Cardiac contractility modulation therapy: Are there superresponders?

Bandar Al-Ghamdi, MD, FACC, FHRS, CCDS, *[†] Azam Shafquat, MD, MBBS, FHRS, *[†] Yaseen Mallawi, MD*

From the *Heart Centre, King Faisal Specialist Hospital and Research Centre, Riyadh, Saudi Arabia, and [†]Alfaisal University, Riyadh, Saudi Arabia.

Introduction

Heart failure (HF) is a common cardiovascular disease, and it is associated with high morbidity and mortality despite advances in medical and device-related management.

Patients with HF with low left ventricular ejection fraction (LVEF) and narrow QRS complexes represent a special challenge, as they are not candidates for cardiac resynchronization therapy (CRT). These patients may benefit from cardiac contractility modulation (CCM) as shown by several studies.^{1,2} Patients with HF New York Heart Association (NYHA) functional class III and an LVEF of $\geq 25\%$ respond exceptionally well to CCM therapy as suggested by subgroup analysis.³

We are presenting a case of dilated cardiomyopathy with significant improvement in HF symptoms and LVEF after initiating CCM therapy.

Case report

We are reporting the case of a 35-year-old male patient with a history of dilated cardiomyopathy and an LVEF of 25% for >2 years. He has an 11-year history of smoking but has no other risk factors for coronary artery disease. His cardiac workup for coronary artery disease was found to be negative using the nuclear stress test.

An implantable cardioverter-defibrillator (ICD) was implanted for primary prevention of sudden cardiac death 8 months previously with no complications. Despite being on guideline-directed medical therapy (GDMT), he was still complaining of shortness of breath on exertion (NYHA functional class III). There was no history of orthopnea or paroxysmal nocturnal dyspnea. His 12-lead electrocardiogram revealed a sinus rhythm at 70 beats/min with a narrow QRS complex. Echocardiogram at baseline and 12 months on GDMT showed severely reduced left ventricular systolic

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function with an LVEF of 25%. There were no regional wall motion abnormalities. The right ventricular systolic function was moderately reduced, but no significant valvular lesion was noted, and pulmonary artery pressure was normal.

The option of CCM therapy was discussed with the patient, and he accepted the procedure. The device was implanted into the right pectoral region. Three pacemaker leads were inserted via the right subclavian vein; 1 lead was placed in the right atrium (at right atrial appendage), and 2 leads were placed at the right ventricular septum about 2 cm apart. During the procedure, a crosstalk test with ICD was performed while the CCM device was active and no significant interaction between the 2 devices was identified. The procedure was uneventful, and the postprocedure chest radiograph is shown in Figure 1. At 3-month follow-up, he reported feeling better with improvement of his shortness of breath on exertion (NYHA functional class I). A 6-minute walk test showed improvement in walking distance from 363 to 528 m.

On cardiopulmonary exercise testing, his maximal oxygen consumption/maximal oxygen uptake ($\dot{V}O_{2max}$) improved from 15.9 to 19.7 mL/(kg•min).

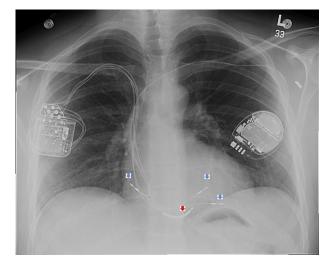


Figure 1 Posteroanterior chest radiograph showing the implantable cardioverter-defibrillator (ICD) on the left upper chest with an ICD lead (red arrow) and the cardiac contractility modulation device on the right upper chest with its leads (blue arrows).

Address reprint requests and correspondence: Dr Bandar Al-Ghamdi, Heart Centre, King Faisal Specialist Hospital and Research Centre, MBC-16, PO Box 3354, Riyadh 11211, Saudi Arabia. E-mail address: balghamdi@kfshrc.edu.sa.

KEY TEACHING POINTS

- Heart failure is a common health care problem.
- Patients with heart failure and narrow QRS complexes who are refractory to medical therapy represent a management challenge, as they are not candidates for cardiac resynchronization therapy.
- Cardiac contractility modulation (CCM) therapy represents a promising therapeutic modality for these patients.
- CCM is associated with increased maximal oxygen consumption and improved quality of life in patients with heart failure.
- CCM may lead to left ventricular reverse remodeling, and it is expected to increase left ventricular ejection fraction (LVEF) by $\sim 5\%$.
- Patients with heart failure New York Heart Association functional class III and an LVEF of ≥25% respond exceptionally well to CCM therapy as suggested by subgroup analysis of large studies in this field.

His echocardiogram showed improvement in LVEF from 20%–25% to 40%–45%. The right ventricle was normal in size and function, and the left ventricular internal diameter at end-diastole and end-systole improved from 7.2 and 6.6 to 6.5 and 5.3 cm, respectively (Figure 2).

Discussion

In patients with symptomatic HF despite GDMT, CRT has proven to be an effective treatment with improvement in 6-minute walk distance, NYHA functional class, quality of life, \dot{VO}_{2max} , reduced left ventricular volumes and mitral regurgitation, and reduced all-cause mortality or hospitalization.⁴ However, CRT is generally recommended for patients in sinus rhythm and prolonged QRS complex (\geq 120 ms) with left bundle branch block or a QRS complex width of \geq 150 ms in the absence of left bundle branch block. In patients with symptomatic HF with narrow QRS complex despite GDMT, CCM may represent an attractive alternative therapy.

We are presenting a case with an exceptionally good response to CCM therapy in a patient with dilated cardiomyopathy and narrow QRS complex who had been on GDMT for at least 1 year with no significant improvement in his symptoms and LVEF.

The CCM device is a cardiac implantable electronic device that enhances ventricular contractile strength by delivering high-voltage, nonexcitatory, biphasic waveform, electrical impulses during the absolute refractory period of the cardiac muscle cells.⁵ These signals do not affect cardiac activation sequence or initiate a new contraction.⁵

To date, the only clinically available system for CCM delivery is the Optimizer IVs system (Impulse Dynamics Inc., Orangeburg, NJ). The device is similar to a pacemaker and consists of 4 components: implantable pulse generator, leads (1 atrial and 2 ventricular leads), battery charger, and a programming unit (Figure 3). The CCM device is implanted into the pectoral region, and 3 bipolar pacemaker leads are introduced into the right side of the heart (commonly Tendril ST, St. Jude Medical, Inc., St. Paul, MN). Two leads are

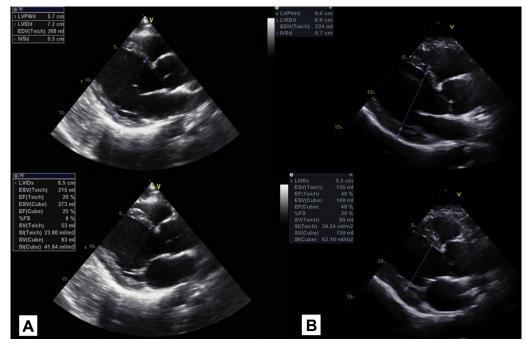


Figure 2 Echocardiogram parasternal long-axis views with left ventricular internal diameter end-diastole (LVIDd) and end-systole (LVIDs), pre (A) and post (B) cardiac contractility modulation implantation.

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