Epidemiological data from the Framingham heart study indicate that the cumulative incidence of atrial fibrillation (AF) over a 22 year follow up was 2.1% in men and 1.7% in women. The prevalence of AF increases with age, doubling with each successive decade, and 70% of people with AF are between 65–85 years old. AF is associated with a three- to fivefold increased risk of stroke, a threefold increased risk of congestive heart failure, and a significant 1.5- to 1.9-fold mortality risk even after adjusting for underlying cardiovascular conditions. Pacemaker follow up physicians often have to deal with AF as a co-morbidity. AF may also be associated with brady–tachy syndrome. A high incidence of AF will be present when we use pacemaker therapy after atrioventricular (AV) nodal ablation for medically refractory AF.

PHARMACOTHERAPY FOR AF: HOW SUCCESSFUL ARE WE?

Conventional pharmacological treatments includes rate control with AV nodal blockers, maintenance of sinus rhythm, and anticoagulation. While rate control and anticoagulation is a recognised treatment strategy, proarrhythmia using class I antiarrhythmic agents to maintain sinus rhythm remains a concern. A recent study has shown that low dose amiodarone, when compared to either sotalol or propafenone, is more efficacious in maintaining sinus rhythm. However, amiodarone had to be discontinued for cardiac and non-cardiac side effects in 18% of patients, while 35% of patients still developed AF at 16 months. While newer antiarrhythmic agents may enhance our success in these refractory cases, the current experience underscores the difficulties of long term pharmacological treatment to maintain sinus rhythm. Indeed, the preliminary results of the AFFIRM (atrial fibrillation following investigation of rhythm management) trial did not show the superiority of rhythm maintenance using drugs over rate control alone (late breaking news, American College of Cardiology annual meeting, 2002). Thus, the use of pacing, either alone or in a hybrid fashion with other treatments, has recently gained favour for treating AF.

MECHANISMS OF PACING FOR PREVENTING AF

AF develops as a result of the interaction between the triggers (atrial premature beat (APB)), the substrate (atrial effective refractory period (AERP), and conduction velocity), and mediation by the autonomic nervous system. Several groups have examined the changes in sinus rhythm and APB prematurity in patients developing AF (table 1). Most data in patients without sinus node disease (sick sinus syndrome (SSS)) suggest that the prevailing sinus rate before the onset of AF is normal or only slightly faster than normal. Thus a single rate support algorithm to prevent bradycardia is unlikely to be effective in suppressing AF in the majority of cases. There are three patterns of APB induced AF onset: APBs that initiate AF after a pause, and APBs that trigger AF with a closely coupled interval, or after a short-long-short cycle. The majority of AF episodes are triggered by closely coupled APBs that have a coupling interval shorter than those APBs that do not induce AF (table 1). Apart from initiating AF, APBs arising from the pulmonary veins may also act as a perpetuator of AF. As AF episodes frequently recur within minutes of termination, high rate overdrive pacing after AF termination may be useful to suppress AF reinitiation.

Atrial electrical remodelling occurs when AF is sustained, leading to a shortening of AERP and slowing of conduction velocity that promotes AF (AF begets AF). Atrial remodelling is inhomogeneous, with more shortening of AERP in the left atrium than the lower right atrium (RA). There is also prolonged internodal conduction time and suppressed sinus node function. Pacing, particularly delivered at multi-sites, may homogenise electrical conduction properties of the atrium and promote sinus rhythm. For example, distal coronary sinus (CS) pacing has been shown to suppress APBs from inducing AF by limiting their prematurity at the triangle of Koch, which is a region of local conduction delay and re-entry. Simultaneous RA and distal CS pacing reduced atrial conduction delay and increased electrogram width at this region and could prevent AF. By overdrive atrial pacing after AF, pacing may avoid AERP dispersion mediated by abrupt cycle length changes, thereby allowing time for reverse atrial remodelling to occur before AF is reinitiated.
Very little is written on the role of the autonomic nervous system on AF mediation. A vagally mediated type of AF has been described, and overdrive pacing suppresses AF by countering the bradycardia. A vagolytic effect of pacing the carotid sympathetic chain has been suggested to suppress certain types of AF or to control the ventricular rate in animals. The role of extracardiac stimulation to control AF in humans remains to be determined.

