Mechanical Dyssynchrony is Similar in Different Patterns of Left Bundle-Branch Block

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Abstract

Background: Left bundle-branch block (LBBB) and the presence of systolic dysfunction are the major indications for cardiac resynchronization therapy (CRT). Mechanical ventricular dyssynchrony on echocardiography can help identify patients responsive to CRT. Left bundle-branch block can have different morphologic patterns.

Objective: To compare the prevalence of mechanical dyssynchrony in different patterns of LBBB in patients with left systolic dysfunction.

Methods: This study assessed 48 patients with ejection fraction (EF) < 40% and LBBB consecutively referred for dyssynchrony analysis. Conventional echocardiography and mechanical dyssynchrony analysis were performed, interventricular and intraventricular, with ten known methods, using M mode, Doppler and tissue Doppler imaging, isolated or combined. The LBBB morphology was categorized according to left electrical axis deviation in the frontal plane and QRS duration > 150 ms.

Results: The patients’ mean age was 60 ± 11 years, 24 were males, and mean EF was 29% ± 7%. Thirty-two had QRS > 150 ms, and 22, an electrical axis between −30° and +90°. Interventricular dyssynchrony was identified in 73% of the patients, while intraventricular dyssynchrony, in 37%-98%. Patients with QRS > 150 ms had larger left atrium and ventricle, and lower EF (p < 0.05). Left electrical axis deviation associated with worse diastolic function and greater atrial diameter. Interventricular and intraventricular mechanical dyssynchrony (ten methods) was similar in the different LBBB patterns (p = ns).

Conclusion: In the two different electrocardiographic patterns of LBBB analyzed, no difference regarding the presence of mechanical dyssynchrony was observed. (Arq Bras Cardiol. 2013;101(5):449-456)

Keywords: Bundle-Branch Block; Ventricular Dysfunction; Cardiac Resynchronization Therapy; Stroke Volume.

Introduction

Heart failure, a clinical syndrome resulting from structural and/or functional cardiac dysfunction, is known to be the end stage of several cardiopathies. Electrocardiographic alterations, such as left bundle-branch block (LBBB), are common findings in patients with heart failure, mainly in the presence of systolic dysfunction.2

Currently, there are several treatment options for heart failure. One efficient alternative is cardiac resynchronization therapy (CRT). The indication for implantation of a resynchronizing pacemaker is based on clinical and electrocardiographic criteria, and echocardiographic data. On the electrocardiogram, QRS complex enlargement, as observed in LBBB, is the most frequent indication for that treatment.4-6 However, treatment failure has been reported in approximately 30% of the cases in several series.3 In addition to the already known classic information, such as left ventricular dimension and ejection fraction, echocardiography allows the analysis of interventricular and intraventricular synchronism, which is the focus of CRT. Different methods, using several echocardiographic techniques, have been used to detect and stratify dyssynchrony, enabling predicting those who will have good results with a certain treatment.

Left bundle-branch block can have different characteristics related to higher morbidity and mortality.9,10 The relationship between different characteristics of LBBB and dyssynchrony assessed on echocardiography is yet to be established, which might contribute to the lack of success of that therapy.

Objectives

This study aimed at comparing conventional echocardiographic findings and those of ventricular synchrony related to different LBBB morphologies in patients with left ventricular systolic dysfunction.

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Methods

This study was approved by the Committee on Ethics and Research of the Instituto Dante Pazzanese de Cardiología.

Study population

This study assessed individuals followed up on an outpatient basis for heart failure treatment, who were referred to the echocardiography section with systolic dysfunction characterized by ejection fraction below 40%, according to the Simpson’s method. All patients had sinus rhythm and LBBB\(^1\). Patients with the following characteristics were excluded: under the age of 18 years; wearing a pacemaker; and those who had undergone previous valvular surgery or had any degree of aortic valvulopathy. The clinical data concerning functional class, history and medications used were also assessed.

Electrocardiogram

Twelve-lead electrocardiography was performed. The PR intervals and QRS complexes were measured, and the frontal axis characteristics were assessed. The patients were classified into two groups according to the presence of QRS interval > 150 ms or left electrical axis deviation in the frontal plane, i.e., frontal axis values < −30°.

