [0.201, 3.683]	-1.2 [-15.6, 13.1]	1.000
		Irbesartan	41	4 (9.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.130
Male	Double-blind period	Sparsentan	139	17 (12.2)	1.943 [0.897, 4.212]	2.075 [0.892, 4.827]	5.9 [-1.5, 13.4]	0.101
		Irbesartan	143	9 (6.3)				
Female	Double-blind period	Sparsentan	63	1 (1.6)	0.312 [0.033, 2.918]	0.301 [0.030, 2.979]	-3.5 [-11.5, 4.5]	0.353
		Irbesartan	59	3 (5.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
 Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.106
<= 45 years	Double-blind period	Sparsentan	96	5 (5.2)	0.737 [0.242, 2.241]	0.722 [0.221, 2.359]	-1.9 [-9.6, 5.9]	0.768
		Irbesartan	99	7 (7.1)				
> 45 years	Double-blind period	Sparsentan	106	13 (12.3)	2.526 [0.934, 6.834]	2.740 [0.940, 7.985]	7.4 [-1.0, 15.9]	0.083
		Irbesartan	103	5 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.404
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.200 + [0.010, 4.166]	0.158 + [0.005, 4.691]	-20.0 [-70.6, 30.6]	0.357
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	10 (9.8)	1.781 [0.672, 4.724]	1.866 [0.653, 5.335]	4.3 [-3.8, 12.4]	0.301
		Irbesartan	109	6 (5.5)				
> 40 years	Double-blind period	Sparsentan	91	8 (8.8)	1.547 [0.526, 4.548]	1.600 [0.503, 5.094]	3.1 [-5.6, 11.8]	0.567
		Irbesartan	88	5 (5.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.353
North America	Double-blind period	Sparsentan	35	5 (14.3)	1.643 [0.476, 5.672]	1.750 [0.433, 7.067]	5.6 [-11.1, 22.3]	0.490
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	10 (10.2)	2.347 [0.830, 6.634]	2.500 [0.824, 7.582]	5.9 [-2.1, 13.9]	0.112
		Irbesartan	115	5 (4.3)				
Asia Pacific	Double-blind period	Sparsentan	69	3 (4.3)	0.594 [0.126, 2.808]	0.576 [0.111, 2.996]	-3.0 [-14.2, 8.3]	0.669
		Irbesartan	41	3 (7.3)				

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N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
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p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
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eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.282
< 27 kg/m**2	Double-blind period	Sparsentan	83	6 (7.2)	0.971 [0.340, 2.773]	0.968 [0.312, 3.006]	-0.2 [-9.0, 8.6]	1.000
		Irbesartan	94	7 (7.4)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	12 (10.1)	2.158 [0.786, 5.925]	2.288 [0.779, 6.723]	5.4 [-2.2, 13.0]	0.138
		Irbesartan	107	5 (4.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.720
eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	1.668 [0.573, 4.856]	1.752 [0.545, 5.637]	4.5 [-6.2, 15.2]	0.394
		Irbesartan	74	5 (6.8)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	2 (3.6)	1.000 [0.146, 6.848]	1.000 [0.136, 7.364]	0.0 [-8.8, 8.8]	1.000
		Irbesartan	55	2 (3.6)				
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	0.973 [0.263, 3.598]	0.970 [0.223, 4.212]	-0.3 [-17.4, 16.8]	1.000
		Irbesartan	36	4 (11.1)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	4 (10.3)	3.795 [0.444, 32.404]	4.114 [0.438, 38.653]	7.6 [-5.9, 21.0]	0.359
		Irbesartan	37	1 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.219
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	10 (7.9)	1.451 [0.570, 3.694]	1.490 [0.549, 4.044]	2.4 [-4.4, 9.3]	0.463
		Irbesartan	129	7 (5.4)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	7 (14.3)	3.429 [0.750, 15.680]	3.833 [0.754, 19.490]	10.1 [-3.3, 23.5]	0.159
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.321 [0.036, 2.880]	0.293 [0.028, 3.029]	-8.2 [-26.8, 10.5]	0.350
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.370
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	8 (9.8)	1.561 [0.533, 4.569]	1.622 [0.507, 5.186]	3.5 [-6.1, 13.1]	0.565
		Irbesartan	80	5 (6.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	2 (4.4)	1.089 [0.160, 7.410]	1.093 [0.147, 8.102]	0.4 [-10.0, 10.7]	1.000
		Irbesartan	49	2 (4.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	7 (14.3)	3.429 [0.750, 15.680]	3.833 [0.754, 19.490]	10.1 [-3.3, 23.5]	0.159
		Irbesartan	48	2 (4.2)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.321 [0.036, 2.880]	0.293 [0.028, 3.029]	-8.2 [-26.8, 10.5]	0.350
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.540
<= 1.75 g/day	Double-blind period	Sparsentan	98	6 (6.1)	1.139 [0.360, 3.605]	1.148 [0.338, 3.897]	0.7 [-6.9, 8.4]	1.000
		Irbesartan	93	5 (5.4)				
> 1.75 g/day	Double-blind period	Sparsentan	104	12 (11.5)	1.797 [0.736, 4.387]	1.901 [0.718, 5.033]	5.1 [-3.5, 13.7]	0.233
		Irbesartan	109	7 (6.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.181
Yes	Double-blind period	Sparsentan	90	9 (10.0)	2.933 [0.821, 10.478]	3.148 [0.823, 12.042]	6.6 [-1.8, 15.0]	0.133
		Irbesartan	88	3 (3.4)				
No	Double-blind period	Sparsentan	112	9 (8.0)	1.018 [0.420, 2.470]	1.019 [0.389, 2.671]	0.1 [-7.8, 8.1]	1.000
		Irbesartan	114	9 (7.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
 Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.135
<= 5 years	Double-blind period	Sparsentan	113	9 (8.0)	1.012 [0.426, 2.400]	1.013 [0.396, 2.588]	0.1 [-7.6, 7.8]	1.000
		Irbesartan	127	10 (7.9)				
> 5 years	Double-blind period	Sparsentan	89	9 (10.1)	3.792 [0.845, 17.013]	4.106 [0.859, 19.632]	7.4 [-1.0, 15.9]	0.067
		Irbesartan	75	2 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AELN\_SSIM: Incidence of AESI hepatic disorders during double-blind period - non-severe by subgroup  
 Safety Set

AESI hepatic disorders - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.358
Yes	Double-blind period	Sparsentan	155	14 (9.0)	1.818 [0.785, 4.211]	1.899 [0.773, 4.663]	4.1 [-2.2, 10.3]	0.187
		Irbesartan	161	8 (5.0)				
No	Double-blind period	Sparsentan	47	4 (8.5)	0.872 [0.233, 3.269]	0.860 [0.201, 3.683]	-1.2 [-15.6, 13.1]	1.000
		Irbesartan	41	4 (9.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AELC\_SSIM: Incidence of AESI hepatic disorders during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AELS\_SSIM: Incidence of AESI hepatic disorders during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.750
Male	Double-blind period	Sparsentan	139	12 (8.6)	1.029 [0.479, 2.212]	1.031 [0.447, 2.381]	0.2 [-7.0, 7.5]	1.000
		Irbesartan	143	12 (8.4)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	1.405 [0.243, 8.113]	1.425 [0.230, 8.844]	1.4 [-7.3, 10.0]	1.000
		Irbesartan	59	2 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.484
<= 45 years	Double-blind period	Sparsentan	96	7 (7.3)	1.444 [0.474, 4.393]	1.479 [0.453, 4.830]	2.2 [-5.5, 10.0]	0.564
		Irbesartan	99	5 (5.1)				
> 45 years	Double-blind period	Sparsentan	106	8 (7.5)	0.864 [0.347, 2.152]	0.853 [0.316, 2.303]	-1.2 [-9.6, 7.2]	0.804
		Irbesartan	103	9 (8.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.563
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	3.000 + [0.171, 52.527]	3.667 + [0.145, 92.653]	22.2 [-20.5, 64.9]	0.505
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	8 (7.8)	1.221 [0.459, 3.247]	1.240 [0.433, 3.552]	1.4 [-6.5, 9.3]	0.791
		Irbesartan	109	7 (6.4)				
> 40 years	Double-blind period	Sparsentan	91	5 (5.5)	0.691 [0.228, 2.095]	0.673 [0.205, 2.205]	-2.5 [-10.9, 6.0]	0.562
		Irbesartan	88	7 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.765
North America	Double-blind period	Sparsentan	35	2 (5.7)	0.657 [0.128, 3.386]	0.636 [0.110, 3.690]	-3.0 [-16.7, 10.7]	0.694
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	9 (9.2)	1.320 [0.530, 3.291]	1.353 [0.501, 3.651]	2.2 [-6.1, 10.5]	0.617
		Irbesartan	115	8 (7.0)				
Asia Pacific	Double-blind period	Sparsentan	69	4 (5.8)	1.188 [0.228, 6.205]	1.200 [0.210, 6.859]	0.9 [-9.6, 11.5]	1.000
		Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.366
< 27 kg/m**2	Double-blind period	Sparsentan	83	8 (9.6)	1.510 [0.546, 4.173]	1.564 [0.520, 4.711]	3.3 [-5.9, 12.4]	0.578
		Irbesartan	94	6 (6.4)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	7 (5.9)	0.787 [0.295, 2.097]	0.773 [0.271, 2.210]	-1.6 [-9.0, 5.8]	0.790
		Irbesartan	107	8 (7.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.190
eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	1.668 [0.573, 4.856]	1.752 [0.545, 5.637]	4.5 [-6.2, 15.2]	0.394
		Irbesartan	74	5 (6.8)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5)	0.600 [0.151, 2.389]	0.577 [0.131, 2.542]	-3.6 [-15.1, 7.9]	0.716
		Irbesartan	55	5 (9.1)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)	6.816 + [0.365, 127.440]	7.406 + [0.369, 148.677]	8.1 [-3.4, 19.6]	0.240
		Irbesartan	36	0 (0.0)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)	0.237 [0.028, 2.025]	0.217 [0.023, 2.040]	-8.2 [-22.0, 5.6]	0.194
		Irbesartan	37	4 (10.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.356
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	11 (8.7)	0.931 [0.427, 2.032]	0.925 [0.392, 2.179]	-0.6 [-8.4, 7.1]	1.000
		Irbesartan	129	12 (9.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	3.918 [0.454, 33.802]	4.178 [0.450, 38.818]	6.1 [-4.6, 16.8]	0.362
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.321 + [0.014, 7.528]	0.308 + [0.012, 7.927]	-4.0 [-15.6, 7.6]	0.490
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.564
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	5 (6.1)	0.976 [0.294, 3.241]	0.974 [0.271, 3.502]	-0.2 [-8.8, 8.5]	1.000
		Irbesartan	80	5 (6.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	6 (13.3)	0.933 [0.339, 2.569]	0.923 [0.285, 2.987]	-1.0 [-17.0, 15.1]	1.000
		Irbesartan	49	7 (14.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	3.918 [0.454, 33.802]	4.178 [0.450, 38.818]	6.1 [-4.6, 16.8]	0.362
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.321 + [0.014, 7.528]	0.308 + [0.012, 7.927]	-4.0 [-15.6, 7.6]	0.490
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.283
<= 1.75 g/day	Double-blind period	Sparsentan	98	5 (5.1)	0.678 [0.223, 2.061]	0.661 [0.202, 2.159]	-2.4 [-10.4, 5.5]	0.560
		Irbesartan	93	7 (7.5)				
> 1.75 g/day	Double-blind period	Sparsentan	104	10 (9.6)	1.497 [0.592, 3.786]	1.550 [0.567, 4.238]	3.2 [-5.0, 11.4]	0.454
		Irbesartan	109	7 (6.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.896
Yes	Double-blind period	Sparsentan	90	8 (8.9)	1.117 [0.423, 2.950]	1.129 [0.391, 3.258]	0.9 [-8.3, 10.2]	1.000
		Irbesartan	88	7 (8.0)				
No	Double-blind period	Sparsentan	112	7 (6.3)	1.018 [0.369, 2.808]	1.019 [0.345, 3.006]	0.1 [-7.1, 7.3]	1.000
		Irbesartan	114	7 (6.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.316
<= 5 years	Double-blind period	Sparsentan	113	7 (6.2)	0.787 [0.310, 1.998]	0.773 [0.284, 2.102]	-1.7 [-9.0, 5.6]	0.802
		Irbesartan	127	10 (7.9)				
> 5 years	Double-blind period	Sparsentan	89	8 (9.0)	1.685 [0.528, 5.378]	1.753 [0.506, 6.069]	3.7 [-5.4, 12.7]	0.549
		Irbesartan	75	4 (5.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEP\_SSIM: Incidence of AESI acute pancreatitis during double-blind period by subgroup  
Safety Set

