

Supplemental Material

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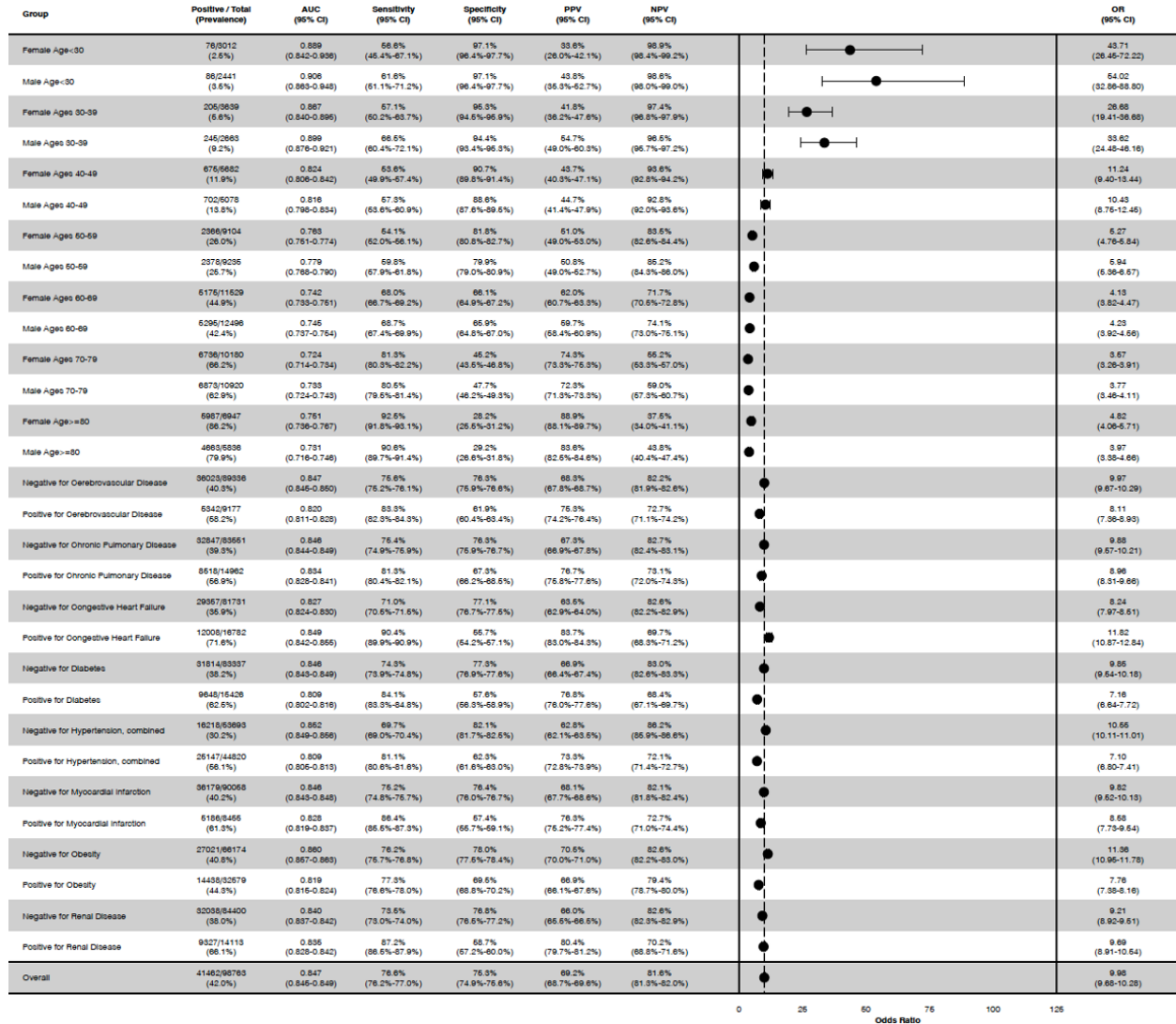
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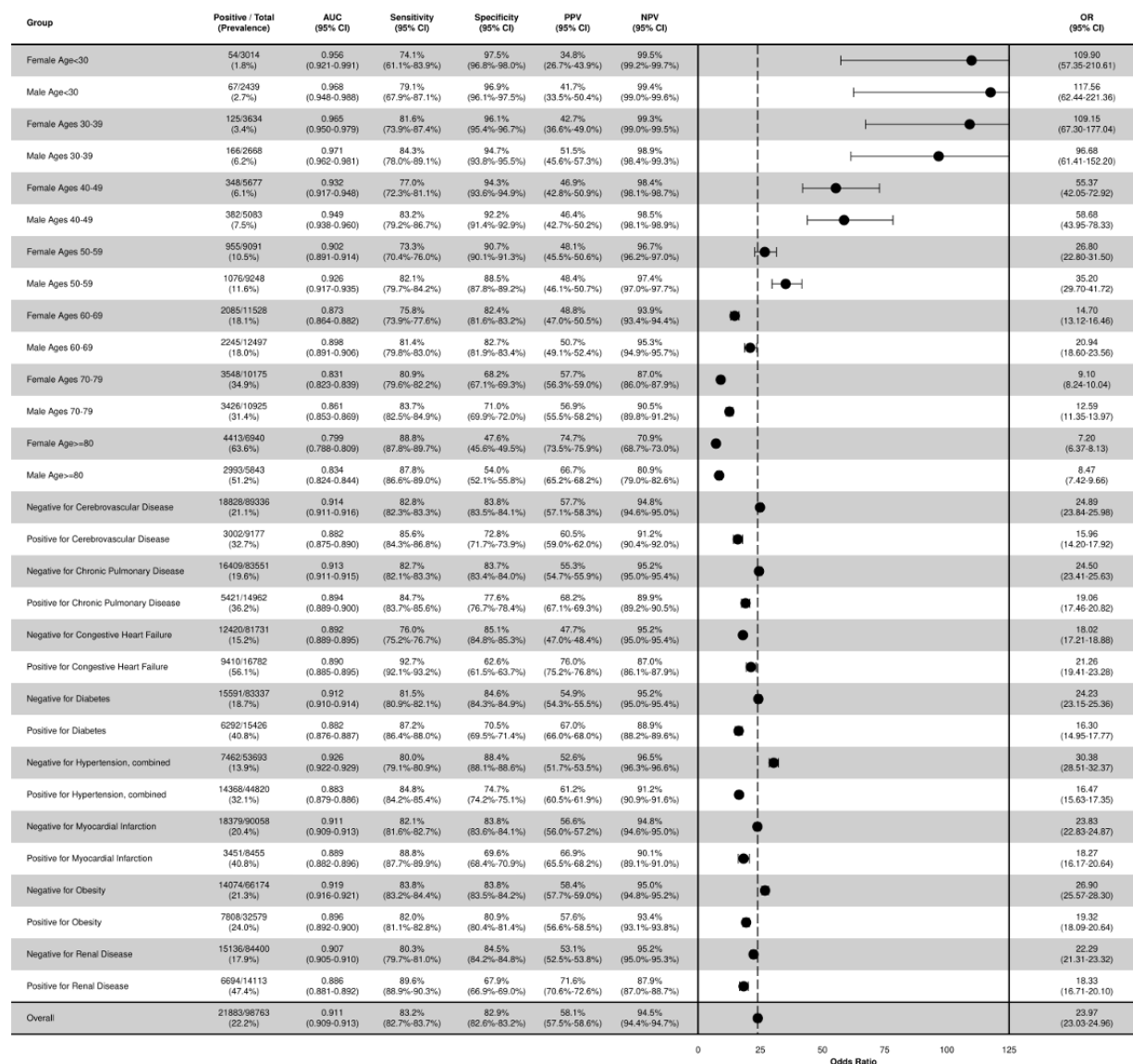
1. Figure