Echocardiogram

Echocardiogram was performed on a Vivid\(^7\)\(^8\) device (GE Vingmed, System VII, Horten, Norway). The images were acquired as digital clips. Then, linear and two-dimensional measures were taken according to the American Society of Echocardiography guidelines, using a mean of three consecutive cycles on an EchoPAC PC workstation, version 6.0.1\(^9\) (GE Vingmed Ultrasound). Diastolic function was also characterized according to the American Society of Echocardiography guidelines, and mitral valve regurgitation was quantified\(^10\),\(^11\).

Interventricular dyssynchrony was assessed as the difference between pre-ejection intervals, i.e., from the beginning of the QRS complex to the beginning of the ventricular ejection into the aortic and pulmonary valves, using pulsed Doppler; interventricular dyssynchrony was considered to exist when that value exceeded 40 ms\(^12\),\(^13\). According to the literature, mechanical intraventricular dyssynchrony has been assessed by use of several methodologies, whose cutoff points have been described as markers of successful CRT. The analysis was performed according to the following criteria: 1) septal-to-posterior wall motion delay, in M mode, > 130 ms, as reported by Pitzalis et al\(^14\); 2) greater interval between maximum systolic motion of six left ventricular basal segments > 110 ms, measured on tissue Doppler, as demonstrated by Notabartolo et al\(^15\); 3) maximum systolic motion interval between the septum and lateral wall on tissue Doppler > 65 ms, as reported by Gorcsan et al\(^16\); 4) presence of positive criterion of the Saint Mary Hospital score, United Kingdom, as reported by Lane et al\(^17\); 5) positive criteria for the presence of dyssynchrony, as established by Cleland et al\(^18\), in the CARE-HF study; 6) standard deviation of the maximum motion times, measured on tissue Doppler imaging, in 12 left ventricular segments > 32 ms, proposed by Yu et al\(^19\); 7) interval values > 60 ms of the onset of mitral ring systolic motion in four segments measured by use of tissue Doppler imaging, as reported by De Sutter et al\(^20\); 8) interval values > 100 ms of the end of mitral ring systolic motion in four segments measured on tissue Doppler imaging, as reported by Perez de Isla et al\(^21\), in the Spanish Ventricular Asynchrony Registry - RAVE; 9) interval between the maximum contraction of the anteroseptal and posterior segments > 130 ms measured by use of two-dimensional strain associated with the interval of the septal-to-lateral wall maximum systolic motion on tissue Doppler > 60 ms, as demonstrated by Gorcsan et al\(^22\).

Statistical analysis

The quantitative variables were described as mean ± standard deviation, and the qualitative ones, as percentages. For comparing the different LBBB presentations, the following tests were used: Student t test; Wilcoxon test; chi-square test; and Fisher exact test. The JMP8.0\(^\circ\) software (Institute Inc., Carry, North Carolina) was used for calculation. The significance level of 5% was adopted.

Results

Table 1 shows the clinical characteristics of the 48 patients studied, with approximately 90% of them on beta-blockers and angiotensin-converting-enzyme inhibitors/angiotensin-receptor blockers. Table 2 shows their electrocardiographic findings. Table 3 shows their echocardiographic variables, with varied degrees of intraventricular dyssynchrony according to the criteria used.

When patients were compared according to their different LBBB morphologies, QRS duration > 150 ms and electrical axis in the frontal plane −30° did not relate to differences concerning sex, age, history, functional class or medication used, as shown in Tables 4 and 5.

Regarding echocardiographic findings, patients with QRS duration > 150 ms showed greater left ventricular linear dimensions and volumes, greater left atrial diameters and lower ejection fraction, as shown in Table 4.

Patients with left electrical axis deviation in the frontal plane, < −30°, showed greater left atrial diameters associated with higher grades of left ventricular diastolic dysfunction, and greater left ventricular diameter, as evidenced in Table 5.

The presence of interventricular and intraventricular dyssynchrony was similar in the two groups of LBBB (longer QRS interval duration and left electrical axis deviation in the frontal plane) (Tables 4 and 5).

Discussion

The different LBBB presentations assessed do not allow identifying a dyssynchrony pattern, and their prevalences did not differ in the different echocardiographic methodologies assessed.
However, the LBBB patterns relate to left ventricular morphologic and functional alterations, in which longer QRS complex durations associate with greater left ventricular dimensions, and the left electrical axis deviation on electrocardiogram relates to greater diastolic dysfunction and greater left atrial dimension.