AESI acute pancreatitis								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.104
Yes	Double-blind period	Sparsentan	155	14 (9.0)	1.454 [0.666, 3.175]	1.499 [0.645, 3.485]	2.8 [-3.7, 9.3]	0.399
		Irbesartan	161	10 (6.2)				
No	Double-blind period	Sparsentan	47	1 (2.1)	0.218 [0.025, 1.874]	0.201 [0.022, 1.877]	-7.6 [-19.9, 4.6]	0.180
		Irbesartan	41	4 (9.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.750
Male	Double-blind period	Sparsentan	139	12 (8.6)	1.029 [0.479, 2.212]	1.031 [0.447, 2.381]	0.2 [-7.0, 7.5]	1.000
		Irbesartan	143	12 (8.4)				
Female	Double-blind period	Sparsentan	63	3 (4.8)	1.405 [0.243, 8.113]	1.425 [0.230, 8.844]	1.4 [-7.3, 10.0]	1.000
		Irbesartan	59	2 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
 Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.484
<= 45 years	Double-blind period	Sparsentan	96	7 (7.3)	1.444 [0.474, 4.393]	1.479 [0.453, 4.830]	2.2 [-5.5, 10.0]	0.564
		Irbesartan	99	5 (5.1)				
> 45 years	Double-blind period	Sparsentan	106	8 (7.5)	0.864 [0.347, 2.152]	0.853 [0.316, 2.303]	-1.2 [-9.6, 7.2]	0.804
		Irbesartan	103	9 (8.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.563
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	3.000 + [0.171, 52.527]	3.667 + [0.145, 92.653]	22.2 [-20.5, 64.9]	0.505
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	8 (7.8)	1.221 [0.459, 3.247]	1.240 [0.433, 3.552]	1.4 [-6.5, 9.3]	0.791
		Irbesartan	109	7 (6.4)				
> 40 years	Double-blind period	Sparsentan	91	5 (5.5)	0.691 [0.228, 2.095]	0.673 [0.205, 2.205]	-2.5 [-10.9, 6.0]	0.562
		Irbesartan	88	7 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.765
North America	Double-blind period	Sparsentan	35	2 (5.7)	0.657 [0.128, 3.386]	0.636 [0.110, 3.690]	-3.0 [-16.7, 10.7]	0.694
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	9 (9.2)	1.320 [0.530, 3.291]	1.353 [0.501, 3.651]	2.2 [-6.1, 10.5]	0.617
		Irbesartan	115	8 (7.0)				
Asia Pacific	Double-blind period	Sparsentan	69	4 (5.8)	1.188 [0.228, 6.205]	1.200 [0.210, 6.859]	0.9 [-9.6, 11.5]	1.000
		Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
 Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.366
< 27 kg/m**2	Double-blind period	Sparsentan	83	8 (9.6)	1.510 [0.546, 4.173]	1.564 [0.520, 4.711]	3.3 [-5.9, 12.4]	0.578
		Irbesartan	94	6 (6.4)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	7 (5.9)	0.787 [0.295, 2.097]	0.773 [0.271, 2.210]	-1.6 [-9.0, 5.8]	0.790
		Irbesartan	107	8 (7.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.190
eGFR Low and UP High	Double-blind period	Sparsentan	71	8 (11.3)	1.668 [0.573, 4.856]	1.752 [0.545, 5.637]	4.5 [-6.2, 15.2]	0.394
		Irbesartan	74	5 (6.8)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5)	0.600 [0.151, 2.389]	0.577 [0.131, 2.542]	-3.6 [-15.1, 7.9]	0.716
		Irbesartan	55	5 (9.1)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)	6.816 + [0.365, 127.440]	7.406 + [0.369, 148.677]	8.1 [-3.4, 19.6]	0.240
		Irbesartan	36	0 (0.0)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)	0.237 [0.028, 2.025]	0.217 [0.023, 2.040]	-8.2 [-22.0, 5.6]	0.194
		Irbesartan	37	4 (10.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.356
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	11 (8.7)	0.931 [0.427, 2.032]	0.925 [0.392, 2.179]	-0.6 [-8.4, 7.1]	1.000
		Irbesartan	129	12 (9.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	3.918 [0.454, 33.802]	4.178 [0.450, 38.818]	6.1 [-4.6, 16.8]	0.362
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.321 + [0.014, 7.528]	0.308 + [0.012, 7.927]	-4.0 [-15.6, 7.6]	0.490
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.564
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	5 (6.1)	0.976 [0.294, 3.241]	0.974 [0.271, 3.502]	-0.2 [-8.8, 8.5]	1.000
		Irbesartan	80	5 (6.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	6 (13.3)	0.933 [0.339, 2.569]	0.923 [0.285, 2.987]	-1.0 [-17.0, 15.1]	1.000
		Irbesartan	49	7 (14.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	3.918 [0.454, 33.802]	4.178 [0.450, 38.818]	6.1 [-4.6, 16.8]	0.362
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.321 + [0.014, 7.528]	0.308 + [0.012, 7.927]	-4.0 [-15.6, 7.6]	0.490
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.283
<= 1.75 g/day	Double-blind period	Sparsentan	98	5 (5.1)	0.678 [0.223, 2.061]	0.661 [0.202, 2.159]	-2.4 [-10.4, 5.5]	0.560
		Irbesartan	93	7 (7.5)				
> 1.75 g/day	Double-blind period	Sparsentan	104	10 (9.6)	1.497 [0.592, 3.786]	1.550 [0.567, 4.238]	3.2 [-5.0, 11.4]	0.454
		Irbesartan	109	7 (6.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.896
Yes	Double-blind period	Sparsentan	90	8 (8.9)	1.117 [0.423, 2.950]	1.129 [0.391, 3.258]	0.9 [-8.3, 10.2]	1.000
		Irbesartan	88	7 (8.0)				
No	Double-blind period	Sparsentan	112	7 (6.3)	1.018 [0.369, 2.808]	1.019 [0.345, 3.006]	0.1 [-7.1, 7.3]	1.000
		Irbesartan	114	7 (6.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.316
<= 5 years	Double-blind period	Sparsentan	113	7 (6.2)	0.787 [0.310, 1.998]	0.773 [0.284, 2.102]	-1.7 [-9.0, 5.6]	0.802
		Irbesartan	127	10 (7.9)				
> 5 years	Double-blind period	Sparsentan	89	8 (9.0)	1.685 [0.528, 5.378]	1.753 [0.506, 6.069]	3.7 [-5.4, 12.7]	0.549
		Irbesartan	75	4 (5.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEPN\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - non-severe by subgroup  
Safety Set

