Distribution of Cardiac Geometric Patterns on Echocardiography in Essential Hypertension. Impact of Two Criteria of Stratification

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Purpose - To evaluate 2 left ventricular mass index (LV-MI) normality criteria for the prevalence of left ventricular geometric patterns in a hypertensive population (HT).

Methods - 544 essential hypertensive patients were evaluated by echocardiography, and different left ventricular hypertrophy criteria were applied: 1 - classic: men - 134 g/m² and women - 110 g/m²; 2 - obtained from the 95th percentil of LVMI from a normotensive population (NT).

Results - The prevalence of 4 left ventricular geometric patterns, respectively for criteria 1 and 2, were: normal geometry - 47.7% and 39.3%; concentric remodeling - 25.4% and 14.3%; concentric hypertrophy - 18.4% and 27.7% and eccentric hypertrophy - 8.8% and 16.7%, which conferred abnormal geometry to 52.6% and 60.7% of hypertensive. The comparative analysis between NT and normal geometry hypertensive group according to criteria 1, detected significative structural differences (*p < 0.05): LVMI - 78.4 ± 1.50 vs 85.9 ± 0.95 g/m²; posterior wall thickness - 8.5 ± 0.1 vs 8.9 ± 0.05 mm; left atrium - 33.3 ± 0.41 vs 34.7 ± 0.30 mm. With criteria 2, significative structural differences between the 2 groups were not observed.

Conclusion - The use of a reference population based criteria, increased the abnormal left ventricular geometry prevalence in hypertensive patients and seemed more appropiate for left ventricular hypertrophy detection and risk stratification.

Keywords - essential hypertension, left ventricular hypertrophy, criteria of hypertrophy

In recent years, the use of echocardiography made possible a better characterization of the cardiac morphometric and functional changes in essential hypertension.

The prevalence of left ventricular hypertrophy, which is estimated in approximately 5% according to electrocardiographic criteria, has ranged from around 20% to 40% in hypertensive populations and from 0% to 10% in normotensive populations assessed on echocardiography.

The use of more sophisticated tools may also detect more accurately the early cardiac structural changes that precede left ventricular hypertrophy.

Recent studies, which have assessed in a global way the spectrum of cardiac geometric changes on echocardiography in hypertensive patients, have established the concept of early cardiac remodeling. This concept along with prospective studies of morbidity and mortality have shown a greater cardiac risk for hypertensive individuals with concentric remodeling and hypertrophy, as compared with patients with normal geometry on echocardiography. This has, therefore, created a greater need for stratifying hypertensive patients in earlier phases of cardiac structural changes. Until then, these patients, unlike those patients with left ventricular hypertrophy, were considered as a low cardiovascular risk group.

The diagnosis of left ventricular hypertrophy has been based on pre-established criteria, whose limits for the cardiac mass index have been obtained in referred normotensive populations.

Therefore, adequate stratification of hypertensive patients requires pre-established limits of cardiac mass index.

Some studies, including that of Cornell University and the Framingham Study, have established limit values of ventricular mass index, which were calculated based on percentiles or standard deviations of the average of the cardiac mass index in populations of normotensive males and females. These limit values were, respectively, 134 and 110g/m² at Cornell University study, and 131 and 100g/m² in the
Framingham Study. Later on, other studies 7,19,20 established new referential values for correcting the mass by body surface area. More recent studies 21,22 have proposed the use of limit values based on indexing by height, square height, and height to the power of 2.13 or 2.7.

However, the application of these criteria in different populations should be carefully performed. Considering the diversity of the populations studied, changes in sensitivity and specificity may occur when pre-established criteria are applied for left ventricular hypertrophy, and, therefore, inadequate stratification in cardiac geometric patterns may happen.

Up to the present time, only a few studies 7,11,23-26 have assessed the impact of different limit values of ventricular mass indexing on the prevalence of cardiac geometric patterns. Likewise, only a few studies 7 have proposed assessing the accuracy of applying criteria usually accepted in their local populations.

The objective of the present study was to assess comparatively the prevalence of cardiac geometric patterns on echocardiography in a population of individuals with essential hypertension using one criterion widely applied (criterion 1) and a second criterion obtained from a referential normotensive population (criterion 2).

