

Effects of Gender and Type 2 Diabetes Mellitus on left Ventricular Structure in a Group of Hypertensive Patients in Erbil-Iraq

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Abstract

Background: The effects of diabetes mellitus and gender on left ventricular structure in hypertension are still controversial.

Objective: To evaluate the effects of diabetes mellitus and gender on left ventricular structure in a sample of hypertensive patients in Erbil-Iraq.

Patients and Methods: This cross sectional study was conducted in Rizgary hospital, Erbil-Iraq, between April 2015 and April 2016. A convenient sample of 200 patients (100 males and 100 females), aged ≥ 18 years were enrolled. Half of the sample (Group I) had diabetes in addition to hypertension, while the other half (Group II) had hypertension only. The effects of diabetes mellitus and gender on left ventricular structure were evaluated in both groups.

Results: In Group I, left ventricular diastolic and systolic dimensions were significantly higher among male patients ($P=0.023$, 0.022 respectively) than female patients. In group II, the mean thickness of posterior wall, left ventricular mass, left ventricular mass index and relative wall thickness was significantly higher among male patients ($P=0.004$, 0.016 , 0.035 and 0.045 respectively) than female patients. The mean thickness of interventricular septum, posterior wall and relative wall thickness ($P=0.0013$, 0.007 and 0.003 respectively) was significantly higher in diabetic females than non-diabetic females. No significant differences were found between diabetic and non-diabetic males.

Conclusion: There is an obvious effect of gender and diabetes on left ventricular structure. In the hypertensive non-diabetic group, the left ventricular structural changes were significantly seen among male patients than females. In contrary, these changes were significantly seen in diabetic females than non-diabetic females.

Keywords: Diabetes mellitus, Gender, Hypertension, Left ventricular structure.

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Introduction

Hypertension is a major modifiable risk factor for cardiovascular diseases [1]. It is associated with a high mortality rate, accounting for ~ 14% of cardiovascular deaths worldwide [2]. According to World Health Statistics 2012, 40 % of the Iraqi adult populations have elevated blood pressure [3]. Echocardiographic studies in hypertension have shown greater left ventricular (LV) mass in men than in women [4].

Diabetes Mellitus (DM) is a cardiovascular risk factor associated with arterial hypertension and accelerated atherosclerosis [5]. Several studies have suggested that diabetes may be associated with LV structural and functional abnormalities [6]. Other studies mentioned that diabetes was associated with higher LV mass in women but not in men [7].

Patients with diabetes and hypertension are exposed to an exceptionally high risk of cardiovascular death, and are generally thought to need a more intensive risk factor control [8]. Both hypertension and DM may synergistically affect LV structure and function. Left ventricular hypertrophy (LVH) is mediated not only by the mechanical stress of pressure overload, but also by various neurohormonal substances (like insulin) that independently exert trophic effects on myocytes in the heart [9]. The relationships between glucose metabolism abnormalities and LV structure have been described in several reports [10]. However, the findings were controversial [11]. The gender differences were not specifically investigated in this issue. Up to our knowledge, there was

no previous study done regarding the same subject in Erbil city. The objective of this study was to find out the effect of gender and type 2 diabetes mellitus on left ventricular structure in a sample of hypertensive patients in Erbil city.

Patients and Methods

This cross-sectional study was conducted in Rizgary teaching hospital between April 2015 and April 2016. A convenient sample of 200 patients presenting with essential hypertension, aged ≥ 18 years were enrolled in the study. Those found to have diabetes in addition to hypertension were assigned to group I of the study (100 patients), while group II (100 patients) consisted of hypertensive patients only. All patients were assessed by a detailed history, physical examination, echocardiographic examinations, and any other necessary investigations to document presence of hypertension, DM and/or left ventricular structure abnormalities.

The exclusion criteria were patients with secondary hypertension (polycystic kidney disease, renovascular hypertension, and Cushing syndrome), chronic renal failure, liver disease and type 1 DM.

Based on recommendations of the Eighth Report of the Joint National Committee on Prevention, Detection, Evaluation, and Treatment of High Blood Pressure (JNC 8) [12], hypertension was defined as systolic blood pressure (SBP) ≥ 140 mmhg and (DBP) diastolic blood pressure ≥ 90 mmhg for adults aged 18 years and less than 60 years, and systolic blood pressure ≥ 150 or diastolic

≥ 90 in general population ≥ 60 years. Blood pressure measurements were taken with a mercury sphygmomanometer. Measurements were made to the nearest 2 mm Hg, in the sitting position with the arm supported, provided that the patient was at rest in the previous 5 minutes. The mean of two BP measurements was taken at each visit.

