

# Chronic heart failure: different treatment methods and their 6-month outcomes



## Abstract

Chronic Heart Failure (CHF) remains one of the most important problems in cardiology, despite the availability of various modern diagnostic methods and a number of advances in treatment. This is due to its widespread use, lowering the quality of life of patients, as well as high rates of recurrent decompensation and death. Despite the optimal use of modern treatments based on proven medical principles, the disease still has a high morbidity and mortality rate [1-6].

**The aims:** The aim of our study was to evaluate the efficacy (activity, BNP levels, etc.) of a pathogenetically complementary conservative treatment with the inclusion of sacubitril/valsartan in the treatment of patients with chronic heart failure in comparison with device-based CRT.

**Materials and methods:** The study included 64 patients over the age of 38 suffering from Chronic Heart Failure (CHF) (45 men, 19 women,  $59.5 \pm 0.9$  years of age). Patients were divided into basic and control groups. 33 patients were included in the main group. In the main group, patients received sacubitril/valsartan twice daily in addition to the classic conservative treatment of CHF. The control group included 31 patients who underwent CRT surgery. During the study, the clinical performance of patients before and after 6 months of treatment, the results of BNP tests, the results of a 6 minute walking test and EcoKG were compared.

**Conclusion:** Evaluation of the results of examinations of patients after 6 months revealed positive changes in the clinical indicators of the majority of patients in both groups compared to 6 months ago. However, better results were obtained in the main group than in the control group.

**Keywords:** CHF, BNP, sacubitril/valsartan, CRT

## Introduction

In the treatment of patients with CHF, our main goal is to improve the clinical condition of patients, increase their functional capacity and quality of life, prevent re-hospitalization and, most importantly, reduce the number of deaths [3,4,7,8]. Many new drugs and devices are currently being used to treat patients with chronic heart failure [9].

Modern principles of existing pharmacological treatments are based on the pathogenetic concept of CHF, which develops as a result of long-term activation of the neurohumoral system. These include, first of all, renin-angiotensin-aldosterone and sympathetic-adrenal systems, which are considered pathognomonic in patients with chronic heart failure with poor prognosis. Theoretically, the combined use of different groups of neurohumoral modulators may provide additional benefits in the treatment of patients with chronic heart failure as a result

of a more complete blockade of neurohormones. The essence of such a concept is very simple, so the higher the level of different levels of neurohumoral regulation, the better the result [9].

In recent years, a new pharmacological drug has been used in the conservative treatment of patients with chronic heart failure with a reduced ejection fraction. This pharmacological drug is a pharmacological agent that can provide simultaneous blockade of both the angiotensin system and neprilysin. Recently, a number of studies have been conducted on this drug, and a series of studies are ongoing.

In addition to drug treatment, the device is widely used in modern therapies. Of these, resynchron heart therapy is the most widely used treatment in recent years in all countries of the worlds.

In patients with moderate to severe heart failure, CRT treatment may improve quality of life in

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two-thirds of patients and prolong life in one-third [3].

However, not all patients receiving this treatment respond positively to the CRT method. A number of features can affect the course of the disease after this treatment and the mortality rate. For example, in patients with ischemic etiology, left ventricular function develops less positively after this treatment due to scar tissue of the myocardium. This reduces the likelihood of favorable remodeling during the use of CRT in such patients [10].

In previous years, according to the guidelines of the European Society of Cardiology, in patients preparing for implantation of a sinus rhythm SRT device, a QRS width of more than 130 ms could be considered a SRT [3]. However, a number of studies published in recent years in 2018-2019, as well as the recommendations of the American Heart Association for 2018, require that this figure be strictly higher than 150 ms [11].

## Materials and methods

The study included 64 patients over the age of 38 who were treated at the Eurasia Hospital with a diagnosis of CHF. The diagnosis of CHF was confirmed on the basis of anamnesis, objective and instrumental examination methods.

Eligibility criteria: History of chronic heart failure; circulatory failure (functional class II-IV, NHYA); left ventricular ejection fraction <40%.

Exclusion criteria: Acute myocardial infarction; hypertrophic cardiomyopathy; congenital heart defects; Patients under 25 years of age; heart failure in oncology patients.

According to the admission criteria, a total of 64 patients were included in the study, 45 men (70.3% ± 5.7%) and 19 women (29.7% ± 5.7%). The mean age of the patients was 59.5 ± 0.9. Of the 64 patients, 3 (4.7%) had 2<sup>nd</sup> f.c., 54 (84.4%) had 3<sup>rd</sup> f.c., and 7 (10.9%) had 4 f.c. suffers from CHF **FIGURE 1**.

During the study, each patient in the main group was given a combination of sacubitril/valsartan twice a day for 6 months in addition to the traditional conservative treatment of CHF (antiarrhythmic, hypolipidemic, anticoagulant, diuretic, hypotensive). Patients in the control group underwent CRT surgery in addition to receiving classic conservative treatment (excluding sakubitril/valsartan). Demographic and clinical characteristics of the patients included in the study are given in **TABLE 1**.