Figure 1. AUC, sensitivity, specificity, PPV, NPV, and odd ratio (OR) with 95% confidence intervals across age, gender, and comorbidity subsets. a. grade 1 or above. b. grade 2 or above (increased filling pressure) c. grade 3.

a.



b.



C.

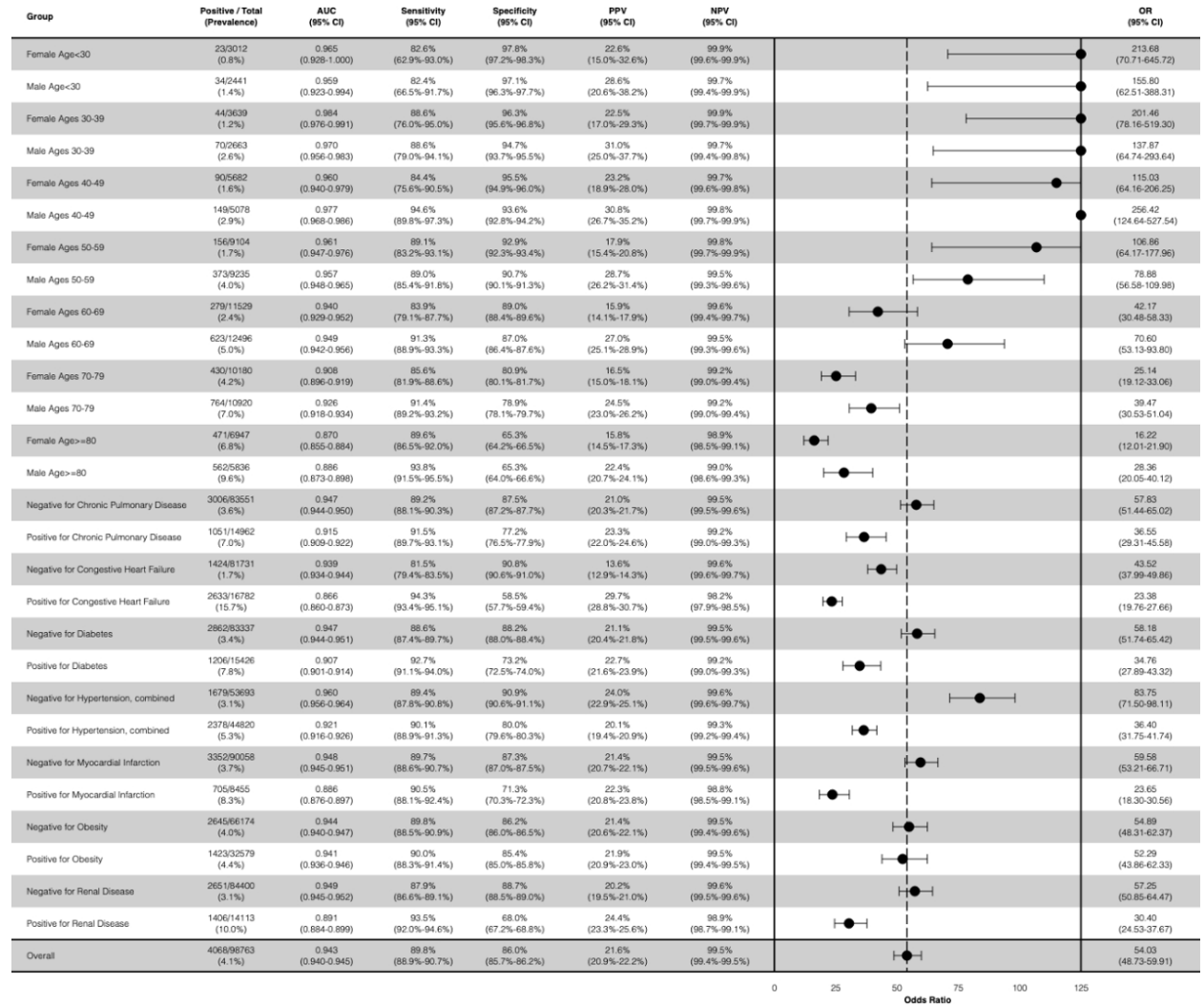


Figure 2. The distribution of left atrial volume index, e' , E/e' , tricuspid regurgitation velocity, and E/A according to filling pressure by our AI-ECG and echocardiography in test set. Each point represents the mean value with 95% confidence interval. Normal filling pressure includes normal and grade1, and increased filling pressure includes grade 2 and grade 3.

