Scoring system based on electrocardiogram features to predict the type of heart failure in patients with chronic heart failure

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ABSTRACT

Heart failure is divided into heart failure with reduced ejection fraction (HFrEF) and heart failure with preserved ejection fraction (HFpEF). Additional studies are required to distinguish between these two types of HF. A previous study showed that HFrEF is less likely when ECG findings are normal. This study aims to create a scoring system based on ECG findings that will predict the type of HF. We performed a cross-sectional study analyzing ECG and echocardiographic data from 110 subjects. HFrEF was defined as an ejection fraction ≤40%. Fifty people were diagnosed with HFpEF and 60 people suffered from HFrEF. Multiple logistic regression analysis revealed certain ECG variables that were independent predictors of HFrEF i.e., LAH, QRS duration > 100 ms, RBBB, ST-T segment changes and prolongation of the QT interval. Based on ROC curve analysis, we obtained a score for HFpEF of -1 to +3, while HFrEF had a score of +4 to +6 with 76% sensitivity, 96% specificity, 95% positive predictive value, an 80% negative predictive value and an accuracy of 86%. The scoring system derived from this study, including the presence or absence of LAH, QRS duration > 100 ms, RBBB, ST-T segment changes and prolongation of the QT interval can be used to predict the type of HF with satisfactory sensitivity and specificity.

ABSTRAK

Latar belakang: Gagal jantung dapat dibagi ke dalam 2 tipe yakni gagal jantung dengan penurunan ejeksi fraksi dan gagal jantung dengan normal ejeksi fraksi. Penelitian lebih lanjut diperlukan untuk membedakan kedua tipe gagal jantung. Penelitian yang pernah dilakukan sebelumnya menunjukkan bahwa hasil EKG pada penderita gagal jantung dengan penurunan ejeksi fraksi kebanyakan menunjukkan hasil yang normal. Tujuan dari penelitian ini adalah untuk membuat system skoring berdasarkan temuan EKG yang dapat memprediksi tipe gagal jantung. Metode penelitian ini menggunakan desain potong lintang dimana akan dilakukan analisis hasil temuan EKG dan Ekokardiografi dari 110 subjek. Gagal jantung dengan penurunan ejeksi fraksi didefinisikan sebagai nilai EF < 40%. Lima puluh pasien terdiagnosis sebagai gagal jantung dengan ejeksi fraksi normal, dan sisanya 60 pasien menderita gagal jantung dengan penurunan ejeksi fraksi. Regresi logistik multiple menunjukkan beberapa variable EKG yang dapat menjadi penunjuk terjadinya penurunan ejeksi fraksi, yakni LAH, QRS > 100 MS, RBBB, perubahan segmen ST-T dan pemanjangan interval QT. Berdasarkan analisis menggunakan kurva ROC, didapatkan skor untuk ejeksi fraksi normal -1 sampai +3, sedangkan untuk penurunan ejeksi fraksi fraksi

sampai +6 dengan sensitivitas 76%, spesifisitas 96%, nilai prediksi positif 95%, nilai prediksi negative 80% dan akurasi 86%. Simpulan skoring system yang didapatkan dari penelitian ini, yang menunjukkan ada atau tidaknya LAH, QRS >100ms, RBBB, perubahan segmen ST-T, pemanjangan interval QT dapat digunakan untuk memprediksi tipe gagal jantung kronik dengan nilai sensitivias dan spesifisitas yang memuaskan.