Transthoracic echocardiographic examination was performed by an expert cardiologist in the left lateral position. Standard M-mode, 2-Dimensional and Doppler echocardiographies were performed using (GE brand, Vivid E9, model 2012) echocardiography machine. All dimensions were measured according to established standards of the American Society of Echocardiography [13]. LV mass (LVM) was calculated according to the Devereux formula [14]: $LVM = 1.04 [(LVDd + IVSt + PWT)^3 - (LVDd)^3] - 13.6$. Thereafter, LV mass index (LVMI) was obtained by the following formula: $LVM / \text{body surface area (g/ m}^2)$ [15]. LVH was defined when LVMI exceeds 115 grams in men and 95 grams in women per meter square body surface area (m^2 BSA).

The data were collected by interviewing the patients using a questionnaire designed by the researchers. The questionnaire included information about socio-demographic data, hypertension, risk factors (like

hyperlipidemia, IHD, obesity, family history), and history of smoking and alcoholism.

Ethical considerations: The study protocol was approved by the ethics committee of the College of Medicine of Hawler Medical University. This study was conducted by using an informed verbal consent from the patients prior to participation in the study. The purpose of the study was carefully explained to each patient.

Statistical analysis

Data were analyzed using the statistical package for social sciences (SPSS, version 19). Student's t test for two independent samples was used to compare means. A 'P' value of ≤ 0.05 was considered as statistically significant.

Results

A total of 200 hypertensive patients were enrolled in this study. Half of the sample (Group I) had diabetes mellitus in addition to hypertension, while the other half (Group II) had hypertension only. Basic characteristics, clinical variables and LV echocardiographic parameters of both groups are shown in Table (1). There were no significant statistical differences between the means of the studied parameters of both groups ($p > 0.05$).

Table (1): Comparison between the means of the two study groups regarding basic characteristics, clinical variables, and echocardiographic parameters

	Group I (DM + HTN)		Group II (HTN only)		p
	Mean	SD	Mean	SD	
Age	55.25	11.36	53.95	10.56	.403
BMI	28.80	5.13	27.73	4.94	.133
SBP	153.20	16.68	149.55	16.71	.124
DBP	94.95	14.85	92.61	9.24	.182
HT duration	4.57	4.78	3.59	3.25	.092
EF	66.12	7.84	65.24	7.59	.42
LVH	58%		52%		.11
IVS	12.46	2.61	11.79	2.32	.06
PW	11.74	2.58	11.39	2.27	.31
Left atrium	33.84	4.57	33.61	5.04	.74
LVM	220.42	69.38	212.93	72.27	.46
LVMi	118.83	36.68	116.01	38.49	.60
RWT	.51	.14	.48	.12	.17

*BMI=Body mass index, EF=Ejection fraction, IVS=Interventricular septum, PW= Posterior wall, RWT=Relative wall thickness.

In the diabetic group Group (I), no inter-gender statistical differences were found regarding the LV echocardiographic parameters (representing LV mass and geometry) as shown in Table (2).

Table (2): Comparison between the means echocardiographic parameters of males and females among group I only

Echocardiographic parameters	Males		Females		p
	Mean	SD	Mean	SD	
EF	64.98	6.95	67.26	8.56	.147
IVS	12.33	2.62	12.59	2.61	.620
PW	11.50	2.49	11.98	2.67	.352
Left atrium	33.40	4.14	34.28	4.97	.338
LVM	223.18	66.13	217.66	73.06	.693
LVMi	120.22	35.52	117.44	38.11	.707
RWT	.48	.14	.53	.15	.088

Echocardiographic parameters of males and females (Group II only) are shown in Table (3). The mean thickness of posterior wall, LV mass, LVM index and relative wall thickness was significantly higher among male patients (P=0.004, 0.016, 0.035 and 0.045 respectively) than female patients.

Table (3): Comparison between the means echocardiographic parameters of males and females among group II only

Echocardiographic parameters	Males		Females		p
	Mean	SD	Mean	SD	
EF	64.74	7.30	65.74	7.91	.513
IVS	12.19	2.47	11.39	2.11	.086
PW	12.04	2.53	10.74	1.76	.004
Left atrium	33.94	4.77	33.28	5.31	.515
LVM	230.18	81.19	195.68	57.92	.016
LVMI	124.08	44.90	107.94	29.06	.035
RWT	.51	.15	.46	.09	.045

In intra-gender comparisons, no significant statistical differences were found in the means of echocardiographic parameters of diabetic and non-diabetic hypertensive male patients Table (4), but diabetic female patients show significant statistical differences regarding the mean thickness of

interventricular septum (IVS), posterior wall and RWT (P=0.013, 0.007, 0.003 respectively) as shown in Table 5, where it is evident that the means of the mentioned parameters were higher among diabetic females than the non-diabetic females.