The differences between the study groups were not statistically significant. Thus, for all indicators it was  $p > 0.05$ . Clinical findings of patients before and 6 months after treatment (anamnesic data, SAH and DAH, pulse rate and fullness, physical examination), duration of initial compensation in both groups and the number of recurrent decompensations, SaO<sub>2</sub>, BNP analysis results, results of the 6-minute walking test and EcoKG were evaluated comparatively. Statistical analysis included variance analysis (ANOVA Test), Wilcoxon Signed Ranks test, Cross analysis (Pearson Chi-Square Test) and Mann-Whitney test **TABLE 2 and TABLE 3**.

## Result

We concluded that the addition of sacubitril/valsartan complex to the treatment of patients

**FIGURE 1. The mean age of the patients suffers from CHF.**

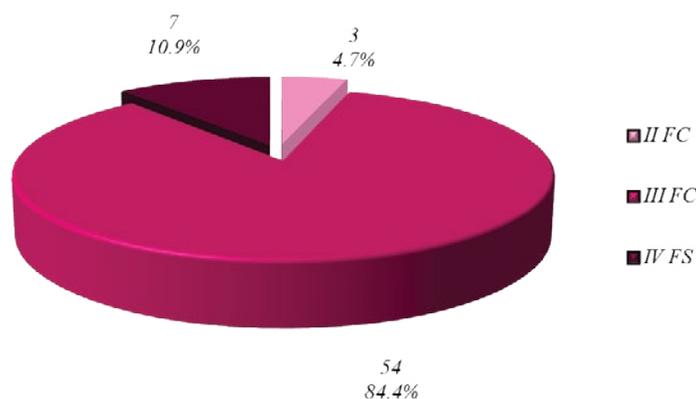


TABLE 1. Demographic and clinical characteristics of patients.

Characteristics		Groups		P <sub>0</sub>	
		I group	II group		
		(n=33)	(n=31)		
Age. years		59.6% ± 1.3% (38-70)	59.5% ± 1.4% (39-73)	0.909	
Sex	Male	25 75.8%	20 64.5%	0.329	
	Female	8 24.2%	11 35.5%		
BMI. kg/m <sup>2</sup>		36.9% ± 0.5% (31.6-43.6)	35.9% ± 0.3% (32.1-38.7)	0.375	
Obesity	I grade	8 24.2% ± 7.5%	6 19.4% ± 7.1%	0.310	
	II grade	18 54.5% ± 8.7%	25 80.6% ± 7.1%		
	III grade	7 21.2% ± 7.1%	-		
Action	Active	4 12.1% ± 5.7%	4 12.9% ± 6.0%	0.925	
	Inactive	29 87.9% ± 5.7%	27 87.1% ± 6.0%		
Smoking	Doesn't smoke	8 24.2% ± 7.5%	11 35.5% ± 8.6%	0.685	
	Less Smoking	8 24.2% ± 7.5%	4 12.9% ± 6.0%		
	Smoking a lot	17 51.5% ± 8.7%	16 51.6% ± 9.0%		
Diabetes mellitus		27 81.8% ± 6.7%	26 83.9% ± 6.6%	0.829	
Arterial hypertension		21 63.6% ± 8.4%	20 64.5% ± 8.6%	0.942	
Family	Mother	5 15.2% ± 6.2%	7 22.6% ± 7.5%	0.975	
	Father	8 24.2% ± 7.5%	4 12.9% ± 6.0%		
	Both	20 60.6% ± 8.5%	20 64.5 ± 8.6%		
Circulatory failure; NYHA	II FC	2 6.1% ± 4.2%	1 3.2% ± 3.2%	0.983	
	III FC	27 81.8% ± 6.7%	27 87.1% ± 6.0%		
	IV FC	4 12.1% ± 5.7%	3 9.7% ± 5.3%		

Note: The difference between the groups is not statistically significant.

led to a decrease in the level of BNP in the blood of patients to lower levels and, consequently, to further regulation of the pathogenetic system. Repeated Echocardiography of patients showed

a positive dynamics in the process of remodeling of the left ventricle in the main group. The 6-minute walking test proved to have a better effect on patients' mobility than other treatments.

TABLE 2. Wilcoxon signed ranks test.

