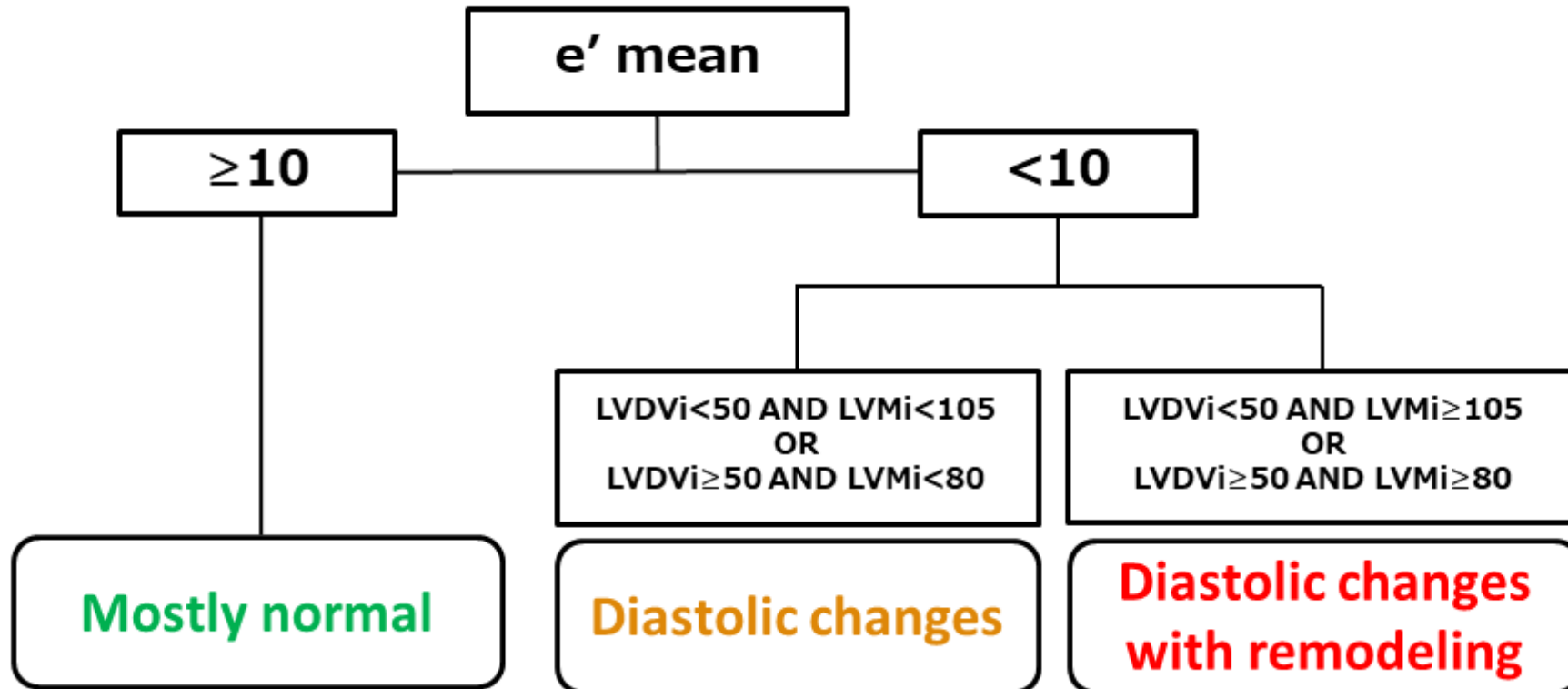


Supplemental Figure 1. e'VM algorithm



Supplemental Table 1. Age-specific limits for e' mean, LVEDVi and LVMi

	e' mean (cm/sec)	LVEDVi (ml/m²)	LVMi (g/m²)	LVMi2 (g/m²)
40 years	12	56	72	97
50 years	11	53	76	101
60 years (reference)	10	50	80	105
70 years	9	47	84	109
80 years	8	44	88	113

Reference limits at 60 years of age were defined by original thresholds of the e'VM algorithm, while age-specific cut-off values were determined according to previous reports^{16, 17}.