HOW CAN PACING BE DELIVERED?

Pacing can be delivered either in a passive or an active manner at a variety of sites (table 2). “Passive pacing” is conventional pacing to prevent or alter the response to AF. For example, different types of pacing modes have been compared to minimise the development of AF. Pacing is used in patients who have a clear bradycardia induced AF. If AF develops in a dual chamber pacemaker (DDD), rapid ventricular response up to the programmed maximum rate can occur as atrial activities are tracked. This is handled in modern DDD pacemakers by an algorithm known as automatic mode switching. When AF is detected or diagnosed, the pacemaker changes automatically to a non-atrial tracking mode (for example, DDI or VVI), so that a rapid ventricular rate does not occur. A clinically proven strategy to treat medically refractory AF is the use of AV nodal ablation followed by permanent DDD(R) or VVI(R) pacing. Both treatments are effectively a type of AF rate control and pacing does not act on the AF itself. Pacing is often used as an adjunct to drug treatment: antibradycardia pacing in the case of successful drug treatment causing sinus bradycardia, partial control with drugs and bradycardia that requires backup pacing so that larger doses of drugs can be used, and intermittent slow rate during AF from drugs used for rate control. An irregular ventricular rate in AF contributes to adverse symptoms and haemodynamics, and ventricular pacing delivered at a rate slightly faster than the average ventricular pacing rate in AF (known as ventricular regularisation pacing) can be used to achieve rate regularisation.

“Active pacing” involves either fixed or dynamic (based on the current sinus or a sensor mediated rate) overdrive of the normal sinus rhythm. Active pacing intervention using algorithms to counteract the mode of APB onset have been developed. These algorithms can be triggered by the onset of APBs, with treatment that aims at minimising the changes of atrial rate.

“Alternative atrial pacing sites” different from the conventional RA appendage or high lateral RA have been evaluated to modify the underlying substrate. Pacing has also been delivered from more than one site in the atrium. It is intuitive that some form of overdrive rate rather than the standard pacing rate will be necessary to maximise the “dose” of pacing to these sites, making them a form of active pacing therapy.

Pacing for AF prevention has been applied to the following patient populations: (1) pacing after AV nodal ablation; (2) vagally/bradycardia related AF; (3) patients with SSS; (4) patients with AF with or without sinus bradycardia; and (5) AF after cardiac surgery. Additionally, pacing methods have now been used to terminate AF precursors, and to control the ventricular rate irregularity once AF develops.

### PACING AFTER AV NODAL ABLATION

One of the most effective ways to treat the fast and irregular rate of AF is to use catheter ablation to interrupt the normal AV conduction system, and leave the patient’s rhythm to be controlled by a pacemaker. Several studies have documented the use of this “ablate and pace” strategy in improving symptoms, heart failure, and well being of patients over conventional drug treatment. In the North American registry that prospectively collected 156 patients followed up for one year, sustained improvement in quality of life was observed. Also, left ventricular ejection fraction was improved in those with a low ejection fraction (<45%).

A disadvantage of this strategy is pacemaker dependency, with the need for replacement and associated morbidity. In addition, there is a high incidence of progression to permanent AF, likely to be caused by withdrawal of antiarrhythmic agents. For example, in one study, AF developed in 24% of patients within six months after “ablate

