High-Grade Atrioventricular Block Requiring Pacemaker Implantation after Cardiac Transplantation: An Unusual Complication

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ECG was suggestive of the right bundle branch block (RBBB) with normal PR interval. The posttransplant patient was started on triple immunosuppressive therapy including tacrolimus, mycophenolate mofetil, and prednisolone as per institutional protocol. Epicardial backup pacing was kept in place as a part of the protocol. The patient was noted to have increasing pacing requirements starting from day 7 in postoperative period in view of 2:1 AV block [Figure 1] with intermittent complete heart block. After ruling out reversible causes (hyperkalemia and AV-blocking drugs) and procedural causes (prolonged ischemia time and perioperative injury to AV node), the evaluation for early rejection was planned. Endomyocardial biopsy and electrophysiological (EP) study

**Case Report**
A 37-year-old patient underwent bicalval OHT for dilated cardiomyopathy. Pretransplant evaluation of donor heart was negative for any structural heart disease or conduction system disease. Baseline electrocardiogram (ECG) of donor heart was normal. The transplant procedure was uneventful with total ischemia time of <3 h. Immediately after the transplant, the ECG was suggestive of the right bundle branch block (RBBB) with normal PR interval. The posttransplant patient was started on triple immunosuppressive therapy including tacrolimus, mycophenolate mofetil, and prednisolone as per institutional protocol. Epicardial backup pacing was kept in place as a part of the protocol. The patient was noted to have increasing pacing requirements starting from day 7 in postoperative period in view of 2:1 AV block [Figure 1] with intermittent complete heart block. After ruling out reversible causes (hyperkalemia and AV-blocking drugs) and procedural causes (prolonged ischemia time and perioperative injury to AV node), the evaluation for early rejection was planned. Endomyocardial biopsy and electrophysiological (EP) study

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were done on postoperative day 19. EP study revealed 2:1 AV conduction with prolonged HV interval (130 ms) suggestive of high-grade infra-Hisian AV block [Figure 2]. Histopathological examination and immunohistochemistry of the biopsy specimens were suggestive of mild acute cell-mediated rejection (the International Society for Heart and Lung Transplantation, Grade 1R) and were negative for pathologic antibody-mediated rejection [Figure 3]. Serum tacrolimus levels were found to be low, and with early cell-mediated rejection in consideration, the dose of tacrolimus and steroids was increased. However, the patient continued to have intermittent high-grade AV block. Repeat tacrolimus levels were in range. A permanent pacemaker implantation (single chamber, VVIR) was done on postoperative day 22. The patient was then discharged on immunosuppressive therapy in stable condition. On follow-up after 1 week, the patient was in sinus rhythm with RBBB [Figure 4] and pacemaker interrogation revealed minimal pacing requirement.

**DISCUSSION**

Bradyarrhythmias after OHT are variably reported, but the most common cause remains sinus node dysfunction. Conduction system abnormalities leading to bradyarrhythmia are distinctly uncommon. Incomplete RBBB is the most common conduction abnormality seen in up to 70% of patients following OHT.[1] The conduction abnormality may persist and may be causally associated with prolonged donor ischemic time and episodes of rejection.[2] However, high-grade AV block remains uncommon, particularly in the early postoperative period. According to a study of 1047 posttransplant patients, first-degree, Mobitz Type I, Mobitz Type II, and complete heart blocks occurred in 8.3%, 0.6%, 0.1%, and 1.8% of patients, respectively.[3]

Whenever AV block occurs after cardiac transplantation, first reversible causes (e.g., hyperkalemia and AV-blocking drugs) should be ruled out and corrected, if any. The preoperative use of amiodarone in the recipient is an important cause of reversible posttransplant bradycardia that should not be missed. Periprocedural injury to AV node and prolonged ischemia time are also associated with postoperative AV node dysfunction and should better be avoided. Rarely, intraoperative coronary air embolism may also cause postoperative conduction disturbances. In a study by Leonelli et al., the duration of graft ischemia time and severity of rejection were the major determinants of conduction abnormality after OHT.[4] A total ischemia time >160 min was associated with persistence of conduction abnormality. Ice preservation of donor heart has been proposed as the likely cause of injury to conduction system. However, postoperative AV block is usually transient.

**Figure 1:** A 12-lead electrocardiogram done on posttransplant day 7 showing 2:1 atrioventricular block with wide QRS complex conduction (left anterior hemiblock and right bundle branch block) suggestive of infra-Hisian block

**Figure 2:** Electrophysiological study trace demonstrating infra-Hisian atrioventricular block (HV interval of 130 ms)

**Figure 3:** (a) Histopathological examination suggestive of mild acute cellular rejection (International Society for Heart and Lung Transplantation, 2004 Grade 1R) and (b) CD68 positivity on immunohistochemistry

**Figure 4:** Normal sinus rhythm with right bundle branch block on follow-up
and only rarely requires permanent pacemaker implantation. In a contemporary case series on 1179 transplant patients, prolonged operative time and biatrial anastomosis were identified as independent risk factors for permanent pacemaker implantation. Cardiac conduction tissue is also a specific target related to allograft rejection. The arteries to sinoatrial and AV tissue are frequently involved in both acute cellular reaction and chronic intimal fibrocellular rejection reaction, causing conduction disturbances. Cardiac rejection should be strongly suspected and ruled out with certainty in cases of postoperative AV blocks. Early deceleration of immunosuppressive therapy and underdosing may precipitate early rejection and should be avoided. Cardiac endomyocardial biopsy is the gold standard for diagnosing cardiac rejection and must be done whenever rejection is suspected. If the AV block persists even after treatment of rejection or EP studies are suggestive of high-risk features, implantation of permanent pacemaker may be the right approach. In the present case, EP study helped to guide the approach and tilted the balance toward advocating permanent pacemaker implantation in view of infra-Hisian block.

In a retrospective study of 114 heart transplant recipients, 14 patients (12%) required temporary pacing and 4 patients (3.5%) required permanent pacing. The indication for permanent pacemaker implantation was sinus node dysfunction in 3 patients (75%) and AV block in 1 patient (25%). They observed transplant rejection in 3 patients (21.4%) requiring temporary pacing and in 2 patients (50%) requiring permanent pacing. Although pacemaker implantation is performed in approximately 10–15% of cardiac transplant recipients, AV block accounts for <20% of these cases. Median time to pacemaker implantation is much later for AV block than for sinus node dysfunction (1511 vs. 27 days in a study by Wellmann et al.).

To sum up, high-grade AV blocks are uncommon in cardiac transplant recipients (especially in early postoperative period), and if noted, acute rejection and other reversible causes should be carefully ruled before considering for permanent pacemaker implantation.

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Conflicts of interest
There are no conflicts of interest.

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