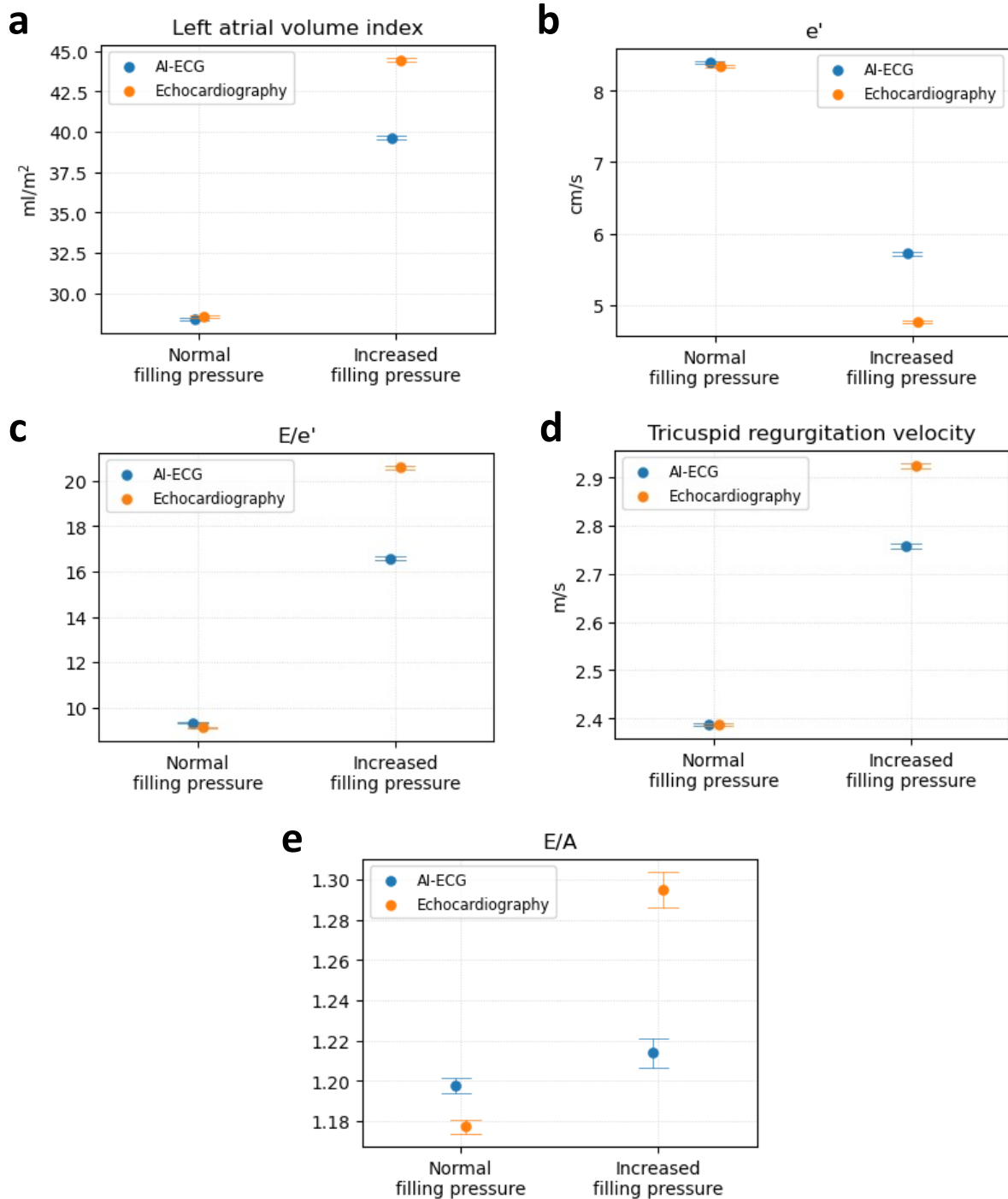


Figure 3. The distribution of left atrial volume index, e' , E/e' , tricuspid regurgitation velocity, and E/A according to filling pressure by our AI-ECG in indeterminate set. Each point represents the mean value with 95% confidence interval. Normal filling pressure includes normal and grade1, and increased filling pressure includes grade 2 and grade 3. Student's t-test is used for p -value.

