

Supplementary Online Content

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This supplementary material has been provided by the authors to give readers additional information about their work.

eAppendix. Definitions, Analyses, and Trial Group Members

Definition of diastolic dysfunction

Diastolic dysfunction (DD) was prospectively identified and graded by an pre-specified algorithm defined in the study protocol with measurements of peak velocities of early (E), late (A) diastolic mitral inflow, peak early (e') tissue Doppler velocity, and peak systolic (S) and diastolic (D) pulmonary venous flow velocity: normal diastolic function ($E/A \geq 1$ and at least two of the following criteria: $E/e' < 10$, $S/D \geq 1$, E/A with Valsalva maneuver ≥ 1), mild DD/ Grade I° ($E/A < 1$), moderate DD/ Grade II° ($1 \leq E/A < 2$ and at least two of the following: $E/e' \geq 10$, $S/D < 1$, E/A Valsalva < 1) and severe DD/ Grade III/IV° ($E/A \geq 2$ and at least one of the following: $E/e' \geq 10$, $S/D < 1$). To further separate severe DD into reversible (Grade III°) and non-reversible (Grade IV°) restrictive filling pattern Valsalva maneuver was needed.¹⁵

Definition of diagnostic criteria of HF with normal EF

Among all included patients the diagnosis criteria of HF with normal EF were analyzed according to current ESC recommendations.¹⁷ Criteria were met (Paulus positive) when E/e' was ≥ 15 , or when $8 < E/e' < 15$ and additional criteria were met (NT-proBNP > 220 pg/mL, LAVI ≥ 40 mL/m², $E/A < 0.5$, E wave deceleration time > 280 msec, A wave duration minus duration of atrial reverse flow of the pulmonary venous flow > 30 ms, LV mass index > 122 g/m² in females and > 149 g/m² in males, or atrial fibrillation).

Sensitivity analysis by imputation for the missing values of the two primary endpoints

In order to analyze how missing values might have affected the effect estimates for the primary endpoints, the data were re-analyzed with several imputations for missing values. Table S4 first displays the effect estimates obtained from the pure non-missing 12-month follow-up data (no imputation).

Secondly, last observation carried forward (LOCF) was applied, inserting the 6-month follow-up values (as far as available) for missing values in the 12-month examination, thus

including cases who had at least one follow-up value but not those who had both follow-up values missing.

Finally, three analyses using multiple imputation (MI) were carried out, including all subjects who were randomized. The imputation was performed as follows. First, two analysis of covariance models were computed from the patients who took part in the final assessment, predicting the 12-month value (1) from the treatment group and the baseline value and (2) from the treatment group, the baseline and the 6-month value. Imputed values for missing 12-month values were then computed from model 1 for subjects who had no follow-up value and from model 2 for those who had the 6-month value (thus using the maximum amount of information available). Each particular imputed value was the sum of the (deterministic) predicted value for the individual subject and a randomly chosen residual of the respective model.

The first MI analysis assumed that missing values were independent of the prognosis for the respective endpoint (“non-informative MI”). The predicted values were computed using the randomized treatment, and the random components were selected from the set of all residuals.

The second MI analysis assumed that subjects who had no 12-month measurement might have skipped the study drug and thus, such patients performed like those in the placebo group (“placebo MI”). Predicted values were computed like for subjects in the placebo group, regardless of the randomized treatment, and a random selection from the set of all residuals was added.

The third MI analysis assumed that missing values in the spironolactone group might have been associated with poor performance, while no such relationship was assumed in the placebo group (“pessimistic MI”). Again, predicted values were computed like for subjects in the placebo group, regardless of the randomized treatment, and the random component was selected from the worst quartile of residuals in the spironolactone group, and from the set of all residuals in the placebo group.

Ten sets of imputed random values were generated in each of the three MI analyses. Computations were performed as outlined in Schafer JL, Multiple imputation: a primer, *Stat Methods Med Res* 1999; 8: 3-15.

Committees and Collaborators (AT: Austria; GE: Germany)

Principal Investigator: Burkert Pieske, Graz (AT).

Study Coordinator: Frank Edelmann, Göttingen (GE).

Steering Committee: Burkert Pieske (Graz, AT); Frank Edelmann (Göttingen, GE); Rolf Wachter (Göttingen, GE); Gerd Hasenfuss (Göttingen, GE); Christiane E Angermann (Würzburg, GE); Michael Böhm (Homburg/Saar, GE); Markus Löffler (Leipzig, GE)

Data Safety Monitoring Board: Stefan D Anker (Berlin, GE); Vera Regitz-Zagrosek (Berlin, GE); Hanjörg Just (Freiburg, GE).

Biometry, Data Collection and Management, Monitoring: Götz Gelbrich (Würzburg/ Leipzig, GE); Christiane Prettin (Leipzig, GE).

Sponsor: Georg-August-University Göttingen, GE.

Reference Centers: Martin Halle (Munich, GE) for Cardiopulmonary exercise testing; Frank Weidemann (Würzburg, GE) and Albrecht Schmidt (Graz, AT) for Echocardiography; Michael Oellerich and Lutz Binder (Göttingen, GE) for Laboratory Analyses; Christoph Herrmann-Lingen (Göttingen, GE) for Psychometric assessments.

