Ablation of ischemic ventricular tachycardia: evidence, techniques, results, and future directions

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Purpose of review
This article summarizes current understanding of the arrhythmia substrate and effect of catheter ablation for infarct-related ventricular tachycardia, focusing on recent findings.

Recent findings
Clinical studies support the use of catheter ablation earlier in the course of ischemic disease with moderate success in reducing arrhythmia recurrence and shocks from implantable defibrillators, although mortality remains unchanged. Ablation can be lifesaving for patients presenting with electrical storm. Advanced mapping systems with image integration facilitate identification of potential substrate, and several different approaches to manage hemodynamically unstable ventricular tachycardia have emerged. Novel ablation techniques that allow deeper lesion formation are in development.

Summary
Catheter ablation is an important therapeutic option for preventing or reducing episodes of ventricular tachycardia in patients with ischemic cardiomyopathy. Present technologies allow successful ablation in the majority of patients, even when the arrhythmia is hemodynamically unstable. Failure of the procedure is often because of anatomic challenges that will hopefully be addressed with technological progress.

Keywords
catheter ablation, ischemic heart disease, ventricular tachycardia

INTRODUCTION
Infarct-related monomorphic ventricular tachycardia typically arises from regions of poorly coupled surviving myofibers within scar tissue characterized by slow electrical conduction. In ischemic cardiomyopathy, such arrhythmia substrates are most commonly located subendocardially but can also occur intramurally and subepicardially. Areas of slow conduction can be large or widely separated within scar tissue, giving rise to multiple and potentially interconnected circuits that can result in multiple ventricular tachycardia morphologies. Progressive fibrosis and ventricular remodeling can lead to onset of ventricular arrhythmias late after myocardial infarction. A small proportion (approximately 10%) of monomorphic ventricular tachycardia in ischemic cardiomyopathy is because of automaticity or reentry involving the Purkinje system.

The occurrence of sustained monomorphic ventricular tachycardia is a marker for increased mortality in patients with structural heart disease. Implantable cardioverter defibrillators (ICDs) are effective for prevention of sudden death, but recurrent shocks reduce quality of life, cause post-traumatic stress syndrome, and are associated with worsened cardiac outcomes [1,2,3]. Interrupting areas of slow conduction by catheter ablation aims to decrease the risk for ventricular tachycardia recurrence and thereby seeks to improve quality of life and potentially reduce the risk of death. Ablation for ventricular tachycardia storm (defined as three or more episodes in 24 h) or incessant ventricular tachycardia can be lifesaving [4].

Effective ablation requires identification of the ventricular tachycardia origin during mapping. Ablation is typically performed during ventricular tachycardia if hemodynamic stability can be maintained. If induced ventricular tachycardias are unstable or the clinical ventricular tachycardia...
cannot be induced, ablation aims to modify the arrhythmia substrate identified during stable sinus or paced rhythms (substrate-guided ablation). Recent studies have advanced both strategies. The review will address the recent advances in techniques, clinical experience, and outcomes for ablation for monomorphic ventricular tachycardia in ischemic cardiomyopathy. Polymorphic ventricular tachycardia and ventricular fibrillation that occur more often in the acute or subacute period of myocardial ischemia or infarction are not discussed.

### Current Evidence, Recent Developments, and Ongoing Trials

#### Indications and timing for postinfarct ventricular tachycardia ablation

With increasing experience and expertise, catheter ablations are being increasingly employed earlier in the course of ventricular tachycardia without waiting to exhaust all pharmacological options. Table 1 summarizes current recommendations for the use of catheter ablation according to the 2009 European Heart Rhythm Association/Heart Rhythm Society (EHRA/HRS) Expert Consensus on Catheter Ablation of Ventricular Arrhythmias [5].