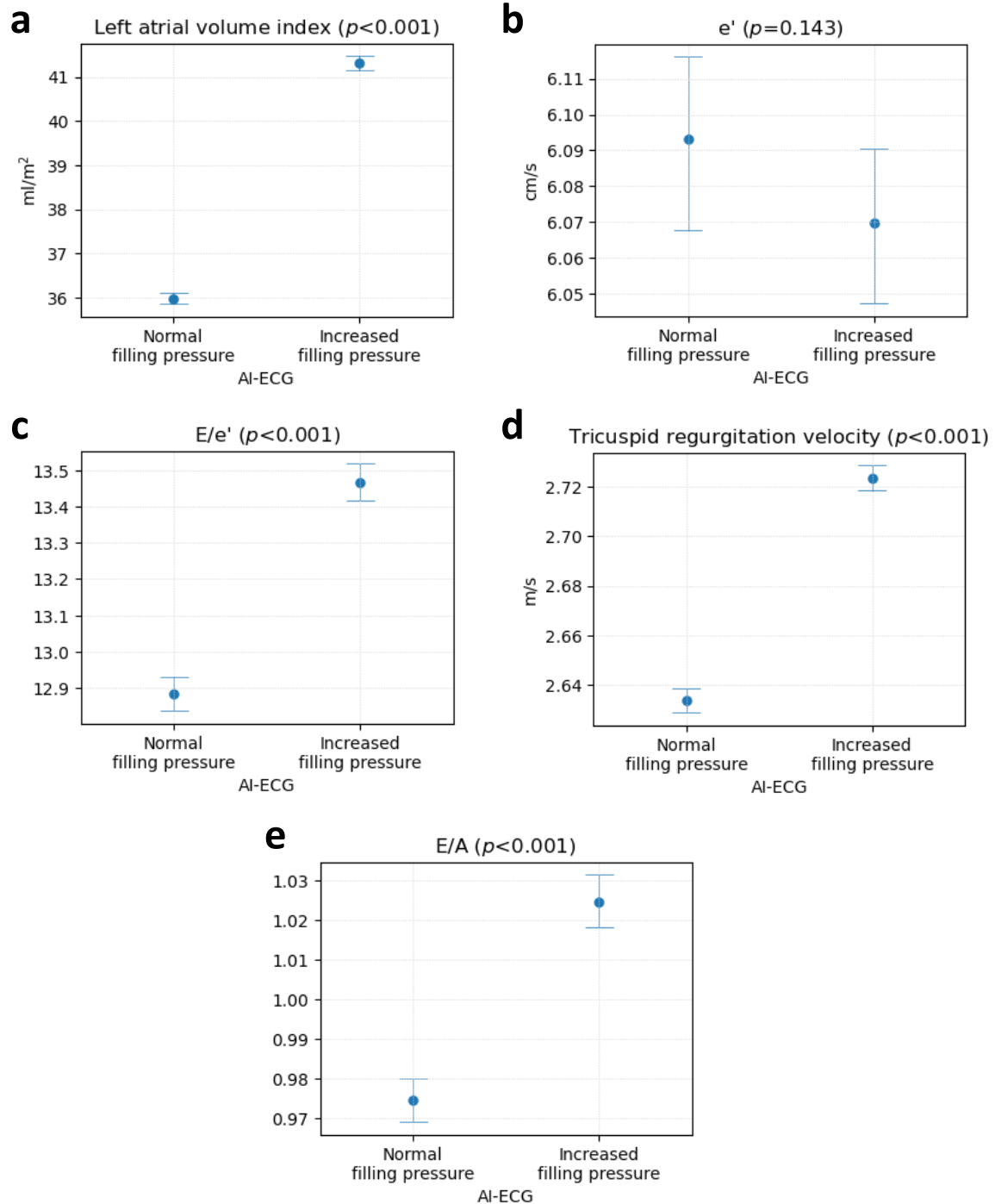


Figure 4. The ROC curves of AI-ECG between before and after the median year of the echocardiography exam in the test set. ROC plots for detecting diastolic function grades using ordinal scale.

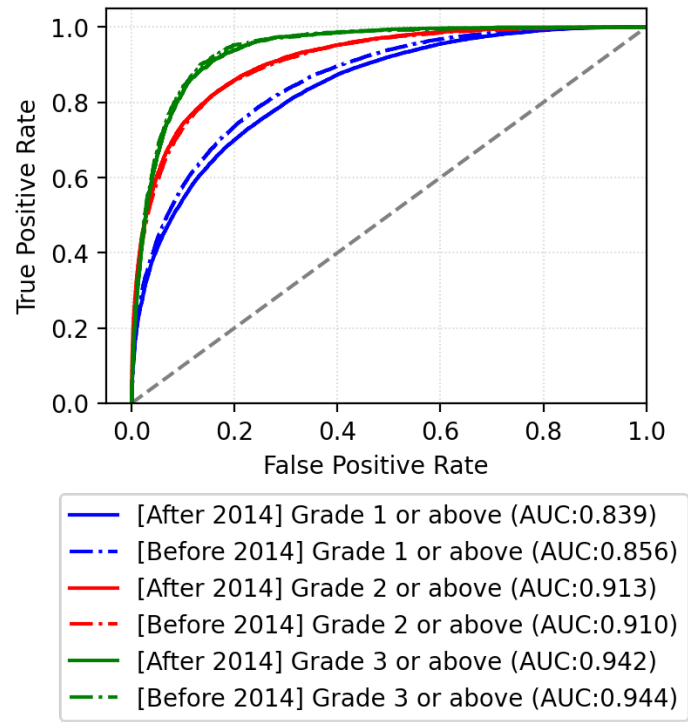


Figure 5. All-cause mortality using a Kaplan-Meier curve with 95% confidence intervals and p-value from a log-rank test by true positive (TP), false positive (FP), false negative (FN), and true negative (TN) for grade 1 or above, grade 2 or above, and grade 3.

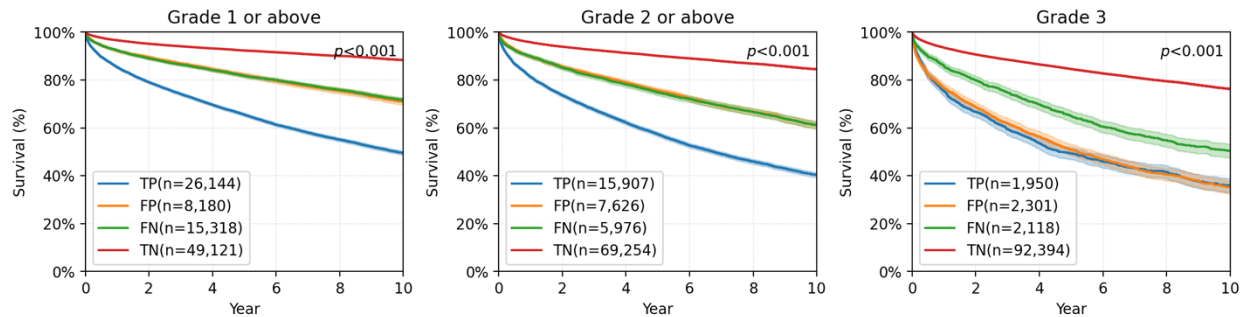


Figure 6. All-cause mortality using a Kaplan-Meier curve with 95% confidence intervals and p-value from a log-rank test by age. a. Kaplan-Meier curve for test group of patients (age \leq 50) according to the filling pressure by our AI-ECG (HR 1.413, 95% CI 1.329-1.503). b. Kaplan-Meier curve for test group of patients (50< age <70) according to the filling pressure by our AI-ECG (HR 1.35, 95% CI 1.313-1.388). c. Kaplan-Meier curve for test group of patients (age \geq 70) according to the filling pressure by our AI-ECG (HR 1.235, 95% CI 1.211-1.26). HRs were calculated with four grades after adjusting by age, sex, and comorbidities.

