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IMPACT OF SACUBITRIL/VALSARTAN COMBINATION ON PREVALENCE OF ARRHYTHMIA IN HEART FAILURE PATIENTS WITH REDUCED EJECTION FRACTION

By

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ABSTRACT

Background: In the PARADIGM-HF trial, Heart failure with reduced ejection fraction (HFrEF) patients on (sacubitril/valsartan) had a substantially lower rate of hospitalization for HF and mortality compared to Enalapril. Only very few data exist regarding the impact of sacubitril/valsartan on arrhythmia in those patients.

Objective: To assess the effect of Sacubitril/Valsartan combination on prevalence of arrhythmias in HFrEF patients and compare it with patients on Angiotensin converting enzyme inhibitor (ACEI) or Angiotensin receptors blockers (ARBs).

Patients and Methods: Sixty patients with heart failure with reduced ejection fraction were classified into: Group A on Sacubitril\Valsartan combination therapy, and group B on ACEI or ARBs for at least 3 months with follow up as regard the burden of arrhythmia by 48 hours holter monitoring at 3 and 6 months.

Results: There was no significant difference in both groups as regard demographic and medical data Group A was characterized by significant decrease in ventricular ectopics and significant improvement in ejection fraction (EF), Left ventricular internal dimension (IVIDd) and right ventricular systolic pressure (RVSP) (p=0.00) in comparison to group B.

Conclusion: Sacubitril/Valsartan combination therapy was superior to ACEIs or ARBs in reducing ventricular ectiopic as a mechanism for preventing of sudden cardic death (SCD).

Keywords: HFrEF, ACEI, ARBs-SCD, Left atrium, LVIDd, RVSP.

INTRODUCTION

The treatment of chronic heart failure with reduced LVEF (HFrEF) using angiotensin- converting enzyme (ACEI) inhibitors is well established, with angiotensin receptor blockers (ARBs) as a safe and proven alternative (Europian society of cardiology guidelines 2012& 2016). New therapeutic class of agents acting on the renin angiotensin

aldosterone system (RAAS) and neutral endopeptidase system has been developed, known as angiotensin receptor neprilysin inhibitors (ARNI) (McMurray et al., 2014). In the PARADIGM-HF trial, HFrEF patients on (sacubitril/valsartan) had a substantially lower rate of hospitalization for HF and mortality compared to Enalapril (King et al., 2015).

A subanalysis of the PARADIGM-HF trial also showed a reduction in sudden

cardiac death by 20% in relation to Enalapril, which does not differ among patients with or without an implantable cardioverter defibrillator (Mangiafico and Costello-Boerrigter 2013). Several potential mechanisms have been linked to this antiarrhythmic effect. Whether this reduction is caused by reverse remodeling, the reduction in myocardial fibrosis, wall stretch sympathetic nervous system activation is not fully understood (de Diego and Gonzalez-Torres 2018 and Martens and Nuyens, 2019).

The authors in this study assess the effect of valsartan \ sacubitril combination on incidence of arrhythmias in HFrEF patients in comparison to patients on ACEI or ARBs only.

Study population: This study conducted at Ain Shams university and Nasr city insurance hospitals including 60 patients with heart failure patients with reduced ejection fraction less than 40% either ischemic (proved by history, metabolic imaging coronary or angiography) or dilated. This was on either Sacubitril\Valsartan combination therapy or on ACEI or ARBs for at least 3 months. Patients with renal impairment, hyperkalemia, pace maker or CRT were excluded.

Ethics approval and informed consent:

The study protocol was approved by Ain Shams Faculty of Medicine scientific and ethical committee. Data confidentiality and privacy were maintained. All patients were informed about the registry and written consent were taken.

PATIENTS AND METHODS

This was an observational prospective study with random sampling of patients with collection of full data including medical histoty, ECG, echocardiography and laboratory data. Follow up the patients at 3, 6 months after initiation of Valsartan\ Sacubitril combination therapy, or ACEIs (ARBs) at maximum tolerated dose by up titration of the dose according the tolerance of patients assessment of blood pressure and also guided by blood investigations including serum creatinine and potassium level. The follow up included the medical status, compliance, ECG. echocardiography, laboratory investigation and holter monitoring.