Those findings are in accordance with the study by Das et al., who have shown that left ventricular ejection fraction is more impaired when the QRS duration is increased in patients with LBBB, but it is not associated with left electrical axis deviation. However, according to Dhingra et al., the higher incidence of events in patients with LBBB and left electrical axis deviation should be associated with greater diastolic dysfunction, which is known to relate to mortality.

Although not all patients meet the criteria proposed by the last guidelines for implantation of resynchronizing pacemakers, those indications have been modified, and most patients studied constitute a group candidate for CRT, including patients with ejection fraction < 40%. Findings might indicate lack of relationship between longer QRS intervals in LBBB and the response to that type of treatment, because the prevalence of mechanical dyssynchrony is similar regardless of the echocardiographic method used.

Despite the limitations of using echocardiography as the method for selecting candidates for pacemaker implantation with evidenced capacity of resynchronization on the PROSPECT study, most of the methods used proved to distinguish patients who would benefit from that therapy. Single center studies have reported several echocardiographic methods that evidenced a better response to that therapy.

Sweeney et al. have shown that the conventional electrocardiographic report in patients with LBBB, such as QRS duration and the presence of left electrical axis deviation, are not enough to predict individuals who will have echocardiographic improvement after cardiac resynchronization by use of pacemaker. However, electrocardiographic evidence of longer left ventricular activation time and smaller scar volume characterizes the group of patients with a better response to the resynchronizer. Such measures were not assessed in the present study, and the study by Sweeney et al. has not compared those electrocardiographic findings with the echocardiographic assessment of mechanical synchrony. That relationship can be tested in a further study.

Conclusion
In the two different electrocardiographic patterns of LBBB analyzed with ten echocardiographic methods, no difference regarding the presence of mechanical dyssynchrony was observed. They can, however, be associated with known risk patterns, such as a reduced ejection fraction and greater diastolic dysfunction grades.

Author contributions
Conception and design of the research: Barreto RBM, Piegas LS, Moreira DA, França FF; Acquisition of data: Barreto RBM, Melo Neto JF, Resende TU; Analysis and interpretation of the data: Barreto RBM, Piegas LS, Assaf JE, Melo Neto JF, Resende TU, Moreira DA, França FF; Statistical analysis and Obtaining funding: Barreto RBM; Writing of the manuscript: Barreto RBM, Piegas LS, Assaf JE; Critical revision of the manuscript for intellectual content: Barreto RBM, Assaf JE, LeBihan DC, Meneghelo RS, Sousa AGMR.

Table 1 – Clinical characteristics of the patients

| Clinical characteristics | Value |
|--------------------------|-------|
| Age (years)              | 59.9 ± 11.1 |
| Male sex                 | 50%   |
| History                  |       |
| Arterial hypertension    | 62.4% |
| Coronary artery disease  | 26.0% |
| Previous myocardial infarction | 22.5% |
| Diabetes mellitus        | 16.5% |
| Chagas’ disease          | 6.1%  |

Functional class (NYHA)

| Class | Percentage |
|-------|------------|
| I     | 9.5%       |
| II    | 47.6%      |
| III   | 42.8%      |

Medications used

| Medication               | Percentage |
|--------------------------|------------|
| Digitalis               | 39.6%      |
| Loop diuretics          | 81.3%      |
| Spironolactone          | 79.2%      |
| Angiotensin-converting-enzyme inhibitor | 56.3% |
| Angiotensin receptor blocker | 31.3% |
| Beta-blocker            | 89.6%      |
| Calcium channel blocker | 6.3%       |
| Nitrate                 | 12.7%      |

NYHA: New York Heart Association.