AESI acute pancreatitis - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.104
Yes	Double-blind period	Sparsentan	155	14 (9.0)	1.454 [0.666, 3.175]	1.499 [0.645, 3.485]	2.8 [-3.7, 9.3]	0.399
		Irbesartan	161	10 (6.2)				
No	Double-blind period	Sparsentan	47	1 (2.1)	0.218 [0.025, 1.874]	0.201 [0.022, 1.877]	-7.6 [-19.9, 4.6]	0.180
		Irbesartan	41	4 (9.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEPC\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEPS\_SSIM: Incidence of AESI acute pancreatitis during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
 Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.714
Male	Double-blind period	Sparsentan	139	21 (15.1)	1.080 [0.613, 1.903]	1.094 [0.564, 2.123]	1.1 [-7.8, 10.1]	0.866
		Irbesartan	143	20 (14.0)				
Female	Double-blind period	Sparsentan	63	15 (23.8)	1.277 [0.639, 2.552]	1.364 [0.569, 3.271]	5.2 [-10.9, 21.3]	0.515
		Irbesartan	59	11 (18.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.813
<= 45 years	Double-blind period	Sparsentan	96	19 (19.8)	1.225 [0.670, 2.238]	1.280 [0.615, 2.666]	3.6 [-8.2, 15.4]	0.577
		Irbesartan	99	16 (16.2)				
> 45 years	Double-blind period	Sparsentan	106	17 (16.0)	1.101 [0.581, 2.087]	1.121 [0.527, 2.382]	1.5 [-9.2, 12.2]	0.849
		Irbesartan	103	15 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.958
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	17 (16.7)	1.135 [0.607, 2.126]	1.163 [0.553, 2.445]	2.0 [-8.8, 12.8]	0.709
		Irbesartan	109	16 (14.7)				
> 40 years	Double-blind period	Sparsentan	91	18 (19.8)	1.160 [0.625, 2.156]	1.200 [0.562, 2.561]	2.7 [-9.7, 15.2]	0.702
		Irbesartan	88	15 (17.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.650
North America	Double-blind period	Sparsentan	35	5 (14.3)	2.190 [0.561, 8.553]	2.389 [0.530, 10.764]	7.8 [-8.4, 23.9]	0.282
		Irbesartan	46	3 (6.5)				
Europe	Double-blind period	Sparsentan	98	22 (22.4)	1.122 [0.668, 1.886]	1.158 [0.599, 2.237]	2.4 [-9.5, 14.4]	0.737
		Irbesartan	115	23 (20.0)				
Asia Pacific	Double-blind period	Sparsentan	69	9 (13.0)	1.070 [0.385, 2.974]	1.080 [0.336, 3.475]	0.8 [-13.9, 15.6]	1.000
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
 Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.498
< 27 kg/m**2	Double-blind period	Sparsentan	83	11 (13.3)	0.958 [0.454, 2.022]	0.952 [0.401, 2.257]	-0.6 [-11.8, 10.7]	1.000
		Irbesartan	94	13 (13.8)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	25 (21.0)	1.322 [0.757, 2.310]	1.408 [0.713, 2.781]	5.1 [-5.8, 16.1]	0.392
		Irbesartan	107	17 (15.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.895
eGFR Low and UP High	Double-blind period	Sparsentan	71	14 (19.7)	1.216 [0.604, 2.446]	1.269 [0.542, 2.971]	3.5 [-10.4, 17.4]	0.667
		Irbesartan	74	12 (16.2)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	11 (20.0)	1.375 [0.599, 3.155]	1.469 [0.541, 3.989]	5.5 [-10.5, 21.4]	0.615
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	6 (16.2)	0.834 [0.310, 2.243]	0.802 [0.241, 2.668]	-3.2 [-23.5, 17.1]	0.768
		Irbesartan	36	7 (19.4)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	5 (12.8)	1.186 [0.345, 4.079]	1.213 [0.299, 4.916]	2.0 [-15.1, 19.1]	1.000
		Irbesartan	37	4 (10.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
 Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.188
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	24 (18.9)	1.283 [0.740, 2.223]	1.349 [0.698, 2.608]	4.2 [-5.8, 14.1]	0.406
		Irbesartan	129	19 (14.7)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	0.544 [0.197, 1.506]	0.492 [0.152, 1.595]	-8.5 [-24.5, 7.4]	0.261
		Irbesartan	48	9 (18.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	7 (26.9)	2.244 [0.652, 7.723]	2.702 [0.612, 11.932]	14.9 [-10.3, 40.1]	0.291
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.265
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	12 (14.6)	1.064 [0.499, 2.271]	1.075 [0.445, 2.601]	0.9 [-11.1, 12.9]	1.000
		Irbesartan	80	11 (13.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	12 (26.7)	1.633 [0.735, 3.627]	1.864 [0.682, 5.092]	10.3 [-8.3, 29.0]	0.313
		Irbesartan	49	8 (16.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	0.544 [0.197, 1.506]	0.492 [0.152, 1.595]	-8.5 [-24.5, 7.4]	0.261
		Irbesartan	48	9 (18.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	7 (26.9)	2.244 [0.652, 7.723]	2.702 [0.612, 11.932]	14.9 [-10.3, 40.1]	0.291
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.593
<= 1.75 g/day	Double-blind period	Sparsentan	98	15 (15.3)	1.017 [0.520, 1.989]	1.020 [0.462, 2.249]	0.3 [-11.0, 11.5]	1.000
		Irbesartan	93	14 (15.1)				
> 1.75 g/day	Double-blind period	Sparsentan	104	21 (20.2)	1.295 [0.725, 2.313]	1.369 [0.677, 2.771]	4.6 [-6.6, 15.8]	0.474
		Irbesartan	109	17 (15.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
 Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.180
Yes	Double-blind period	Sparsentan	90	17 (18.9)	0.875 [0.488, 1.570]	0.846 [0.407, 1.759]	-2.7 [-15.6, 10.2]	0.711
		Irbesartan	88	19 (21.6)				
No	Double-blind period	Sparsentan	112	19 (17.0)	1.612 [0.821, 3.162]	1.737 [0.800, 3.771]	6.4 [-3.4, 16.3]	0.179
		Irbesartan	114	12 (10.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.089
<= 5 years	Double-blind period	Sparsentan	113	24 (21.2)	1.587 [0.900, 2.798]	1.745 [0.883, 3.449]	7.9 [-2.6, 18.3]	0.123
		Irbesartan	127	17 (13.4)				
> 5 years	Double-blind period	Sparsentan	89	12 (13.5)	0.722 [0.356, 1.465]	0.679 [0.293, 1.575]	-5.2 [-17.7, 7.4]	0.397
		Irbesartan	75	14 (18.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEO\_SSIM: Incidence of AESI fluid retention during double-blind period by subgroup  
Safety Set

AESI fluid retention								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.750
Yes	Double-blind period	Sparsentan	155	31 (20.0)	1.150 [0.725, 1.823]	1.188 [0.674, 2.093]	2.6 [-6.6, 11.8]	0.567
		Irbesartan	161	28 (17.4)				
No	Double-blind period	Sparsentan	47	5 (10.6)	1.454 [0.370, 5.714]	1.508 [0.337, 6.739]	3.3 [-10.8, 17.5]	0.719
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.800
Male	Double-blind period	Sparsentan	139	21 (15.1)	1.137 [0.640, 2.021]	1.161 [0.594, 2.269]	1.8 [-7.0, 10.7]	0.734
		Irbesartan	143	19 (13.3)				
Female	Double-blind period	Sparsentan	63	15 (23.8)	1.277 [0.639, 2.552]	1.364 [0.569, 3.271]	5.2 [-10.9, 21.3]	0.515
		Irbesartan	59	11 (18.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.706
<= 45 years	Double-blind period	Sparsentan	96	19 (19.8)	1.306 [0.706, 2.419]	1.382 [0.656, 2.909]	4.6 [-7.0, 16.3]	0.452
		Irbesartan	99	15 (15.2)				
> 45 years	Double-blind period	Sparsentan	106	17 (16.0)	1.101 [0.581, 2.087]	1.121 [0.527, 2.382]	1.5 [-9.2, 12.2]	0.849
		Irbesartan	103	15 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.961
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	17 (16.7)	1.211 [0.639, 2.296]	1.253 [0.590, 2.663]	2.9 [-7.7, 13.6]	0.571
		Irbesartan	109	15 (13.8)				
> 40 years	Double-blind period	Sparsentan	91	18 (19.8)	1.160 [0.625, 2.156]	1.200 [0.562, 2.561]	2.7 [-9.7, 15.2]	0.702
		Irbesartan	88	15 (17.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.433
North America	Double-blind period	Sparsentan	35	5 (14.3)	3.286 [0.677, 15.949]	3.667 [0.667, 20.156]	9.9 [-5.6, 25.5]	0.230
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	22 (22.4)	1.122 [0.668, 1.886]	1.158 [0.599, 2.237]	2.4 [-9.5, 14.4]	0.737
		Irbesartan	115	23 (20.0)				
Asia Pacific	Double-blind period	Sparsentan	69	9 (13.0)	1.070 [0.385, 2.974]	1.080 [0.336, 3.475]	0.8 [-13.9, 15.6]	1.000
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.616
< 27 kg/m**2	Double-blind period	Sparsentan	83	11 (13.3)	1.038 [0.484, 2.227]	1.044 [0.434, 2.510]	0.5 [-10.6, 11.6]	1.000
		Irbesartan	94	12 (12.8)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	25 (21.0)	1.322 [0.757, 2.310]	1.408 [0.713, 2.781]	5.1 [-5.8, 16.1]	0.392
		Irbesartan	107	17 (15.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.874
eGFR Low and UP High	Double-blind period	Sparsentan	71	14 (19.7)	1.327 [0.646, 2.724]	1.407 [0.591, 3.348]	4.9 [-8.8, 18.5]	0.512
		Irbesartan	74	11 (14.9)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	11 (20.0)	1.375 [0.599, 3.155]	1.469 [0.541, 3.989]	5.5 [-10.5, 21.4]	0.615
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	6 (16.2)	0.834 [0.310, 2.243]	0.802 [0.241, 2.668]	-3.2 [-23.5, 17.1]	0.768
		Irbesartan	36	7 (19.4)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	5 (12.8)	1.186 [0.345, 4.079]	1.213 [0.299, 4.916]	2.0 [-15.1, 19.1]	1.000
		Irbesartan	37	4 (10.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
 Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.176
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	24 (18.9)	1.354 [0.774, 2.371]	1.437 [0.737, 2.801]	4.9 [-4.9, 14.8]	0.314
		Irbesartan	129	18 (14.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	0.544 [0.197, 1.506]	0.492 [0.152, 1.595]	-8.5 [-24.5, 7.4]	0.261
		Irbesartan	48	9 (18.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	7 (26.9)	2.244 [0.652, 7.723]	2.702 [0.612, 11.932]	14.9 [-10.3, 40.1]	0.291
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.276
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	12 (14.6)	1.171 [0.536, 2.556]	1.200 [0.487, 2.958]	2.1 [-9.6, 13.9]	0.819
		Irbesartan	80	10 (12.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	12 (26.7)	1.633 [0.735, 3.627]	1.864 [0.682, 5.092]	10.3 [-8.3, 29.0]	0.313
		Irbesartan	49	8 (16.3)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	0.544 [0.197, 1.506]	0.492 [0.152, 1.595]	-8.5 [-24.5, 7.4]	0.261
		Irbesartan	48	9 (18.8)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	7 (26.9)	2.244 [0.652, 7.723]	2.702 [0.612, 11.932]	14.9 [-10.3, 40.1]	0.291
		Irbesartan	25	3 (12.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.508
<= 1.75 g/day	Double-blind period	Sparsentan	98	15 (15.3)	1.017 [0.520, 1.989]	1.020 [0.462, 2.249]	0.3 [-11.0, 11.5]	1.000
		Irbesartan	93	14 (15.1)				
> 1.75 g/day	Double-blind period	Sparsentan	104	21 (20.2)	1.376 [0.761, 2.487]	1.471 [0.720, 3.005]	5.5 [-5.6, 16.6]	0.366
		Irbesartan	109	16 (14.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
 Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.132
Yes	Double-blind period	Sparsentan	90	17 (18.9)	0.875 [0.488, 1.570]	0.846 [0.407, 1.759]	-2.7 [-15.6, 10.2]	0.711
		Irbesartan	88	19 (21.6)				
No	Double-blind period	Sparsentan	112	19 (17.0)	1.758 [0.877, 3.524]	1.913 [0.865, 4.231]	7.3 [-2.4, 17.0]	0.119
		Irbesartan	114	11 (9.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.069
<= 5 years	Double-blind period	Sparsentan	113	24 (21.2)	1.686 [0.944, 3.010]	1.871 [0.937, 3.735]	8.6 [-1.7, 19.0]	0.084
		Irbesartan	127	16 (12.6)				
> 5 years	Double-blind period	Sparsentan	89	12 (13.5)	0.722 [0.356, 1.465]	0.679 [0.293, 1.575]	-5.2 [-17.7, 7.4]	0.397
		Irbesartan	75	14 (18.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEON\_SSIM: Incidence of AESI fluid retention during double-blind period - non-severe by subgroup  
Safety Set