Supplemental Table 2. Change in echocardiographic and biomarker variables in response to spironolactone across the e'VM phenotype strata

	MN phenotype				D phenotype				D/S phenotype			
	Spironolactone	placebo	mean \pm SD	adj-p	Spironolactone	placebo	mean \pm SD	adj-p	Spironolactone	placebo	mean \pm SD	adj-p
LVEDVi change, ml/m²	-0.6 \pm 5.5	-1.0 \pm 5.2	0.4 \pm 1.5	0.87	-1.2 \pm 4.5	-1.0 \pm 4.4	-0.2 \pm 0.6	0.95	-1.8 \pm 5.2	-1.6 \pm 5.3	-0.4 \pm 0.9	0.71
LVMi change, g/m²	-0.7 \pm 15.4	0.1 \pm 9.6	-0.4 \pm 3.4	0.65	-0.5 \pm 8.9	0.7 \pm 10.4	-1.2 \pm 1.4	0.32	-6.7 \pm 13.9	-0.9 \pm 14.7	-6.4 \pm 2.7	0.012
LAVi change, ml/m²	-3.3 \pm 8.4	-1.2 \pm 4.6	-1.8 \pm 1.6	0.29	-1.0 \pm 5.0	-0.9 \pm 6.2	-0.2 \pm 0.7	0.70	-2.1 \pm 5.2	-0.0 \pm 5.3	-2.3 \pm 1	0.028
E/e' change, /unit	0.1 \pm 2.2	0.9 \pm 1.5	-0.7 \pm 0.5	0.04	-0.3 \pm 2.1	-0.4 \pm 2.4	0.2 \pm 0.3	0.70	-1.0 \pm 2.5	0.0 \pm 3.2	-1.5 \pm 0.5	0.004
SBP change, mmHg	-9.1 \pm 15.4	0.2 \pm 14.3	-12.2 \pm 3.1	0.002	-13.0 \pm 19.8	-3.5 \pm 18.5	-8.3 \pm 2.2	0.002	-15.9 \pm 17.0	-7.0 \pm 21.8	-10.7 \pm 3	<0.001
PIIINP change, μg/L	-0.1 \pm 2.2	-0.2 \pm 1.4	0.1 \pm 0.5	0.63	-0.0 \pm 1.7	-0.3 \pm 1.7	0 \pm 0.2	0.93	-0.2 \pm 1.7	-0.3 \pm 2.3	-0.2 \pm 0.3	0.45
PICP change, μg/L	-9.0 \pm 24.8	1.5 \pm 19.6	-6.9 \pm 4.6	0.12	-6.6 \pm 26.5	-1.9 \pm 18.5	-6.3 \pm 3	0.032	-9.9 \pm 22.9	0.2 \pm 26.3	-12.6 \pm 4	0.003
NTproBNP change, μg/L	-8.0 \pm 179.3	68.5 \pm 202.5	-23.5 \pm 49.3	0.64	-25.4 \pm 163.5	-10.4 \pm 196.9	-17.9 \pm 21.9	0.33	-18.9 \pm 155.4	119.3 \pm 613.6	-134.2 \pm 84.9	0.06
BNP change	-0.1 \pm 1.2	0.5 \pm 1.1	-0.4 \pm 0.3	0.15	-0.0 \pm 1.0	-0.1 \pm 0.9	0 \pm 0.1	0.67	-0.3 \pm 1.0	0.3 \pm 1.0	-0.6 \pm 0.2	0.002

* Adjusted for age, sex, body mass index, a prevalence of hypertension, diabetes mellitus, glomerular filtration rate and baseline echocardiographic/proteomic variables.

**Echocardiographic and biomarker variables were assessed by changes from baseline to month 9.