Two randomized trials [ventricular tachycardia ablation in coronary heart disease (VTACH) and Substrate Mapping and Ablation in Sinus Rhythm to Halt Ventricular Tachycardia (SMASH-VT)] explored the use of catheter ablation in ICD recipients with prior myocardial infarction, ventricular tachycardia, and impaired left ventricular function [6,7]. Both studies found a significant reduction in ventricular tachycardia recurrence during follow-up of approximately 2 years. However, neither of these trials was sufficiently powered to examine mortality. In a retrospective study, Bunch et al. [8] reported a lower risk of death and heart failure hospitalizations in patients treated with ventricular tachycardia ablation after an ICD shock compared with patients managed medically only. Recent publications support an early invasive approach.

#### Table 1. Indications for catheter ablation of ventricular tachycardia in patients with structural heart disease (adapted from [5])

<table>
<thead>
<tr>
<th>Catheter ablation of VT is recommended for</th>
</tr>
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<tbody>
<tr>
<td>1  Patients with SMVT, including VT terminated by an ICD, that recurs despite antiarrhythmic drug therapy or when antiarrhythmic drugs are not tolerated or not desired</td>
</tr>
<tr>
<td>2  Control of incessant SMVT or VT storm that is not because of a transient reversible cause</td>
</tr>
<tr>
<td>3  Patients with frequent PVCs, NSVTs, or VT that is presumed to cause ventricular dysfunction</td>
</tr>
<tr>
<td>4  Bundle branch reentrant or interfascicular VTs</td>
</tr>
<tr>
<td>5  Recurrent sustained polymorphic VT and VF that is refractory to antiarrhythmic therapy when there is a suspected trigger that can be targeted for ablation</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>Catheter ablation should be considered for</th>
</tr>
</thead>
<tbody>
<tr>
<td>1  Patients who have one or more episodes of SMVT despite therapy with one or more class I or III antiarrhythmic drugs</td>
</tr>
<tr>
<td>2  Patients with recurrent SMVT because of prior MI who have LV ejection fraction &gt;0.30 and expectation for 1 year of survival, and it is an acceptable alternative to amiodarone therapy</td>
</tr>
<tr>
<td>3  Patients with hemodynamically tolerated SMVT due to prior MI who have reasonably preserved LV ejection fraction (&gt;0.35) even if they have not failed antiarrhythmic drug therapy</td>
</tr>
</tbody>
</table>

<table>
<thead>
<tr>
<th>VT catheter ablation is contraindicated</th>
</tr>
</thead>
<tbody>
<tr>
<td>1  In the presence of a mobile ventricular thrombus (epicardial ablation may be considered)</td>
</tr>
<tr>
<td>2  For asymptomatic PVCs and/or NSVT that are not suspected of causing or contributing to ventricular dysfunction</td>
</tr>
<tr>
<td>3  When VT is because of transient, reversible causes, such as acute ischemia, hyperkalemia, or drug-induced torsade de pointes</td>
</tr>
</tbody>
</table>

ICD, implantable cardioverter defibrillator; LV, left ventricle; MI, myocardial infarction; NSVT, nonsustained ventricular tachycardia; PVCs, premature ventricular contractions; SMVT, symptomatic sustained monomorphic ventricular tachycardia; VF, ventricular fibrillation; VT, ventricular tachycardia.
observational study, Dinov et al. [9**] reported better acute and long-term success if catheter ablation of scar-related ventricular tachycardia was performed within 30 days after the first documented ventricular tachycardia. Once again, a mortality benefit was not evident in patients who underwent early ventricular tachycardia ablation compared with those who had their ablation late (>1 year) after their first presentation with ventricular tachycardia.