<p>| Table 1 Characteristics of atrial fibrillation (AF) onset |
|----------------|-----------------|-----------------|</p>
<table>
<thead>
<tr>
<th>References</th>
<th>Number of episodes (patients)</th>
<th>Preceding sinus rate (%)</th>
<th>APB coupling interval (ms)</th>
</tr>
</thead>
<tbody>
<tr>
<td>Kilip (1965)</td>
<td>18 (14)</td>
<td>NA</td>
<td>0.48*</td>
</tr>
<tr>
<td>Bernett (1970)</td>
<td>32 (8)</td>
<td>NA</td>
<td>0.68*</td>
</tr>
<tr>
<td>Capucci (1992)</td>
<td>168 (20)</td>
<td>15</td>
<td>0.30</td>
</tr>
<tr>
<td>Murgatroyd (1993)</td>
<td>1126 (78)</td>
<td>8.5</td>
<td>0.40</td>
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<tr>
<td>Mehra (1996)</td>
<td>193 (80)</td>
<td>12</td>
<td>0.35</td>
</tr>
<tr>
<td>Tse (1999)</td>
<td>58 (53)</td>
<td>0</td>
<td>0.33</td>
</tr>
</tbody>
</table>

*Ratio of atrial premature beat (APB) coupling interval to proceeding sinus cycle length.

<table>
<thead>
<tr>
<th>Table 2 Types of pacing intervention for atrial fibrillation</th>
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<tr>
<td><strong>“Passive pacing”</strong></td>
</tr>
<tr>
<td>1. Pacemakers for AF prevention in sinus node disease</td>
</tr>
<tr>
<td>AAI(R) v VVI(R) DDD(R) v VVI(R) DDD(R) v AAI(R)</td>
</tr>
<tr>
<td>2. Ventricular rate control</td>
</tr>
<tr>
<td>Automatic mode switching</td>
</tr>
<tr>
<td>AV node ablation and pacing</td>
</tr>
<tr>
<td>Support pacing for concomitant drug treatment</td>
</tr>
<tr>
<td>Ventricular rate stabilisation pacing</td>
</tr>
<tr>
<td><strong>“Active pacing”</strong></td>
</tr>
<tr>
<td>Automatic atrial overdrive pacing</td>
</tr>
<tr>
<td>Post-ectopic atrial overdrive pacing</td>
</tr>
<tr>
<td>Post-AF atrial/overdrive pacing</td>
</tr>
<tr>
<td>AF termination</td>
</tr>
<tr>
<td><strong>“Alternative sites”</strong></td>
</tr>
<tr>
<td>Bachmann’s bundle region pacing</td>
</tr>
<tr>
<td>Low interatrial septal pacing (outside coronary sinus)</td>
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<tr>
<td>Biatrial endocardial pacing (right atrial and coronary sinus)</td>
</tr>
<tr>
<td>Biatrial epicardial pacing (right and left atrium)</td>
</tr>
<tr>
<td>Dual site atrial pacing (right atrial appendage and low interatrial septum)</td>
</tr>
</tbody>
</table>
and pace", but in none of the controls in the continued drug arm. A mortality rate of 15%, with 3% of patients dying suddenly, was reported. Recent evidence suggests that in the absence of previous myocardial infarction, congestive heart failure, and the use of cardiac medications after pacing, patients after ablate and pace had similar survival to age and sex matched populations when followed up for three years. These factors should be considered when prescribing this treatment for an individual patient.

A DDDR device is commonly prescribed after AV nodal ablation, together with automatic mode switching to avoid rapid ventricular tracking of AF. Arguably, because of poor long term sinus maintenance in this resistant group of patients, and in patients with persistent AF before ablation, a VVIR device may be an alternative.

VAGALLY/BRADYCARDIA MEDIATED AF
In some patients, AF episodes typically occur after meals or exercise, or during sleep and after prolonged sinus pauses. The use of class Ic agents such as propafenone is the recommended treatment. If Blockers should be avoided as they may aggravate the associated bradycardia. An early study by Coumel and colleagues showed in 4/6 patients that atrial pacing prevented these AF episodes during 5.5 years of follow up, although almost all patients took disopyramide. This group subsequently used DDD pacing in 10 patients with SSS in whom AF onset was related to bradycardia, and 7/10 patients also exhibited interatrial conduction delay that was reversed with higher rate atrial pacing. By overdrive pacing at a rate slightly above the mean diurnal rate, this group reported successful control of AF. These studies suggest that in a small group of patients in whom AF was clearly related to bradycardia, atrial based pacing could be effective in preventing AF episodes.