Scientific review and editorial contributions to the manuscript: Wendy Gattis Stough, PharmD (Cary, North Carolina)

Trial Sites: Frank Edelmann, Rolf Wachter, Kathleen Durstewitz, André Duvinage, & Raoul Stahrenberg, Göttingen; Burkert Pieske, Caterina Colantonio, Albrecht Schmidt, Elisabeth Kraigher-Krainer, Stephanie Walther & Sascha Pätzold (Graz, AT); Wolfram Kamke, Burg; Hans-Dirk Düngen & Simone Inkrot, Berlin; Carsten Tschöpe & Dirk Westermann, Berlin. Stephan Gielen & Marco Sandri, Leipzig; Rüdiger Braun-Dullaeus & Alexander Schmeißer, Magdeburg; Christiane E Angermann & Frank Weidemann, Würzburg; Wolfgang von Scheidt, Augsburg; Till Neumann, Essen.

Collaborators: Regine Gebauer, Jürgen Wolf, Silja Schwarz, Otto Zelger, Melanie Wätzold, Friederike Polzin, Manuela Knoke, Inke Bartels, Lars Platschek, Inga Roesler, Wiebke Aufderheide, Anneke Behrens, Caroline Pasedach, Annkatrin Pieper, Ingmar Rahn, Fabian Gabriel, Ines Unkelbach, Hannes Fricke, Lutz Binder, Sabine Dreyer, Doris von Grünhagen, Jeanette Kühn, Volker Holzendorf, Meinhard Mende, Birgit Binder, Birgit Saumer, Cornelia Kos.

eTable 1. Baseline Characteristics*

Variable	All patients	Placebo	Spironolactone
Number of subjects	422	209	213
History - no. (%)			
Smoking			
Never smoked	223 (53)	108 (52)	115 (54)
Former smoker	172 (41)	88 (42)	84 (39)
Current smoker	27 (6)	13 (6)	14 (7)
Previous myocardial infarction	67 (16)	29 (14)	38 (18)
Previous coronary artery bypass graft surgery	31 (7)	16 (8)	15 (7)
Stroke or TIA	45 (11)	22 (11)	23 (11)
Peripheral arterial disease	17 (4)	10 (5)	7 (3)
History of depression	47 (11)	25 (12)	22 (10)
Physical examination			
Mean arterial pressure- mmHg	98±12	98±12	98±11
Puls pressure - mmHg	56±15	56±16	56±15
Laboratory			
Total cholesterol - mg/dL	195±44	196±45	195±44
Uric acid - mg/dL	6.1±1.6	6.2±1.5	6.1±1.7
Echocardiography			
IV septum thickness – mm	12.2±1.8	12.2±2.0	12.1±1.7
Posterior wall thickness - mm	11.5±1.5	11.5±1.5	11.6±1.9
TEI index	0.44±0.23	0.43±0.18	0.44±0.25
Left ventricular volume (end-diastolic) – mL	77±29	77±28	77±30
Left ventricular volume (end-systolic) – mL	27±13	26±12	27±14
Left atrial area, 4-CH – cm ²	22.0±4.6	22.1±4.5	21.9±4.8
Left atrial area, 2-CH – cm ²	22.0±4.5	22.2±4.2	21.9±4.8
LA _{ES} – mm	44.2±5.7	44.3±5.4	44.1±6.0
A wave velocity - cm/s	83±18	83±18	83±18
A wave duration – ms	154±32	156±33	152±32
Medial a' wave velocity - cm/s	9.3±1.8	9.3±1.9	9.2±1.8
PVF systolic - cm/s	58±12	58±12	58±12
PVF diastolic - cm/s	46±14	47±15	46±13

eTable 1. Baseline Characteristics* (continued)

Variable	All patients	Placebo	Spironolactone
PVA velocity - cm/s	33±11	33±11	33±11
PVA duration – ms	125±30	122±31	127±30
Flow propagation time - cm/s	31±9	31±9	32±9
Maximal workload - W	100±29	101±29	99±29
BP systolic at rest - mmHg	122±18	123±18	121±18
BP systolic at maximal stress - mmHg	169±28	171±30	167±27
BP diastolic at rest – mmHg	79±12	80±12	79±12
BP diastolic at maximal stress - mmHg	85±18	86±18	85±17
Heart rate at rest – bpm	70±13	68±12	71±13
Heart rate at maximal stress - bpm	117±21	116±22	117±20
V _E at rest - L/min	8.4±2.4	8.3±2.4	8.4±2.4
V _E at maximal stress - L/min	45.7±12.9	46.5±13.4	44.9±12.4
Anaerobic threshold – W	64±25	63±23	65±27
RER at rest	0.84±0.07	0.84±0.07	0.84±0.08
RER _{max}	1.11±0.12	1.12±0.12	1.10±0.12
RER _{max} post exercise	1.38±0.19	1.39±0.18	1.37±0.19
Six-minute walk test			
Terminated before 6 min – n (%)	8 (2)	4 (2)	4 (2)
Borg scale	3.1±1.8	3.2±2.0	3.0±1.5
Current medication – no. (%)			
Anti-platelet agent	221 (52)	108 (52)	113 (53)
Anticoagulant	58 (14)	32 (15)	26 (12)
Lipid lowering drug	230 (55)	118 (56)	112 (53)
Allopurinol	40 (10)	21 (10)	19 (9)
Antidepressant	30 (7)	13 (6)	17 (8)