The benefits of early catheter ablation and its superiority to medical therapy are the subject of ongoing randomized trials. The PARTITA trial (NCT01547208) is randomizing patients who experience an appropriate ICD shock to immediate catheter ablation or waiting until the occurrence of a ventricular tachycardia storm. The Substrate Targeted Ablation Using the FlexAbility™ Ablation Catheter System for the Reduction of Ventricular Tachycardia (STAR-VT) trial (NCT02130765) aims to show a benefit of ventricular tachycardia ablation as first line therapy compared with medical therapy in patients with documented monomorphic ventricular tachycardia (either spontaneous or induced by programmed stimulation) who have an implanted ICD. The Ventricular Tachycardia (VT) Ablation Versus Enhanced Drug Therapy (VANISH) trial (NCT00905853) aims to compare aggressive antiarrhythmic therapy to catheter ablation for ventricular tachycardia recurrence after receiving ICD therapy despite antiarrhythmic drugs.

Mapping strategies and image integration
Electroanatomic mapping systems are routinely used in most centers for voltage and activation mapping and for navigation of catheters within the cardiac chambers. Automated multipoint acquisition from multipolar catheters permit fast, high-resolution mapping to identify abnormal tissue that typically demonstrates low voltages, fractionation, and late potentials [10]. The judicious use of unipolar and bipolar voltage maps enables the identification of border zones. When combined with analysis of electrograms and pacing maneuvers, conducting channels that support reentry circuits within scars can be identified during stable sinus or paced rhythm [11,12].

Mapping systems also allow integration of real-time intracardiac ultrasound images, fluoroscopy, and previously acquired MRI or computer tomography. Contrast enhanced MRI may be used to identify ventricular tachycardia substrate and areas that are likely to contain critical isthmus sites [13,14]. Prospective studies are required to assess whether such imaging guidance will improve the efficiency and efficacy of ventricular tachycardia ablation. Image integration is potentially useful to minimize risks of phrenic nerve and coronary artery injury during epicardial ventricular tachycardia ablation [15]. MRI-guided interventions with intra-procedural imaging may facilitate substrate identification, navigation, lesion formation, and early detection of complications [16]. However, the incorporation of an MRI scanner in the electrophysiology laboratory is expensive and raises multiple logistical issues.

Contact force-sensing catheters
Radiofrequency ablation lesion creation is critically dependent on adequate contact force between the catheter and tissue [17]. Newer ablation catheters incorporate contact force sensors [18]. In an ovine model, contact force monitoring improved lesion formation in the ventricles when ablating both from the endocardium and the epicardium [19]. Contact force monitoring during mapping increases specificity for identifying low-voltage area and abnormal electrograms [20]. The clinical benefits of contact force mapping and ablation for ventricular tachycardia are yet to be fully elucidated.

Substrate-based ablation
The critical isthmus of a specific monomorphic ventricular tachycardia can be identified by activation or entrainment mapping during ventricular tachycardia, if the ventricular tachycardia is hemodynamically stable or can be induced and terminated reproducibly to allow mapping during short episodes [21–23]. In clinical practice, the majority of induced ventricular tachycardias are hemodynamically unstable and multiple morphologies of ventricular tachycardia, often reflecting multiple reentry circuits, are induced with repeated attempts to initiate ventricular tachycardia. Therefore, ablation is often limited to targeting the arrhythmia substrate during sinus or paced rhythm. Areas of slow conduction and late potentials within areas of low voltage have been correlated with critical isthmus sites for reentry [24–26].

Based on these observations, several strategies for modifying the ventricular tachycardia substrate have been described recently (Table 2). These have included the targeting of sites with late potentials or fractionated electrograms that can be shown to be poorly coupled to the surrounding myocardium [27,28], circumferential ablation of the scar area [29], circumferential core isolation around critical ventricular tachycardia circuit elements [30], or ‘homogenization’ of the scar with extensive overlapping ablation lesions [31]. de Chillou et al. [32]
located the critical isthmus by pace-mapping in sinus rhythm in a selected population with hemodynamically stable ventricular tachycardia, and Berruezo et al. [33] recently suggested that 'dechannealing' of the scar specifically targets channels relevant to the ventricular tachycardia. These investigators targeted the earliest of all recorded 'late' potentials in the scar border zone during sinus rhythm under the assumption that these areas represent entrance sites of conduction channels. Ablation at the entrance sites eliminated the conducting channels, thereby limiting the total ablations necessary for abolishing conducting channels in the scar. Tung et al. demonstrated that local ablation can modify electrical activity in scar regions remote from the ablation site [34]. However, the optimal strategy resulting in effective ablation but at the same time minimizing unnecessary radiofrequency application and procedure time is still not known. Anatomically ablating unexcitable scar might not be necessary, and abnormal electrograms may arise from tissue remote from the ablation catheter [35].