SICK SINUS SYNDROME
AF occurs in a significant proportion of patients with SSS after pacing. Retrospective studies suggested the use of atrial pacing in these patients was associated with a lower incidence of AF compared to ventricular pacing (6.8% v 2.6% annually). In three prospective randomised studies, AF was also reduced by atrial pacing. In the Danish study, 225 patients with SSS were randomised to either single chamber atrial (AAI) pacing or ventricular (VVI) pacing, with a follow up of eight years. The relative risk for AF (0.35 v 0.54), thromboembolic events, and heart failure were lower with AAI pacing. Similarly, in a Canadian study involving 1474 patients randomised to VVI(R) pacing and 1094 to atrial based pacing, the annual rate of AF was reduced from 6.6% to 5.5% with physiological pacing, with a relative risk reduction of AF of 18% by three years.

The effect on AF was only apparent after two years. In the elderly population, DDD systems also improved quality of life and reduced the progression to chronic AF. However, in all of these studies, conventional pacing is used and a control group is not possible as all of the involved patients required pacing therapy. It can be argued that atrial pacing does not actually suppress AF, rather it is ventricular pacing that is proarrhythmogenic.

Nevertheless, both retrospective and prospective data teach us that when prescribing pacemaker therapy for SSS, an atrial based pacing mode is preferred to ventricular pacing to minimise the incidence of AF. The impact of pacemaker prescription on AF incidence in patients with complete AV block is less certain, and is the subject of several ongoing studies (for example, UK PACE). As a high percentage of patients developed AF even with atrial pacing, some would advocate prophylactic use of additional strategies (hardware and software) to combat future AF episodes. These are described below.

AF WITH OR WITHOUT ASSOCIATED BRADYCARDIA
There are several situations in which a pacemaker is used in patients with AF. Paroxysmal AF (PAF) is present in about half of patients with SSS and a third of patients with AV block at the time of pacing implantation. Antiarrhythmic medications can depress sinus node function that requires pacing backup, and this is now increasingly an indication for pacing in many centres. In refractory cases, some would argue for implanting a pacemaker first, and delay or avoid AV nodal ablation if AF can be controlled with a device. Finally, a device to treat AF in patients without bradycardia has been tested in several clinical studies. The above categories of patients represent the largest body of data on which pacing therapy has been tested, either alone or more often in combination with antiarrhythmic medications. Data are also emerging for AF as a co-morbidity in patients receiving implantable cardioverter-defibrillators (ICDs) or cardiac resynchronisation devices for heart failure.

Conventional pacing
In patients with medically refractory PAF pending AV nodal ablation, the PA’ (atrial pacing periablation for paroxysmal AF) study randomised patients to either no pacing (DDI at 30 beats/min (bpm)) or to DDDR pacing at a lower rate of 70 bpm, with continuation of antiarrhythmic drugs. Unexpectedly, pacing did not prolong the time to the first AF recurrence (1.9 days v 4.2 days with no pacing, p = NS). In fact, pacing was associated with a trend for higher AF burden. Potential limitations in this study are the use of a pacing mode (DDDR) that did not guarantee AV synchrony, the lack of an overdrive algorithm to ensure a high percentage of atrial pacing (the atrium was paced in only 67% in this study), the use of atrial pacing at the conventional single site at the RA appendage, and the relatively short follow up (10 weeks). The PA’ study suggests that in patients with medically refractory AF who do not have bradycardia, conventional atrial pacing at 70 bpm in the short term is not effective in preventing AF.