**Methods**

We retrospectively studied 544 patients with essential hypertension (173 males and 371 females), who underwent Doppler echocardiography in the hypertension clinic of the nephrology department at Unifesp (Federal University of São Paulo). All patients had a previous diagnosis of hypertension, which was established by assessment of medical records. Most patients (85.4%) were under medicamentous treatment, and the remaining patients were out of medication for at least 8 weeks prior to echocardiography.

The following exclusion criteria were considered: previous diagnosis of severe hypertension; pressure levels: systolic blood pressure (SBP) ≥ 180mmHg or diastolic blood pressure (DBP) ≥ 110mmHg, or both, on the day of echocardiography; secondary hypertension; a pre-established diagnosis of diabetes or fasting glycemia levels ≥ 140mg/dL, or both; chronic renal failure defined as serum creatinine ≥ 2.0mg/dL; coronary heart disease diagnosed by angiography, history of myocardial infarction, angina, or positive exercise test; and clinical signs of congestive heart failure.

Out of the local referential population, a group of 106 normotensive individuals was chosen to undergo Doppler echocardiography assessment as a control group.

On the day of the examination, demographic data and measurements of blood pressure were taken from the patients and control group. For means of analysis, only the first examination of each patient during the period studied was considered. All patients with an inadequate echocardiographic window or with valvar lesions and hemodynamic repercussion evidenced on echocardiography were excluded from the analysis.

On Doppler echocardiography, the following structural parameters were assessed: thickness of the posterior wall, of the interventricular septum, and left ventricular diameter during systole and diastole. The ventricular mass was calculated with the modified Devereux formula 27: 0.80[(1.04 x DIVS + DLVPW)3 / LVDD] + 0.6, and the ventricular mass index obtained through correction of mass by the body surface area. Measurements of the relative thickness of the wall and relative thickness of the septum were obtained as follows: 2 x DLVPW/LVDD 28 and 2 x DIVS/LVDD 29, where DLVPW, DIVS, and LVDD correspond, respectively, to measurements of the thickness of the posterior wall, of the interventricular septum, and the ventricular diameter during diastole.

All measurements of the septum and the posterior wall were performed at the end of diastole, including the endocardial thickness, according to recommendations of the American Society of Echocardiography 30, justifying the use of its formula modified by Devereux 27. This formula approximates the values of ventricular mass obtained with the formula initially validated by the American Society of Echocardiography 31, to the values of mass obtained with the equation of the convention of Penn 32. This latter, despite being more accurate, applies a method of measurements less used, which excludes from the analysis the endocardial thickness of the septum and wall.

Two criteria were used for the definition of hypertrophy: 1) the classical criterion, whose limits of ventricular mass index are 134g/m² for males and 110g/m² for females 17 (criterion 1); 2) the criterion obtained from the 95th percentile of the measurements of the ventricular mass index obtained in males and females of the referential normotensive population (criterion 2), which resulted in the respective limits of 110g/m² and 96g/m².

According to these criteria (1 and 2), the patients were classified into 4 ventricular geometry groups 9,10,29, as follows: normal geometry – index of normal ventricular mass and relative thickness of the wall and relative thickness of the septum <0.45 (NG1 – criterion 1; NG2 – criterion 2); concentric remodeling (CR) – normal ventricular mass index and relative wall thickness or relative septum thickness >0.45, or both, (CR1 and CR2); hypertrophy – ventricular mass index ≥ pre-established limits, concentric hypertrophy (CH) if the relative wall thickness was >0.45, and eccentric hypertrophy (EH) if the relative wall thickness was <0.45.

Direct measurements of ventricular diameter and volume of the chambers obtained on echocardiography allowed direct determination of parameters of systolic function and hemodynamic derivatives, such as: systolic volume (SV) = end-diastolic volume (EDV) − end-systolic volume (ESV), where EDV = LVDD 3, ESV = LVSD 3 and LVSD is the ventricular diameter obtained during systole; cardiac output (CO) = SV x HR (heart rate) and corrected for body surface for obtainment of the cardiac index (CI); ejection fraction (EF) = (EDV – ESV)/EDV x 100 34; percentage of fractional shortening (FS%) obtained through the expected FS values divided by the FS values obtained, where FS = (DVD – SVD)/DVD x 100 35 and expected FS = 99.9 - (35.4 log 10 end-systolic stress); end-systolic stress (ESS) = SBP x LVSD/4 x SLVPW 36,37.
Diastolic function was assessed by mitral Doppler, using the ratios of the E and A wave (cm/s) velocities and the ratio of these waves (E/A).\(^{38}\)