Keywords: chronic heart failure - scoring system - electrocardiogram features - type of heart failure

BACKGROUND

Heart failure has a high incidence, is one of the major causes of mortality from cardiovascular diseases in the world and is a major health problem in society. Heart failure is divided into heart failure with reduced ejection fraction (HFrEF) and heart failure with preserved ejection fraction (HFpEF).2 In the literature, HFrEF and HFpEF are also referred to as systolic and diastolic heart failure, respectively. Patients with systolic dysfunction can also exhibit diastolic dysfunction, particularly in late stage heart failure.3 The mortality from heart failure is inversely related to left ventricular systolic function. Ejection fraction is considered one of the strongest prognosistic factors that influences a poor outcome for heart failure patients.4

Echocardiography is considered the gold standard for assessing diastolic or systolic dysfunction in patients with heart failure. However, an expert is required to conduct the examination and not all health facilities provide an echocardiography machine. Thus, a simple device that can measure both systolic and diastolic function can help a physician determine the diagnosis.⁵ Electrocardiogram (ECG) examination is inexpensive tool that is accessible in almost all primary health care setting. A normal or minor change in the ECG is consistent with a low likelihood

of left ventricle dysfunction. Conversely, left ventricle systolic dysfunction is usually accompanied by major ECG changes.⁶ The ECG is useful because it can serve as an initial investigative tools that physicians can use to determine the presence of systolic and diastolic dysfunction in patients with chronic heart failure, though it cannot replace echocardiography.

A scoring system is a simple method for diagnosing disease.⁷ Several scoring systems based on ECG findings have been studied to estimate left ventricular function. However, additional studies have suggested that scoring systems have limitations or are less accurate in estimating left ventricular function in patients with coronary heart disease.^{8,9} To our knowledge, there no previous study has evaluated a scoring system based of ECG results to predict HFpEF or HFrEF in patients with heart failure.

METHODS

We used a cross-sectional research design to determine a scoring system based on ECG findings to predict the type of heart failure (HFpEF or HFrEF). The population consisted of patients with chronic heart failure that were hospitalized or seen as outpatients in Dr. Sardjito General Hospital between April and July 2015 and in whom both echocardiography and ECG had been conducted. The inclusion

criteria were patients with heart failure diagnosed based on ESC (2012) or AHA (2013) guidelines, the presence of sinus rhythm, age > 18 years and agreement to participate in the study. The exclusion criteria were patients with congenital heart disease, primary valve disease, massive pericardial effusion, patients with acute coronary syndrome, severe pulmonary disease (cor pulmonale, pneumothorax) or who had pacemakers. The independent variable in this study was the ECG findings in patients with chronic heart failure. The dependent variable was chronic heart failure (HFrEF or HFpEF). The confounding variable was patient medications.

Subjects were enrolled in the cardio-vascular clinic, the hospital ward, and the echocardiography clinic at Dr. Sardjito General Hospital Subjects who met the inclusion criteria was included in the study and were enrolled consecutively. The collected data included demographic information, clinical examination, ECG and echocardiography results. Demographic data included age, sex and medication history. Clinical data included the NYHA class determination.

Twelve lead ECGs were obtained by a nurse or a cardiology resident with patients in the supine position at a speed of 25 mm/s. The ECGs were read by 3 expert physicians (who were blinded to the results of the echocardiogram). The ECG report included an evaluation of heart rhythm and rate, heart axis, presence or absence of: chamber enlargement, intraventricular block, and ST-T segment changes, and the duration of the QRS complex, QT and QTc intervals. A wide QRS complex defined as QRS duration >100 ms. The echocardiogram was supervised by a cardiologist and 2 expert examiners (who were blinded to the ECG result). Echocardiography data included a determination of the ejection fraction (calculated by Simpson's method) and the presence or absence of diastolic dysfunction. Heart failure was divided into HFrEF and HFpEF. The criterion for a diagnosis of HFrEF and HFpEF was an ejection fraction ≤ 40% and > 40%, respectively. Subjects were divided into two groups: HFrEF group and HFpEF group.

Bivariate analysis was used to analyze the relationship between the ECG results and HFrEF or HFpEF. Inter-variable bivariate analysis was analyzed with the Chi-square test followed by Fisher's test. Further, variables in the bivariate analysis with a p <0.25 were tested using a multivariate analysis, specifically logistic regression with a backward stepwise method. Each variable from the multivariate analysis was scored by using B and SE values (from SPSS statistical analysis program). There were 2 scoring systems based on the probability and cut-off point from the ROC curve. Next, the scoring system was validated in several samples to obtain the diagnostic value. The entire data analysis was statistically conducted using software. Additional information about the research subjects that was deemed necessary for the study was collected from the medical record or via a direct interview. Sample data collection was non-probability sampling.

