- 1 Supplemental Appendix
- 3 The role and prognostic value of growth differentiation factor 15 in patient of
- 4 heart failure with preserved ejection fraction: Insights from the PURSUIT-HFpEF
- 5 Registry

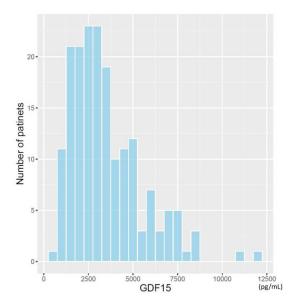
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- 7 Supplemental Figure
- **8** Supplemental Table
- 9 Collaborators list

#### 1 **Supplemental Figure 1. Distribution of GDF15**



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- 6 The distribution of GDF15 is shown using a histogram. In the histogram, the x-axis
- 7 shows GDF15 level, and the y-axis shows the number of patients. Abbreviations:
- 8 GDF15, growth differentiation factor 15.

## 1 Supplemental Table 1. Comparison of background of patients included and those

# 2 not included for analysis of GDF15

Characteristic	Patients included	Patients not included	P value
	(N = 181)	(N = 1050)	
Age (years)	81.0 [75.0, 85.0]	83.0 [77.0, 87.0]	<0.01
Male sex	86 (47.5)	465 (44.3)	0.47
Body mass index (kg/m <sup>2</sup> )	21.5 [19.3, 24.3]	21.4 [18.9, 24.3]	0.71
NYHA≥II	116 (64.1)	658 (64.0)	>0.99
Clinical frailty scale $\geq 5$	32 (17.7)	348 (33.2)	<0.01
Smoking history	75 (41.7)	385 (37.3)	0.30
Systolic blood pressure	126.0 [112.0, 136.0]	117.0 [105.0, 129.0]	<0.01
Heart rate	70.0 [60.0, 76.0]	70.0 [61.0, 80.0]	0.39
Past medical history			
Atrial fibrillation	70 (38.7)	561 (53.4)	<0.01
Coronary artery disease	35 (19.6)	177 (17.1)	0.48
Hypertension	162 (89.5)	877 (83.8)	0.07
Diabetes mellitus	67 (37.2)	336 (32.2)	0.22

Dyslipidemia	87 (48.3)	436 (41.8)	0.12
COPD	18 (10.1)	70 (7.0)	0.20
Dialysis	6 (3.3)	12 (1.1)	0.06
Medication at discharge			
ACE-Is/ARBs	90 (49.7)	562 (53.9)	0.34
β blockers	128 (70.7)	541 (51.9)	<0.01
Diuretics	155 (85.6)	840 (80.5)	0.13
MRAs	70 (38.7)	415 (39.8)	0.84
SGLT2 inhibitors	18 (9.9)	63 (6.1)	0.07
Biguanides	7 (3.9)	31 (3.0)	0.67
Laboratory data			
Albumin (g/dL)	3.3 [3.1, 3.6]	3.4 [3.1, 3.7]	0.27
Hemoglobin (g/dL)	11.6 [10.0, 13.1]	11.2 [10.1, 12.6]	0.39
Sodium (mEq/L)	140.0 [138.0, 141.0]	139.0 [137.0, 141.0]	0.27
eGFR (mL/min/1.73m <sup>2</sup> )	43.9 [28.1, 58.4]	40.9 [29.5, 53.7]	0.51
Cholinesterase (U/L)	208.0 [172.0, 255.0]	207.0 [165.0, 254.0]	0.28
C-reactive protein (mg/dL)	0.6 [0.2, 1.6]	0.2 [0.1, 0.8]	<0.01
Hemoglobin A1c	6.0 [5.6, 6.5]	6.0 [5.6, 6.5]	0.70

LDL cholesterol	92.0 [73.0, 113.5]	91.0 [73.0, 109.0]	0.38		
NT-proBNP (pg/mL)	1280.0 [529.2, 2995.0]	1058.5 [492.0, 2370.8]	0.17		
Echo cardiography at discharge					
LVDd (mm)	46.4 [42.0, 51.2]	45.0 [41.0, 49.9]	0.04		
LVEF (%)	61.0 [55.0, 65.0]	61.0 [55.1, 65.6]	0.60		
LAVI (mL/m <sup>2</sup> )	50.0 [37.0, 64.8]	51.0 [38.0, 67.0]	0.42		
LVMI (g/m <sup>2</sup> )	117.9 [98.9, 146.3]	99.5 [82.3, 119.6]	<0.01		
TAPSE (mm)	17.4 [15.1, 20.4]	17.3 [14.6, 20.2]	0.57		
Mean E/e'	12.7 [10.0, 17.2]	12.4 [9.7, 16.7]	0.48		
TRPG (mmHg)	25.9 [20.4, 32.0]	27.0 [22.0, 33.0]	0.02		
Nutritional status					
CONUT score	3.0 [2.0, 5.0]	3.0 [2.0, 5.0]	0.57		
GNRI	91.2 [84.5, 98.3]	92.0 [84.5, 99.2]	0.67		
PNI	40.6 [37.4, 44.4]	41.1 [37.2, 44.8]	0.36		

<sup>2</sup> This comparison summarizes the difference between patients included and those not

<sup>3</sup> included. Data are shown as median [interquartile range] or number (percentage).

<sup>4</sup> Abbreviations: GDF15, Growth differentiation factor 15; NYHA, New York Heart

1 Association; COPD, chronic obstructive pulmonary disease; ACE-Is, 2 angiotensin-converting enzyme inhibitors; ARBs, angiotensin II receptor blockers; 3 MRAs, mineralocorticoid receptor antagonists; SGLT2 inhibitors, sodium glucose 4 cotransporter2 inhibitors; eGFR, estimated glomerular filtration rate; LDL, low density 5 lipoprotein; NT-proBNP, N-terminal pro-brain natriuretic peptide; LVDd, left 6 ventricular diastolic diameter; LVEF, left ventricular ejection fraction; LAVI, left atrial 7 volume index; LVMI, left ventricular mass index; TAPSE, tricuspid annular plane 8 systolic excursion; E, early diastolic velocity on transmitral doppler; e', early diastolic 9 velocity of the mitral valve annulus; TRPG, tricuspid regurgitation pressure gradient; 10 CONUT, controlling nutritional status; GNRI, geriatric nutritional risk index; PNI, 11 prognostic nutritional index.

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