For statistical analysis, demographic data were recorded with Dbase III software, and the Sigma Stat software was used. Demographic and pressure parameters, as well as echocardiographic analysis, were presented as mean ± standard error. The student \(t\) test was used for comparative analysis of demographic and pressure parameters between the normotensive and hypertensive groups. The variance test (ANOVA) was used for comparing these parameters between groups NT, NG1, and NG2, and separately for males and females. Comparative analysis of the means of the structural and hemodynamic parameters, as well as the systolic and diastolic functions, between the groups NT x HT and NT x NG1 x NG2 was performed through the covariance test (ANCOVA), after adjusting for age, sex, and body mass index. Comparison of prevalence of the parameters of ventricular geometry in the groups of males and females, according to both criteria, was obtained through the chi-square test.

**Results**

Table I shows the prevalence of cardiac geometric patterns obtained according to criteria 1 (classical criterion) and 2 (based on a normotensive population). The percentage of patients with cardiac structural changes (CR + CH + EH) was significantly higher when criterion 2 was used (60.7%) as compared with criterion 1 (52.6%). In addition, prevalence of cardiac hypertrophy (CH + EH) increased from 27.2% (criterion 1) to 45.7% when criterion 2 was used. On the other hand, a reduction in the number of patients with normal geometry on echocardiography occurred (47.4% with criterion 1, and 39.3% with criterion 2).

When assessing prevalence of different patterns of ventricular geometry for both sexes and according to criteria 1 and 2, we observed a distribution similar to that obtained with the analysis of the total population. The distribution of abnormalities according to criteria 1 and 2 in the male group was, respectively, 50.3% and 57.7%, and in the female group, respectively, 53.6% and 62.0%.

Table II shows pressure and demographic data of the normotensive and essentially hypertensive population. The subgroups of individuals with normal geometry obtained using the two criteria were also assessed. It is worth noting that hypertensive patients with normal geometry did not differ from the total group of hypertensive individuals in regard to age, body mass index, and blood pressure levels. These subgroups of hypertensive individuals, however, showed a significant increase in age and body mass index as compared with the normotensive individuals. In addition, a higher proportion of males was found among the hypertensive individuals.
Table III illustrates the demographic data obtained separately with the evaluation of males and females in the subgroups of normotensive individuals and hypertensive individuals with normal geometry, according to the 2 criteria employed.

As expected, cardiac structural parameters were significantly higher in hypertensive individuals as compared with the normotensive population as follows: ventricular mass index – 78.8±1.2 vs 103.8±1.3 *; left atrium – 33.4±0.41 vs 35.2±0.2 *; LVDD – 47.2±0.34 vs 46.8±0.23; LVPW – 8.5±0.10 vs 10.2±0.08 *; IVS – 8.8±0.10 vs 10.7±0.09 * (* p<0.05).

Table IV shows that the structural parameters were also significantly higher in the group of hypertensive individuals with normal geometry obtained with the classical criterion as compared with the group of normotensive individuals. However, when using the criterion based on the local normal population, no differences were observed in the assessment of the 2 groups.

Table IV also shows the structural evaluation in subgroups of males and females. It is worth noting that in the male subgroup, no structural differences were observed in the NG2 group as compared with the group of normotensive individuals (as in the global evaluation). In the female subgroup, significant differences were observed between the 2 groups, despite the fact that by applying criterion 2 the structural parameters of the NG2 group were more close to those of the normotensive group.

Cardiac functional alterations resulting from hypertension were not influenced by the use of one or the other criterion. Patients with normal geometry obtained according to both criteria showed an increase in the peripheral vascular resistance, end-systolic stress, cardiac index, fractional shortening, and contractility index, when compared with the normotensive individuals (Table V). In regard to the diastolic function, similar reductions in the E wave/A wave ratio were observed in both groups as compared with the normotensive individuals. This was also observed when males and females were separately evaluated.
Table V – Hemodynamic parameters (systolic and diastolic function) in normotensive individuals (NT), individuals with essential hypertension (HT), and hypertensive individuals with normal cardiac geometry (HT - NG1; HT - NG2).