Atrial overdrive
In patients with conventional pacing with a DDDR pacemaker, it is simple to just increase the backup rate to suppress AF. Ward and colleagues’ randomised 18 patients with PAF and SSS to a backup rate of 60, 75, and 90 bpm, each for a two month period to test this hypothesis. While the percentage of atrial pacing increased by 44%, 57.5%, and 73.5%, respectively,

Abbreviations

<table>
<thead>
<tr>
<th>Abbreviation</th>
<th>Description</th>
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<tbody>
<tr>
<td>AERP</td>
<td>atrial effective refractory period</td>
</tr>
<tr>
<td>AF</td>
<td>atrial fibrillation</td>
</tr>
<tr>
<td>APB</td>
<td>atrial premature beat</td>
</tr>
<tr>
<td>ATP</td>
<td>antitachycardia pacing</td>
</tr>
<tr>
<td>AV</td>
<td>atroventricular</td>
</tr>
<tr>
<td>CS</td>
<td>coronary sinus</td>
</tr>
<tr>
<td>ERAF</td>
<td>early reinitiation of atrial fibrillation</td>
</tr>
<tr>
<td>ICD</td>
<td>implantable cardioverter-defibrillator</td>
</tr>
<tr>
<td>PAF</td>
<td>paroxysmal atrial fibrillation</td>
</tr>
<tr>
<td>RA</td>
<td>right atrium</td>
</tr>
<tr>
<td>SSS</td>
<td>sick sinus syndrome</td>
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www.heartjnl.com
Nevertheless, acute testing suggests that pacing at the anterior Bachmann’s bundle region or interatrial septal pacing有益，这些益处与心房率的变化无关。患者中，心房率在60%至45%抑制的基础上显著下降（37%），并改善了心房率的幅度。该算法被很好地耐受。因此，如果存在AF，与起搏器的患者，可以自动抑制AF心房率。所有患者具有SSS和阵发性AF的历史，2年随访后，AF发生率的近似值为75%。Bachmann’s bundle区域被定位在主动固定电极的位置，可以在左前心房（使用荧光左前心房的观察视图）和心房尖部之间。Bachmann’s bundle区域的电极位置对抑制APBs和ERAf具有显著延迟作用（75%）。AF发生率在RA尖端中为0，然后在永久AF发展的两个月后心房内。Bachmann’s bundle区域的电极位置对抑制APBs和ERAf具有显著延迟作用（75%）。

**Algorithms specific to APB/AF triggers**

经验与那些能够抑制心房率的算法有关。当APBs和心房率在起搏率中出现时，这些算法在抑制APBs的频率方面非常有效，但不是所有的情况。自发性或除颤/终止性心房率衰减的AF终止，心房率可能会在34%的患者中出现。近端和远端心房电极是早期重新开始的AF（ERAf），它限制了长期的心房率的持续时间。Tse和同事们测试了，在随机化的方式下，使用心房起搏和ERAf的百分比在12例患者中的百分比显著升高。Pacing in RA尖端预防了AF。Pacing in CS9–10）。在左房尖端（最早的心房记录区）和CS9–10）中。B）心房起搏在500 ms以上预防了AF。从起搏开始。APB的持续时间为210 ms到240 ms，没有引起AF。CS，冠状窦；Hls，His bundle；RA，右房；RV，右心室。来源：Tse et al.13的许可。

**Alternative and multiple site atrial pacing**

这些包括Bachmann’s bundle区域/心房内隔膜、左心房起搏（RA appendage and distal CS），和心房尖端起搏（RA appendage and low atrial septum）。Bachmann’s bundle区域或心房内隔膜起搏

Bachmann’s bundle区域的电极位置是争议的。Bachmann’s bundle区域的电极位置是争议的。
might have influenced the outcome of this study. The proximity of the site to the aortic arch is a potential concern, although no complication related to the aorta was observed in this study. AF burden was not measured. Several studies are now underway to test the incremental benefit of automatic atrial overdrive in suppressing AF burden in the high septal region.