AESI fluid retention - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.788
Yes	Double-blind period	Sparsentan	155	31 (20.0)	1.193 [0.748, 1.901]	1.241 [0.701, 2.196]	3.2 [-5.9, 12.4]	0.471
		Irbesartan	161	27 (16.8)				
No	Double-blind period	Sparsentan	47	5 (10.6)	1.454 [0.370, 5.714]	1.508 [0.337, 6.739]	3.3 [-10.8, 17.5]	0.719
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEOC\_SSIM: Incidence of AESI fluid retention during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEOS\_SSIM: Incidence of AESI fluid retention during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
 Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.580
Male	Double-blind period	Sparsentan	139	15 (10.8)	1.715 [0.776, 3.788]	1.801 [0.761, 4.264]	4.5 [-2.7, 11.7]	0.204
		Irbesartan	143	9 (6.3)				
Female	Double-blind period	Sparsentan	63	6 (9.5)	2.810 [0.590, 13.376]	3.000 [0.581, 15.495]	6.1 [-4.1, 16.4]	0.275
		Irbesartan	59	2 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.074
<= 45 years	Double-blind period	Sparsentan	96	8 (8.3)	1.031 [0.403, 2.637]	1.034 [0.372, 2.876]	0.3 [-8.5, 9.0]	1.000
		Irbesartan	99	8 (8.1)				
> 45 years	Double-blind period	Sparsentan	106	13 (12.3)	4.211 [1.236, 14.345]	4.659 [1.287, 16.872]	9.4 [1.4, 17.3]	0.017 *
		Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.735
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.600 + [0.014, 26.475]	0.579 + [0.010, 33.502]	0.0 [NE, NE]	NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	9 (8.8)	1.603 [0.591, 4.345]	1.661 [0.570, 4.845]	3.3 [-4.6, 11.2]	0.426
		Irbesartan	109	6 (5.5)				
> 40 years	Double-blind period	Sparsentan	91	12 (13.2)	2.321 [0.853, 6.317]	2.522 [0.850, 7.483]	7.5 [-2.1, 17.1]	0.125
		Irbesartan	88	5 (5.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.681
North America	Double-blind period	Sparsentan	35	2 (5.7)	2.629 [0.248, 27.835]	2.727 [0.237, 31.357]	3.5 [-7.7, 14.8]	0.575
		Irbesartan	46	1 (2.2)				
Europe	Double-blind period	Sparsentan	98	9 (9.2)	2.112 [0.732, 6.094]	2.225 [0.720, 6.876]	4.8 [-2.9, 12.6]	0.175
		Irbesartan	115	5 (4.3)				
Asia Pacific	Double-blind period	Sparsentan	69	10 (14.5)	1.188 [0.437, 3.235]	1.220 [0.386, 3.857]	2.3 [-12.7, 17.3]	1.000
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.737
< 27 kg/m**2	Double-blind period	Sparsentan	83	13 (15.7)	1.840 [0.803, 4.220]	1.996 [0.783, 5.088]	7.2 [-3.6, 17.9]	0.166
		Irbesartan	94	8 (8.5)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	8 (6.7)	2.398 [0.653, 8.807]	2.498 [0.645, 9.672]	3.9 [-2.4, 10.3]	0.222
		Irbesartan	107	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.647
eGFR Low and UP High	Double-blind period	Sparsentan	71	9 (12.7)	1.563 [0.587, 4.167]	1.645 [0.554, 4.887]	4.6 [-6.7, 15.9]	0.422
		Irbesartan	74	6 (8.1)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	5 (9.1)	1.250 [0.354, 4.409]	1.275 [0.324, 5.025]	1.8 [-10.2, 13.9]	1.000
		Irbesartan	55	4 (7.3)				
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	3.892 [0.457, 33.169]	4.242 [0.451, 39.943]	8.0 [-6.1, 22.1]	0.358
		Irbesartan	36	1 (2.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	3 (7.7)	6.650 + [0.355, 124.506]	7.192 + [0.359, 144.166]	7.7 [-3.3, 18.7]	0.241
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.461
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	14 (11.0)	1.580 [0.709, 3.519]	1.652 [0.688, 3.966]	4.0 [-3.7, 11.8]	0.282
		Irbesartan	129	9 (7.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	5.878 [0.735, 47.011]	6.558 [0.759, 56.698]	10.2 [-1.9, 22.3]	0.111
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.567
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	10 (12.2)	1.951 [0.698, 5.457]	2.083 [0.679, 6.393]	5.9 [-4.1, 16.0]	0.279
		Irbesartan	80	5 (6.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	1.089 [0.289, 4.099]	1.098 [0.258, 4.675]	0.7 [-12.7, 14.2]	1.000
		Irbesartan	49	4 (8.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	5.878 [0.735, 47.011]	6.558 [0.759, 56.698]	10.2 [-1.9, 22.3]	0.111
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.448
<= 1.75 g/day	Double-blind period	Sparsentan	98	9 (9.2)	2.847	3.034	6.0	0.135
		Irbesartan	93	3 (3.2)	[0.795, 10.193]	[0.795, 11.575]	[-1.8, 13.8]	
> 1.75 g/day	Double-blind period	Sparsentan	104	12 (11.5)	1.572	1.647	4.2	0.351
		Irbesartan	109	8 (7.3)	[0.670, 3.690]	[0.644, 4.208]	[-4.6, 13.0]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
 Safety Set

AESI anemia					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.480
Yes	Double-blind period	Sparsentan	90	9 (10.0)	1.467	1.519	3.2	0.591
		Irbesartan	88	6 (6.8)	[0.545, 3.948]	[0.517, 4.461]	[-6.1, 12.4]	
No	Double-blind period	Sparsentan	112	12 (10.7)	2.443	2.616	6.3	0.082
		Irbesartan	114	5 (4.4)	[0.890, 6.708]	[0.890, 7.688]	[-1.4, 14.1]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
Safety Set

AESI anemia					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.555
<= 5 years	Double-blind period	Sparsentan	113	14 (12.4)	1.748	1.854	5.3	0.191
		Irbesartan	127	9 (7.1)	[0.787, 3.884]	[0.770, 4.465]	[-3.1, 13.7]	
> 5 years	Double-blind period	Sparsentan	89	7 (7.9)	2.949	3.116	5.2	0.182
		Irbesartan	75	2 (2.7)	[0.632, 13.774]	[0.627, 15.476]	[-2.7, 13.1]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEA\_SSIM: Incidence of AESI anemia during double-blind period by subgroup  
 Safety Set

AESI anemia					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.661
Yes	Double-blind period	Sparsentan	155	16 (10.3)	2.077	2.201	5.4	0.090
		Irbesartan	161	8 (5.0)	[0.915, 4.715]	[0.914, 5.303]	[-1.1, 11.8]	
No	Double-blind period	Sparsentan	47	5 (10.6)	1.454	1.508	3.3	0.719
		Irbesartan	41	3 (7.3)	[0.370, 5.714]	[0.337, 6.739]	[-10.8, 17.5]	