*Plus - minus values are means ± SD; TIA, transient ischemic attack; IV, interventricular; BMI, body-mass index (weight in kilograms divided by the square of the height in meters); LA_{ES}, left atrial endsystolic; A, peak atrial transmitral ventricular filling velocity; E/A, peak early transmitral ventricular filling velocity to peak atrial transmitral ventricular filling velocity ratio; IVRT, isovolumetric relaxation time; a', atrial diastolic tissue Doppler velocity; PVF, pulmonary venous flow; PVA, pulmonary venous atrial reverse flow; BP, blood pressure; V_E, minute ventilation; RER, respiratory exchange ratio.

Higher values indicate better performance: Flow propagation time; Maximal workload; V_E; Anaerobic threshold

Lower values indicate better performance: Puls pressure; Total cholesterol; Uric acid; LA_{ES}; PVA velocity; PVA duration; Borg Scale

eTable 2. Echocardiography, Cardiopulmonary Exercise Testing, and Clinical and Laboratory Results After 12 Months

Echocardiography^a				
Variable	Placebo	Spirolactone	Difference Spirolactone–Placebo	
	(n=195)	(n=203)	Difference (95% CI)	P-value
IV septum thickness - mm	12.1 (11.8-12.4)	11.9 (11.9-12.1)	-0.2 (-0.4 to +0.1)	0.21
Posterior wall thickness - mm	11.4 (11.2-11.6)	11.3 (11.1-11.5)	-0.1 (-0.4 to +0.1)	0.29
TEI index	0.44 (0.40 to 0.47)	0.44 (0.41 to 0.48)	0.00 (-0.05 to +0.04)	0.84
Left ventricular volume (end-diastolic) – mL	79 (74 to 83)	73 (69 to 77)	-5 (-10 to 0)	0.06
Left ventricular volume (end-systolic) – mL	29 (27 to 32)	25 (23 to 27)	-4 (-7 to -1)	0.005
Left atrial area, 4-CH – cm ²	22.2 (21.5 to 22.8)	21.8 (21.0 to 22.5)	-0.2 (-0.9 to +0.5)	0.60
Left atrial area, 2-CH – cm ²	22.2 (21.6 to 22.8)	21.7 (21.0 to 22.4)	-0.3 (-1.0 to +0.5)	0.49
LA _{ES} - mm	43.9 (43.1-44.6)	43.2 (42.4-44.0)	-0.4 (-1.1 to +0.2)	0.17
A wave velocity - cm/s	82 (79-85)	80 (78-83)	-2 (-4 to +1)	0.25
A wave duration - ms	156 (152-161)	153 (148-158)	-2 (-9 to +4)	0.52
Medial a' wave velocity - cm/s	9.2 (8.9-9.5)	9.3 (9.0-9.5)	+0.2 (-0.1 to +0.5)	0.23
PVF systolic - cm/s	59 (57-61)	58 (57-60)	-1 (-3 to +2)	0.62
PVF diastolic - cm/s	47 (45-49)	46 (44-48)	0 (-3 to +2)	0.85
PVA velocity - cm/s	33 (32-35)	34 (32-36)	+1 (-2 to +3)	0.48
PVA duration - ms	123 (118-127)	123 (119-127)	-2 (-8 to +4)	0.45
Flow propagation time - cm/s	31 (30-32)	32 (30-33)	0 (-2 to +2)	0.74

eTable 2. Echocardiography, Cardiopulmonary Exercise Testing, and Clinical and Laboratory Results After 12 Months (continued)

Cardiopulmonary Exercise Testing^b				
Variable	Placebo	Spironolactone	Difference Spironolactone – Placebo	
	(n=187)	(n=187)	Difference (95% CI)	P-value
Maximal workload – W	101 (97-106)	100 (95-105)	+2 (–2 to +5)	0.35
BP systolic at rest – mmHg	122 (120-124)	116 (114-118)	–5 (–8 to –2)	<0.001
BP systolic at maximal stress - mmHg	171 (167-175)	166 (162-170)	–3 (–8 to +2)	0.20
BP diastolic at rest – mmHg	76 (75-78)	74 (73-76)	–2 (–4 to 0)	0.09
BP diastolic at maximal stress - mmHg	84 (82-87)	82 (80-85)	–2 (–5 to +1)	0.20
Heart rate at rest – bpm	69±13	71±12	0 (–2 to +2)	0.80
Heart rate at maximal stress - bpm	118 (115-122)	120 (117-124)	+1 (–3 to +4)	0.64
V _E at rest - L/min	9.1 (8.7-9.4)	8.8 (8.4-9.2)	–0.2 (–0.7 to +0.2)	0.27
V _E at maximal stress - L/min	50.1 (47.8-52.4)	48.7 (46.5-50.9)	+0.6 (–1.7 to +3.0)	0.58
Anaerobic threshold – W	66 (62-69)	65 (61-69)	–1 (–5 to +4)	0.79
RER at rest	0.84 (0.82-0.85)	0.84 (0.82-0.85)	0 (–0.02 to +0.02)	0.97
RER _{max}	1.13 (1.11-1.15)	1.12 (1.10-1.14)	–0.01 (–0.03 to +0.02)	0.60
RER _{max} post exercise	1.38 (1.35-1.41)	1.38 (1.36-1.41)	+0.01 (–0.02 to +0.05)	0.52
Clinical/Laboratory^c				
Variable	Placebo	Spironolactone	Difference Spironolactone–Placebo	
	(n=196)	(n=204)	Difference (95% CI)	P-value
<i>Clinical variables</i>				
Mean arterial pressure [mmHg] – mean±SD	99 (98-101)	94 (92-95)	–5 (–7 to –3)	<0.001

eTable 2. Echocardiography, Cardiopulmonary Exercise Testing, and Clinical and Laboratory Results After 12 Months (continued)