Statistical analysis: Using computer software statistical package for the social sciences (SPSS, version 20, SPSS Inc., Chicago, Illinois, USA) Description of quantitative (numerical) variables was performed in the form of mean \pm SD. Description of qualitative (categorical) data was performed in the form of number of cases and percent. Appropriate test of associations was performed using Chisquare test, Paired t-test. Wilcoxon signed-rank repeated test, measure ANOVA test and Friedman test the significance level was set at p-value of less than 0.05.

RESULTS

The number of samples that fulfilled the inclusion and exclusion criteria in this study were 60 patients that were classified into 2 groups: 30 patients on Sacubitril\Valsartan combination therapy

(group A), and 30 patients on ACEIs or ARBs (group B).

There was no significant difference in demographic and clinical data of both groups (**Table 1**).

Table (1): Demographic and clinical data of patients

	Group A	Grou	ıp A	Gr	oup B	D volvo	
Parameters		No. = 30		No. = 30		P-value	
Λ σο	Mean ± SD	58.23	± 4.80	56.3	7 ± 4.49	0.125	
Age	Range	48 - 65		49	0-64	0.125	
Gender	Female	8 (26	.7%)	6 (2	20.0%)	0.542	
Gender	Male	22 (73	3.3%)	24 (80.0%)	0.342	
		No	%	No.	%		
HTN	No	5	16.7%	5	16.7%	1.000	
пін	Yes	25	83.3%	25	83.3%	1.000	
Controlled	No	1	4.0%	3	12.5%	0.277	
Controlled	Yes	24	96.0%	21	87.5%	0.277	
DM	No	15	50.0%	11	36.7%	0.207	
DM	Yes	15	50.0%	19	63.3%	0.297	
Com (11	No	4	26.7%	10	52.6%	0.107	
Controlled	Yes	11	73.3%	9	47.4%	0.127	
C 1	No	9	30.0%	9	30.0%	1 000	
Smoker	Yes	21	70.0%	21	70.0%	1.000	
шь	No	4	13.3%	3	10.0%	0.600	
IHD	Yes	26	86.7%	27	90.0%	0.688	
Alcoholic	No	30	100.0%	30	100.0%	NA	
, , , , , , , , , , , , , , , , , , ,	No	27	90.0%	28	93.3%	0.640	
VHD	Yes	3	10.0%	2	6.7%		
arr.	No	29	96.7%	26	86.7%		
CKD	Yes	1	3.3%	4	13.3%	0.161	
CV.disease	e No	30	100.0%	30	100.0%	NA	
D 11 11	. No	7	23.3%	12	40.0%	0.165	
Dyslipidem	1a Yes	23	76.7%	18	60.0%	0.165	
	No	10	33.3%	15	50.0%	0.400	
FH (IHD/SC	(D) Yes	20	66.7%	15	50.0%	0.190	
	NYHA I	0	0.0%	1	3.3%		
NTX/TT A 1	NVHAII	21	70.0%	21	70.0%		
NYHA clas	NYHA III	9	30.0%	8	26.7%	0.589	
	NYHA IV	0	0	0	0		
Prior MI	No	6	20.0%	4	13.3%	0.488	
FIIOT IVII	Yes	24	80.0%	26	86.7%	0.488	
Drice DCI	No	10	33.3%	6	20.0%	0.242	
Prior PCI	Yes	20	66.7%	24	80.0%	0.243	
Prior CAB	G No	22	73.3%	27	90.0%	0.095	
THUI CAD	Yes	8	26.7%	3	10.0%	0.093	
Cardiomyopa	Non-ICM	4	13.3%	3	10.0%	0.687	
Caruioniyopa	ICM	26	86.7%	27	90.0%	0.687	

Through six months of follow up, Group A was characterized by nonsignificant reduction in QRS width and non-significant shortening in QTc while

Group B was associated with non-significant increasing QTc values through the follow up period (**Tables 2 & 3**).