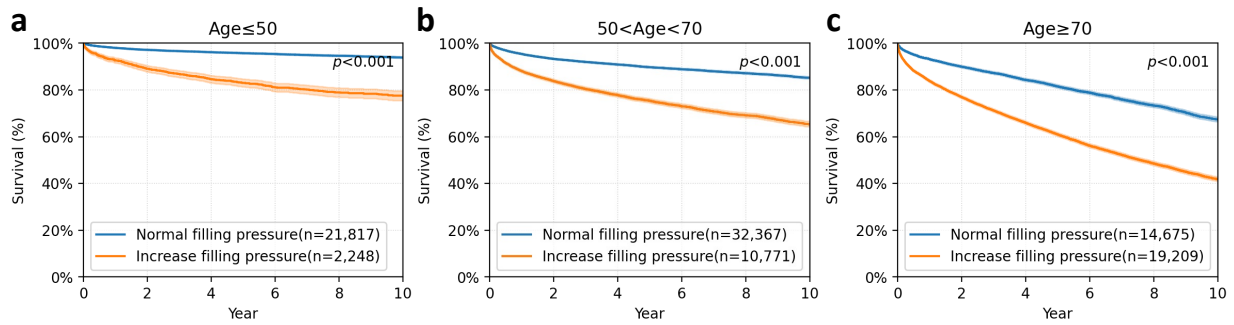


Figure 7. The distribution of left atrial volume index, e' , E/e' , tricuspid regurgitation velocity, and E/A according to diastolic filling pressure by our AI-ECG in grade 1 by echocardiography. Each point represents the mean value with 95% confidence interval. Normal filling pressure includes normal and grade1, and increased filling pressure includes grade 2 and grade 3. Student's t-test is used for p -value.

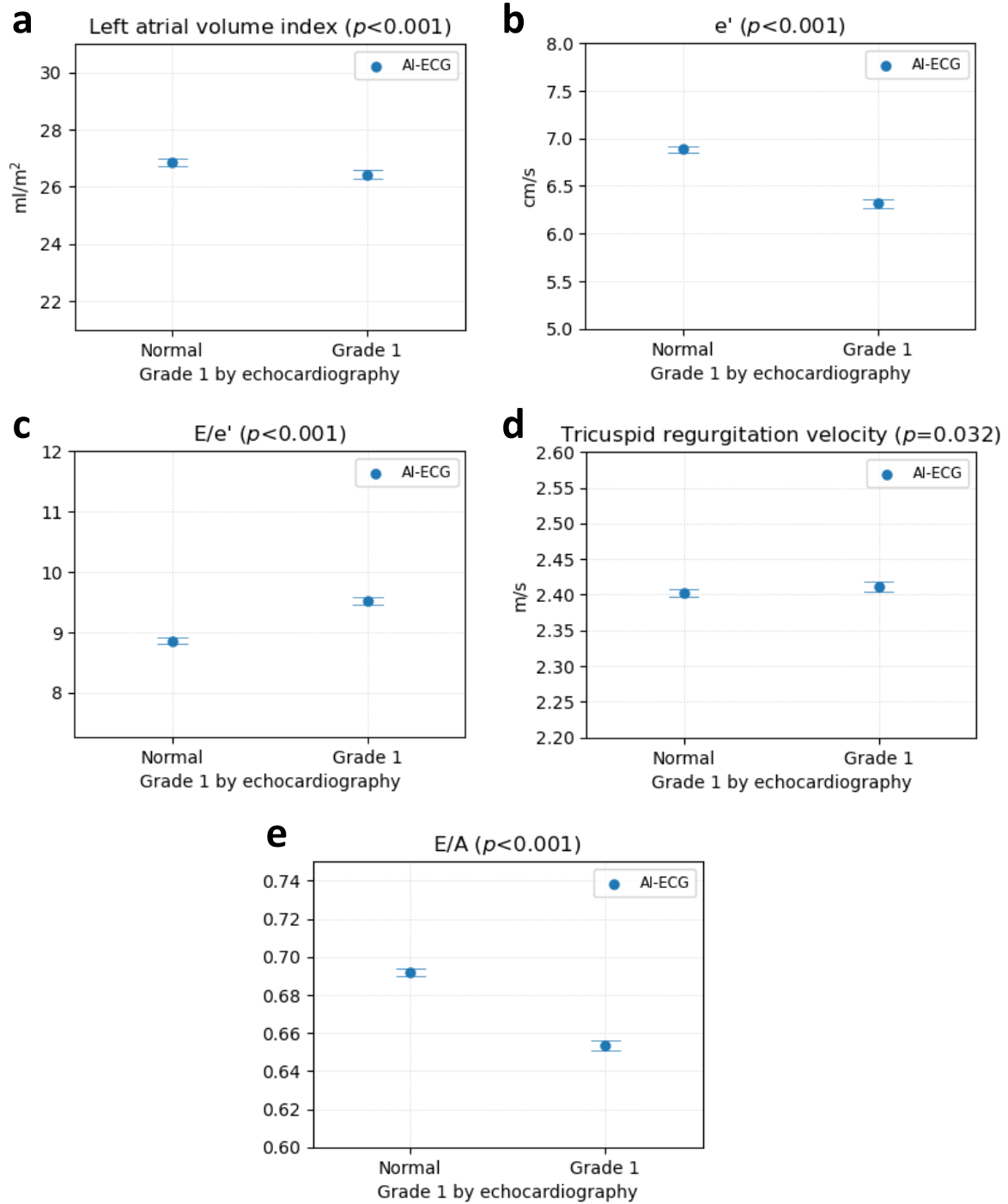
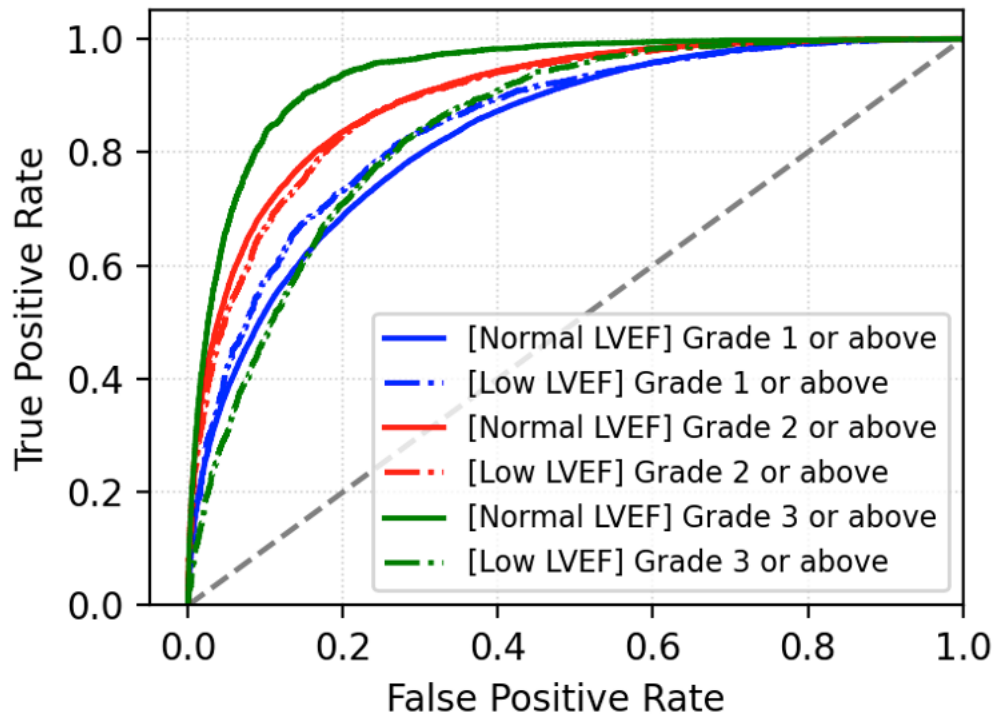