Table (4): Comparison between the means of echocardiographic parameters of the two study groups, among males only

Echocardiographic parameters	Group I (DM+HTN)		Group II (HTN only)		p
	Mean	SD	Mean	SD	
EF	64.980	6.950	64.740	7.303	.867
IVS	12.334	2.624	12.190	2.472	.778
PW	11.500	2.495	12.036	2.535	.289
Left atrium	33.400	4.135	33.940	4.774	.547
LVM	223.180	66.126	230.180	81.194	.637
LVMI	120.220	35.523	124.080	44.898	.635
RWT	.483	.136	.506	.147	.423

Table (5): Comparison between the means of echocardiographic parameters of the two study groups, among females only

Echocardiographic parameters	Group I (DM+HTN)		Group II (HTN only)		p
	Mean	SD	Mean	SD	
EF	67.260	8.561	65.740	7.912	.359
IVS	12.594	2.610	11.392	2.111	.013
PW	11.984	2.673	10.744	1.760	.007
Left atrium	34.280	4.965	33.280	5.311	.333
LVM	217.660	73.064	195.680	57.916	.099
LVMI	117.440	38.108	107.940	29.061	.164
RWT	.532	.147	.456	.091	.003

Discussion

In the present study, LVM was significantly higher in male patients than females in Group (II). Effects of gender on LVM are somewhat indefinite [7]. Several studies have identified sex-related differences in the response to chronic pressure overload, with men having higher LV mass than women [4, 16, 17]. Regarding other echocardiographic parameters and despite the fact that LVH was not statistically significant but the means of posterior wall thickness, LVM index and relative wall thickness (RWT) were significantly higher among hypertensive non-diabetic Group (II) male patients than female patients. In the determination of left ventricular adaptive process that takes place in the over-loaded ventricle, fundamental components namely wall thickness must be taken into account [18]. Parietal thickness and its relation to LV chamber size has been recognized as a measure of hypertrophy for more than 30 years [19]. RWT (which depends on the posterior wall thickness) provides information regarding LV geometry independent of other calculations [20]. In the hypertensive diabetic group, no inter-gender statistical differences were found regarding the above parameters but only left ventricular diastolic and systolic dimensions were significantly higher among male patients.

The prevalence of LVH in patients with essential hypertension is ranging from 12 to 70%, depending on clinical characteristics of the population studied and largely on the measurement technique used [21].

In the present study, LVH was present in 52% of patients with hypertension alone (non-diabetic group). This result is within the above-mentioned range.

In intra-gender comparisons, no significant differences were found in diabetic and non-diabetic hypertensive male patients, but diabetic females show significant differences regarding interventricular septum, posterior wall and RWT. Despite being not statistically significant, but other parameters like LVM and LVM index were higher in diabetic women. Many studies have shown that DM affects the cardiac function and structure independent of hypertension or any other known risk factors [22]. Diabetes has been implicated as an important determinant of left ventricular mass in many studies. Myocardial and systemic mechanisms, as an increased extra-cellular matrix, vascular hypertrophy, and vasoconstriction have been attributed to this hypertrophic response [5-8]. It is now well recognized that insulin directly promotes myocyte hypertrophy independent of its effects on systemic arterial pressure [10]. In addition to circulating insulin, insulin growth factor-1 is also an independent determinant of LVM and geometry in essential hypertension [9]. However, Galvin *et al* [11] concluded that insulin resistance and hyperinsulinemia are not independent predictors of LVM in humans. In this study, the prevalence of LVH was higher (58%) in patients with both hypertension and DM than in patients with hypertension alone, although it was statistically not significant. The high

prevalence of LVH in our study is comparable with previously reported data. Somaratne *et al* [23] demonstrated that LVH was common (56%) among patients with type2 DM with no known cardiac, cerebrovascular or peripheral vascular disease. Echocardiographic LVH was found in (22-51%) of Australian diabetes clinic attendees [24]. A higher prevalence of echocardiographic LVH (71%) was noted in diabetes clinic attendees in Dundee, Scotland [25].

Conclusions

There is an obvious effect of gender and diabetes on left ventricular structure. In the hypertensive non-diabetic group, the left ventricular structural changes were significantly seen among male patients than females. In contrary, these changes were significantly seen in diabetic females than non-diabetic females.

Recommendations

- 1- Large scale prospective studies are needed in the future to evaluate the impact of gender and diabetes mellitus in hypertensive patients on left ventricular structure and function, and the development of LVH.
- 2- More studies are needed to evaluate the effect of type 1 DM on left ventricular structure and function, and to reveal if there is any gender difference to effect.

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