Table (2): ECG changes in group A through the follow up period

	Group A	Initial	3 months	6 months	D volue	
Parameters		No. = 30	No. = 30	No. = 30	P-value	
HR	Mean ± SD	75.60 ± 8.51	73.17 ± 6.75	72.57 ± 7.20	0.168	
ПK	Range	63 - 100	65 – 90	60 – 96	0.100	
Dhydhm	AF	5 (16.7%)	7 (23.3%)	7 (23.3%)	0.766	
Rhythm	Sinus	25 (83.3%)	23 (76.7%)	23 (76.7%)	0.700	
DD interval	Mean ± SD	125.60 ± 19.81	129.57 ± 31.98	127.39 ± 27.67	0.623	
PR interval Range		100 - 160	80 - 240	80 - 220	0.023	
ODC: 14h	Mean ± SD	111.00 ± 19.18	109.67 ± 17.12	109.66 ± 17.21	0.416	
QRS width	Range	80 - 160	80 - 130	80 - 130	0.410	
ОТа	Mean ± SD	425.20 ± 26.94	425.70 ± 27.06	424.27 ± 22.16	0.774	
QTc	Range	382 - 480	382 - 477	382 – 453	0.774	
	Inverted t	23 (76.7%)	24 (80.0%)	23 (76.7%)	0.938	
T wave	Bihasic t	9 (30.0%)	6 (20.0%)	7 (23.3%)	0.656	
morphology	Flat t	4 (13.3%)	6 (20.0%)	5 (16.7%)	0.787	
	Norml T	2 (6.7%)	2 (6.7%)	2 (6.7%)	1.000	
PVCs	No	27 (90.0%)	26 (86.7%)	28 (93.3%)	0.690	
FVCS	Yes	3 (10.0%)	4 (13.3%)	2 (6.7%)	0.090	

Table (3): ECG changes in group B through the follow up period

	Group B	Initial	3 months	6 months		
Parameters	Oroup D	No. = 30	No. = 30	No. = 29	P-value	
ш	Mean ± SD	72.50 ± 6.68	72.27 ± 6.76	71.38 ± 6.33	0.713	
HR	Range	60 – 85	55 – 90	65 – 95	0.712	
Dl41	AF	3 (10.0%)	5 (16.7%)	8 (27.6%)	0.207	
Rhythm	Sinus	27 (90.0%)	25 (83.3%)	21 (72.4%)	0.207	
DD :41	Mean ± SD	137.41±18.73	134.40 ± 19.38	134.29 ± 17.77	0.251	
PR interval	Range	120 – 180	100 – 180	120 – 180	0.251	
ODC: 44h	Mean ± SD	118.17 ± 12.21	120.17 ± 11.48	119.66 ± 12.39	0.007	
QRS width	Range	100 – 140	100 – 140	100 – 140	0.087	
ОТо	Mean ± SD	418.37 ± 22.18	419.43 ± 23.85	419.21 ± 23.00	0.729	
QTc	Range	384 – 463	369 – 462	381 – 464	0.738	
	Inverted t	26 (86.7%)	26 (86.7%)	24 (80.0%)	0.887	
T wave	Bihasic t	3 (10.0%)	2 (6.7%)	2 (6.7%)	0.867	
morphology	Flat t	2 (6.7%)	2 (6.7%)	2 (6.7%)	1.000	
	Norml T	4 (13.3%)	4 (13.3%)	4 (13.3%)	1.000	
PVCs	No	25 (83.3%)	23 (79.3%)	24 (82.8%)	0.911	

Through six months of follow up, group A show significant improvement in EF (P=<0.001), RVSP (P=<0.001) and SWMA (P=0.005) with maximal improvement after 6 months While LA diameter increased after 3 month then declined again after the 6 month (p=0.02)

while in group B Mean left atrium diameter was found to be increasing through the follow up period with highest values after six months. And RVSP was found to be improving through the follow up period with lowest values after six months (**Tables 4, 5, 6 & 7**).