Figure 8. The ROC curves and performance of AI-ECG in normal ejection fraction and low ejection fraction, defined as left ventricular ejection fraction <50% in the test set. ROC plots for detecting diastolic function grades using ordinal scale.



Ordinal grade	LVEF	AUROC	Sensitivity	Specificity	PPV	NPV
Grade 1 or above	Normal	0.83	0.58	0.87	0.73	0.77
	Low	0.85	0.87	0.64	0.89	0.58
Grade 2 or above (Increased filling pressure)	Normal	0.90	0.67	0.91	0.63	0.92
	Low	0.89	0.90	0.69	0.82	0.82
Grade 3 or above	Normal	0.94	0.36	0.99	0.40	0.98
	Low	0.84	0.62	0.85	0.51	0.90

Figure 9. An illustrative case of a heart failure patient. Video for the echocardiograms is attached separately (Supplemental Video 1).

30-year-old female with heart failure due to post-partum cardiomyopathy Baseline and Follow-up Echocardiography and ECG

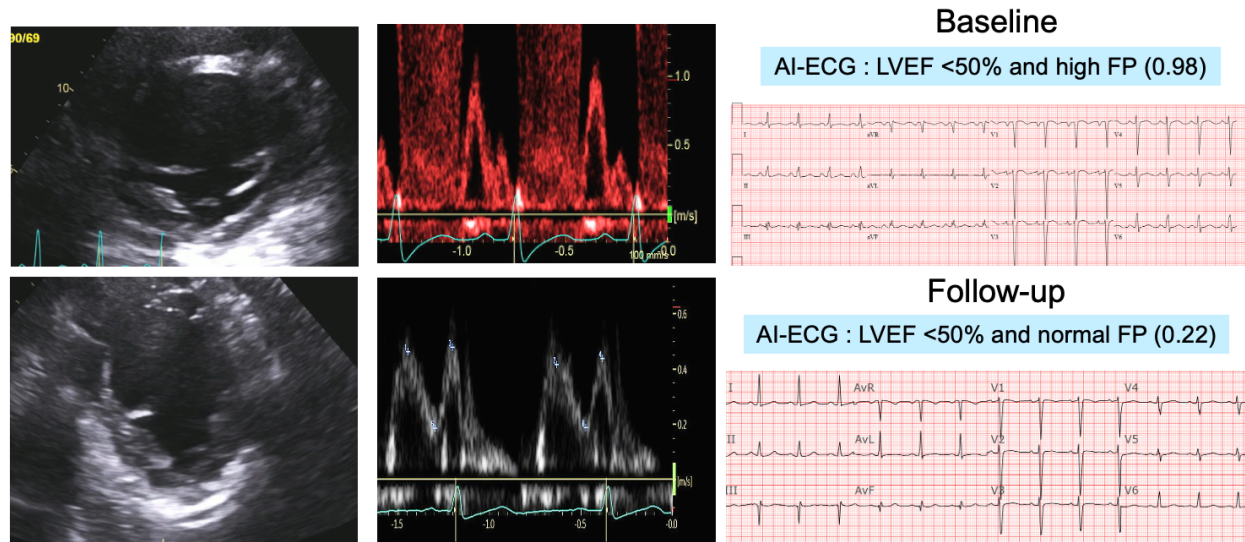


Figure 10. Flow chart of dataset construction and patient distribution. a. A flow chart demonstrating patient selection. b. A pie chart for dataset split.