Padeletti and colleagues reported the result of pacing at the low interatrial septum. This site was chosen as it is near the triangle of Koch (an area of slow conduction), and was approached by using a screw-in lead above the CS os. They studied 46 patients with paroxysmal AF, randomised to either RA appendage or low septal pacing. Either pacing mode reduced AF compared to pre-implantation frequency, but low interatrial septal pacing was superior to RA appendage pacing in reducing AF burden over a three month period. Again, P wave duration was significantly reduced compared to that in sinus rhythm.

Taken together, these studies suggest that pacing at the interatrial septum (high or low) shortens the P wave duration, and reduces the incidence of AF compared to the RA appendage site. At least in the short and medium term, the right interatrial septal site appears to be as stable and safe as the conventional appendage position. These sites may be an alternative pacing site for AF control in patients with SSS and AF if the issues of complexity in implantation and long term lead stability can be solved.

Biatrial pacing
Daubert and his colleagues pioneered biatrial pacing by using a CS bipolar lead to achieve left atrial pacing simultaneously with conventional RA pacing. They tested the efficacy of biatrial pacing in patients with either prolonged P wave duration (> 120 ms) or interatrial conduction time (> 100 ms). In a group of 86 patients with atrial tachyarrhythmias, they were able to reduce P wave duration (from mean (SD) 187 (29) ms to 160 (14) ms), and maintained sinus rhythm in 64% (with 33% free from any episode of AF). However, in a multicentre European trial (the SYNBIPACE (synchronous biatrial pacing for reduction of paroxysmal/permanent AF) study), such a benefit in AF suppression was not reproduced. Thus this technique may be applicable to select patients with long interatrial conduction delay, and can also possibly contribute to better left heart AV interval programming and haemodynamic benefits. However, double sensing of A and V electrograms in the CS can be a problem, and special blanking is required. There is concern (as with biventricular pacing for heart failure) over the stability of the CS lead, and the ease with which lead extraction can be effected.

Dual site atrial pacing
Delfaut and colleagues pioneered the use of RA appendage to RA low septal pacing (just outside the CS os) in suppressing AF (fig 2B). Thirty patients with drug refractory symptomatic AF and documented primary or drug induced bradyarrhythmia underwent a crossover study to assess: (1) if pacing was useful to prevent AF compared to pre-implant history; (2) if single site (RA appendage or CS os pacing) were different; and (3) if dual site pacing had additional benefit to single site pacing. A fixed rate overdrive was used and event recorder documented first AF recurrence was used as the primary end point. The mean arrhythmia-free interval was increased from mean (SD) of 9 (10) days before implant, to 143 (110) days during single site periods, and to 195 (96) days during dual site crossover period. The authors did not find any difference between single site pacing at the RA appendage or CS os pacing in suppressing AF. Significantly, this study also documented long term safety of dual site pacing up to three years, with no case of CS os lead dislodgement after patient discharge from hospital, compared to a rate of dislodgement of up to 8% in dual site pacing. Although uncontrolled, the long term efficacy of maintaining sinus rhythm was 78% at one year and 56% at three years, which was remarkable in a very refractory group of patients. The limitations of this study were the lack of an unpaced controlled group, frequent crossover with potential carryover effect, and the need for antiarrhythmic medications to maintain sinus rhythm.