Table (4): Echocardiography changes in group A through the follow up period

	Group A	Initial	3 months	6 months	P-value	
Paramete	ers	No. = 30	No. = 30	No. = 30	P-value	
EF	Mean \pm SD	29.77 ± 4.09	31.57 ± 4.49	33.83 ± 4.83	< 0.001	
ЕГ	Range	22 - 38	22 - 40	24 - 45	<0.001	
LA	Mean \pm SD	4.67 ± 0.53	4.82 ± 0.55	4.75 ± 0.50	0.029	
LA	Range	3.95.8	4.1 - 5.9	4 - 5.7	0.029	
LVIDd	Mean \pm SD	6.11 ± 0.48	6.01 ± 0.57	6.05 ± 0.56	0.068	
LVIDa	Range	5.57.3	4.8 - 7.2	5.3 - 7.5	0.008	
LVIDs	Mean ± SD	4.61 ± 0.57	4.61 ± 0.49	4.64 ± 0.47	N 969	
LVIDS	Range	3.35.8	3.3 - 5.9	3.3 - 5.7	0.868	
RVSP	Mean \pm SD	47.40 ± 10.21	42.27 ± 7.53	39.73 ± 7.05	< 0.001	
KVSF	Range	25 - 65	28 - 60	25 - 55	<0.001	
SWMA	Median (IQR)	7(3-7)	3 (2 –7)	3 (2 –7)	0.005	
SWIM	Range	2 - 7	2 –7	2 - 7	0.003	
	Mild	13 (43.3%)	13 (43.3%)	16 (53.3%)		
MR	Moderate	16 (53.3%)	17 (56.7%)	14 (46.7%)	0.605	
	Trivial	1 (3.3%)	0 (0.0%)	0 (0.0%)		
	Mild	5 (16.7%)	7 (23.3%)	8 (26.7%)		
TR	Moderate	15 (50.0%)	18 (60.0%)	20 (66.7%)	0.173	
1 K	Severe	8 (26.7%)	5 (16.7%)	2 (6.7%)	0.175	
	Moderate to severe	2 (6.7%)	0 (0.0%)	0 (0.0%)		

Table (5):Post Hoc analysis of significant echocardiography changes in group A through the follow up period

Post Hoc analysis	Initial Vs 3 months	Initial Vs 6 months	3 months Vs 6 months
EF	0.001	0.000	0.005
LA	0.028	0.084	0.081
RVSP	0.001	0.000	0.001
SWMA	0.014	0.010	0.085

Table (6): Echocardiography changes in group B through the follow up period

	Group B	Initial	3 months	6 months	D l	
Parameters		No. = 30	No. = 30	No. = 29	P-value	
EF	Mean ± SD	32.43 ± 4.58	33.10 ± 4.68	32.72 ± 5.71	0.338	
EF	Range	22 - 40	24 - 40	22 - 45	0.338	
LA	Mean ± SD	4.39 ± 0.44	4.40 ± 0.39	4.48 ± 0.36	0.026	
LA	Range	3.7 - 5.7	3.8 - 5.6	3.9 - 5.5	0.026	
LVIDd	Mean ± SD	6.34 ± 0.44	6.31 ± 0.47	6.32 ± 0.51	0.548	
LVIDa	Range	5.5 - 7.4	5.5 - 7.2	5.4 - 7.5	0.348	
LVIDs	Mean \pm SD	4.57 ± 0.69	4.55 ± 0.60	4.50 ± 0.50	0.670	
LVIDS	Range	3.9 - 6.9	3.7 - 6.3	3.8 - 6.1	0.670	
RVSP	Mean \pm SD	45.63 ± 9.89	41.97 ± 9.00	40.79 ± 8.55	0.000 *	
KVSP	Range	30 - 65	25 - 60	25 - 60	0.000 **	
SWMA	Median (IQR)	7 (2 –7)	5 (2 –7)	3 (2 –7)	0.076	
SWMA	Range	1 –7	1 –7	1 –7	0.076	
	No	0 (0.0%)	0 (0.0%)	1 (3.4%)		
MR	Mild	13 (43.3%)	11 (36.7%)	9 (31.0%)	0.579	
	Moderate	17 (56.7%)	19 (63.3%)	19 (65.5%)		
	Mild	5 (16.7%)	6 (20.0%)	6 (20.7%)		
TR	Moderate	16 (53.3%)	13 (43.3%)	13 (44.8%)	0.949	
	Severe	9 (30.0%)	11 (36.7%)	10 (34.5%)		

Table (7): Post Hoc analysis of significant echocardiography changes in group B through the follow up period

Post Hoc analysis		Initial Vs 6 months	3 months Vs 6 months
LA	0.672	0.017	0.015
RVSP	0.001	0.001	0.232

There was no significant difference through the follow up period in group A populations regarding serum creatinine and potassium level while significant elevation in serum creatinine in group B (p=0.001) (**Tables 8, 9 & 10**).