At our center, substrate modification of the scar is performed, targeting regions of slow conduction evidenced by fractionated potentials. We perform pace-mapping in sinus rhythm to identify sites where pace-mapping approaches the morphology of an induced clinical ventricular tachycardia and has a stimulus to QRS delay >40ms (Fig. 1). Ablation is usually performed for 10-20s with a contact-sensing catheter aiming for 10-15g of contact force and impedance drop of 10-15V during ablation. As an acute endpoint, we aim to achieve noninducibility of all sustained monomorphic ventricular tachycardia by rendering the whole scar or scar areas felt to be involved in ventricular tachycardia circuits unexcitable to pacing (10 mA at 2ms pulse width) or to disconnect them from healthy myocardium.

### Table 2. Recently published data on substrate-based ablation approaches for ablation of ischemic ventricular tachycardia

<table>
<thead>
<tr>
<th>Technique</th>
<th>Patient selection</th>
<th>Patients</th>
<th>Men</th>
<th>Age (years)</th>
<th>LVEF (%)</th>
<th>Ischemic heart disease</th>
<th>No. of VTs induced at baseline</th>
<th>Epicardial ablation</th>
<th>RF time (min)</th>
<th>Procedure time (min)</th>
<th>Adverse events</th>
<th>Noninducible</th>
<th>Mean or median follow-up (months)</th>
<th>VT recurrence</th>
<th>Death</th>
</tr>
</thead>
<tbody>
<tr>
<td>Jais et al. [27]</td>
<td>Elimination of LAVA</td>
<td>Unselected</td>
<td>70</td>
<td>90%</td>
<td>67 ± 11</td>
<td>80%</td>
<td>35 ± 10</td>
<td>2 (1–3)</td>
<td>30%</td>
<td>23 ± 11</td>
<td>148 ± 73</td>
<td>9%</td>
<td>70%</td>
<td>22 (14–27)</td>
<td>46%</td>
</tr>
<tr>
<td>Vergara et al. [28]</td>
<td>Late potential abolition</td>
<td>Selected</td>
<td>50</td>
<td>9.4%</td>
<td>66 ± 7</td>
<td>72%</td>
<td>32 ± 9</td>
<td>2.8 ± 1.0</td>
<td>42%</td>
<td>N/A</td>
<td>N/A</td>
<td>N/A</td>
<td>80%</td>
<td>13 ± 4</td>
<td>20%</td>
</tr>
<tr>
<td>Iliz et al. [29]</td>
<td>Scar isolation</td>
<td>Selected</td>
<td>12</td>
<td>100%</td>
<td>54 ± 8</td>
<td>100%</td>
<td>32 ± 13</td>
<td>3 ± 2</td>
<td>0%</td>
<td>35 ± 13</td>
<td>195 ± 64</td>
<td>N/A</td>
<td>92%</td>
<td>16 (10–26)</td>
<td>30%</td>
</tr>
<tr>
<td>Lou et al. [30]</td>
<td>Core isolation</td>
<td>Selected</td>
<td>44</td>
<td>9.5%</td>
<td>63 ± 14</td>
<td>73%</td>
<td>31 ± 13</td>
<td>3 ± 2</td>
<td>11%</td>
<td>N/A</td>
<td>326 ± 121</td>
<td>2%</td>
<td>82%</td>
<td>17 ± 9</td>
<td>14%</td>
</tr>
<tr>
<td>Di Biase et al. [31]</td>
<td>Scar homogenization</td>
<td>Unselected</td>
<td>43</td>
<td>75%</td>
<td>62 ± 8</td>
<td>100%</td>
<td>24 ± 8</td>
<td>2 (2–5)</td>
<td>100%</td>
<td>74 ± 21</td>
<td>288 ± 90</td>
<td>2%</td>
<td>100%</td>
<td>21 (19–25)</td>
<td>19%</td>
</tr>
<tr>
<td>de Chillou et al. [32]</td>
<td>Isthmus localization by pace mapping</td>
<td>Selected</td>
<td>10</td>
<td>80%</td>
<td>71 ± 11</td>
<td>100%</td>
<td>37 ± 14</td>
<td>N/A</td>
<td>0%</td>
<td>10 ± 9</td>
<td>203 ± 62</td>
<td>0%</td>
<td>80%</td>
<td>65 ± 6</td>
<td>33%</td>
</tr>
<tr>
<td>Bemèveau et al. [33]</td>
<td>Scar dechanneling</td>
<td>Unselected</td>
<td>101</td>
<td>91%</td>
<td>65 ± 12</td>
<td>74%</td>
<td>36 ± 13</td>
<td>N/A</td>
<td>27%</td>
<td>28 ± 16</td>
<td>227 ± 69</td>
<td>7%</td>
<td>78%</td>
<td>21 (11–29)</td>
<td>26%</td>
</tr>
</tbody>
</table>