We have specifically addressed the use of dual site atrial pacing in patients with paroxysmal AF without conventional
indication for pacing, using pacemakers with the Continuous Atrial Overdrive algorithm.29 Twenty two patients who had AF recurrence despite sotalol treatment underwent randomised crossover periods of 12 weeks with either pacing on (plus sotalol) or continuation of sotalol only. The end points were event recorder documented AF recurrence and pacemaker memory of AF burden. Dual site atrial pacing increased the percentage of atrial pacing (13 (18)% to 80 (30)%), reduced the number of APBs (from 8265/day to 2740/day), prolonged the time to the first documented AF (symptomatic or asymptomatic), and reduced AF burden (45 (34)% to 22 (29)%). PACing reduced the risk of AF recurrence by 3.2 times. There was significant change in some measures of quality of life, but no overall change in AF symptoms. The DAPPAD (dual site atrial pacing for the prevention of AF) study prospectively randomised and crossed over patients between dual site and RA appendage pacing and support pacing in patients with PAF randomised and crossed over patients between dual site and atrial pacing for the prevention of AF) study prospectively randomised and crossed over patients between dual site and RA appendage pacing and support pacing in patients with PAF and pacing indications. The preliminary results suggested that dual site pacing with overdrive in combination with either class I or class III antiarrhythmic agents was better tolerated and more effective in AF prevention than overdrive RA pacing or support pacing. Several other studies have also reported on the efficacy of dual site pacing.

Taken together, these trials indicate that pacing has an effect on the burden of AF in patients with or without the need of a pacemaker. Automatic atrial overdrive pacing is necessary for pacing to be effective, and multisite pacing may have an incremental benefit. However, the use of multiple leads (with their complexity of implantation and programming) should probably be reserved for those with AF of moderate severity. Use of a β blocker may provide an incremental benefit. Uncertainty remains as to the clinical predictors for a favourable response to dual site atrial pacing. Neither the P wave duration nor interatrial delay before pacing were predictive of an effective outcome.

AF AFTER CARDIAC SURGERY

Approximately 20% of patients developed AF after bypass surgery. The causes are multifactorial, and may include the effects of cardiac bypass, changes in sympathetic tone, and postoperative infection. The development of AF is associated with an increase in stroke, heart failure, and length of hospital stay, which substantially increase management cost. β Blockers and amiodarone have been shown to be effective prophylactic agents for AF, but AF still developed in up to 30% of these patients despite amiodarone pre-treatment. After cardiac surgery, epicardial pacing placement is often routine for support pacing, and it is of interest to assess if pacing may have effects on postoperative AF. Several studies have recently examined the effect of pacing (in addition to β blocker treatment) in suppressing postoperative AF. Fan and colleagues25 randomised 132 patients to biatrial, RA, and left atrial pacing or control. Manual overdrive atrial pacing was performed for five days, at 10 bpm above the intrinsic rate up to 90 bpm. Only biatrial pacing reduced the development of sustained AF (42% in control v 12.5% during pacing), whereas single site pacing had no effect. The beneficial effect was attributable to a larger reduction of P wave dispersion during biatrial pacing compared with single site pacing or control, and only patients with AF prevention had a reduction in P wave duration. Postoperative intensive care stay and the associated cost were reduced. Several studies have since reported the efficacy of pacing, particularly biatrial pacing, in suppressing postoperative AF, with the left atrial lead in the posterior atrium. The convenience and ease of instrumentation of biatrial pacing is a strong incentive to apply this technique to most patients after postoperative cardiac surgery.

ANTITACHYCARDIA PACING

While the short excitable gap during sustained AF does not lend itself to pacing termination, ATP has several potential mechanisms to reduce AF burden. Many episodes of AF degenerate from atrial tachycardia or flutter, and early termination of these precursor rhythms may prevent AF from becoming established. Conversely, after antiarrhythmic agents (especially class Ic drugs), AF may be converted to flutter or a slower atrial tachycardia that can be terminated. It is logical to consider ATP in an implanted device to terminate these AF related rhythms.

Atrial ATP (burst, ramp, and 50 Hz stimulation) are now available in some ICDs and pacemakers. Several groups have reported on the efficacy of ATP, ranging from 33–86%, depending on the organisation of AF (fig 3A). Interestingly, “organised” rhythms were encountered in nearly half of all recorded episodes in patients with a clinical diagnosis of paroxysmal AF, suggesting that ATP may have a role in these patients. Twenty five per cent of patients with ICDs have associated AF. Friedman and colleagues20 randomised 52/269 patients with a combined atrial and ventricular ICD to either ATP, atrial defibrillation, and preventive pacing versus only ventricular ICD function, each for a three month randomised period. Atrial therapies significantly reduced AF burden from