The research was conducted after receiving permission from the faculty of medicine at UGM/RSUP Dr. Sardjito Ethical Committee. Informed consent was obtained from the subjects after they received complete information about the trial.

RESULTS

Data were collected between April and July 2015 from patients seen in the cardiovascular clinic, hospital ward and echocardiography

clinic at Dr. Sardjito General Hospital. A total of 110 heart failure patients met the inclusion criteria. Sixty patient (54.5%) had systolic heart failure (HFrEF) and 50 patients had diastolic heart failure (HFpEF). There

was no significant difference in the average age of patients in the HFpEF and HFrEF group (59.7±9.2 years and 57.9±10.1 years, respectively). All of the subjects met NYHA class II or III criteria (TABLE 1).

TABLE 1. Basic subject characteristics based on heart failure type

Variable	HFpEF $(n = 50)$	HfrEF $(n = 60)$	Total (n = 110)	p
Age, year ±SD	59.7±9.2	57.9±10.1	58.6±9.9	0.29
Sex				0.003
Male, n (%)	28 (56)	49 (81.7)	77 (70)	
Female, n (%)	22 (44)	11 (18.3)	33 (30)	
Risk factor				
Hypertension, n (%)	46 (92)	39 (65)	85 (77.2)	0.001
DM, n (%)	19 (38)	13 (21.7)	32 (29.1)	0.06
Smoker, n (%)	11 (22)	29 (48.3)	40 (36.3)	0.004
Therapy				
ACE-I / ARB, n (%)	50 (100)	60 (100)	110 (100)	1.000
Diuretic, n (%)	37 (74)	57 (95)	94 (85.4)	0.002
Beta Blocker, n (%)	21 (42)	30 (50)	51 (46.3)	0.042
CCB, n (%)	22 (44)	6 (10)	28 (25.4)	< 0.0001

HFpEF: Heart Failure preserved Ejection Fraction; HFrEF: Heart Failure reduced Ejection Fraction; DM: Diabetes mellitus; ACE-I: Angiotensin Converting Enzyme Inhibitor; ARB: Angiotensin Receptor Blocker; CCB: Calcium Channel Blocker

Patients with HFrEF had a larger left atrial diameter, LAVI, LVMI, LVIDd and E/A ratio compare to patients with HFpEF, while IVSD, DT and TAPSE were lower in patients with HFrEF. Kinetic disturbance in the myocardium

was found in all patients with HFrEF patient and in 30% of patients with HFpEF. Characteristics of the echocardiograms are shown in TABLE 2.

TABLE 2. Echocardiography basic characteristics based on heart failure type

77 ' 11	IIE EE (50)	III. EE((0)	
Variable	HFpEF (n = 50)	HFrEF(n = 60)	<i>p</i>
LA Diameter, mm±SD	34.3±5.2	41.5±6.2	< 0.0001
LVIDd, mm±SD	49.2±7.1	64.9±7.9	< 0.0001
IVSd, mm±SD	12.5±2.1	10.3 ± 2.6	< 0.0001
EF Simpson, %±SD	59.2.±8.5	29.4 ± 6.8	< 0.0001
LVMI, g/m2±SD	$133.3.\pm40.2$	170.3 ± 46.7	< 0.0001
Septal e'	$5.09.\pm1.1$	4.3±1.2	< 0.0001
Lateral e'	6.21±1.6	5.6±2.5	0.146
LAVI, ml/m2±SD	34.5±4.3	47.1±14.4	< 0.0001
E/A ratio	0.77 ± 0.2	1.8 ± 1.2	< 0.0001
DT, ms±SD	234±53.3	158.4±52.8	< 0.0001
Diastolic dysfunction			1.000
relaxation, n	39	16	
pseudonormal, n	11	15	
restrictive, n	0	29	
TAPSE, mm±SD	23.2±3.1	18.1±4.3	< 0.0001
RVSP, mmHg±SD	$6.64.\pm 9.1$	26.8 ± 22.6	< 0.0001
Myocard Kinetic disturbance			< 0.0001
yes	15	60	
no	35	0	
Heart rate, x/minute ±SD	71.8±12.8	81.7±15	0.135