<table>
<thead>
<tr>
<th></th>
<th>NT</th>
<th>HT</th>
<th>HT – NG1</th>
<th>HT – NG2</th>
</tr>
</thead>
<tbody>
<tr>
<td>N</td>
<td>106</td>
<td>544</td>
<td>258</td>
<td>214</td>
</tr>
<tr>
<td>PVRi</td>
<td>1306.7 ± 35.9</td>
<td>1672.1 ± 24.3*</td>
<td>1462.1 ± 22.7*</td>
<td>1497.8 ± 25.1*</td>
</tr>
<tr>
<td>ESS</td>
<td>44.2 ± 1.47</td>
<td>48.0 ± 0.63*</td>
<td>50.4 ± 0.73*</td>
<td>49.6 ± 0.77*</td>
</tr>
<tr>
<td>IC</td>
<td>3.34 ± 0.11</td>
<td>3.54 ± 0.05</td>
<td>3.75 ± 0.05*</td>
<td>3.63 ± 0.05*</td>
</tr>
<tr>
<td>EF</td>
<td>0.75 ± 0.06</td>
<td>0.75 ± 0.002</td>
<td>0.76 ± 0.002*</td>
<td>0.76 ± 0.002*</td>
</tr>
<tr>
<td>FS</td>
<td>37.7 ± 0.47</td>
<td>38.1 ± 0.20</td>
<td>38.4 ± 0.23</td>
<td>38.6 ± 0.24*</td>
</tr>
<tr>
<td>FS%</td>
<td>89.3 ± 1.13</td>
<td>92.7 ± 0.48*</td>
<td>95.9 ± 0.61*</td>
<td>96.0 ± 0.64*</td>
</tr>
<tr>
<td>CTI</td>
<td>1.79 ± 0.06</td>
<td>2.07 ± 0.02*</td>
<td>2.05 ± 0.03*</td>
<td>2.13 ± 0.04*</td>
</tr>
<tr>
<td>E/A</td>
<td>1.32 ± 0.03</td>
<td>1.12 ± 0.01*</td>
<td>1.27 ± 0.02*</td>
<td>1.26 ± 0.02*</td>
</tr>
</tbody>
</table>

p<0.05 vs NT; PVRi – peripheral vascular resistance index (din/sec/m²); ESS – systolic stress (10² din/cm²); CI – cardiac index (l/min/m²); EF– ejection fraction; FS- fractional shortening (%); FS%- fractional shortening corrected to the expected FS; CTI- contractility index (10³ din/cm²); E/A- E wave/A wave ratio.

Discussion

The diagnosis of ventricular hypertrophy in hypertension requires pre-established criteria derived from normotensive populations, and these criteria are based on the normal limits of the cardiac mass index.17,18

One of the currently most used criteria is that derived from studies at Cornell University 17, which is based on a normotensive population of New York City (ventricular mass index of 134g/m² for males and 110g/m² for females).

When the limits of thickness of the septum and the left ventricular posterior wall are taken into consideration, stratification in 4 cardiac geometric patterns is possible. This has proven to be useful as it allows the identification of a significant percentage of hypertensive individuals in the population, who, despite their ventricular mass index within the normal range, show an increase in the relative thickness of the septum or the posterior wall, or both, as compared with individuals with normal geometry.10,12,29

These patients, classified as having concentric remodeling, are currently known to have higher cardiovascular morbidity and mortality.11,12

In our study, the different prevalences of ventricular geometric patterns obtained with the use of both criteria (Table I) allow the identification of a substantial number of individuals with hypertrophy (CH and EH) or with some kind of cardiac structural change (CH, EH, and CR) in a population of mild to moderate hypertensive individuals both for the male and female groups.

Using criterion 1, 47.4% of the patients had NG, 25.4% CR, and 27.2% hypertrophy, adding to a total of 52.6% of patients with cardiac structural changes. Similar prevalence was found in the evaluation of males and females separately.

Using the criterion based on the normotensive population (criterion 2), we found 45.7% of individuals with hypertrophy in the total group, 42.7% in the male group, and 47.9% in the female group, constituting along with the percentage of individuals with concentric remodeling a total of, respectively, 60.7%, 57.8%, and 62.0% of hypertensive individuals with some kind of structural alteration. This means a significant increment of 8.1%, 7.5%, and 8.4% of patients with an increased risk in stratification.

On the other hand, patients with ventricular mass index, and septal and wall thickness within normal range have normal geometry and, theoretically, should not differ from normotensive patients in regard to cardiac structural parameters. However, the results of our study have shown that individuals classified as having normal geometry by applying criterion 1 had significantly higher structural changes in mass ventricular index and relative thickness of the septum and wall than the referential population of normotensive individuals.