Variable	Placebo	Spironolactone	Difference Spironolactone–Placebo	
	(n=196)	(n=204)	Difference (95% CI)	P-value
<i>Clinical variables (continued)</i>				
Pulse pressure [mmHg] – mean±SD	56 (54-58)	52 (50-54)	–5 (–7 to –2)	<0.001
Heart rate on ECG – mean±SD	65 (63-66)	66 (65-68)	+1 (–1 to +3)	0.56
<i>Laboratory variables</i>				
Total cholesterol [mg/dL] – mean±SD	197 (191-202)	196 (190-202)	–1 (–7 to +5)	0.85
Uric acid [mg/dL] – mean±SD	6.1 (5.9-6.3)	6.2 (6.0-6.5)	+0.2 (0 to +0.4)	0.10

^aQuantitative variables are presented by group-wise means and 95% CI. Between-group differences are from ANCOVA, adjusting for baseline. IV, interventricular; LA_{ES}, left atrial endsystolic; A, peak atrial transmitral ventricular filling velocity; E/A, peak early transmitral ventricular filling velocity to peak atrial transmitral ventricular filling velocity ratio; IVRT, isovolumetric relaxation time; a', atrial diastolic tissue Doppler velocity; PVF, pulmonary venous flow; PVA, means pulmonary venous atrial reverse flow. Higher values indicate better performance: A wave duration; Flow propagation time

Lower values indicate better performance: TEI index; Left atrial area; LA_{ES}; PVA velocity; PVA duration

^bQuantitative variables are presented by group-wise means and 95% CI (exception: NT-proBNP is described by median and inter-quartile range). Between-group differences are from ANCOVA, adjusting for baseline. BP, blood pressure; V_E, minute ventilation; RER, respiratory exchange ratio. Higher values indicate better performance: Maximal workload; V_E; Anaerobic threshold

^cQuantitative variables are presented by group-wise means and 95% CI. Between-group differences are from ANCOVA, adjusting for baseline. Lower values indicate better performance: Puls pressure; Total Cholesterol; Uric acid

eTable 3. Effect of Spironolactone Treatment on E/e' and peak VO₂ in Subgroups

		E/e'^a	
Subgroups	N in subgroup	Treatment effect (95% CI) in subgroups	Interaction P-value
Subgroup #1	Placebo / Spironolactone		
Subgroup #2	Placebo / Spironolactone		
Age			
<67 years	84 / 93	-1.73 (-2.57 to -0.89)	0.42
≥67 years	111 / 110	-1.27 (-2.02 to -0.52)	
Gender			
Men	95 / 93	-1.42 (-2.24 to -0.61)	0.82
Women	100 / 110	-1.55 (-2.32 to -0.78)	
Body mass index			
<29 kg/m ²	96 / 99	-1.08 (-1.88 to -0.28)	0.17
≥29 kg/m ²	99 / 104	-1.87 (-2.65 to -1.08)	
Systolic blood pressure			
<134 mmHg	87 / 108	-1.41 (-2.21 to -0.60)	0.84
≥134 mmHg	108 / 95	-1.53 (-2.31 to -0.74)	
Heart rate			
<63 bpm	92 / 84	-1.43 (-2.27 to -0.58)	0.95
≥63 bpm	103 / 119	-1.46 (-2.21 to -0.71)	
NYHA class			
II	170 / 171	-1.48 (-2.09 to -0.88)	0.91
III	25 / 32	-1.39 (-2.89 to -0.10)	
Grade of diastolic dysfunction			
I°	139 / 148	-1.54 (-2.20 to -0.88)	0.78
II-III° or atrial fibrillation	56 / 55	-1.37 (-2.42 to -0.31)	
Paulus criteria			
Negative	92 / 97	-1.72 (-2.54 to -0.91)	0.41
Positive	103 / 106	-1.25 (-2.03 to -0.48)	
Estimated glomerular filtration rate			
<73 mL/min/1.73m ²	92 / 104	-1.53 (-2.33 to -0.73)	0.88
≥73 mL/min/1.73m ²	103 / 99	-1.44 (-2.23 to -0.66)	

eTable 3. Effect of Spironolactone Treatment on E/e' and peak VO₂ in Subgroups (continued)