LA: left atrium; LVIDd: left ventricular internal diameter end diastole; IVSd: Interventricular septal end diastole; EF: ejection fraction; LVMI: left ventricular mass index; LAVI: left atrial volume index; DT: deceleration time; TAPSE: tricuspid annular plane systolic excursion; RVSP: right ventricle systolic pressure

The major ECG abnormalities were a prolonged QT interval (62.7%), ST-T segment changes (55.4%) and prolonged QRS duration (48.2%). Thirty eight percent of patients with HFpEF had ST-T segment changes and a

prolonged QT interval, while a prolonged QT interval, prolonged QRS duration and ST-T segment changes were found in 83.3%, 71.7% and 70% of patients with HFrEF, respectively. ECG characteristics are shown in TABLE 3.

ECG Parameter	HFpEF $(n = 50)$	HFrEF(n = 60)	Total(n = 110)	p
LAH, n (%)	6 (12)	22 (36.7)	28 (25.5)	0.003
LVH, n (%)	15 (30)	33 (55)	48 (43.6)	0.008
Poor R wave, n (%)	5 (10)	30 (50)	35 (31.8)	< 0.0001
LAD, n (%)	15 (30)	27 (45)	42 (38.2)	0.107
RAD, n (%)		3 (5)	3 (2.7)	0.249
Q wave, n (%)	9 (18)	17 (28.3)	26 (23.6)	0.204
Wide QRS, n (%)	10 (20)	43 (71.7)	53 (48.2)	< 0.0001
QRS duration, ms±SD	97.3±20.7	124 ± 30.4		
LBBB, n (%)		12 (20)	12 (10.9)	0.001
RBBB, n (%)	7 (14)	3 (5)	10 (9.1)	0.181
ST-T changes, n (%)	19 (38)	42 (70)	61 (55.4)	0.001
Prolong QT, n (%)	19 (38)	50 (83.3)	69 (62.7)	< 0.0001
Interval QTc, ms±SD	453.2±42.8	499±50.9		

LAH: left atrial hypertrophy; LVH: left ventricular hypertrophy; LAD: left axis deviation; RAD: right axis deviation; LBBB: left bundle branch block; RBBB: right bundle branch block

Multivariate analysis (TABLE 4) demonstrated that the ECG variables influencing whether a patient had HFpEF or HFrEF were LAH, a wide QRS complex, RBBB, ST-T changes and a prolonged QT interval. The power of the association from

largest to smallest was a *wide QRS* (OR 12.657), prolonged QT interval (OR 7.401), LAH (OR 4.449), ST-T segment changes (OR 4.35) and RBBB (OR 0.109). Thus, the scores from these variables were used to calculate the scoring system (TABLE 5).

TABLE 4. Multivariate analysis of variable ECG with logistic regression to test the possibility of systolic heart failure or HFrEF (n = 110). * = statistically significant #= n.a

ECG Parameter	OR	IK 95%	p
LVH	1.24	0.284 - 5.407	0.774
Poor R wave	2.004	0.423 - 9.494	0.381
LAD	0.661	0.161 - 2.708	0.565
RAD	#	#	#
Q wave	3.756	0.947 - 14.891	0.060
LBBB	#	#	#
LAH	4.449	1.109 - 17.848	0.035*
Wide QRS	12.657	3.277 - 48.895	< 0.0001*
RBBB	0.109	0.012 - 0.986	0.049*
ST-T changes	4.35	1.277 - 14.817	0.019*
Prolong QT	7.401	2.134 - 25.672	0.002*

LAH: left atrial hypertrophy; LVH: left ventricular hypertrophy; LAD: left axis deviation; RAD: right axis deviation; LBBB: left bundle branch block; RBBB: right bundle branch block

