

## Cardiomyopathy Induced by Artificial Heart Stimulation

Cardiomiopatia Induzida por Estimulação Cardíaca Artificial Silas dos Santos Galvão Filho<sup>1,\*</sup>

With the advent of cardiac pacemakers more than 60 years ago, the era of artificial cardiac pacing began, which changed the natural history of symptomatic bradycardias, significantly increasing the survival especially of patients with complete atrioventricular block. Initially, the pacemakers were ventricular single chamber (VVI), bringing some limitations in patients with bradycardia consequent to sinus dysfunction, which mostly presented ventriculoatrial retrograde conduction, determining atrioventricular dyssynchrony with damage to cardiac function and consequent symptoms, which was called pacemaker syndrome<sup>1</sup>. In the early 1980s, the incorporation of one more electro in the atrium and the development of dual chamber pacemakers (DDD), also called "physiological pacemakers", recovered the atrioventricular synchronism and solved this problem. The success of dual-chamber pacemakers led to the idea that they could further increase the survival of their carriers compared to VVI. However, although DDD pacemakers show benefits in tolerance to effort and improvement in quality of life, no significant increase in survival was evidenced in this comparison.

In the first years of the second millennium, concomitant to the appearance of cardiac resynchronization therapy (CRT) and the concepts of dyssynchronization, it was observed that patients with double chamber devices that presented effective ventricular pacing in more than 40% of the heartbeat presented some problems, such as: appearance or worsening of heart failure, development of atrial fibrillation and even increased mortality<sup>2–4</sup> when compared to those who did not present RV stimulation, being that the worse and the greater the percentage of ventricular stimulation. These problems were attributed to ventricular dyssynchrony caused by RV stimulation and led to the development of programmable algorithms of minimal ventricular pacing in artificial cardiac pacing devices in order to avoid unnecessary ventricular stimulations in patients who had preserved atrioventricular and interventricular conduction.

Already in cases of atrioventricular blocks, where RV stimulation is absolutely necessary, new optional stimulation sites to the classical RV tip stimulation have been sought. In this sense, several ventricular pacing sites were attempted: RV outflow tract, mid-septal, inferoseptal region, without, however, showing significant differences when compared to classical RV tip stimulation<sup>5,6</sup>.

In recent years, the development of special sheaths and electrodes for stimulation of the His bundle enabled this type of cardiac stimulation that preserves the interventricular conduction through the normal conduction system, consequently avoiding ventricular dyssynchrony induced by RV stimulation<sup>7,8</sup>. This modality of stimulation, however, has its indication in cases that present intraventricular conduction preserved through the normal conduction system, because in cases where there are intraventricular conduction disorders such as the left bundle branch block (LBBB), the His bundle stimulation does not bring benefits, since it determines the persistence of conduction disorder and, consequently, dyssynchrony. More recently, using the same system developed for His pacing, the muscle region of the interventricular septum was stimulated, immediately below the fibrotic portion where the trunk of the His bundle

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is located, penetrating the interventricular septum and stimulating the initial portions of the His bundle branches, correcting in many cases conduction disorders such as LBBB. This promising modality of artificial cardiac pacing has been placed as an option of unisite pacing to CRT with biventricular pacing, although it still needs more knowledge of its long-term results.

Our idea that ventricular dyssynchrony could induce dilated myocardiopathy started with a study we presented in 1997 at the *Congresso Brasileiro de Arritmias Cardíacas* (Brazilian Congress of Cardiac Arrhythmias)<sup>10</sup>, where a study was presented with 12 patients with idiopathic dilated cardiomyopathy who had ventricular dyssynchrony (LBBB), reduced left ventricle ejection fraction (LVEF) (< 35%), and CHF refractory to drug therapy, which, after CRT, normalized the cardiac systolic function with LVEF normalization. In three of these patients there was normalization of all echocardiographic parameters with the examination showing a structurally normal heart. We attributed the appearance of dilated cardiomyopathy considered idiopathic to the dyssynchrony determined by LBBB, which we named "dyssyncronyomyopathy".

Conventional RV pacing determines the electrocardiographic pattern of LBBB, with QRS width frequently greater than 150 ms and changes in myocardial ventricular contractility (dyssynchrony) similar to LBBB, with damage to the cardiac systolic function<sup>11</sup>. In some (but not all) patients, these alterations may be responsible for the appearance and/or worsening of heart failure and even structural alterations of the heart, and may therefore be responsible for the development of "dyssyncronyomyopathy". These patients benefit greatly from the CRT upgrade call, with the implantation of another electrode for LV stimulation.

Gage et al.<sup>12</sup> published a study in which they compared the clinical and echocardiographic response to CRT of patients with or without previous conventional cardiac pacing (PCCP), showing a similar positive clinical response between both groups and a better response of patients with PCCP regarding ventricular function evaluated by eco. In addition, a trend of lower mortality of patients with PCCP was demonstrated when compared to those with no PCCP. The authors concluded that patients submitted to CRT upgrade of pacemakers present a similar, if not better, response than CRT in patients without PCCP.

Several studies such as that of Martineli et al.<sup>13</sup> showed better results of CRT implantation when compared to the conventional pacemaker in patients with indication for ventricular pacing and reduced ventricular systolic function. Recently, a systematic review<sup>14</sup> was published involving 679 patients from eight studies that evaluated the impact of what they called physiological cardiac pacing (PCP) (CRT or His/ParaHis) versus conventional cardiac pacing in patients with indication for cardiac pacemakers and moderately impaired ventricular systolic function (> 35%), and significant superiority of PCP in relation to echocardiographic parameters (LVEF, LV end diastolic volume, LV end systolic volume) was demonstrated.

In the European Society of Cardiology (ESC) for the management of CHF 2016<sup>15</sup>, the CRT is class 1 in patients with indication for cardiac pacing who require RV pacing and have EF < 40%, to the detriment of conventional pacemakers. In these guidelines, the CRT upgrade for pacemaker is indicated for patients with systolic dysfunction (LVEF < 40%) who present worsening of CHF after device implantation with RV pacing in more than 40% of the heartbeats.

Although it may cause damage to the cardiac systolic function in some patients, it may even be responsible for the appearance/aggregation of dilated cardiomyopathies called "dyssyncronyomyopathies", conventional unisite pacing RV remains routinely used. The good evolution of the majority of conventional pacemaker patients, the greater complexity of biventricular pacing procedures and even the much higher costs of CRT devices limit the more frequent use of this type of cardiac pacing in patients with indication for pacemakers that require ventricular pacing. Transeptal stimulation of the initial portions of His bundle branches can be an excellent option for unisite stimulation from RV to CRT with biventricular stimulation.

Having knowledge of the possible damage caused by unisite RV stimulation of conventional pacemakers, the carriers of these devices should be evaluated in relation to the structural and functional part of the heart, through imaging methods such as echocardiography, at least annually and, if they show relevant and/or progressive changes, upgrade for CRT should always be considered.

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