Subgroups	N in subgroup	Peak VO ₂ ^b	
		Treatment effect (95% CI) in subgroups [mL/min/kg]	Interaction P-value
Subgroup #1	Placebo / Spironolactone		
Subgroup #2	Placebo / Spironolactone		
Age			
<67 years	80 / 86	-0.02 (-1.08 to +1.04)	0.82
≥67 years	107 / 101	+0.15 (-0.80 to +1.10)	
Gender			
Men	93 / 88	-0.32 (-1.33 to +0.70)	0.26
Women	94 / 99	+0.49 (-0.49 to +1.47)	
Body mass index			
<29 kg/m ²	93 / 92	-0.10 (-1.10 to +0.91)	0.62
≥29 kg/m ²	94 / 95	+0.26 (-0.73 to +1.25)	
Systolic blood pressure			
<134 mmHg	82 / 97	-0.24 (-1.26 to +0.78)	0.49
≥134 mmHg	105 / 90	+0.25 (-0.72 to +1.23)	
Heart rate			
<63 bpm	90 / 81	+0.74 (-0.30 to +1.79)	0.10
≥63 bpm	97 / 106	-0.44 (-1.39 to +0.52)	
NYHA class			
II	163 / 157	+0.26 (-0.50 to +1.01)	0.36
III	24 / 30	-0.69 (-2.55 to +1.17)	
Grade of diastolic dysfunction			
I°	132 / 134	+0.08 (-0.76 to +0.91)	0.96
II-III° or atrial fibrillation	55 / 53	+0.12 (-1.20 to +1.43)	
Paulus criteria			
Negative	88 / 89	-0.16 (-1.19 to +0.86)	0.52
Positive	99 / 98	+0.30 (-0.67 to +1.28)	
Estimated glomerular filtration rate			
<73 mL/min/1.73m ²	88 / 95	+0.43 (-0.58 to +1.44)	0.36
≥73 mL/min/1.73m ²	98 / 92	-0.23 (-1.22 to +0.76)	

^aThe treatment effect is the mean difference in E/e' at 12-month follow-up between the spironolactone and the placebo arm in the respective subgroups, adjusted for baseline. The interaction P-value is for the test of the null hypothesis that the treatment effects in the respective two subgroups is the same. Treatment effects and interaction P-values were computed from analysis of covariance.

^bThe treatment effect is the mean difference in peak VO₂ at 12-month follow-up between the spironolactone and the placebo arm in the respective subgroups, adjusted for baseline. The interaction P-value is for the test of the null hypothesis that the treatment effects in the respective two subgroups is the same. Treatment effects and interaction P-values were computed from analysis of covariance.

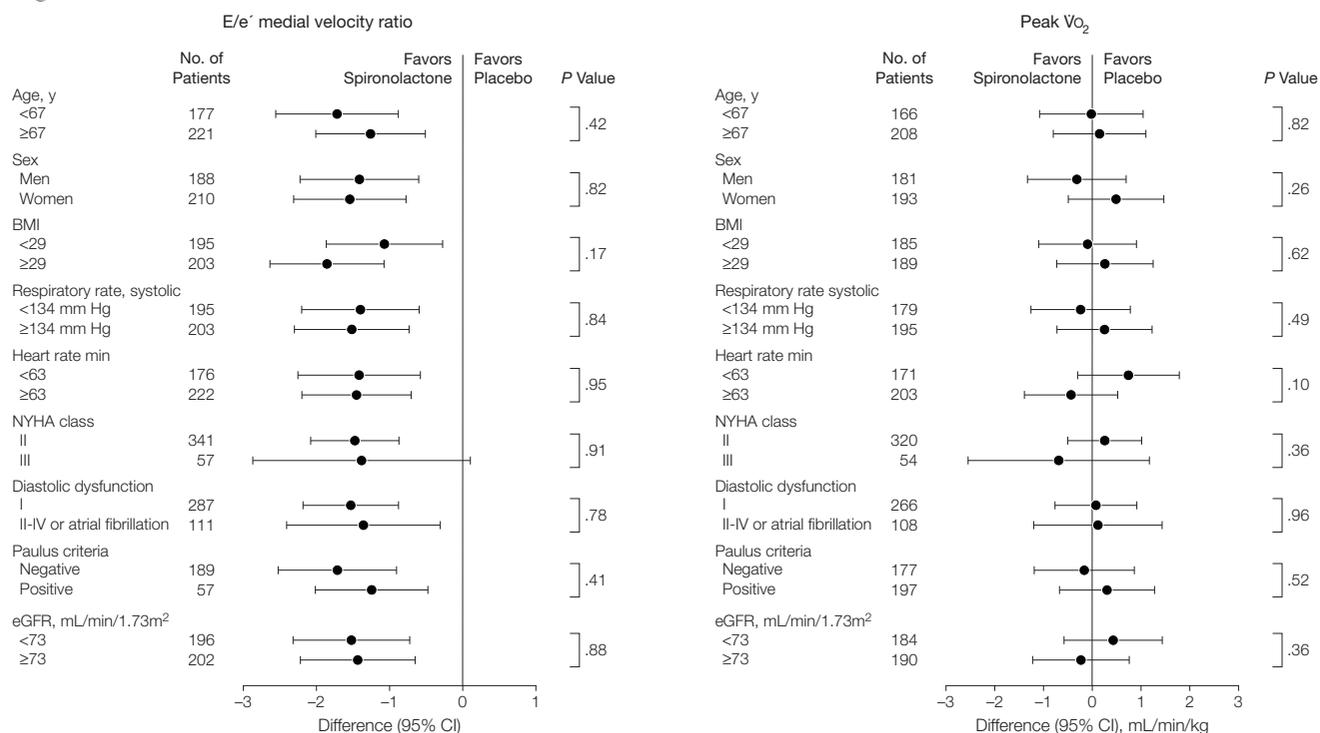
eTable 4. Estimates for the Treatment Effect on the Primary Endpoints When Different Imputation Methods for Missing Values are Applied