Table (8): Kidney function and serum potassium changes in group A and B through the follow up period

	Group A	Initial	3 month	6 month	P-value
Parameters		No. = 30	No. = 30	No. = 30	r-value
C	Median (IQR)	1.1 (0.9 – 1.3)	1.15 (0.8 – 1.3)	1 (0.8 – 1.1)	0.672
Serum cr.	Range	0.5 - 1.6	0.5 - 1.7	0.5 - 1.6	0.672
Corum V	Mean ± SD	4.43 ± 0.56	4.38 ± 0.54	4.28 ± 0.41	0.224
Serum K	Range	3.3 - 5.3	3.3 - 5.2	3.5 - 5	0.234

Table (9): Kidney function and serum potassium changes in group B through the follow up period

	Group B	Initial	3 month	6 month	D volue
parameters		No. = 30	No. = 29	No. = 29	P-value
Comum or	Median (IQR)	1.3 (1 – 1.6)	1.4 (1.1 – 1.7)	1.5 (1.2 – 1.8)	0.001
Serum cr	Range	0.8 - 2.2	0.9 - 2.4	0.9 - 2.5	0.001
Serum K	Mean ± SD	4.35 ± 0.45	4.29 ± 0.46	4.33 ± 0.40	0.457
	Range	3.2 - 5.1	3.2 - 5.1	3.7 - 5.1	0.457

Table (10): Post Hoc analysis of significant kidney function in group B through the follow up period

Periods			
Post Hoc analysis	Initial Vs 3 months	Initial Vs 6 months	3 months Vs 6 months
Serum cr	0.008	0.002	0.056

There was no significant difference through the follow up period in group A populations regarding clinical data while

one mortality was in group B (tables 11& 12).

Table (11): Clinical data of group A through the follow up period

Group A Parameters	Clinical data after 3 months	Clinical data after 6 months	P-value
Hospitalization and decompensation	1 (3.3%)	1 (3.3%)	1.000
SCD	0 (0.0%)	0 (0.0%)	_

Table (12): Clinical data of group B through the follow up period

Group I	Clinical data after	Clinical data after	P-value
Parameters	3 months	6 months	1 -value
Hospitalization and decompensation	3 (10.3%)	4 (13.8%)	0.687
SCD	0 (0.0%)	1 (3.4%)	0.313

Burden of ventricular ectopics was found to be significantly lowered (p=0.01) through the follow up period with lower

incidence and percentage after six months of drug use (Tables 13& 14).

Table (13): Holter study changes through the follow up period in group A

	Group A	3 months	6 months	P-value
Parameters		No. = 30	No. = 30	
Ventricular ectopics	Median (IQR)	2511 (823 – 6312)	2022 (416 – 4216)	0.012
	Range	19 – 16422	10 - 18411	
Percentage	Median (IQR)	2.1 (0.4 – 3.1)	1.09(0.06-2.93)	0.013
	Range	0.01 - 12	0 - 175	
Bigemini	Median (IQR)	126.5 (24 – 584)	82 (3 – 778)	0.674
	Range	1 – 10366	1 - 861	
Trigemini	Median (IQR)	67 (18 – 1011)	53 (2 – 522)	0.062
	Range	1 - 4232	1 - 2206	
Couplets	Median (IQR)	3 (2 – 35)	7 (2.5 – 83)	0.753
	Range	1 - 428	2 - 420	
Triplets	Median (IQR)	3 (1 – 30)	21 (1 – 410)	0.180
	Range	1 – 613	1 - 410	
Runs of nsvt	Median (IQR)	4 (1 – 12)	4 (3 – 10)	0.279
	Range	1 - 24	1 - 26	