Table 2 – Electrocardiographic (ECG) characteristics

| ECG measures | Value |
|--------------|-------|
| Heart rate (bpm) | 72.3 ± 14.2 |
| PR interval (ms)  | 232.8 ± 58.9 |
| QRS width (ms)    | 165.0 ± 28.1 |
| Frontal ECG axis (º) | -2.3 ± 45.8 |

Pattern (%)

| Pattern | Percentage |
|---------|------------|
| QRS > 150 ms | 66.7%     |
| Frontal ECG axis between -30º and +90º | 54.2% |

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Table 3 – Echocardiographic characteristics

| Echocardiographic measures | M mode | Two-dimensional |
|---------------------------|--------|-----------------|
| Echocardiographic measures |        |                 |
| M mode                    |        |                 |
| Left atrium (mm)          | 45.5 ± 7.0 |
| Left ventricle, diastole (mm) | 74.1 ± 9.8 |
| Left ventricle, systole (mm) | 64.3 ± 10.7 |
| Two-dimensional           |        |                 |
| End-diastolic volume (mL) | 203.4 ± 79.8 |
| End-systolic volume (mL)  | 148.5 ± 66.1 |
| Left ventricular ejection fraction (%) | 28.7 ± 7.3 |
| Diastolic pattern         |        |                 |
| Grade IA dysfunction      | 37.5%  |
| Grade II dysfunction      | 31.3%  |
| Grade III/IV dysfunction  | 20.8%  |
| Undetermined              | 10.4%  |
| Mitral regurgitation      |        |                 |
| Absent                    | 14.6%  |
| Mild                      | 54.2%  |
| Moderate                  | 27.1%  |
| Severe                    | 4.2%   |
| Prevalence of interventricular dyssynchrony | 72.9% |
| Interval between pulmonary and aortic pre-ejective periods > 40 ms | 72.9% |
| Prevalence of intraventricular dyssynchrony | 72.9% |
| Criteria                  |        |                 |
| Pitzalis et al[16]        | 50.0%  |
| Notabartolo et al[17]     | 39.6%  |
| Gorcsan et al[18]         | 37.4%  |
| Lane et al[19]            | 97.9%  |
| Cleland et al[20]         | 60.4%  |
| Yu et al[21]              | 60.4%  |
| De Sutter et al[22]       | 85.4%  |
| Perez de Isla et al[23]   |        | Criterion I    |
| Gorcsan et al[24]         |        | 60.4%          |
|                           |        | Criterion II   |
|                           |        | 41.7%          |
|                           |        |                |
| Potential Conflict of Interest |        |                 |
| No potential conflict of interest relevant to this article was reported. |
| Study Association         |        |                 |
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Table 4 – Comparison of echocardiographic data between patients with different QRS intervals

| Echocardiographic measures                  | Variable                  | QRS interval ≤ 150 ms (n = 16) | QRS interval > 150 ms (n = 32) | p value |
|--------------------------------------------|---------------------------|---------------------------------|---------------------------------|---------|
| **M mode**                                 |                           |                                 |                                 |         |
| Left atrium (mm)                           | 42.2 ± 7.1                | 47.1 ± 6.5                      | 0.03                            |         |
| Left ventricle, diastole (mm)              | 71.1 ± 9.5                | 78.4 ± 8.8                      | 0.001                           |         |
| Left ventricle, systole (mm)               | 57.9 ± 10.9               | 64.4 ± 9.26                     | 0.03                            |         |
| **Two-dimensional**                        |                           |                                 |                                 |         |
| End-diastolic volume (mL)                  | 153.3 ± 69.9              | 228.6 ± 73.0                    | 0.002                           |         |
| End-systolic volume (mL)                   | 108.3 ± 58.8              | 168.6 ± 60.8                    | 0.002                           |         |
| Left ventricular ejection fraction (%)     | 31.8 ± 7.4                | 27.2 ± 6.8                      | 0.04                            |         |
| **Diastolic pattern**                      |                           |                                 |                                 |         |
| Grade IA dysfunction                       | 12.5%                     | 6.2%                            | ns                              |         |
| Grade II dysfunction                       | 43.8%                     | 59.4%                           | ns                              |         |
| Grade III/IV dysfunction                   | 37.5%                     | 25.0%                           | ns                              |         |
| Undetermined                               | 6.2%                      | 9.4%                            | ns                              |         |
| **Mitrail regurgitation**                  |                           |                                 |                                 |         |
| Absent                                     | 31.2%                     | 6.2%                            | ns                              |         |
| Mild                                       | 50.0%                     | 56.2%                           | ns                              |         |
| Moderate                                    | 12.5%                     | 34.4%                           | ns                              |         |
| Severe                                     | 6.2%                      | 3.1%                            | ns                              |         |
| **Prevalence of interventricular dyssynchrony** |                       |                                 |                                 |         |
| Interval between pulmonary and aortic pre-ejective periods > 40 ms | 68.8% | 75.0% | ns |
| **Prevalence of intraventricular dyssynchrony** |                       |                                 |                                 |         |
| Criterias                                  |                           |                                 |                                 |         |
| Pitzalis et al16                           | 61.5%                     | 59.3%                           | ns                              |         |
| Notabartolo et al17                        | 56.3%                     | 31.3%                           | ns                              |         |
| Gorcsan et al18                            | 50.0%                     | 31.3%                           | ns                              |         |
| Lane et al19                               | 100.0%                    | 96.9%                           | ns                              |         |
| Cleland et al19                            | 56.3%                     | 62.5%                           | ns                              |         |
| Yu et al20                                 | 87.5%                     | 62.5%                           | ns                              |         |
| De Sutter et al20                          | 87.5%                     | 84.4%                           | ns                              |         |
| Perez de Isla et al21                      | Criterion I               | 62.5%                           | 59.4%                           | ns      |
|                                            | Criterion II              | 43.4%                           | 40.6%                           | ns      |
| Gorcsan et al22                            | 50.0%                     | 74.2%                           | ns                              |         |