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.580
Male	Double-blind period	Sparsentan	139	15 (10.8)	1.715 [0.776, 3.788]	1.801 [0.761, 4.264]	4.5 [-2.7, 11.7]	0.204
		Irbesartan	143	9 (6.3)				
Female	Double-blind period	Sparsentan	63	6 (9.5)	2.810 [0.590, 13.376]	3.000 [0.581, 15.495]	6.1 [-4.1, 16.4]	0.275
		Irbesartan	59	2 (3.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.074
<= 45 years	Double-blind period	Sparsentan	96	8 (8.3)	1.031 [0.403, 2.637]	1.034 [0.372, 2.876]	0.3 [-8.5, 9.0]	1.000
		Irbesartan	99	8 (8.1)				
> 45 years	Double-blind period	Sparsentan	106	13 (12.3)	4.211 [1.236, 14.345]	4.659 [1.287, 16.872]	9.4 [1.4, 17.3]	0.017 *
		Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.735
<= 18 years	Double-blind period	Sparsentan	9	0 (0.0)	0.600 + [0.014, 26.475]	0.579 + [0.010, 33.502]	0.0 [NE, NE]	NE
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	9 (8.8)	1.603 [0.591, 4.345]	1.661 [0.570, 4.845]	3.3 [-4.6, 11.2]	0.426
		Irbesartan	109	6 (5.5)				
> 40 years	Double-blind period	Sparsentan	91	12 (13.2)	2.321 [0.853, 6.317]	2.522 [0.850, 7.483]	7.5 [-2.1, 17.1]	0.125
		Irbesartan	88	5 (5.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.681
North America	Double-blind period	Sparsentan	35	2 (5.7)	2.629 [0.248, 27.835]	2.727 [0.237, 31.357]	3.5 [-7.7, 14.8]	0.575
		Irbesartan	46	1 (2.2)				
Europe	Double-blind period	Sparsentan	98	9 (9.2)	2.112 [0.732, 6.094]	2.225 [0.720, 6.876]	4.8 [-2.9, 12.6]	0.175
		Irbesartan	115	5 (4.3)				
Asia Pacific	Double-blind period	Sparsentan	69	10 (14.5)	1.188 [0.437, 3.235]	1.220 [0.386, 3.857]	2.3 [-12.7, 17.3]	1.000
		Irbesartan	41	5 (12.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.737
< 27 kg/m**2	Double-blind period	Sparsentan	83	13 (15.7)	1.840 [0.803, 4.220]	1.996 [0.783, 5.088]	7.2 [-3.6, 17.9]	0.166
		Irbesartan	94	8 (8.5)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	8 (6.7)	2.398 [0.653, 8.807]	2.498 [0.645, 9.672]	3.9 [-2.4, 10.3]	0.222
		Irbesartan	107	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.647
eGFR Low and UP High	Double-blind period	Sparsentan	71	9 (12.7)	1.563 [0.587, 4.167]	1.645 [0.554, 4.887]	4.6 [-6.7, 15.9]	0.422
		Irbesartan	74	6 (8.1)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	5 (9.1)	1.250 [0.354, 4.409]	1.275 [0.324, 5.025]	1.8 [-10.2, 13.9]	1.000
		Irbesartan	55	4 (7.3)				
eGFR High and UP High	Double-blind period	Sparsentan	37	4 (10.8)	3.892 [0.457, 33.169]	4.242 [0.451, 39.943]	8.0 [-6.1, 22.1]	0.358
		Irbesartan	36	1 (2.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	3 (7.7)	6.650 + [0.355, 124.506]	7.192 + [0.359, 144.166]	7.7 [-3.3, 18.7]	0.241
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.461
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	14 (11.0)	1.580 [0.709, 3.519]	1.652 [0.688, 3.966]	4.0 [-3.7, 11.8]	0.282
		Irbesartan	129	9 (7.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	5.878 [0.735, 47.011]	6.558 [0.759, 56.698]	10.2 [-1.9, 22.3]	0.111
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.567
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	10 (12.2)	1.951 [0.698, 5.457]	2.083 [0.679, 6.393]	5.9 [-4.1, 16.0]	0.279
		Irbesartan	80	5 (6.3)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	1.089 [0.289, 4.099]	1.098 [0.258, 4.675]	0.7 [-12.7, 14.2]	1.000
		Irbesartan	49	4 (8.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	6 (12.2)	5.878 [0.735, 47.011]	6.558 [0.759, 56.698]	10.2 [-1.9, 22.3]	0.111
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.448
<= 1.75 g/day	Double-blind period	Sparsentan	98	9 (9.2)	2.847 [0.795, 10.193]	3.034 [0.795, 11.575]	6.0 [-1.8, 13.8]	0.135
		Irbesartan	93	3 (3.2)				
> 1.75 g/day	Double-blind period	Sparsentan	104	12 (11.5)	1.572 [0.670, 3.690]	1.647 [0.644, 4.208]	4.2 [-4.6, 13.0]	0.351
		Irbesartan	109	8 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.480
Yes	Double-blind period	Sparsentan	90	9 (10.0)	1.467 [0.545, 3.948]	1.519 [0.517, 4.461]	3.2 [-6.1, 12.4]	0.591
		Irbesartan	88	6 (6.8)				
No	Double-blind period	Sparsentan	112	12 (10.7)	2.443 [0.890, 6.708]	2.616 [0.890, 7.688]	6.3 [-1.4, 14.1]	0.082
		Irbesartan	114	5 (4.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.555
<= 5 years	Double-blind period	Sparsentan	113	14 (12.4)	1.748 [0.787, 3.884]	1.854 [0.770, 4.465]	5.3 [-3.1, 13.7]	0.191
		Irbesartan	127	9 (7.1)				
> 5 years	Double-blind period	Sparsentan	89	7 (7.9)	2.949 [0.632, 13.774]	3.116 [0.627, 15.476]	5.2 [-2.7, 13.1]	0.182
		Irbesartan	75	2 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEAN\_SSIM: Incidence of AESI anemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI anemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.661
Yes	Double-blind period	Sparsentan	155	16 (10.3)	2.077 [0.915, 4.715]	2.201 [0.914, 5.303]	5.4 [-1.1, 11.8]	0.090
		Irbesartan	161	8 (5.0)				
No	Double-blind period	Sparsentan	47	5 (10.6)	1.454 [0.370, 5.714]	1.508 [0.337, 6.739]	3.3 [-10.8, 17.5]	0.719
		Irbesartan	41	3 (7.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEAC\_SSIM: Incidence of AESI anemia during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEAS\_SSIM: Incidence of AESI anemia during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.452
Male	Double-blind period	Sparsentan	139	26 (18.7)	1.408 [0.817, 2.424]	1.502 [0.789, 2.860]	5.4 [-3.8, 14.7]	0.256
		Irbesartan	143	19 (13.3)				
Female	Double-blind period	Sparsentan	63	8 (12.7)	0.937 [0.376, 2.334]	0.927 [0.324, 2.654]	-0.9 [-14.5, 12.8]	1.000
		Irbesartan	59	8 (13.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.975
<= 45 years	Double-blind period	Sparsentan	96	17 (17.7)	1.252 [0.654, 2.397]	1.307 [0.604, 2.824]	3.6 [-7.7, 14.9]	0.559
		Irbesartan	99	14 (14.1)				
> 45 years	Double-blind period	Sparsentan	106	17 (16.0)	1.271 [0.651, 2.481]	1.322 [0.607, 2.883]	3.4 [-7.0, 13.9]	0.556
		Irbesartan	103	13 (12.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.692
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	18 (17.6)	1.069 [0.589, 1.937]	1.083 [0.529, 2.220]	1.1 [-10.0, 12.2]	0.856
		Irbesartan	109	18 (16.5)				
> 40 years	Double-blind period	Sparsentan	91	15 (16.5)	1.612 [0.744, 3.490]	1.732 [0.715, 4.195]	6.3 [-4.8, 17.3]	0.274
		Irbesartan	88	9 (10.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.325
North America	Double-blind period	Sparsentan	35	8 (22.9)	2.629 [0.860, 8.031]	3.111 [0.853, 11.347]	14.2 [-4.5, 32.8]	0.114
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	14 (14.3)	1.027 [0.528, 1.996]	1.031 [0.476, 2.236]	0.4 [-10.0, 10.7]	1.000
		Irbesartan	115	16 (13.9)				
Asia Pacific	Double-blind period	Sparsentan	69	12 (17.4)	1.019 [0.436, 2.379]	1.023 [0.367, 2.848]	0.3 [-16.2, 16.8]	1.000
		Irbesartan	41	7 (17.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
 Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.914
< 27 kg/m**2	Double-blind period	Sparsentan	83	14 (16.9)	1.220 [0.609, 2.443]	1.264 [0.557, 2.871]	3.0 [-8.8, 14.8]	0.676
		Irbesartan	94	13 (13.8)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	20 (16.8)	1.285 [0.683, 2.414]	1.342 [0.641, 2.811]	3.7 [-6.4, 13.9]	0.462
		Irbesartan	107	14 (13.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.288
eGFR Low and UP High	Double-blind period	Sparsentan	71	18 (25.4)	1.251 [0.684, 2.286]	1.336 [0.613, 2.912]	5.1 [-9.9, 20.1]	0.553
		Irbesartan	74	15 (20.3)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	11 (20.0)	1.100 [0.509, 2.377]	1.125 [0.434, 2.914]	1.8 [-14.7, 18.3]	1.000
		Irbesartan	55	10 (18.2)				
eGFR High and UP High	Double-blind period	Sparsentan	37	5 (13.5)	10.711 + [0.614, 186.919]	12.354 + [0.657, 232.145]	13.5 [-0.2, 27.3]	0.054
		Irbesartan	36	0 (0.0)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0)	0.190 + [0.009, 3.831]	0.180 + [0.008, 3.872]	-5.4 [-15.3, 4.5]	0.234
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.920
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	29 (22.8)	1.227 [0.758, 1.988]	1.295 [0.706, 2.375]	4.2 [-6.5, 14.9]	0.443
		Irbesartan	129	24 (18.6)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	1.633 [0.413, 6.457]	1.705 [0.384, 7.567]	4.0 [-9.0, 16.9]	0.715
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.915
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	23 (28.0)	1.122 [0.671, 1.876]	1.169 [0.581, 2.352]	3.0 [-11.8, 17.9]	0.723
		Irbesartan	80	20 (25.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	6 (13.3)	1.633 [0.493, 5.416]	1.731 [0.455, 6.582]	5.2 [-9.5, 19.8]	0.512
		Irbesartan	49	4 (8.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	1.633 [0.413, 6.457]	1.705 [0.384, 7.567]	4.0 [-9.0, 16.9]	0.715
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
 Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.676
<= 1.75 g/day	Double-blind period	Sparsentan	98	13 (13.3)	1.122 [0.529, 2.377]	1.140 [0.483, 2.690]	1.4 [-9.0, 11.9]	0.829
		Irbesartan	93	11 (11.8)				
> 1.75 g/day	Double-blind period	Sparsentan	104	21 (20.2)	1.376 [0.761, 2.487]	1.471 [0.720, 3.005]	5.5 [-5.6, 16.6]	0.366
		Irbesartan	109	16 (14.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.095
Yes	Double-blind period	Sparsentan	90	21 (23.3)	1.867 [0.957, 3.640]	2.130 [0.959, 4.735]	10.8 [-1.4, 23.1]	0.079
		Irbesartan	88	11 (12.5)				
No	Double-blind period	Sparsentan	112	13 (11.6)	0.827 [0.417, 1.639]	0.804 [0.367, 1.760]	-2.4 [-12.0, 7.2]	0.692
		Irbesartan	114	16 (14.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.419
<= 5 years	Double-blind period	Sparsentan	113	21 (18.6)	1.475 [0.810, 2.685]	1.584 [0.781, 3.210]	6.0 [-4.1, 16.0]	0.215
		Irbesartan	127	16 (12.6)				
> 5 years	Double-blind period	Sparsentan	89	13 (14.6)	0.996 [0.474, 2.091]	0.995 [0.417, 2.373]	-0.1 [-12.1, 12.0]	1.000
		Irbesartan	75	11 (14.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEU\_SSIM: Incidence of AESI hyperkalemia during double-blind period by subgroup  
Safety Set