Endpoint	Imputation	Effect estimate (95% CI)	P-value
E/e'	None	-1.48 (-2.04 to -0.92)	<0.001
	LOCF	-1.53 (-2.09 to -0.96)	<0.001
	Non-informative MI	-1.53 (-2.08 to -0.98)	<0.001
	Placebo MI	-1.47 (-2.02 to -0.92)	<0.001
	Pessimistic MI	-1.30 (-1.86 to -0.74)	<0.001
Peak VO ₂ [mL/min/kg]	None	+0.08 (-0.62 to +0.79)	0.81
	LOCF	-0.02 (-0.70 to +0.67)	0.96
	Non-informative MI	+0.06 (-0.63 to +0.75)	0.86
	Placebo MI	+0.04 (-0.65 to +0.73)	0.91
	Pessimistic MI	-0.46 (-1.15 to +0.24)	0.20

LOCF- last observation carried forward; MI- multiple imputation

According to these results, the effect estimates are not materially changed when interim follow-up observations were carried forward or MI was applied, assuming that missing data were not related with the outcome or were associated with the absence of an intervention effect. In addition, a substantial effect remained in E/e' and no significant effect occurred in peak VO₂ when MI was applied with the pessimistic assumption that missing data were associated with unfavorable outcome in the spironolactone group but not in the placebo group. We conclude that missing values had no material impact on the primary results of this study.

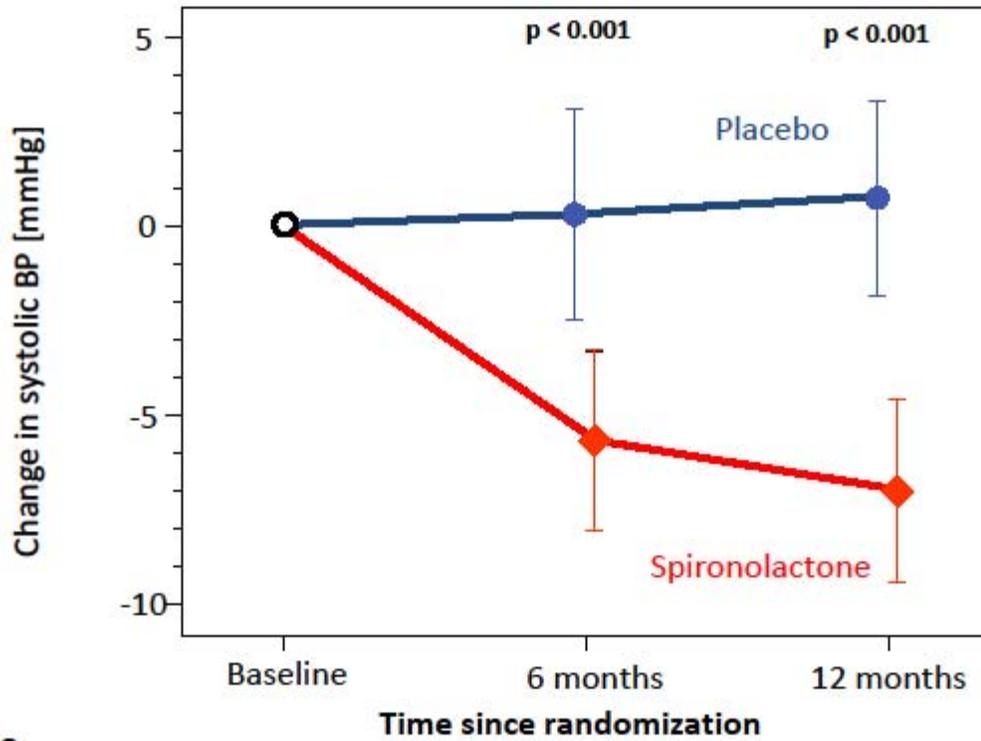
eFigure 1. Predefined Subgroup Analysis of Ratio of Peak Early Transmitral Ventricular Filling Velocity to Early Diastolic Tissue Doppler Velocity (E/e') and Peak Oxygen Consumption (Peak VO_2) According to Assigned Study Treatment



Differences are given in absolute values with 95% CIs for the spironolactone group compared with the placebo group. Body mass index (BMI) is defined as weight in kilograms divided by height in meters squared. eGFR indicates estimated glomerular filtration rate; NYHA, New York Heart Association; Paulus positive/negative, fulfillment/nonfulfillment of criteria for diagnosis of heart failure with preserved ejection fraction proposed by Paulus et al.¹⁷ All values and subgroup numbers shown in Figure 3 are provided in eTable 3A (E/e') and eTable 3B (peak O_2).

eFigure 2. Main Clinical Endpoint, According to the Assigned Study Treatment

Shown are the changes after 6 and 12 months in systolic blood pressure. The bars represent the analysis of covariance estimates of treatment effects within subgroups. P-values describe comparisons of the changes of placebo or spironolactone at the respective time-point vs. Baseline. All values are means (95 % confidence interval).