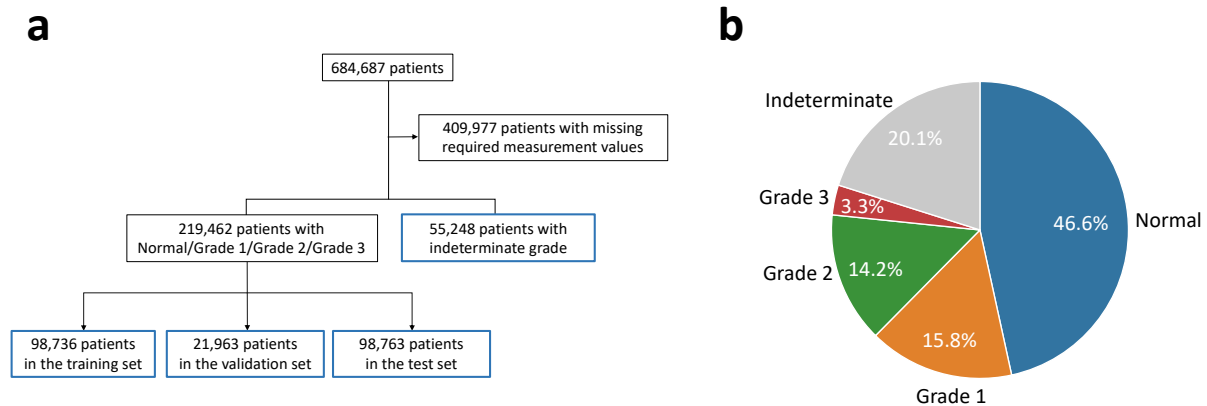
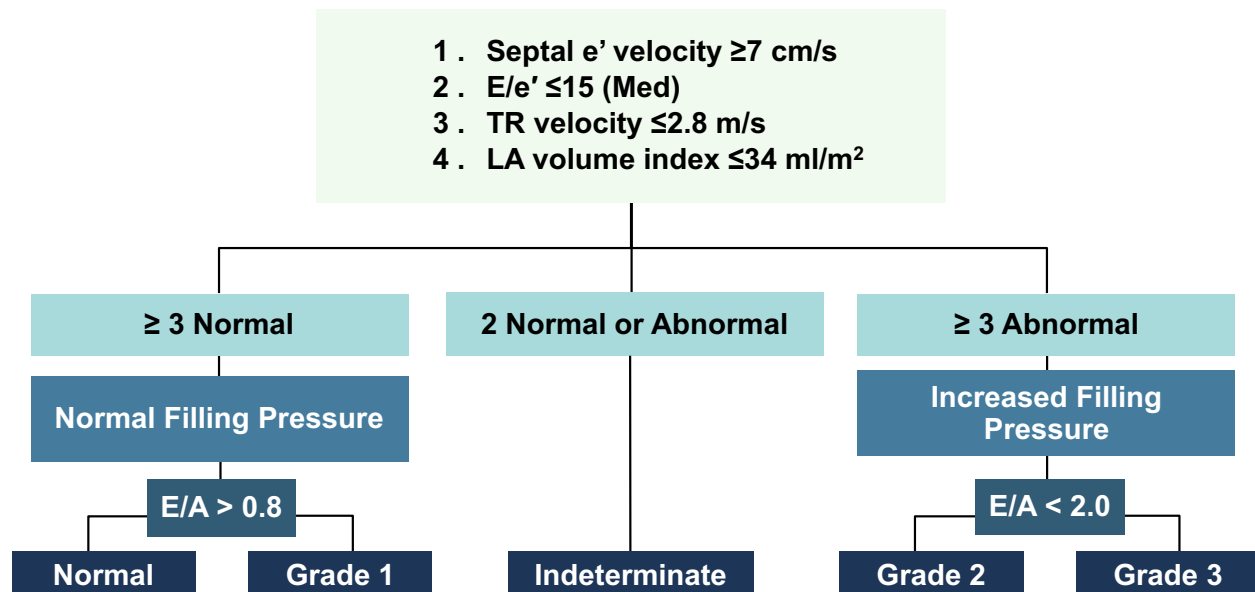


Figure 11. Algorithm for assessment of diastolic filling pressure and function¹.



A=Late mitral inflow velocity with atrial contraction, E=Early diastolic mitral inflow velocity, e'=early medial (Med) or septal mitral annulus diastolic velocity, LA=Left atrium, TR=Tricuspid regurgitation velocity.

¹ Oh JK, Miranda WR, Bird JG, Kane GC, Nagueh SF. The 2016 diastolic function guideline: is it already time to revisit or revise them? JACC: Cardiovascular Imaging 2020;13(1 Part 2):327-335.

2. Table

Table 1. Patient characteristics of training, validation, and testing group. Values are n (%) or mean \pm SD. Obesity was defined as body mass index ≥ 30 . Renal disease includes any stages of chronic kidney disease, hypertensive kidney disease, glomerulonephritis, nephritic syndrome, hereditary nephropathy, end stage renal disease, unspecified kidney failure, dialysis, and kidney transplant status.

	Overall (n=219,462)		
	Training (n=98,736)	Validation (n=21,963)	Test (n=98,763)
Age, y	61.0 \pm 16.8	60.8 \pm 16.8	61.2 \pm 16.8
Female sex, Number (%)	50,574 (51.2%)	11,164 (50.8%)	50,093 (50.7%)
Myocardial infarction, Number (%)	8,513 (8.6%)	1,914 (8.7%)	8,455 (8.6%)
Congestive heart failure, Number (%)	16,797 (17.1%)	3,833 (17.5%)	16,782 (17.0%)
Cerebrovascular disease, Number (%)	9,082 (9.2%)	1,970 (9.0%)	9,177 (9.3%)
Chronic pulmonary disease, Number (%)	14,869 (15.1%)	3,277 (15.0%)	14,962 (15.2%)
Diabetes Mellitus, Number (%)	15,598 (15.8%)	3,504 (16.0%)	15,426 (15.6%)
Renal disease, Number (%)	13,825 (14.0%)	3,086 (14.1%)	14,113 (14.3%)
Hypertension, Number (%)	44,639 (45.3%)	9,888 (45.1%)	44,820 (45.5%)
Obesity, Number (%)	32,696 (33.1%)	7,280 (33.1%)	32,579 (33.0%)

Table 2. Patient characteristics of four diastolic grade groups determined by AI-ECG in test set. Values are n (%).