AESI hyperkalemia								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.277
Yes	Double-blind period	Sparsentan	155	29 (18.7)	1.434 [0.856, 2.404]	1.534 [0.833, 2.827]	5.7 [-3.0, 14.3]	0.217
		Irbesartan	161	21 (13.0)				
No	Double-blind period	Sparsentan	47	5 (10.6)	0.727 [0.239, 2.207]	0.694 [0.195, 2.470]	-4.0 [-20.2, 12.2]	0.749
		Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.452
Male	Double-blind period	Sparsentan	139	26 (18.7)	1.408 [0.817, 2.424]	1.502 [0.789, 2.860]	5.4 [-3.8, 14.7]	0.256
		Irbesartan	143	19 (13.3)				
Female	Double-blind period	Sparsentan	63	8 (12.7)	0.937 [0.376, 2.334]	0.927 [0.324, 2.654]	-0.9 [-14.5, 12.8]	1.000
		Irbesartan	59	8 (13.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.975
<= 45 years	Double-blind period	Sparsentan	96	17 (17.7)	1.252 [0.654, 2.397]	1.307 [0.604, 2.824]	3.6 [-7.7, 14.9]	0.559
		Irbesartan	99	14 (14.1)				
> 45 years	Double-blind period	Sparsentan	106	17 (16.0)	1.271 [0.651, 2.481]	1.322 [0.607, 2.883]	3.4 [-7.0, 13.9]	0.556
		Irbesartan	103	13 (12.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.692
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	18 (17.6)	1.069 [0.589, 1.937]	1.083 [0.529, 2.220]	1.1 [-10.0, 12.2]	0.856
		Irbesartan	109	18 (16.5)				
> 40 years	Double-blind period	Sparsentan	91	15 (16.5)	1.612 [0.744, 3.490]	1.732 [0.715, 4.195]	6.3 [-4.8, 17.3]	0.274
		Irbesartan	88	9 (10.2)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.325
North America	Double-blind period	Sparsentan	35	8 (22.9)	2.629 [0.860, 8.031]	3.111 [0.853, 11.347]	14.2 [-4.5, 32.8]	0.114
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	14 (14.3)	1.027 [0.528, 1.996]	1.031 [0.476, 2.236]	0.4 [-10.0, 10.7]	1.000
		Irbesartan	115	16 (13.9)				
Asia Pacific	Double-blind period	Sparsentan	69	12 (17.4)	1.019 [0.436, 2.379]	1.023 [0.367, 2.848]	0.3 [-16.2, 16.8]	1.000
		Irbesartan	41	7 (17.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.914
< 27 kg/m**2	Double-blind period	Sparsentan	83	14 (16.9)	1.220 [0.609, 2.443]	1.264 [0.557, 2.871]	3.0 [-8.8, 14.8]	0.676
		Irbesartan	94	13 (13.8)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	20 (16.8)	1.285 [0.683, 2.414]	1.342 [0.641, 2.811]	3.7 [-6.4, 13.9]	0.462
		Irbesartan	107	14 (13.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.288
eGFR Low and UP High	Double-blind period	Sparsentan	71	18 (25.4)	1.251 [0.684, 2.286]	1.336 [0.613, 2.912]	5.1 [-9.9, 20.1]	0.553
		Irbesartan	74	15 (20.3)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	11 (20.0)	1.100 [0.509, 2.377]	1.125 [0.434, 2.914]	1.8 [-14.7, 18.3]	1.000
		Irbesartan	55	10 (18.2)				
eGFR High and UP High	Double-blind period	Sparsentan	37	5 (13.5)	10.711 + [0.614, 186.919]	12.354 + [0.657, 232.145]	13.5 [-0.2, 27.3]	0.054
		Irbesartan	36	0 (0.0)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	0 (0.0)	0.190 + [0.009, 3.831]	0.180 + [0.008, 3.872]	-5.4 [-15.3, 4.5]	0.234
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.920
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	29 (22.8)	1.227 [0.758, 1.988]	1.295 [0.706, 2.375]	4.2 [-6.5, 14.9]	0.443
		Irbesartan	129	24 (18.6)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	1.633 [0.413, 6.457]	1.705 [0.384, 7.567]	4.0 [-9.0, 16.9]	0.715
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.915
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	23 (28.0)	1.122 [0.671, 1.876]	1.169 [0.581, 2.352]	3.0 [-11.8, 17.9]	0.723
		Irbesartan	80	20 (25.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	6 (13.3)	1.633 [0.493, 5.416]	1.731 [0.455, 6.582]	5.2 [-9.5, 19.8]	0.512
		Irbesartan	49	4 (8.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	5 (10.2)	1.633 [0.413, 6.457]	1.705 [0.384, 7.567]	4.0 [-9.0, 16.9]	0.715
		Irbesartan	48	3 (6.3)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.676
<= 1.75 g/day	Double-blind period	Sparsentan	98	13 (13.3)	1.122 [0.529, 2.377]	1.140 [0.483, 2.690]	1.4 [-9.0, 11.9]	0.829
		Irbesartan	93	11 (11.8)				
> 1.75 g/day	Double-blind period	Sparsentan	104	21 (20.2)	1.376 [0.761, 2.487]	1.471 [0.720, 3.005]	5.5 [-5.6, 16.6]	0.366
		Irbesartan	109	16 (14.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.095
Yes	Double-blind period	Sparsentan	90	21 (23.3)	1.867 [0.957, 3.640]	2.130 [0.959, 4.735]	10.8 [-1.4, 23.1]	0.079
		Irbesartan	88	11 (12.5)				
No	Double-blind period	Sparsentan	112	13 (11.6)	0.827 [0.417, 1.639]	0.804 [0.367, 1.760]	-2.4 [-12.0, 7.2]	0.692
		Irbesartan	114	16 (14.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.419
<= 5 years	Double-blind period	Sparsentan	113	21 (18.6)	1.475 [0.810, 2.685]	1.584 [0.781, 3.210]	6.0 [-4.1, 16.0]	0.215
		Irbesartan	127	16 (12.6)				
> 5 years	Double-blind period	Sparsentan	89	13 (14.6)	0.996 [0.474, 2.091]	0.995 [0.417, 2.373]	-0.1 [-12.1, 12.0]	1.000
		Irbesartan	75	11 (14.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEUN\_SSIM: Incidence of AESI hyperkalemia during double-blind period - non-severe by subgroup  
 Safety Set

AESI hyperkalemia - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.277
Yes	Double-blind period	Sparsentan	155	29 (18.7)	1.434 [0.856, 2.404]	1.534 [0.833, 2.827]	5.7 [-3.0, 14.3]	0.217
		Irbesartan	161	21 (13.0)				
No	Double-blind period	Sparsentan	47	5 (10.6)	0.727 [0.239, 2.207]	0.694 [0.195, 2.470]	-4.0 [-20.2, 12.2]	0.749
		Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEUC\_SSIM: Incidence of AESI hyperkalemia during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEUS\_SSIM: Incidence of AESI hyperkalemia during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.327
Male	Double-blind period	Sparsentan	139	7 (5.0)	0.450 [0.191, 1.060]	0.421 [0.168, 1.057]	-6.2 [-13.2, 0.9]	0.081
		Irbesartan	143	16 (11.2)				
Female	Double-blind period	Sparsentan	63	5 (7.9)	0.937 [0.286, 3.071]	0.931 [0.255, 3.395]	-0.5 [-11.9, 10.9]	1.000
		Irbesartan	59	5 (8.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
 Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.423
<= 45 years	Double-blind period	Sparsentan	96	5 (5.2)	0.430 [0.157, 1.174]	0.398 [0.135, 1.178]	-6.9 [-15.8, 1.9]	0.127
		Irbesartan	99	12 (12.1)				
> 45 years	Double-blind period	Sparsentan	106	7 (6.6)	0.756 [0.292, 1.954]	0.738 [0.264, 2.063]	-2.1 [-10.3, 6.0]	0.611
		Irbesartan	103	9 (8.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.637
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	0.556 [0.044, 7.095]	0.500 [0.024, 10.251]	-8.9 [-65.1, 47.3]	1.000
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	5 (4.9)	0.411 [0.152, 1.112]	0.381 [0.131, 1.109]	-7.0 [-15.4, 1.3]	0.085
		Irbesartan	109	13 (11.9)				
> 40 years	Double-blind period	Sparsentan	91	6 (6.6)	0.829 [0.290, 2.369]	0.817 [0.263, 2.534]	-1.4 [-10.1, 7.4]	0.780
		Irbesartan	88	7 (8.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.172
North America	Double-blind period	Sparsentan	35	2 (5.7)	0.438 [0.094, 2.041]	0.404 [0.076, 2.136]	-7.3 [-22.2, 7.6]	0.455
		Irbesartan	46	6 (13.0)				
Europe	Double-blind period	Sparsentan	98	8 (8.2)	1.043 [0.418, 2.600]	1.047 [0.388, 2.826]	0.3 [-7.9, 8.6]	1.000
		Irbesartan	115	9 (7.8)				
Asia Pacific	Double-blind period	Sparsentan	69	2 (2.9)	0.198 [0.042, 0.936]	0.174 [0.033, 0.908]	-11.7 [-25.2, 1.7]	0.050
		Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 0.988
< 27 kg/m**2	Double-blind period	Sparsentan	83	5 (6.0)	0.566 [0.202, 1.590]	0.538 [0.176, 1.645]	-4.6 [-13.8, 4.6]	0.295
		Irbesartan	94	10 (10.6)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	7 (5.9)	0.572 [0.230, 1.423]	0.545 [0.203, 1.462]	-4.4 [-12.4, 3.6]	0.325
		Irbesartan	107	11 (10.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.855
eGFR Low and UP High	Double-blind period	Sparsentan	71	5 (7.0)	0.744 [0.248, 2.238]	0.725 [0.219, 2.400]	-2.4 [-12.7, 7.9]	0.765
		Irbesartan	74	7 (9.5)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5)	0.375 [0.105, 1.340]	0.339 [0.085, 1.353]	-9.1 [-22.0, 3.8]	0.202
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)	0.730 [0.176, 3.034]	0.706 [0.146, 3.403]	-3.0 [-19.3, 13.3]	0.711
		Irbesartan	36	4 (11.1)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)	0.474 [0.045, 5.014]	0.461 [0.040, 5.305]	-2.8 [-14.3, 8.6]	0.610
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.754
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	7 (5.5)	0.474 [0.200, 1.124]	0.443 [0.174, 1.127]	-6.1 [-13.7, 1.5]	0.117
		Irbesartan	129	15 (11.6)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.784 [0.224, 2.744]	0.764 [0.192, 3.037]	-2.3 [-15.9, 11.4]	0.740
		Irbesartan	48	5 (10.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.595
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	3 (3.7)	0.293 [0.084, 1.025]	0.266 [0.070, 1.005]	-8.8 [-18.4, 0.7]	0.046 *
		Irbesartan	80	10 (12.5)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	0.871 [0.249, 3.043]	0.859 [0.216, 3.419]	-1.3 [-15.3, 12.7]	1.000
		Irbesartan	49	5 (10.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.784 [0.224, 2.744]	0.764 [0.192, 3.037]	-2.3 [-15.9, 11.4]	0.740
		Irbesartan	48	5 (10.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.457
<= 1.75 g/day	Double-blind period	Sparsentan	98	7 (7.1)	0.738 [0.287, 1.901]	0.718 [0.256, 2.014]	-2.5 [-11.5, 6.4]	0.606
		Irbesartan	93	9 (9.7)				
> 1.75 g/day	Double-blind period	Sparsentan	104	5 (4.8)	0.437 [0.159, 1.197]	0.408 [0.139, 1.202]	-6.2 [-14.3, 1.9]	0.129
		Irbesartan	109	12 (11.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.819
Yes	Double-blind period	Sparsentan	90	8 (8.9)	0.602 [0.262, 1.380]	0.563 [0.221, 1.433]	-5.9 [-16.5, 4.7]	0.252
		Irbesartan	88	13 (14.8)				
No	Double-blind period	Sparsentan	112	4 (3.6)	0.509 [0.158, 1.642]	0.491 [0.143, 1.679]	-3.4 [-10.1, 3.3]	0.375
		Irbesartan	114	8 (7.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
 Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.135
<= 5 years	Double-blind period	Sparsentan	113	6 (5.3)	0.397 [0.162, 0.971]	0.363 [0.138, 0.955]	-8.1 [-16.1, -0.0]	0.047 *
		Irbesartan	127	17 (13.4)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	1.264 [0.370, 4.313]	1.283 [0.348, 4.728]	1.4 [-7.1, 9.9]	0.756
		Irbesartan	75	4 (5.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEY\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period by subgroup  
 Safety Set