Placebo

Number of patients

Systolic BP (mmHg)

Spironolactone

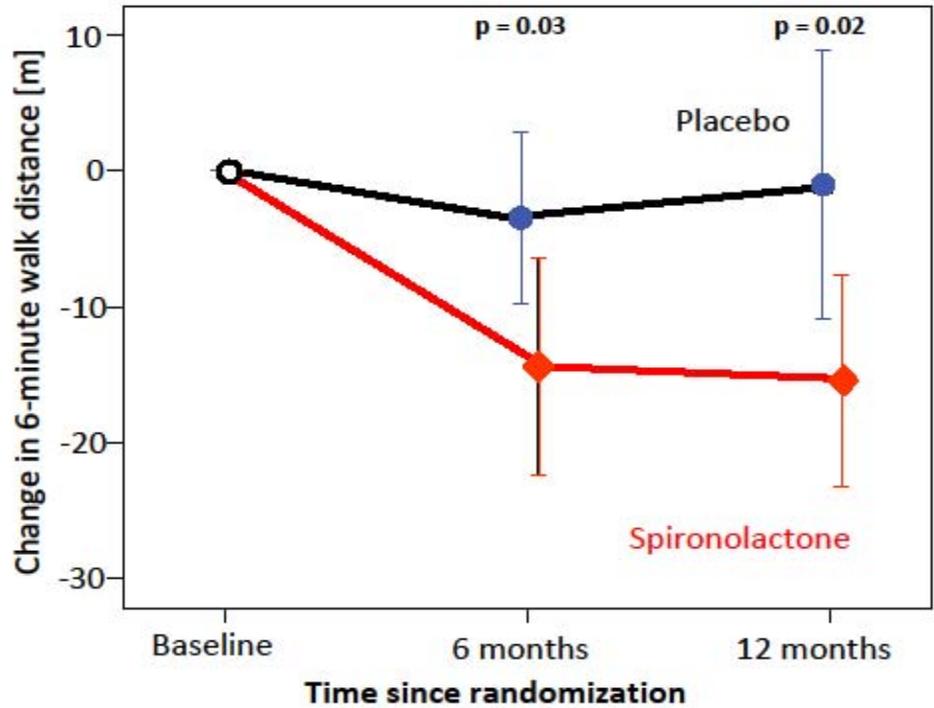
Number of patients

Systolic BP (mmHg)

	Baseline	6 months	12 months
Number of patients	209	198	195
Systolic BP (mmHg)	135 (133-138)	136 (134-139)	137 (135-139)
Number of patients	213	206	202
Systolic BP (mmHg)	135 (132-137)	129 (127-131)	128 (126-130)

eFigure 3. Main Secondary Endpoint, According to the Assigned Study Treatment

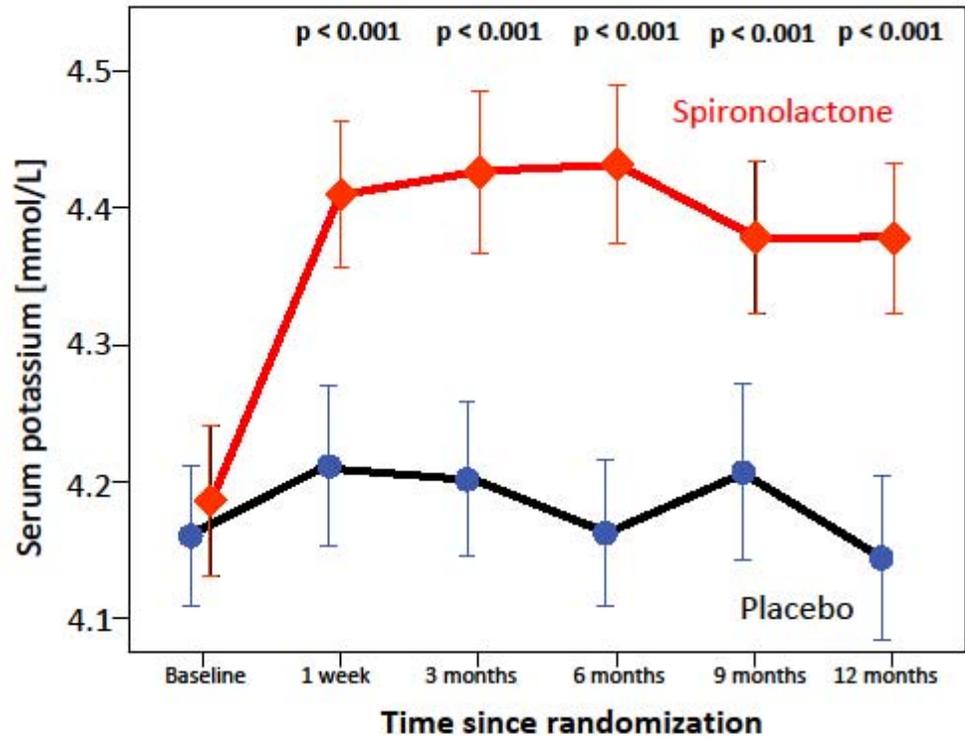
Shown are the changes after 6 and 12 months in 6-minute-walking distance. The bars represent the analysis of covariance estimates of treatment effects within subgroups. P-values describe comparisons of the changes of placebo or spironolactone at the respective time-point vs. Baseline. All values are means (95 % confidence interval).



