Obstruction in Hypertrophic Cardiomyopathy
How Often Does It Occur? Should It be Treated? If So, How?

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Hypertrophic cardiomyopathy (HCM) is the most common monogenic cardiac disorder and has been estimated to occur in 1 of every 500 people in the general population, amounting to a total of ≈600,000 persons in the United States. Its pathophysiology and optimal management have been the subject of conjecture and debate for more than a century. The issues surrounding left ventricular outflow tract (LVOT) obstruction in HCM have evoked the most discussion.

In 1907, a German pathologist, A. Schminke, described 2 hearts from women in their mid-50s. Decades before the development of left heart catheterization, and before any pressure gradients had ever been measured in humans, he wrote the following: “Diffuse muscular hypertrophy of the left ventricular outflow tract causes an obstruction. The left ventricle has to work harder to overcome the obstruction. So, the primary hypertrophy will be accompanied by a secondary hypertrophy, causing an incremental (further) narrowing of the outflow tract.” Thus, Schminke presciently understood that the obstruction can only be explained by muscular hypertrophy of the left ventricular outflow tract.

As open-heart surgery exploded in the early 1960s, patients with this condition (the name HCM had not yet been agreed on) were encountered with increasing frequency by cardiologists and cardiac surgeons around the world; the British surgeon, Sir Russell Brock provided especially useful insights. Indeed, HCM soon became the poster child for the hemodynamic era. Although, by definition, all patients had LVOT obstruction, the primary hypertrophy will be accompanied by a secondary hypertrophy, causing an incremental (further) narrowing of the outflow tract.

Now, to fast-forward to 2006, when the Marons and their colleagues showed by echocardiography that LVOT gradients at rest or during exercise occur in 70% of patients with HCM, are frequently associated with symptoms and adverse clinical outcomes, and concluded that when these obstructions are not responsive to pharmacological therapy, they require mechanical relief. Two approaches to accomplish this are available today; the first is SSM, sometimes referred to as the Morrow procedure, which has gradually become both more extensive and safer over the years. In 1995, alcohol septal ablation (ASA) was introduced, and because it relieved obstruction without requiring open-heart surgery, it quickly became the more frequently used procedure. However, SSM has continued to be carried out, mostly at specialized centers, and it is still considered to be the gold standard.

Three major comparisons are available. Argawal et al performed a comparison of the results in 8 institutions in which both procedures were used; 326 patients received SSM and 380 ASA. Leonardi et al compared the results on 1887 patients who received SSM with those on 2153 patients who received ASA, usually in different hospitals. Several studies found their way into both comparisons. Both analyses concluded that the mortality rates were similarly low with both procedures. However, Argawal et al pointed out that ASA increased the need for implantation of a permanent pacemaker and left patients with low, but significantly higher, LVOT pressure gradients than did SSM.

In this issue of Circulation, the excellent study by Sorajja et al from the Mayo Clinic, Rochester provides the largest...
single-center comparison of 177 patients who underwent ASA with an equal number of age- and sex-matched patients who underwent SSM. These 2 groups were not randomized, and the time that these procedures were carried out differed, from 1998 to 2010 for ASA and from 1983 to 2001 for SSM. At baseline, when compared with the SSM patients, the ASA patients had a significantly greater incidence of New York Heart Association class III/IV, and more frequently had a history of coronary artery disease and of treatment with a β-blocker. Quite remarkably, the 8-year survival estimates were identical, at 79% in both groups. However, the ASA patients had a residual gradient averaging 11 mm Hg and early pacemaker dependency in 20% of patients, compared with 5 mm Hg and 2%, respectively for SSM patients. During prolonged follow-up of ASA patients, 5.6% required subsequent SSM and 2.8% repeat SA. The postablation LVOT gradient in ASA patients was an independent predictor of all-cause mortality and subsequent need for these reinterventions.

The American College of Cardiology/American Heart Association Guidelines for the Treatment of HCM published in 2011 recommended, with a Class I indication, that “septal reduction therapy should be performed only by experienced operators … and only for the treatment of patients with severe drug-refractory symptoms and LVOT obstruction.” They also provided a IIa recommendation that SSM “is the first consideration for the majority of eligible patients with HCM” and that “when surgery is contraindicated, or the risk is considered unacceptable because of serious comorbidity or advanced age, ASA … can be beneficial in … patients with HCM with LVOT obstruction and severe drug-refractory symptoms.” The study by Sorajja et al18 provides further support for these recommendations.

Ordinarily, when there is a question regarding the choice between 2 competing therapeutic approaches, a randomized clinical trial is used to provide the answer. However, Olivotto et al20 have made the case, and I believe quite convincingly, that such a trial presents logistical hurdles that cannot be surmounted in the foreseeable future.

So, where do go from here? The available comparisons between the 2 approaches to septal reduction therapy are largely retrospective, and the risks for bias, both in patient selection and publication, creep in. A prospective multicenter registry that provides detailed baseline characteristics and descriptions of the procedure, as well as careful and systematic follow-up, could be very helpful. Much of the early work on HCM was conducted at the National Heart, Lung, and Blood Institute, which has had extensive favorable experience with prospective registries—on cardiac transplantation, pulmonary hypertension, and assisted circulation, to name a few. It would now be quite appropriate for the institute to organize a comparison of these 2 techniques of septal reduction, which would be relatively inexpensive and cost-effective at a time of fiscal stringency.

**Disclosures**

None.

**References**


**Key Words:** Editorials ■ ablation techniques
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Circulation. 2012;126:2369-2370; originally published online October 17, 2012; doi: 10.1161/CIRCULATIONAHA.112.144667

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