Original Article
Use of a dual-chamber pacemaker to achieve an excellent response to cardiac resynchronization therapy in patients with dilated cardiomyopathy and complete left bundle branch block: a case report

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Abstract: The atrioventricular delay (AVD) in cardiac resynchronization therapy (CRT) is usually programmed to be shorter than the physiological AVD, which can lead to patients non-responsive to CRT. Although the programming of rate-adaptive AV delay (RAAVD) has been recommended for the optimization of the AVD in CRT with a triloculare pacemaker, the response to CRT is disappointing. Moreover, few studies have reported a dual chamber pacemaker to achieve response to CRT based on RAAVD algorithm. This study reported a 58-year-old male patient who was diagnosed with dilated cardiomyopathy and complete left bundle branch block. He achieved an excellent response to CRT with a dual-chamber pacemaker by left univentricular pacing with RAAVD algorithm.

Keywords: Cardiac resynchronization therapy, dilated cardiomyopathy, rate-adaptive atrioventricular delay algorithm, optimal atrioventricular delay

Introduction
Cardiac resynchronization therapy (CRT) has been demonstrated to significantly improve cardiac function and clinical outcomes in patients with New York Heart Association (NYHA) class III or IV congestive heart failure (CHF) and a left ventricular ejection fraction (LVEF) less than 30%. However, approximately 20-30% of patients are non-responsive to CRT [1]. Fixed short atrioventricular delay (AVD) in current CRT [2] has been considered as one of the reasons, which can abolish the physiological AVD of atrioventricular node (AVN), thereby counteracting the benefit of standard biventricular (Biv) pacing. It is well known that physiological AVD plays a key role in achieving optimal atrioventricular (AV) synchrony and atrial contribution to ventricular filling [3]. Therefore, the optimal AVD is important for patients' response to CRT. Currently, the programming of rate-adaptive AV delay (RAAVD) has been recommended for the optimization of the AVD in atrioventricular block pacing [4]. To date, few studies have reported a dual chamber (DDD) pacemaker achieving response to CRT based on RAAVD algorithm. Here, we reported a patient with dilated cardiomyopathy (DCM) and chronic complete left bundle branch block (CLBBB) who was implanted with a DDD pacemaker to achieve an excellent response to CRT by left-univentricular (LUV) pacing with a novel RAAVD algorithm.

Case presentation
A 58-year-old male patient was admitted to the hospital with palpitations and dyspnea after physical activities for more than 1 year (approved by ethical committee of the First affiliated hospital of Kunming medical university). There were no special records in the medical history and neither alcohol nor smoking addiction was recorded. Physical examination showed that cardiac dullness area was extended in the left side and no pathological murmurs were heard in all cardiac auscultation areas. Heart rate (HR) was 64 beats/min without arrhythmia
and LVEF was 34%. Electrocardiograph (ECG) revealed a sinus rhythm and CLBBB. Finally, the patient was diagnosed with DCM, CLBBB and NYHA class III by echocardiography.

Based on the diagnosis of DCM and CLBBB, the patient accepted the pacemaker implantation. After the informed consent to implantation of pacemaker was obtained, a DDD pacemaker (Relia RED01, Medtronic, Inc., Minneapolis, MN, USA) was implanted on October 15, 2013. The surgery of DDD pacemaker implantation was performed according to the standard procedure. X-ray percutaneous transluminal coronary angioplasty (PTCA) wire guided electrode of left ventricular lead to position in the left posterolateral cardiac vein and right atrium electrode in the right atrial appendage. The left ventricular electrode lead and right atrial electrode lead were connected to the jacks on the pulse generator correspondently. The operation was finished successfully without complications.

The patient was followed up after short time of interval optimization with echocardiography. Start and stop rates were set at 60 bpm and 100 bpm according to the lower and upper limit heart rate in holter ECG recording, respectively. Interval from onset of A wave to atrial sensing in intracardiac electrograms was 28 ms. Therefore, atrial sense compensation (ASC) was set at 30 ms, and PR interval at optimized ASC (190 ms - 30 ms) was taken as baseline interval (160 ms). Because a sensed AV (SAV) interval of 150 ms (160 ms - 10 ms) achieved the largest aortic velocity-time integral (AVI), the optimal AVD was set at 150 ms. Then, RAAVD was programmed as following: SAV interval for start rate = optimal AVD + (intrinsic PR interval at start rate - intrinsic PR interval at optimization) = 150 + (220 - 190) = 180 (ms); paced AV (PAV) interval for start rate = SAV interval for start rate + ASC = 180 + 30 = 210 (ms); and maximum offset was set at -40 ms (intrinsic PR interval at stop rate - intrinsic PR interval at start rate). Medications were unchanged during follow-up (oral administration of 40 mg furosemide and 25 mg spironolactone daily). Transthoracic echocardiogram was performed at the follow-up of 6 and 12 months after the device implantation. Cardiac output was assessed using the LV outflow tract velocity-time integral.
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Table 1. NYHA class and echocardiographic parameters of baseline, 6 and 12 months post-implantation

<table>
<thead>
<tr>
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<th>Before CRT</th>
<th>6 months post</th>
<th>12 months post</th>
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<tbody>
<tr>
<td>NYHA class</td>
<td>III</td>
<td>II</td>
<td>I</td>
</tr>
<tr>
<td>LVEF (%)</td>
<td>34</td>
<td>47</td>
<td>58</td>
</tr>
<tr>
<td>LVEDD (mm)</td>
<td>69</td>
<td>58</td>
<td>53</td>
</tr>
<tr>
<td>LAD (mm)</td>
<td>36</td>
<td>24</td>
<td>27</td>
</tr>
<tr>
<td>AVI (cm)</td>
<td>24.9</td>
<td>25.3</td>
<td>28.2</td>
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<tr>
<td>IVMD (ms)</td>
<td>111</td>
<td>74</td>
<td>69</td>
</tr>
<tr>
<td>E/A Pd (ms)</td>
<td>383</td>
<td>300</td>
<td>235</td>
</tr>
<tr>
<td>Ts-SD12 (ms)</td>
<td>92</td>
<td>61</td>
<td>70</td>
</tr>
</tbody>
</table>

