Editorial

Who needs ventricular stimulation studies?

Two randomised studies reported in the New England Journal of Medicine have cast doubt on the value of ventricular stimulation studies in the assessment of antiarrhythmic drug therapy in patients with symptomatic sustained ventricular arrhythmias. In the study by Steinbeck and coworkers 115 patients with inducible ventricular arrhythmias were assigned to treatment with electrophysiologically guided drug therapy or with empirical metoprolol. During a mean follow up of 23 months the incidence of recurrent arrhythmia and sudden death was almost identical in the two groups. Steinbeck et al concluded that electrophysiologically guided drug therapy did not improve the overall outcome of patients with sustained ventricular arrhythmias. In the Electrophysiologic Study Versus Electrocardiographic Monitoring (ESVEM) study 486 patients with both inducible ventricular arrhythmias and frequent ventricular extrasystoles during Holter monitoring were randomly assigned to antiarrhythmic therapy guided by either electrophysiologic study or by Holter monitoring. Over a 6 year follow up there was no significant difference between the two study groups in terms of recurrence of arrhythmia, sudden death, or total mortality. The ESVEM group recommended that treatment with sotalol (the most effective drug in the trial) and assessment of its potential efficacy by Holter monitoring was a reasonable initial strategy in patients with characteristics similar to those of their study population.

A casual reader of the New England Journal of Medicine might be forgiven for thinking that there are now few, if any, indications for ventricular stimulation studies in the management of patients with symptomatic sustained ventricular arrhythmias.

Current practice

Ventricular stimulation studies are widely used for the management of recurrent sustained ventricular tachycardia and cardiac arrest in both Europe and the United States. This widespread use is not, however, based on an assumption that electrophysiologically guided therapy itself improves outcome in these patients. Indeed it has been frequently stated that this may not be so, and the results of CAST in 1989 underlined this possibility. It is increasingly recognised that antiarrhythmic drug therapy, particularly with agents of Vaughan Williams class 1, may have a deleterious effect on overall mortality in several different groups of patients.

Programmed electrical stimulation of the ventricle is performed principally as a means of assessing prognosis in patients with recurrent ventricular arrhythmias, rather than as a means of selecting a particular antiarrhythmic drug. Gone are the days of prolonged hospital admissions with repeated testing of different classes of antiarrhythmic drugs or different drugs from within the same class. More relevant is the ability of the technique to provide prognostic information that, in conjunction with the clinical history (ventricular tachycardia or cardiac arrest) and an index of ventricular function, allows rational selection of those patients most likely to benefit from non-pharmacological therapy. For instance, it has been realised for some time that the prognosis of patients with ventricular arrhythmias that are not inducible by ventricular stimulation is relatively good. This is borne out by the study of Steinbeck et al, in which recurrent arrhythmia and sudden death were significantly less common in patients with non-inducible arrhythmia at baseline than in patients with inducible arrhythmia, irrespective of whether the patients with inducible arrhythmia were assigned to electrophysiologically guided drug therapy or to empirical metoprolol.

Similarly patients with inducible arrhythmias that become non-inducible on drug therapy are known to have a better prognosis than those in whom the arrhythmia remains inducible despite drug therapy. This has been confirmed by several studies with different drugs including that of Steinbeck and coworkers. Persistence of arrhythmia inducibility despite drug therapy has been shown to be an independent predictor of sudden death or recurrent cardiac arrest. This is not to suggest, of course, that the drugs themselves are actually improving prognosis: drug suppression may be merely a marker of a prognosis that is already good.

Era of the transvenous implantable defibrillator

Assessment of prognosis in patients with sustained ventricular arrhythmias has become critically important since the advent of cardioverter defibrillators that can be implanted transvenously without thoracotomy. Despite the conspicuous absence of a randomised controlled trial showing prolonged overall survival with the use of implantable cardioverter defibrillators in high risk patients, even the critics of the devices have conceded that they are effective in preventing sudden death from cardiac tachyarrhythmias. Transvenous cardioverter defibrillators can now be implanted with an operative mortality less than 0.5% and a 1 year mortality from sudden death of less than 1%, suggesting the likelihood of a higher overall survival than with previous epicardial patch systems. These results, in conjunction with the relative inefficacy of most antiarrhythmic drugs, the currently limited role of ablation techniques, and relatively high mortality associated with surgery for ventricular tachycardia, have caused a major shift to device therapy in many centres. It is becoming increasingly clear that patient groups other than those resuscitated from ventricular fibrillation are likely to benefit from these devices. Widespread implantation is now a practical option and cost restraints have come to the fore. Selection of patients for device therapy is increasingly based on cost-benefit issues, with those at highest risk from sudden death having the most to gain.

A large body of information is available on the
prognostic value of ventricular stimulation studies\textsuperscript{7,11-13} and together with the clinical history and assessment of ventricular function, such studies are an integral part of the assessment of risk in patients with sustained ventricular arrhythmias. In many centres the decision to implant a device is based on the inducibility of a ventricular arrhythmia at baseline or in the presence of a single antiarrhythmic drug.\textsuperscript{14} Usually the response to a class 1 agent (often intravenous procaainamide) is used for this purpose but there is evidence that other drugs, including amiodarone, may be as useful in such tests. The likelihood of tachycardia induction is higher after amiodarone (approximately 80\% of patients) than after administration of class 1 agents and the effect of amiodarone on the rate of the induced arrhythmia provides the principal means of prognostic assessment.\textsuperscript{12,15}

Although the ESVEM study showed that there was no difference in terms of arrhythmia recurrence between drug therapy selected by invasive and non-invasive techniques (they seemed equally ineffective) the study was not designed to compare the prognostic value of the two techniques in terms of mortality and did not have the power to do so. There is considerably less information on the prognostic power of Holter monitoring in the setting of recurrent sustained ventricular arrhythmias than on that of programmed electrical stimulation. Newer non-invasive means of risk assessment (heart rate variability, signal averaged electrocardiography) have shown promise in patients assessed after infarction,\textsuperscript{16} but have yet to be fully evaluated in patients with recurrent ventricular arrhythmias. Consequently the suggestion that Holter assessment of drug therapy is a reasonable initial strategy in patients with sustained ventricular arrhythmias cannot be supported.

In summary, it is clear that the issues concerning management of sustained ventricular arrhythmias have changed since the design of the ESVEM study in the middle 1980s. There has been a dramatic trend away from antiarrhythmic drugs and towards device therapy. The priority now is to be able to stratify patients with ventricular arrhythmias prognostically so as to determine those likely to benefit most from a limited resource. At present programmed electrical stimulation of the ventricle together with evaluation of the clinical history and assessment of ventricular function is the best way to do this.

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*Br Heart J* 1994 71: 307-308
doi: 10.1136/hrt.71.4.307

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