AESI cardiac arrhythmias								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.401
Yes	Double-blind period	Sparsentan	155	12 (7.7)	0.656 [0.330, 1.306]	0.627 [0.294, 1.340]	-4.1 [-11.2, 3.1]	0.259
		Irbesartan	161	19 (11.8)				
No	Double-blind period	Sparsentan	47	0 (0.0)	0.175 + [0.009, 3.543]	0.166 + [0.008, 3.567]	-4.9 [-13.8, 4.0]	0.214
		Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.426
Male	Double-blind period	Sparsentan	139	7 (5.0)	0.514 [0.214, 1.236]	0.489 [0.191, 1.250]	-4.8 [-11.5, 2.0]	0.173
		Irbesartan	143	14 (9.8)				
Female	Double-blind period	Sparsentan	63	5 (7.9)	0.937 [0.286, 3.071]	0.931 [0.255, 3.395]	-0.5 [-11.9, 10.9]	1.000
		Irbesartan	59	5 (8.5)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.409
<= 45 years	Double-blind period	Sparsentan	96	5 (5.2)	0.469 [0.169, 1.299]	0.440 [0.147, 1.317]	-5.9 [-14.5, 2.7]	0.192
		Irbesartan	99	11 (11.1)				
> 45 years	Double-blind period	Sparsentan	106	7 (6.6)	0.850 [0.320, 2.260]	0.840 [0.293, 2.406]	-1.2 [-9.1, 6.8]	0.794
		Irbesartan	103	8 (7.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.590
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	0.556 [0.044, 7.095]	0.500 [0.024, 10.251]	-8.9 [-65.1, 47.3]	1.000
		Irbesartan	5	1 (20.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	5 (4.9)	0.445 [0.163, 1.220]	0.417 [0.141, 1.228]	-6.1 [-14.3, 2.1]	0.131
		Irbesartan	109	12 (11.0)				
> 40 years	Double-blind period	Sparsentan	91	6 (6.6)	0.967 [0.324, 2.885]	0.965 [0.299, 3.113]	-0.2 [-8.7, 8.2]	1.000
		Irbesartan	88	6 (6.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
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eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: 0.195
North America	Double-blind period	Sparsentan	35	2 (5.7)	0.657 [0.128, 3.386]	0.636 [0.110, 3.690]	-3.0 [-16.7, 10.7]	0.694
		Irbesartan	46	4 (8.7)				
Europe	Double-blind period	Sparsentan	98	8 (8.2)	1.043 [0.418, 2.600]	1.047 [0.388, 2.826]	0.3 [-7.9, 8.6]	1.000
		Irbesartan	115	9 (7.8)				
Asia Pacific	Double-blind period	Sparsentan	69	2 (2.9)	0.198 [0.042, 0.936]	0.174 [0.033, 0.908]	-11.7 [-25.2, 1.7]	0.050
		Irbesartan	41	6 (14.6)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: 1.000
< 27 kg/m**2	Double-blind period	Sparsentan	83	5 (6.0)	0.629 [0.220, 1.803]	0.605 [0.194, 1.885]	-3.6 [-12.5, 5.4]	0.418
		Irbesartan	94	9 (9.6)				
≥ 27 kg/m**2	Double-blind period	Sparsentan	119	7 (5.9)	0.629 [0.248, 1.595]	0.606 [0.222, 1.654]	-3.5 [-11.3, 4.4]	0.450
		Irbesartan	107	10 (9.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: 0.703
eGFR Low and UP High	Double-blind period	Sparsentan	71	5 (7.0)	1.042 [0.315, 3.447]	1.045 [0.289, 3.778]	0.3 [-9.3, 9.9]	1.000
		Irbesartan	74	5 (6.8)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5)	0.375 [0.105, 1.340]	0.339 [0.085, 1.353]	-9.1 [-22.0, 3.8]	0.202
		Irbesartan	55	8 (14.5)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1)	0.730 [0.176, 3.034]	0.706 [0.146, 3.403]	-3.0 [-19.3, 13.3]	0.711
		Irbesartan	36	4 (11.1)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6)	0.474 [0.045, 5.014]	0.461 [0.040, 5.305]	-2.8 [-14.3, 8.6]	0.610
		Irbesartan	37	2 (5.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.858
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	7 (5.5)	0.547 [0.226, 1.326]	0.521 [0.201, 1.351]	-4.6 [-11.9, 2.8]	0.244
		Irbesartan	129	13 (10.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.784 [0.224, 2.744]	0.764 [0.192, 3.037]	-2.3 [-15.9, 11.4]	0.740
		Irbesartan	48	5 (10.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: 0.768
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	3 (3.7)	0.366 [0.101, 1.330]	0.342 [0.087, 1.338]	-6.3 [-15.3, 2.6]	0.129
		Irbesartan	80	8 (10.0)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)	0.871 [0.249, 3.043]	0.859 [0.216, 3.419]	-1.3 [-15.3, 12.7]	1.000
		Irbesartan	49	5 (10.2)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	0.784 [0.224, 2.744]	0.764 [0.192, 3.037]	-2.3 [-15.9, 11.4]	0.740
		Irbesartan	48	5 (10.4)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	1 (3.8)	0.962 [0.064, 14.551]	0.960 [0.057, 16.232]	-0.2 [-14.7, 14.4]	1.000
		Irbesartan	25	1 (4.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.441
<= 1.75 g/day	Double-blind period	Sparsentan	98	7 (7.1)	0.830 [0.314, 2.199]	0.817 [0.284, 2.351]	-1.5 [-10.2, 7.2]	0.791
		Irbesartan	93	8 (8.6)				
> 1.75 g/day	Double-blind period	Sparsentan	104	5 (4.8)	0.476 [0.171, 1.324]	0.450 [0.151, 1.343]	-5.3 [-13.2, 2.6]	0.195
		Irbesartan	109	11 (10.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.879
Yes	Double-blind period	Sparsentan	90	8 (8.9)	0.652 [0.280, 1.517]	0.618 [0.240, 1.594]	-4.7 [-15.1, 5.6]	0.351
		Irbesartan	88	12 (13.6)				
No	Double-blind period	Sparsentan	112	4 (3.6)	0.582 [0.175, 1.932]	0.566 [0.161, 1.990]	-2.6 [-9.0, 3.9]	0.539
		Irbesartan	114	7 (6.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.185
<= 5 years	Double-blind period	Sparsentan	113	6 (5.3)	0.450 [0.181, 1.119]	0.419 [0.157, 1.119]	-6.5 [-14.3, 1.3]	0.108
		Irbesartan	127	15 (11.8)				
> 5 years	Double-blind period	Sparsentan	89	6 (6.7)	1.264 [0.370, 4.313]	1.283 [0.348, 4.728]	1.4 [-7.1, 9.9]	0.756
		Irbesartan	75	4 (5.3)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEYN\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - non-severe by subgroup  
 Safety Set

AESI cardiac arrhythmias - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.363
Yes	Double-blind period	Sparsentan	155	12 (7.7)	0.733 [0.362, 1.484]	0.711 [0.328, 1.542]	-2.8 [-9.8, 4.2]	0.439
		Irbesartan	161	17 (10.6)				
No	Double-blind period	Sparsentan	47	0 (0.0)	0.175 + [0.009, 3.543]	0.166 + [0.008, 3.567]	-4.9 [-13.8, 4.0]	0.214
		Irbesartan	41	2 (4.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEYC\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEYS\_SSIM: Incidence of AESI cardiac arrhythmias during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
 Safety Set

AESI acute kidney injury								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.813
Male	Double-blind period	Sparsentan	139	10 (7.2)	2.572 [0.826, 8.008]	2.694 [0.824, 8.802]	4.4 [-1.4, 10.2]	0.105
		Irbesartan	143	4 (2.8)				
Female	Double-blind period	Sparsentan	63	2 (3.2)	1.873 [0.174, 20.118]	1.902 [0.168, 21.541]	1.5 [-5.6, 8.6]	1.000
		Irbesartan	59	1 (1.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
 Safety Set

AESI acute kidney injury								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.079
<= 45 years	Double-blind period	Sparsentan	96	10 (10.4)	5.156 [1.160, 22.925]	5.640 [1.202, 26.456]	8.4 [0.7, 16.1]	0.017 *
		Irbesartan	99	2 (2.0)				
> 45 years	Double-blind period	Sparsentan	106	2 (1.9)	0.648 [0.111, 3.798]	0.641 [0.105, 3.917]	-1.0 [-6.1, 4.1]	0.680
		Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
Safety Set

AESI acute kidney injury								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.971
<= 18 years	Double-blind period	Sparsentan	9	2 (22.2)	3.000 + [0.171, 52.527]	3.667 + [0.145, 92.653]	22.2 [-20.5, 64.9]	0.505
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	8 (7.8)	2.137 [0.664, 6.883]	2.234 [0.652, 7.659]	4.2 [-3.1, 11.4]	0.240
		Irbesartan	109	4 (3.7)				
> 40 years	Double-blind period	Sparsentan	91	2 (2.2)	1.934 [0.179, 20.950]	1.955 [0.174, 21.955]	1.1 [-3.8, 5.9]	1.000
		Irbesartan	88	1 (1.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
Safety Set

<u>AESI acute kidney injury</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: NE
North America	Double-blind period	Sparsentan	35	3 (8.6) all n<10				NE
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	7 (7.1) all n<10				NE
		Irbesartan	115	2 (1.7)				
Asia Pacific	Double-blind period	Sparsentan	69	2 (2.9) all n<10				NE
		Irbesartan	41	1 (2.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
 Safety Set

<u>AESI acute kidney injury</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: NE
< 27 kg/m**2	Double-blind period	Sparsentan	83	6 (7.2) all n<10				NE
		Irbesartan	94	2 (2.1)				
>= 27 kg/m**2	Double-blind period	Sparsentan	119	6 (5.0) all n<10				NE
		Irbesartan	107	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
 Safety Set

<u>AESI acute kidney injury</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	5 (7.0) all n<10				NE
		Irbesartan	74	3 (4.1)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5) all n<10				NE
		Irbesartan	55	1 (1.8)				
eGFR High and UP High	Double-blind period	Sparsentan	37	3 (8.1) all n<10				NE
		Irbesartan	36	1 (2.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6) all n<10				NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
Safety Set

AESI acute kidney injury								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.792
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	8 (6.3)	2.031 [0.627, 6.578]	2.101 [0.616, 7.160]	3.2 [-2.8, 9.2]	0.253
		Irbesartan	129	4 (3.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)	3.918 [0.454, 33.802]	4.178 [0.450, 38.818]	6.1 [-4.6, 16.8]	0.362
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
 Safety Set

<u>AESI acute kidney injury</u>					RR	OR	RD	
Subgroup	Time	Treatment	N	n (%)	[95 % CI]	[95 % CI]	[95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	4 (4.9)				NE
		Irbesartan	80	3 (3.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	4 (8.9)				NE
		Irbesartan	49	1 (2.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	4 (8.2)				NE
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)				NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
 Safety Set