NYHA, New York heart association; CRT, cardiac resynchronization therapy; LVEF, left ventricular ejection fraction; LVEDD, left ventricular end diastolic diameter; LAD, left anterior descending; AVI, aortic velocity-time integral; IVMD, interventricular mechanical delay; E/A Pd, E/A procedure duration; Ts-SD12, standard deviation of time intervals of the 12 LV segments.

Discussion

In this study, for the first time, we implanted a dual chamber pacemaker and developed a novel algorithm to preserve the physiological AVD. The results showed that QRS duration was shortened after LUV pacing with RAAVD, as well as the echocardiographic parameters such as NYHA class, LVEF, LVDD, Ts-SD12, AVI, IVMD, and E/A Pd were all improved after DDD pacemaker implantation for 6 and 12 months, which suggested that the patient with DCM and CLBBB achieved an excellent response to CRT by LUV pacing with this novel RAAVD algorithm.

Our results showed that the LUV pacing with RAAVD algorithm shortened the QRS duration from 200 ms to 137 ms in patients with DCM and CLBBB. Because the BiV pacing activation was slowly conducted retrograde through cardiomyocytes via His-Purkinje system, the time for RV activation was longer under the standard BiV pacing than that for physiological activation, which manifested as the longer QRS duration. Previous research had proved that the abnormality of intraventricular conduction was associated with the extension of QRS duration, which increased the percentage of mechanical dyssynchrony, thereby leading to the lower cardiac function and the higher mortality [5]. Therefore, narrow QRS wave under LUV pacing with RAAVD algorithm may obtain more benefit than standard BiV pacing and LUV pacing with fixed AVD.

The first study of the RAAVD in patients with CRT was reported by Scharf et al. and the results showed that the optimal AVD increased during exercise and a progressive fusion at longer AVD might increase ventricular resynchronization [6]. Therefore, the recommendation by Scharf et al. was suggested to be used with extreme caution. In addition, determined the optimal AVD by simultaneous measurement of electrocardiographic and doppler-echocardiographic parameters, and then programmed the RAAVD in CRT [1]. The results showed that optimal AVD was shortened with increased HR and the RAAVD parameters in CRT patients was advised to be turned off. Excitingly, our study showed that QRS duration and morphology did not change with varying HR, which demonstrated that RAAVD algorithm could track physiological AVD and enable fusion of intrinsic right bundle conduction with paced LV conduction,
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thereby solving the problem that the changes of fusion and QRS duration resulted from fixed AVD and intrinsic conduction. This may partly explain the results of the previous studies which showed that fusion was not established and LV pacing was not superior to BiV pacing [7, 8]. Furthermore, the left-sided AVD automatically tracked physiological AVD by RAAV algorithm to allow fusion with intrinsic conduction coming from the normal-conducting right bundle branch, which averted the deleterious effects from right ventricular (RV) pacing, then increased device longevity due to unnecessary RV pacing. Higher percentage of ventricular pacing in patients with intact AVD had been found to be associated with increased incidence of atrial fibrillation and heart failure [9]. Our previous study showed that preserving intrinsic conduction via the right bundle branch potentially and pacing LV simultaneously could improve acute hemodynamic effect than standard right BiV pacing and avert deleterious effects from RV pacing [10].

AV and interventricular interval optimization were time-consuming in conventional BiV pacing, and it was difficult to achieve individualization and dynamic optimization [11]. However, under LUV pacing with RAAV algorithm, the intrinsic PR interval was the physiological and optimal AVD, it was unnecessary to optimize. Therefore, LUV pacing RAAV algorithm could realize dynamic optimization of AVD, which adapted to physiological alterations during exercise and sympathetic tone changes, and thereby preserving optimal atrial contribution to ventricular filling. However, a randomized, multicenter, double-blind, controlled clinical research is needed to confirm our finding.

The algorithm tracking physiological AVD in LUV pacing will contribute to the research and development of new pacemaker. Previous study had shown that the numerous proposed methods, including predefined formulas, iterative methods and automatic methods, were used to obtain optimized AVD. Recent study had shown that adaptive CRT algorithm (Medtronic, Inc.) significantly reduces RV pacing and improves response rates to CRT [12]; however, adaptive CRT algorithm was based on three chamber pacemaker.

In conclusion, our LUV pacing with RAAV algorithm could be used in both dual and three-chamber pacemaker. This novel algorithm will cause dispute on preserve or abolish of the physiological AVD of AVN in CRT. The patient with DCM and CLBBB who use dual-chamber pacemaker achieved superior response to CRT by LUV pacing with RAAV algorithm.

Figure 2. Chest X-ray before and 1 year after PM implantation. A: Before PM implantation, chest X-ray showed LV dilated. CTR was 56%. B: 12 months after PM implantation, great reduction in ventricular size, the CTR decreased from 56 to 49%.
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Figure 3. Ts-SD12 at base and papillary level were also improved 12 months after RAAV LUV pacing. A: TS-SD12 was 92 ms before RAAV LUV pacing; B: TS-SD12 was 70 ms 12 months post-implantation.
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Disclosure of conflict of interest

None.

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