TABLE 5. Calculated score for each ECG variable resulting from the multivariate analysis

ECG Parameter	В	SE	B/SE	[B/SE] / x	Score
LAH	1.493	0.709	2.105	1.068	1
Wide QRS	2.538	0.69	3.678	1.865	2
RBBB	-2.214	1.123	-1.971	-1	-1
ST-T changes	1.47	0.625	2.352	1.192	1
Prolong QT	2.002	0.635	3.152	1.599	2

LAH: left atrial hypertrophy; RBBB: right bundle branch block

The first scoring model was based on a subject's probability of having HFrEF. The minimum score was -1, and the maximum score was +6 (see TABLE 5). A regression tool was used to calculate the probability of each subject having systolic heart failure (see TABLE 6).

The second scoring model was based on a cut-off point derived from the ROC (Receiver Operating Characteristic) curve analysis. The ROC method is based on a calculation between sensitivity and specificity values surrounding various cut-off points represented in the graph. Results of the ROC curve analysis are shown in FIGURE 1.

TABLE 6. Scoring system based on the probability of systolic heart failure (HFrEF)

Score	Probability (%)	
-1	0.9	
0	3.16	
1	10.4	
2	29.3	
3	59.6	
4	84	
5	94.9	
6	98.5	

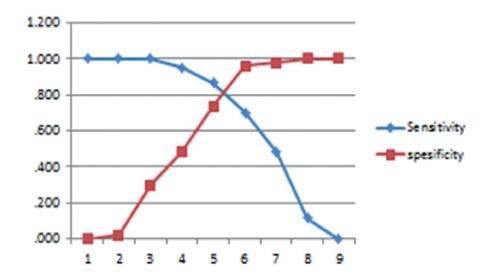


FIGURE 1. This figure shows the sensitivity and specificity based on a cut-off point derived from the ROC (Receiver Operating Characteristic) curve analysis

From the cut-off analysis in Figure 1, the cut-off value was 4. A score of +4 to +6 indicated the possibility of systolic heart failure (HFrEF), and a score of -1 to +3 indicated the possibility diastolic heart failure (HFpEF). The scoring system was validated

with additional samples to better determine the diagnostic value of the result and demonstrated 76% sensitivity, 96% specificity, 95% positive predictive value, 80% negative predictive value and an accuracy of 86% (TABLE 7).

TABLE 7. Validity test based on the scoring system of heart failure patients (n = 50)

		ECHO (gold standard)	
		HfrEF (EF \leq 40%) HfpEF (EF $>$ 40°	
Scoring system	HfrEF (+4 sd +6)	19	1
	HfpEF $(-1 \text{ sd } +3)$	6	24

HFpEF: Heart Failure preserved Ejection Fraction; HFrEF: Heart Failure reduced Ejection Fraction

DISCUSSION

In this study, hypertension was the major risk factor for patients with HFpEF (92%), while HFrEF was associated with multiple risk factors. This result is in agreement with Paulus and Tschope,10 and Tsutsui et al.,11 who found that hypertension is comorbid in HFpEF, while HFrEF is mostly associated with ischemic heart disease with multiple risk factors such as hypertension, DM or smoking. 10,11 Patients with HFrEF had evidence of morphological changes such as an increase in the size of the left ventricle accompanied by an increase in end-systolic and end-diastolic volume, reduced wall thickness and increased left ventricular mass. The morphology in patients with HFrEF was opposite to that observed in patients with HFpEF (TABLE 2).12 Most of the differences in echocardiography parameters between HFrEF and HFpEF were significant (p < 0.05) i.e., LVID size $(64.9\pm7.9 \text{ vs } 49.2\pm7.1)$ mm) and IVSd (12.5±2.1 vs 10.3±2.6 mm).

ECG changes are usually found in patients with heart failure. This finding is supported by Karaye and Sani who reported ECG abnormalities in 98.2% of patients

with heart failure. Further, the majority of these patients (65.5%) had at least 3 types of ECG abnormalities.¹³ If the ECG was normal, the probability that a patient had systolic dysfunction (HFrEF) was small.^{5,6} Multivariate analysis showed that LAH, a wide QRS complex, RBBB, ST-T segment changes and a prolonged QT interval were independent predictors of systolic heart failure (HFrEF) (TABLE 4).