Table (14): Holter study changes through the follow up period in group B

	Group B	3 months	6 months	P-value
Parameters		No. = 29	No. = 29	P-value
ventricular ectopics	Median (IQR)	1221 (689 – 7624)	1878.5 (632.5 – 5237)	
	Range	17 - 34582	201 - 28674	0.891
Percentage	Median (IQR)	0.9(0.3-4.1)	0.9(0.3-4.1)	
	Range	0 - 17	0 - 17	1.000
Bigemini	Median (IQR)	19.5 (9 – 42.5)	19 (3 – 23)	
	Range	2 - 408	1 - 475	0.158
Trigemini	Median (IQR)	17 (6 – 45)	15 (3 – 45)	
	Range	2 - 1203	2 - 1400	0.972
Couplets	Median (IQR)	2 (1 – 13)	2.5 (2 – 7)	
	Range	1 - 25	1 – 31	0.443
Triplets	Median (IQR)	2(2-3)	2 (2 – 13)	
	Range	1 - 4	1 - 14	1.000
Runs of NSVT	Median (IQR)	7 (3 – 14)	5 (3 – 11)	
	Range	1 - 22	1 - 22	0.017

DISCUSSION

In the PARADIGM-HF trial, sudden cardiac death (SCD) more decreased in the sacubitril/valsartan as compared with ACEIs. A further prospective study recruited patients with HFrEF, treated with sacubitril/valsartan, and compared them with the treatment data on ACEI and/or angiotensin receptor blocker

(ARBs) (de Diego and Gonzalez-Torres, 2018).

However, published data presented six cases of ventricular arrhythmic storm shortly after initiating sacubitril/valsartan that required drug withdrawal *Vicent et al.*, (2019).

No systematic analysis of the incidence of ventricular tachyarrhythmia in patients treated with sacubitril/valsartan in a sufficient number of patients with longterm follow-up has been conducted yet.

We conducted a prospective observational study on 60 patients with failure with reduced ejection fraction, divided into two groups; 30 patients Sacubitril/Valsartan on combination, and 30 patients on ACEIs or ARBs to evaluate the incidence of ventricular arrhythmia in each group by with follow Holter. up Electrocardiogram and Echocardiography within 6 months after drug use. To assess to what extent the patient can benefit from medications to reduce hospitalization and prevent SCD. There was no impact of risk factors among our patients on incidence of arrhythmias in both groups. We missed one of our patients in ACEIs group due to sudden cardiac death in home. But in Sacubitril/Valsartan group we did not lose any of our patients. There was no significant difference regarding hospitalization on top of decomposition between both groups.

They have found that the functional capacity of Sacubitril/Valsartan groups was improving through the follow up period by documenting their quality of life and according to NYHA classification of HF. Also there was an antiarrhythmic effect of Sacubitril/Valsartan combination therapy, compared to ACEIs characterized by reduced burden of ventricular ectopics through the follow up period by Holter monitoring. The frequency of ectopics after the first three months was 2511 and became 2022 ectopics after 6 months that represent a significant difference with P value 0.012. These results consistent with the study of Martens and Nuyens (2019) that concluded a decrease in the burden of ventricular arrhythmias, as assessed by ICD monitoring.

In contrast, ACEIs groups did not show reduced frequencies of ventricular ectopics through the follow up period and in contrast the ectopics increased from mean 1221 ectopics in the first three months to become 1878.5 ectopics after 6 months of initiations of medications which represents non-significant difference with P value 0.891 and it was consistent with de Diego and Gonzalez-(2018)who found that Torres Angiotensin-neprilysin inhibition decreased ventricular arrhythmias and appropriate ICD shocks in HFrEF patients under home monitoring compared to angiotensin inhibition Sacubitril/Valsartan group.

We found non-significant decrease in OTc values which reached the lowest values after 6 months. In contrast ACEIs group was associated with non-significant increase in QTc values through the follow up period to reach the highest values after six months. It was consistent with Valentim Gonçalves (2019) who found that QTc interval were significantly reduced by 5.7% in Sacubitril/Valsartan combination therapy in HFrEF. Runs of NSVT in ACEIs group but not in Sacubitril/Valsartan combination decreased during the follow up; it was against the study of Martens and Nuyens (2019) that concluded that runs of NSVT are higher in **ACEIs** than Sacubitril/Valsartan combination.