AESI acute kidney injury								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: 0.300
<= 1.75 g/day	Double-blind period	Sparsentan	98	6 (6.1)	5.694 [0.699, 46.400]	6.000 [0.708, 50.825]	5.0 [-1.2, 11.3]	0.119
		Irbesartan	93	1 (1.1)				
> 1.75 g/day	Double-blind period	Sparsentan	104	6 (5.8)	1.572 [0.457, 5.413]	1.607 [0.440, 5.866]	2.1 [-4.5, 8.7]	0.531
		Irbesartan	109	4 (3.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
 Safety Set

AESI acute kidney injury								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: 0.382
Yes	Double-blind period	Sparsentan	90	7 (7.8)	1.711 [0.519, 5.640]	1.771 [0.500, 6.277]	3.2 [-4.9, 11.4]	0.536
		Irbesartan	88	4 (4.5)				
No	Double-blind period	Sparsentan	112	5 (4.5)	5.089 [0.604, 42.874]	5.280 [0.607, 45.936]	3.6 [-1.5, 8.7]	0.118
		Irbesartan	114	1 (0.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024



Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
 Safety Set

AESI acute kidney injury								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: 0.594
<= 5 years	Double-blind period	Sparsentan	113	8 (7.1)	2.997 [0.815, 11.024]	3.149 [0.815, 12.174]	4.7 [-1.5, 11.0]	0.121
		Irbesartan	127	3 (2.4)				
> 5 years	Double-blind period	Sparsentan	89	4 (4.5)	1.685 [0.317, 8.947]	1.718 [0.306, 9.650]	1.8 [-5.0, 8.7]	0.689
		Irbesartan	75	2 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEI\_SSIM: Incidence of AESI acute kidney injury during double-blind period by subgroup  
 Safety Set

AESI acute kidney injury								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.647
Yes	Double-blind period	Sparsentan	155	10 (6.5)	2.077 [0.727, 5.940]	2.152 [0.718, 6.445]	3.3 [-2.0, 8.7]	0.192
		Irbesartan	161	5 (3.1)				
No	Double-blind period	Sparsentan	47	2 (4.3)	4.375 + [0.216, 88.579]	4.560 + [0.213, 97.787]	4.3 [-3.8, 12.3]	0.497
		Irbesartan	41	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
 Safety Set

AESI acute kidney injury - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Sex	Double-blind period	Sparsentan						Interaction test: 0.670
Male	Double-blind period	Sparsentan	139	7 (5.0)	1.800 [0.539, 6.014]	1.843 [0.527, 6.441]	2.2 [-3.0, 7.5]	0.372
		Irbesartan	143	4 (2.8)				
Female	Double-blind period	Sparsentan	63	1 (1.6)	0.937 [0.060, 14.634]	0.935 [0.057, 15.304]	-0.1 [-6.3, 6.0]	1.000
		Irbesartan	59	1 (1.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
Safety Set

AESI acute kidney injury - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age	Double-blind period	Sparsentan						Interaction test: 0.045 #
<= 45 years	Double-blind period	Sparsentan	96	8 (8.3)	4.125 [0.899, 18.933]	4.409 [0.912, 21.323]	6.3 [-0.9, 13.5]	0.056
		Irbesartan	99	2 (2.0)				
> 45 years	Double-blind period	Sparsentan	106	0 (0.0)	0.139 + [0.007, 2.655]	0.135 + [0.007, 2.643]	-2.9 [-7.1, 1.3]	0.118
		Irbesartan	103	3 (2.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
 Safety Set

AESI acute kidney injury - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Age at IgAN diagnosis	Double-blind period	Sparsentan						Interaction test: 0.596
<= 18 years	Double-blind period	Sparsentan	9	1 (11.1)	1.800 + [0.086, 37.493]	1.941 + [0.066, 56.760]	11.1 [-25.0, 47.2]	1.000
		Irbesartan	5	0 (0.0)				
> 18 to 40 years	Double-blind period	Sparsentan	102	7 (6.9)	1.870 [0.564, 6.199]	1.934 [0.549, 6.815]	3.2 [-3.8, 10.2]	0.362
		Irbesartan	109	4 (3.7)				
> 40 years	Double-blind period	Sparsentan	91	0 (0.0)	0.322 + [0.013, 7.811]	0.319 + [0.013, 7.930]	-1.1 [-4.5, 2.2]	0.492
		Irbesartan	88	1 (1.1)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
Safety Set

<u>AESI acute kidney injury - non-severe</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Geographic region	Double-blind period	Sparsentan						Interaction test: NE
North America	Double-blind period	Sparsentan	35	3 (8.6) all n<10				NE
		Irbesartan	46	2 (4.3)				
Europe	Double-blind period	Sparsentan	98	4 (4.1) all n<10				NE
		Irbesartan	115	2 (1.7)				
Asia Pacific	Double-blind period	Sparsentan	69	1 (1.4) all n<10				NE
		Irbesartan	41	1 (2.4)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
 Safety Set

<u>AESI acute kidney injury - non-severe</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline BMI	Double-blind period	Sparsentan						Interaction test: NE
< 27 kg/m**2	Double-blind period	Sparsentan	83	4 (4.8)				NE
		Irbesartan	94	2 (2.1)				
>= 27 kg/m**2	Double-blind period	Sparsentan	119	4 (3.4)				NE
		Irbesartan	107	3 (2.8)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
Safety Set

<u>AESI acute kidney injury - non-severe</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Randomization strata	Double-blind period	Sparsentan						Interaction test: NE
eGFR Low and UP High	Double-blind period	Sparsentan	71	3 (4.2) all n<10				NE
		Irbesartan	74	3 (4.1)				
eGFR Low and UP Low	Double-blind period	Sparsentan	55	3 (5.5) all n<10				NE
		Irbesartan	55	1 (1.8)				
eGFR High and UP High	Double-blind period	Sparsentan	37	1 (2.7) all n<10				NE
		Irbesartan	36	1 (2.8)				
eGFR High and UP Low	Double-blind period	Sparsentan	39	1 (2.6) all n<10				NE
		Irbesartan	37	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024



Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
Safety Set

AESI acute kidney injury - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 1	Double-blind period	Sparsentan						Interaction test: 0.953
< 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	127	6 (4.7)	1.524 [0.440, 5.271]	1.550 [0.427, 5.627]	1.6 [-3.9, 7.2]	0.538
		Irbesartan	129	4 (3.1)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)	1.959 [0.184, 20.900]	2.000 [0.175, 22.815]	2.0 [-6.9, 10.9]	1.000
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)	0.963 + [0.020, 46.760]	0.962 + [0.018, 50.349]	0.0 [NE, NE]	NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
Safety Set

<u>AESI acute kidney injury - non-severe</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline eGFR Group 2	Double-blind period	Sparsentan						Interaction test: NE
< 45 mL/min/1.73 m**2	Double-blind period	Sparsentan	82	3 (3.7)				NE
		Irbesartan	80	3 (3.8)				
45 to < 60 mL/min/1.73 m**2	Double-blind period	Sparsentan	45	3 (6.7)				NE
		Irbesartan	49	1 (2.0)				
60 to < 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	49	2 (4.1)				NE
		Irbesartan	48	1 (2.1)				
≥ 90 mL/min/1.73 m**2	Double-blind period	Sparsentan	26	0 (0.0)				NE
		Irbesartan	25	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High ≥ 60 ml/min/1.73m\*\*2, UP Low = ≤ 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
 Safety Set

<u>AESI acute kidney injury - non-severe</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline urine protein excretion	Double-blind period	Sparsentan						Interaction test: NE
<= 1.75 g/day	Double-blind period	Sparsentan	98	6 (6.1) all n<10				NE
		Irbesartan	93	1 (1.1)				
> 1.75 g/day	Double-blind period	Sparsentan	104	2 (1.9) all n<10				NE
		Irbesartan	109	4 (3.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
Safety Set

<u>AESI acute kidney injury - non-severe</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Baseline use of antihypertensives	Double-blind period	Sparsentan						Interaction test: NE
Yes	Double-blind period	Sparsentan	90	4 (4.4)				NE
		Irbesartan	88	4 (4.5)				
No	Double-blind period	Sparsentan	112	4 (3.6)				NE
		Irbesartan	114	1 (0.9)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
 Safety Set

<u>AESI acute kidney injury - non-severe</u>								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
Time since renal biopsy	Double-blind period	Sparsentan						Interaction test: NE
<= 5 years	Double-blind period	Sparsentan	113	5 (4.4)				NE
		Irbesartan	127	3 (2.4)				
> 5 years	Double-blind period	Sparsentan	89	3 (3.4)				NE
		Irbesartan	75	2 (2.7)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
 N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
 95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
 p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
 RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
 eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
 eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
 Source Data: aae, created on: 30JAN2024

Table PT2AEIN\_SSIM: Incidence of AESI acute kidney injury during double-blind period - non-severe by subgroup  
Safety Set

AESI acute kidney injury - non-severe								
Subgroup	Time	Treatment	N	n (%)	RR [95 % CI]	OR [95 % CI]	RD [95 % CI]	p-value
History of hypertension	Double-blind period	Sparsentan						Interaction test: 0.731
Yes	Double-blind period	Sparsentan	155	7 (4.5)	1.454 [0.472, 4.485]	1.476 [0.458, 4.752]	1.4 [-3.4, 6.3]	0.567
		Irbesartan	161	5 (3.1)				
No	Double-blind period	Sparsentan	47	1 (2.1)	2.625 + [0.110, 62.727]	2.677 + [0.106, 67.539]	2.1 [-4.3, 8.5]	1.000
		Irbesartan	41	0 (0.0)				

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEIC\_SSIM: Incidence of AESI acute kidney injury during double-blind period - severe by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024

Table PT2AEIS\_SSIM: Incidence of AESI acute kidney injury during double-blind period - serious by subgroup  
Safety Set

Not done. Less than 10 patients with events in main analysis.

Notes: Due to different lengths of observation periods, incidences between primary and final analysis are not comparable.  
N = number of patients in analysis set. n = number of patients with event. AESI = adverse event of special interest.  
95% CI = 95% confidence interval (RR/OR: Wald, RD: Newcombe method).  
p-value = exact test of Fisher. \* = significant treatment effect. p-value interaction = Cochran's Q. # = significant interaction.  
RR = relative risk. OR = odds ratio. RD = risk difference. + = adding of 0.5 to each cell. NE = not evaluable.  
eGFR Low = 30 - < 60 ml/min/1.73m\*\*2, eGFR High >= 60 ml/min/1.73m\*\*2, UP Low = <= 1.75 g/day, UP High = > 1.75 g/day.  
eGFR = estimated glomerular filtration rate. UP = urine protein. IgAN = immunoglobulin A nephropathy. BMI = body mass index.  
Source Data: aae, created on: 30JAN2024