![Image](http://heart.bmj.com/Heart/firstpublishedas10.1136/heart.89.1.106on1January2003.Downloadedfromhttp://heart.bmj.com/onJune10,2022byguest.Protectedbycopyright.)
58.5 to 7.8 h/month. The efficacy of ATP for terminating slower and faster atrial tachyarrhythmias were 62% and 49%, respectively. While encouraging, these studies are limited by frequent patient exclusion and insufficient randomisation, and it is uncertain if burden reduction was related to pacing prevention or to ATP itself. Thus, in the absence of a well controlled randomised trial, it is uncertain whether ATP works or may be proarrhythmogenic.

VENTRICULAR RATE STABILISATION
Apart from a rapid rate, irregularity in AF contributes to abnormal cardiac haemodynamics. By pacing the RV at a rate slightly faster than the mean ventricular rate of AF, it is possible to suppress shorter cycles and regularise the rate. This has been attributed to retrograde concealed activation in the AV node. Acute testing suggested that rate regularisation pacing can regularise AF at rest and to some extent during exercise. Ventricular rate stabilisation algorithms have been developed by several manufacturers. Clinical benefit in ambulatory patients remains to be confirmed, and the long term effect on left ventricular function because of pacing needs to be considered.

FUTURE PERSPECTIVES
Pacing either in the treatment or prevention of AF should not be an isolated therapy. AF is a heterogeneous disease, and a hybrid approach is the standard. For example, concomitant antithrombotic medications are commonly used with pacing. Radiofrequency ablation can eliminate pulmonary vein ectopic foci for AF or modify the atrial substrate, and atrial defibrillation can further enhance the maintenance of sinus rhythm. The various types of hybrid therapy are under evaluation.

An important development in device therapy for AF is the ability to measure the total amount of AF (AF burden) that can be confirmed with stored atrial electrograms (fig 3B). This is a more accurate assessment of AF than the time to the first recurrence of AF itself. In addition, device based AF recording gives the clinician the possibility of objectively measuring the severity of AF, and may become a useful guide to assess interventional procedures, the need for anticoagulation, and to understand the symptomatology of AF itself.

CONCLUSION
Ablation and pacing for medically refractory AF is clinically proven, and is an effective symptomatic therapy. In patients with SSS, an atrial based pacemaker should be prescribed to reduce future episodes of AF. An automatic atrial overdrive algorithm appears to be effective in reducing symptomatic AF. Dual site right atrial pacing, in the presence of overdrive and β blocker, confers additional benefit to single site pacing. Epicardial biastral pacing is a useful technique to reduce the incidence of AF complicating cardiac surgery. While automatic mode switching and ventricular rate stabilisation will become programmable features of modern pacemakers, the role of ATP in patients with AF remains to be confirmed. It is likely that pacing efficacy will be enhanced when combined with strategies such as ablation, pharmacotherapy, and defibrillation.

REFERENCES
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11 An important study which shows the lack of efficacy of conventional pacing alone in preventing drug refractory AF.
15 This is the only randomised study to show that pacing can suppress early re-initiation of AF from atrial ectopic discharges.
17 A prospective study which demonstrates the efficacy of pacing the Bachmann’s bundle region in preventing persistent AF.
19 This randomised study investigated the use of low right atrial septal pacing in preventing AF.
21 One of several studies by this group which pioneered the use of biastral pacing in preventing atrial tachyarrhythmias in patients with interstitial conduction delay.
23 The most important of several studies by this group which compared the relative efficacy of drug treatment, single site pacing, and dual site pacing on the development of AF.
25 A multicentre, randomised, controlled study that shows the efficacy of dual site pacing in preventing both symptomatic and asymptomatic AF over sotalol in patients without bradycardia.
27 One of several studies recently published which demonstrates the role of multisite pacing in reducing the incidence of postoperative AF.
Web-Only References