Ranks	Pf	Pu	Px <sup>2</sup>	Pw	
				Gr 1	Gr 2
Shortness of breath		0.890	0.99		
Shortness of breath a		0.012	0.043	0	0
Heartbeat		0.325	0.107		
Heartbeat a		0.298	0.076	0	0
Cough		0.713	0.068		
Cough a		0.021	0.063	0	0.001
pulmonary auscultation		0.056	0.054		
pulmonary auscultation a		0.707	0.17	0	0
edema in the legs		0.399	0.395		
edema in the legs a		0.000	0.001	0	0
Pulse fullness		0.877	0.876		
Pulse fullness a		0.424	0.42	0.317	0.046
Pulse rate	0.921	0.845			
Pulse rate a	0.931	0.772		0	0
SaO <sub>2</sub>	0.104	0.138			
SaO <sub>2</sub> a	0.325	0.439		0	0
QRS	0.583	0.655			
Decompensation	0.579	0.373			
Decompensation a	0.668	0.859		0	0
BNP	0.766	0.687			
BNP a	0	0		0	0
6 min. walk test		0.189	0.185		
6 min. walk test a		0.002	0.001	0	0
FC		0.983	0.813		
FC a		0.017	0.083	0	0
Initial compensation period		0.050	0.065		
SAH	0.46	0.620			
SAH a	0.529	0.937		0	0
DAH	0.526	0.718			
DAH a	0.567	0.573		0	0

TABLE 3. ExoKG indicators of groups.

Sings.	Groups	Count	M	± m	min	max	P <sub>F</sub>	P <sub>U</sub>	P <sub>w</sub>
ESD	Group 1	33	41.4	0.9	33	50	0.158	0.068	
	Group 2	31	43.3	0.9	30	55			
ESD a	Group 1	33	39.8	1	32	50	0.01	0.006	<0.001
	Group 2	31	43.5	1	29	56			0.197
EDD	Group 1	33	60.5	0.5	57	66	0.161	0.133	
	Group 2	31	61.6	0.6	55	67			
EDD a	Group 1	33	58.3	0.6	53	66	<0.001	0.001	<0.001
	Group 2	31	61.8	0.7	55	67			0.142
EFLW	Group 1	33	26.9	0.9	15	33	0.957	0.651	
	Group 2	31	26.8	1.1	15	38			
EFLW a	Group 1	33	32.8	1.1	20	44	0.257	0.184	<0.001
	Group 2	31	31	1.2	19	45			<0.001
EAD	Group 1	33	39.2	0.7	32	51	0.056	0.055	
	Group 2	31	41	0.7	34	52			
SPA <sub>h</sub>	Group 1	33	28.8	1.1	20	41	0.824	0.951	
	Group 2	31	29.1	0.8	23	40			
SPA <sub>h</sub> a	Group 1	33	26.5	0.9	18	39	0.158	0.354	0.001
	Group 2	31	28.1	0.6	23	36			0.002

## Discussion

As can be seen from the tables above, all

patients' pulse and blood pressure, anamnesis, physical examination results, 6 minute walking test results, as well as EcoGq results, and B-type

natriuretic peptide levels in the blood were examined in detail by statistical analysis. Both qualitative and quantitative tests were used in statistical analysis.

During the Wilxson test, statistically significant differences were obtained in the results of other indicators after 6 months, except for pulse saturation in group 1  $p < 0.05$ . The calculation of pulse saturation results before and after 6 months did not make a statistically significant difference ( $p = 0.317$ ). There are also positive changes in the comparison of pre- and post-treatment outcomes of patients in group 2. Thus, statistically significant changes were obtained on most indicators during the statistical analysis in this group as well.  $p < 0.05$  only the Echocardiogram did not show a statistically significant difference in the size of the left ventricle before and after 6 months  $p = 0.197$ ,  $p = 0.142$ . However, in this group there was a statistically significant difference in the results of LVEF  $p < 0.001$ . That is, positive results were obtained from the treatments performed in both groups separately. 6 months later intergroup analysis of patients' results was performed with the Mann-Whitney Test, ANOVA Test, and Pearson Chi-Square Test. Although these analyzes did not show statistically significant differences in all indicators, statistically significant changes were obtained in some indicators (history of shortness of breath, cough, lower peripheral

edema during physical examination, number of recurrent decompensations, etc.). The most important of these were BNP levels in blood, initial compensation period, 6-minute walking test and echocardiography. In the intergroup analyzes, the Mann-Whitney test showed statistically significant differences in BNP levels, initial compensation duration, 6-minute walking test and some EcoGq indicators  $p = 0.000$ ,  $p = 0.50$ ,  $p = 0.002$ ,  $p = 0.006$ ,  $p = 0.001$ . The Pearson Chi-Square Test, a qualitative analysis, did not show a statistically significant difference in most indicators, in the initial compensation period either ( $p = 0.065$ ). However, a statistically significant difference was obtained in the indicators of 6-minute walking test.  $p = 0.001$ . Although the ANOVA test, a dispersion analysis, did not show statistically significant differences in other parameters, a statistically significant difference was obtained in BNP and left ventricular remodeling (ESD and EDD).  $p = 0.000$ ,  $p = 0.010$ ,  $p < 0.001$ .

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## Conclusion

Evaluation of the results of examinations of patients after 6 months revealed positive changes in the clinical indicators of the majority of patients in both groups compared to 6 months ago. However, better results were obtained in the main group than in the control group.

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