	Baseline	6 months	12 months
Placebo			
Number of patients	207	189	185
Six-minute walk distance (m)	531 (520-543)	530 (517-543)	536 (521-550)
Spironolactone			
Number of patients	213	199	186
Six-minute walk distance (m)	529 (517-541)	517 (503-531)	517 (504-531)

eFigure 4. Main Safety Endpoint, According to the Assigned Study Treatment

Shown are the potassium values at baseline, after 1 week, and after 3, 6, 9 and 12 months. The bars represent the analysis of covariance estimates of treatment effects within subgroups. P-Values describe comparisons between placebo and spironolactone at the respective time-point, and the bars denote the 95 % confidence interval. All values are means (95 % confidence interval).



Placebo

Number of patients
Potassium (mmol/L)

209	206	202	197	190	195
4.16	4.21	4.20	4.16	4.21	4.14
(4.11-4.21)	(4.15-4.27)	(4.15-4.26)	(4.11-4.22)	(4.14-4.27)	(4.08-4.20)

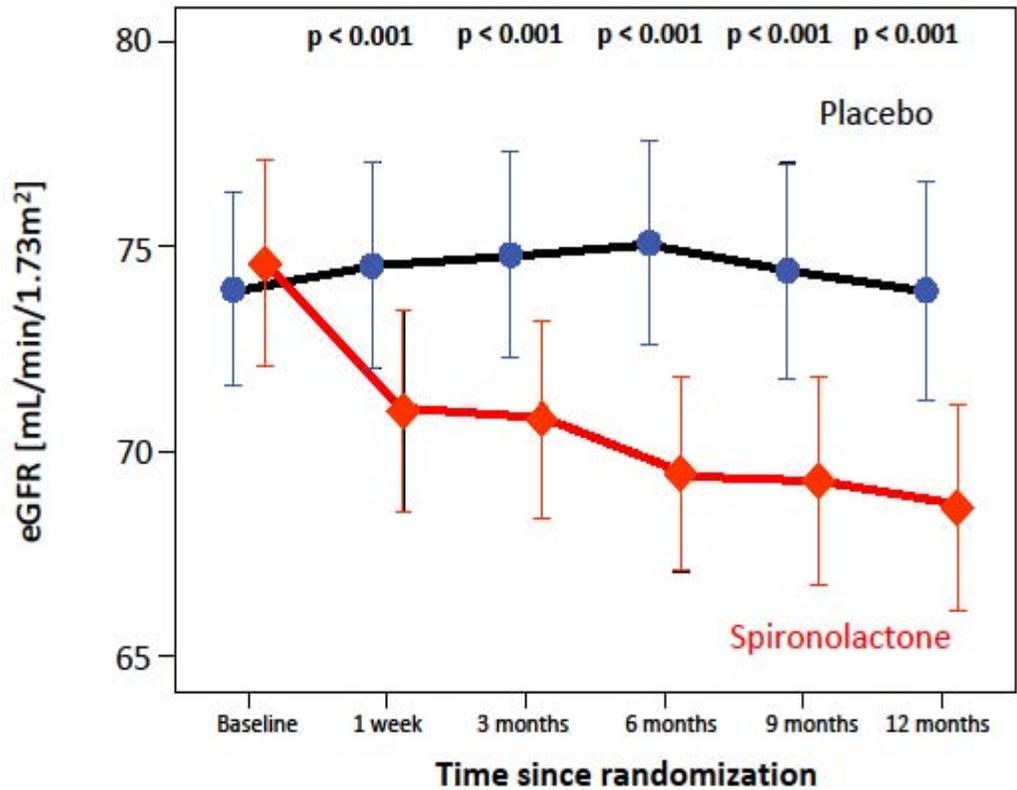
Spironolactone

Number of patients
Potassium (mmol/L)

211	210	207	204	195	203
4.19	4.41	4.43	4.43	4.38	4.38
(4.13-4.24)	(4.36-4.46)	(4.37-4.49)	(4.37-4.49)	(4.32-4.43)	(4.32-4.43)

eFigure 5. Main Safety Endpoint, According to the Assigned Study Treatment

Shown are the estimated glomerular filtration rate values at baseline, after 1 week, and after 3, 6, 9 and 12 months. The bars represent the analysis of covariance estimates of treatment effects within subgroups. P-Values describe comparisons between placebo and spironolactone at the respective time-point, and the bars denote the 95 % confidence interval. All values are means (95 % confidence interval).



Placebo

Number of patients
eGFR (mL/min/1.73 m²)

206	205	202	197	190	194
74 (72-76)	75 (72-77)	75 (72-77)	75 (73-78)	74 (72-77)	74 (71-77)

Spironolactone

Number of patients
eGFR (mL/min/1.73 m²)

210	210	207	204	195	203
75 (72-77)	71 (69-73)	71 (68-73)	69 (67-72)	69 (67-72)	69 (66-71)