AI-ECG diastolic function grade	Test (n=98,763)			
	Normal (n=64,439)	Grade 1 (n=10,791)	Grade 2 (n=19,282)	Grade 3 (n=4,251)
Age, year	55.6 ± 16.1	70.0 ± 10.5	73.0 ± 12.7	69.8 ± 14.7
Female sex, Number (%)	32,801 (50.9%)	5,338 (49.5%)	10,340 (53.6%)	1,614 (38.0%)
Myocardial infarction, Number (%)	3,674 (5.7%)	1,071 (9.9%)	2,897 (15.1%)	813 (19.2%)
Congestive heart failure, Number (%)	5,425 (8.4%)	1,414 (13.1%)	7,012 (36.5%)	2,931 (69.1%)
Cerebrovascular disease, Number (%)	4,513 (7.0%)	1,341 (12.4%)	2,793 (14.5%)	530 (12.5%)
Chronic pulmonary disease, Number (%)	7,848 (12.2%)	1,760 (16.3%)	4,164 (21.7%)	1,190 (28.1%)
Diabetes, Number (%)	6,961 (10.8%)	1,901 (17.6%)	5,188 (26.9%)	1,376 (32.4%)
Renal disease, Number (%)	5,694 (8.9%)	1,427 (13.2%)	5,329 (27.7%)	1,663 (39.2%)
Hypertension, Number (%)	23,310 (36.3%)	6,151 (57.1%)	12,825 (66.7%)	2,534 (59.7%)
Obesity, Number (%)	20,356 (31.6%)	4,060 (37.6%)	6,847 (35.5%)	1,316 (31.0%)

Table 3. Model performance for filling pressure and diastolic function grade from the AI-ECG in test set by hypertension, obesity, and diabetes. The threshold from the entire validation were used for each class.

Comorbidity	Class	Prevalence	AUCROC	Sensitivity	Specificity	PPV	NPV
Hypertension	Grade 1 or above	Y	0.81	0.8	0.65	0.72	0.74
		N	0.86	0.67	0.86	0.6	0.89
	Grade 2 or above	Y	0.88	0.84	0.76	0.6	0.92
		N	0.94	0.8	0.91	0.5	0.98
	Grade 3	Y	0.93	0.9	0.81	0.21	0.99
		N	0.97	0.9	0.93	0.25	1
Obesity	Grade 1 or above	Y	0.82	0.77	0.7	0.67	0.79
		N	0.86	0.76	0.78	0.7	0.83
	Grade 2 or above	Y	0.9	0.82	0.81	0.58	0.93
		N	0.92	0.84	0.84	0.58	0.95
	Grade 3	Y	0.94	0.9	0.85	0.22	0.99
		N	0.94	0.9	0.86	0.21	1
Diabetes	Grade 1 or above	Y	0.81	0.83	0.61	0.75	0.71
		N	0.85	0.74	0.78	0.66	0.84
	Grade 2 or above	Y	0.89	0.86	0.73	0.65	0.9
		N	0.91	0.81	0.85	0.54	0.95
	Grade 3	Y	0.91	0.92	0.76	0.22	0.99
		N	0.95	0.88	0.89	0.21	1

Table 4. Clinical characteristics of normal or grade 1 group by AI-ECG in grade 1 patients by echocardiography. Values are n (%) or mean \pm SD. Student t-test and chi-squared test were used.

	Normal (n=10,712)	Grade 1 (n=5,669)	<i>p</i>-value
Age, y	66.1 \pm 10.9	71.6 \pm 9.1	<0.001
Female sex, Number (%)	5,194 (48.5%)	2,823 (49.8%)	0.861
Myocardial infarction, Number (%)	799 (7.5%)	512 (9.0%)	<0.001
Congestive heart failure, Number (%)	1,063 (10.0%)	747 (13.2%)	<0.001
Cerebrovascular disease, Number (%)	1,064 (10.0%)	737 (13.0%)	<0.001
Chronic pulmonary disease, Number (%)	1,608 (15.1%)	935 (16.5%)	0.015
Diabetes Mellitus, Number (%)	1,643 (15.3%)	1,001 (17.7%)	<0.001
Renal disease, Number (%)	1,185 (11.1%)	788 (13.9%)	<0.001
Hypertension, Number (%)	5,319 (49.8%)	3,350 (59.1%)	<0.001
Obesity, Number (%)	3,565 (33.3%)	2,044 (36.1%)	<0.001

Table 5. The number of patients for hypertrophic cardiomyopathy (HCM), cardiac amyloidosis, moderate to severe mitral valve regurgitation (MR) and aortic stenosis (AS), and reduced left ventricular ejection fraction (LVEF) according to diastolic function grade by AI-ECG. Percent in each column indicates distribution of a particular condition within the same diastolic function category.

AI-ECG diastolic function grade	Test (n=98,763)			
	Normal (n=64,439)	Grade 1 (n=10,791)	Grade 2 (n=19,282)	Grade 3 (n=4,251)
HCM	1,103 (1.7%)	233 (2.2%)	1,267 (6.6%)	852 (20%)
Amyloidosis	2,590 (4%)	373 (3.5%)	1,571 (8.1%)	2,225 (52.3%)
Moderate to severe MR	1,070 (1.7%)	144 (1.3%)	1,958 (10.2%)	1,017 (23.9%)
Moderate to severe AS	588 (<1%)	242 (2.2%)	1,374 (7.1%)	265 (6.2%)
Reduced LVEF (<50%)	2,252 (3.5%)	755 (7%)	3,813 (19.8%)	2,252 (53%)