LAH on ECG was found in 25.5% of patients (28 out of the total number of patients). Of these, 78.5 % (22 cases; p=0.003) were from patients with HFrEF. This result is similar to that reported in a study by Karaye and Sani,13 in which LAH was more commonly found in patients with an ejection fraction <50% compared to those with an ejection fraction $\geq 50\%$ (77.5% vs 22.4%; p=0.001).¹³ In this study, a prolonged QRS duration >100 ms was predominantly found in patients with HFrEF compare to those with HFpEF (81.1% vs 18.9%), with values of 124±30.4 vs, 97.3±20.7 ms, respectively. Murkofsky et al. 14 showed that a prolonged QRS duration (> 0.10 second) is a very specific, though not a very

sensitive, indicator of left ventricular systolic function. A QRS duration of >0.10 seconds has a high likelihood of being associated with an ejection fraction <45 %.¹⁴

RBBB was mostly found in patients with HFpEF compared to those with HFrEF (70% vs 30%), while LBBB was found in 20% of patients with HFrEF and in no patients with HFpEF. This result is similar to the findings by Lee et al. 15 RBBB is caused by myocardial ischemia, infarction, inflammation (myocarditis), chronic increase in right ventricular pressure (cor pulmonale) and a sudden dilatation of the right ventricle (observed in acute pulmonale secondary to emboli). Less common causes of RBBB include hypertension, cardiomyopathy and congenital heart disease. 16 RBBB is also found in subjects without any underlying disease (isolated RBBB).¹⁷ In our study, the presence of RBBB in patients with HFpEF might have been related to age, coronary artery disease, hypertension or due to isolated RBBB.

ST-T segment changes were evident in 61 patients (55.4%). Of these, 68.8% (42 patients) were found in patients with HFrEF and 19 patients with HFpEF. This finding is supports similar findings by Basnet et al.,18 who reported that ST-T segment changes are a common ECG finding that is present in 48.57% of patients with left ventricular systolic dysfunction. Nielsen et al., 19 also found that ECGs with significant Q wave abnormalities, LBBB and ST-T segment changes (p <0.012) are associated with left ventricle systolic dysfunction. Strain pattern ST-T segment changes have a strong association with a left ventricle internal dimension (LVIDd) >55 mm compared to a posterior wall thickness >12 mm. This observation explains the strong association between strain pattern and ventricular hypertrophy with eccentric vs. concentric remodelling.20

A prolonged QTc interval was found in 69 patients (62.7%). Of these, 72.4% (50 patients) were patients with HFrEF. The average QTc interval in patients with HFrEF was also longer than that of patients with HFpEF (499±50.9 vs 453.2±42.8 ms; p=0.000). Wilcox et al., 21 showed that patients with grade II or III diastolic dysfunction have longer QTc intervals compare to patients with non-diastolic or grade I diastolic dysfunction (QTc 461±34 vs 432±32 ms; p=0.0003). In our study, most of the patients with HFrEF has grade III diastolic dysfunction (restrictive type) (48.3%), while 78% of patients with HFpEF had grade I diastolic dysfunction (relaxation type).

A limitation of our study is variable duration of heart failure therapy for each patient, though it is difficult to determine the impact this would have on our results.

CONCLUSION

Our study suggests that a scoring system based on ECG findings that include the presence or absence of LAH, a wide QRS duration, RBBB, ST-T segment changes and a prolonged extended QTc interval can be used to predict the type of heart failure (HFpEF and HFrEF) in patients with chronic heart failure. This scoring system has a sensitivity of 76%, a specificity of 96%, a positive predictive value of 95%, a negative predictive value of 80% and an accuracy of 86%. A score of -1 to +3 suggests the possibility of HFpEF, while a score of +4 to +6 suggests the possibility of HFrEF.

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