ns: non-significant, p > 0.05.
Table 5 – Comparison of echocardiographic data between patients with different axis orientation in the frontal plane

| Variable                        | Axis in the frontal plane | p value |
|---------------------------------|----------------------------|---------|
|                                 | between – 30° and +90°     | < – 30° |
|                                 | (n = 26)                   | (n = 22) |
| **M mode**                      |                            |         |
| Left atrium (mm)                | 42.1 ± 6.3                 | 49.4 ± 5.7 | 0.0001 |
| Left ventricle, diastole (mm)   | 71.45 ± 10.5               | 77.2 ± 8.1  | ns  |
| Left ventricle, systole (mm)    | 61.7 ± 11.6                | 67.4 ± 8.7  | ns  |
| **Two-dimensional**             |                            |         |
| End-diastolic volume (mL)       | 191.2 ± 92.4               | 217.8 ± 80.8 | ns  |
| End-systolic volume (mL)        | 137.9 ± 74.9               | 161.1 ± 52.7 | ns  |
| Left ventricular ejection fraction (%) | 30.4 ± 7.3      | 26.8 ± 6.9  | ns  |
| **Diastolic pattern**           |                            |         |
| Grade IA dysfunction            | 53.9%                      | 18.2%   | 0.01 |
| Grade II dysfunction            | 23.1%                      | 40.9%   |      |
| Grade III/IV dysfunction        | 7.7%                       | 36.4%   |      |
| Undetermined                    | 15.4%                      | 4.6%    |      |
| **Mitral regurgitation**        |                            |         |
| Absent                          | 23.1%                      | 4.6%    |      |
| Mild                            | 50.0%                      | 59.1%   |      |
| Moderate                        | 23.1%                      | 31.8%   |      |
| Severe                          | 3.8%                       | 4.6%    |      |
| **Prevalence of interventricular dyssynchrony** | | |
| Interval between pulmonary and aortic pre-ejective periods > 40 ms | 76.9% | 68.2% | ns |
| **Prevalence of intraventricular dyssynchrony** | | |
| Criteria                        |                            |         |
| Pitzalis et al16                | 63.7%                      | 55.6%   | ns  |
| Notabartolo et al17             | 34.6%                      | 27.3%   | ns  |
| Gorcsan et al18                 | 38.4%                      | 36.4%   | ns  |
| Lane et al19                    | 96.2%                      | 100.0%  | ns  |
| Cleland et al14                 | 61.5%                      | 59.1%   | ns  |
| Yu et al30                      | 76.9%                      | 63.6%   | ns  |
| De Sutter et al22               | 84.6%                      | 83.4%   | ns  |
| Perez de Isla et al23           | Criterion I 53.9%          | 66.2%   | ns  |
|                                 | Criterion II 42.3%         | 40.9%   | ns  |
| Gorcsan et al27                 | 60.0%                      | 72.7%   | ns  |

ns: non-significant, p > 0.05.