When we applied the criterion based on the 95th percentile of the mean of the ventricular mass indices obtained from a local referential population instead of the criterion of Cornell University, significant differences were no longer observed in the structural parameters of individuals with normal geometry and those of the normotensive population. When analyzing males and females separately, however, the results obtained showed that in the female group, despite the approximation of the structural parameters between the NG group and the normotensive group, when using criterion 2, significant differences in the assessment of the ventricular mass index and LVPW occurred. In addition, no significant differences were observed in most morphometric evaluations of NG1 and NG2 subgroups.

Nevertheless, the application of criterion 2 seemed more appropriate for detecting cardiac hypertrophy in this population, because a greater approximation of the structural parameters of the NG and normotensive groups occurred.

The fact that pre-established criteria obtained from specific populations might not be applied with the same accuracy for other populations had already been observed and motivated the studies at Cornell University. This showed that the application of the criteria obtained in the Framingham Study18 to a population of New York City ended up by overestimating the prevalence of hypertrophy in groups of normotensive individuals and groups of individuals with borderline and sustained hypertension in this population.7 On that occasion, 9.4% of the normotensive individuals, who supposedly had normal cardiac geometry, were classified as having left ventricular hypertrophy, and a prevalence of 19.6% of hypertrophy in borderline hyperten-
sive individuals was detected; on the other hand, with the use of local criteria, this prevalence was 12.4%. The low specificity (negative predictive value) of the criteria of the Framingham Study for the population of New York City (90.6%) was attributed to population changes in lifestyle, prevalence of obesity, sedentary lifestyle, etc.

Differences in body constitution and life habits between our population and that of New York City may perhaps explain the better accuracy obtained when a criterion based on the local population was used. This was stressed when we applied criterion 1 in our normotensive population, which provided a prevalence of 0.9% of individuals with hypertrophy in a population that supposedly had no hypertrophy. Therefore, we obtained a 99.1% specificity for criterion 1, which is extremely high when one considers the ideal 97% specificity², with no loss in sensitivity (positive predictive value).

A recent study²⁶ assessed the use of different criteria of hypertrophy in a normotensive population and in a previously selected subpopulation based on the presence of hypertrophy on the electrocardiography (subpopulation of the LIFE Study). This study could well analyze the inverse relation existing between specificity and sensitivity for the criterion of ventricular mass correction used (body geometric patterns depending on the population sub-entails ends up in modifying the prevalence of ventricular hypertrophy, which ranged from 42% to 72% in this study, depending on the criterion used.

Other studies have shown that the application of different criteria ends up in modifying the prevalence of ventricular geometric patterns depending on the population subgroups analyzed (according to sex, age, body mass index), on the criterion of ventricular mass correction used (body surface, height, square height), or on the covariables present in the population being studied (level of hypertension, medication use, blood pressure control, etc).

In the VITAE Study²⁵, the echocardiographic assessment of a large population of essential hypertension obtained from reference centers in Spain provided a prevalence of ventricular hypertrophy and of concentric remodeling that ranged from 59.2% to 72.2% and from 6.5% to 11.4%, respectively, depending on the criterion used.

The use of the same criteria applied in our study (110g/m² and 134g/m²) in 510 participants of the HOT Study²⁰ provided a prevalence of hypertrophy of 62%. In another study²⁹, a prevalence of hypertrophy of 25% for males and 26% for females was found with the criterion of 134g/m² and 102g/m², which is very similar to that used in our study.

Considering the prognostic implications associated with the diagnosis of ventricular hypertrophy and remodeling changes in hypertensive individuals⁴¹, the standardization of criteria has been emphasized, confirming the need for establishing specific population criteria.

As expected, the use of a more adequate criterion of normality for the ventricular mass index did not alter accuracy in detecting early functional changes that accompany the development of left ventricular hypertrophy³⁰,²⁹,⁴²,⁴³. In fact, functional (systolic and diastolic) and hemodynamic changes in hypertensive patients with normal cardiac geometry were equally demonstrated when both criteria were used.

In conclusion, the findings of the present study suggest that the use of a criterion of normality based on a referential normotensive population may perhaps be more appropriate for detecting ventricular hypertrophy and stratification of the hypertensive population in different cardiac geometric patterns. Obtaining these criteria based on the assessment of a larger referential normotensive population in our country is required.

References