Ejection fraction was found to be increased in Sacubitril/Valsartan group through the follow up period with high significant difference between its values before drug use and 6 months after

treatment, its initial mean EF was 29.77±4.09% and became 31.57±4.49% after 3 months and reached 33.83±4.83% after 6 months of treatment and it represents highly significant difference, it was consistent with Bayard and Decosta (2020)who found that under sacubitril/valsartan, **LVEF** improvred from 32.6 ± 5 to $36 \pm 6\%$.In contrast in ACEIs group, EF has no significant difference through follow up period.

LA diameter was found to has significant difference in Sacubitril/Valsartan group after six months of drug use and declined after three months to become $(4.75\pm0.50\text{cm})$ after six months; In ACEIs group, mean LA diameter was significantly increasing from its initial values after 6 months and it was consistent with Landolfo and Piani (2020) who found also a significant reduction of LA diameter and RVSP was found to have a significant difference in both groups through the 6 months of follow up with highest values before initiation of the drug and lowest values after 6 months and it was also consistent with Martens and Nuyens (2019) who stated that a trend toward reduction in RVSP was noted Martens et al 2019.

CONCLUSION

Sacubitril/Valsartan combination therapy was superior to ACEIs in reducing ventricular ectopics and decrease in QTc values which reflects the role of this combination to reduce ventricular arrhythmias.

Ejection fraction in Sacubitril/Valsartan combination group was improving through the follow up period as well as LA dimension, LVID and RVSP which can be explains in reverse remodeling.

Functional capacity in Sacubitril/Valsartan combinations group was improving through follow up period which means this drug reduces hospitalization.

LIMITATIONS

The sample was not large enough and was only conducted on 60 patients. Also assessment of functional capacity was subjective without qualitative assessment method to evaluate our patients.

Conflict of interest statement:

The authors have no conflicts of interest to declare.

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تأثير عقار ساكيوبتريل/فالسارتان علي معدل حدوث الإضطرابات في ضربات القلب في مرضي فشل القلب الناتج عن ضعف كفاءة العضلة

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الهدف من البحث: در اسه تأثير عقار ساكيوبتريل/فالسارتان علي معدل حدوث اضطرابات في ضربات القلب في مرضي فشل القلب الناتج عن ضعف كفاءه العضله مع مقارنته بمثبطات انزيم تحويل الانجيوتنسين.

المرضي و طرق البحث: التحق بهذه الدراسية ستون مريضا تم تقسيمهم إلي: مجموعة تعالج بعقار ساكيوبتريل فالسارتان والأخري بمثبطات انزيم تحويل الأنجيوتنسين أو مضادات الأنجيوتنسن لمده لاتقل عن 3 شهور و يتم متابعتهم بعد ثلاث و ست شهور عن طريق مسجل ضربات القلب لمدة ثمان و أربعين ساعة.

نتائج البحث: أوضحت الدراسه التأثير الهام للعقار علي المجموعة الاولي وذلك في تقصير مدة QRS، وتقليل معدل حدوث الضربات البطينية و تحسن كفاءة عضلة القلب و تقليل الحجم الانبساطي للقلب و تخفيض ضغط الشريان الرئوي و ذلك بالمقارنة مع المجموعه الأخرى.

الاستنتاج: عقار ساكيوبتريل /فالسارتان أكثر فاعليه من مثبطات انزيم تحويل الأنجيوتنسين. او مضادات الأنجيوتنسن في تقليل الضربات البطينية و تقصير QTc مما يؤدي لمنع الموت المفاجئ.

الكلمات الدالة: فشل القلب الناتج عن ضعف كفاءة العضلة، ساكيوبتريل، مثبطات انسزيم تحويل الأنجيوتنسين، مضادات الأنجيوتنسن، الموت المفاجئ، الأذين الايسر و ضغط الشريان الرئوي.