LAVA, local abnormal ventricular activity; LVEF, left ventricular ejection fraction; N/A, not available; RF, radiofrequency; VT, ventricular tachycardia.
for relatively longer periods of time and were terminated by radiofrequency ablation more often compared with a control group without hemodynamic support or with an intraaortic balloon pump. However, there was no difference in acute procedural success or ventricular tachycardia recurrence rates during follow-up. In addition, the magnetic motor of the Impella system can interfere with the electroanatomic mapping, especially during mapping in the ventricular outflow tracts. Extracorporeal membrane oxygenation offers biventricular support and does not interfere with the mapping systems. Extracorporeal membrane oxygenation does not unload the left ventricle as compared with pLVAD but offers better hemodynamic support (4–6 l/min).

**Epicardial ablation**

When endocardial ablation fails, it is often because of the location of critical components of a ventricular tachycardia circuit deep in the myocardium or in the subepicardium. Epicardial ablation has become an essential component of ablation strategies, especially for ventricular tachycardia because of nonischemic cardiomyopathies, although proximity of major coronary vessels or the phrenic nerve.
to critical sites may preclude adequate ablation [39]. In ischemic heart disease, epicardial ablation is usually performed after failed endocardial ablation. An initial endo–epicardial approach might be beneficial in selected patients. A recent small single-center study showed an association between an initial combined approach and less hospitalization and reablation during follow-up, but failed to show any differences in ventricular tachycardia recurrence or mortality [40].

Techniques for ablation of intramural substrate

Although the advent of cooled tip radiofrequency catheters and contact force monitoring has improved lesion formation, safe delivery of deep intramyocardial lesions remains a problem. Bipolar radiofrequency energy between two separate ablation catheters positioned on the septum from both ventricles has proven to be beneficial in isolated cases and in preliminary studies [41–43].

Transcoronary ethanol ablation is an approach that is used in the setting of malignant refractory ventricular tachycardia when standard approaches fail [44]. Important considerations including coronary anatomy, risk of heart block, and risk of hemodynamic deterioration from the loss of functioning myocardium.

Needle electrodes for the creation of deeper intramural lesions show promise in early human studies where conventional catheter ablation techniques have been unsuccessful [45,46]. The use of a warm saline-enhanced needle electrode where radiofrequency energy is applied through the saline stream into deep myocardial layers shows promise in ablation of viable tissue in large myocardial infarct scars [47].

Procedural endpoints and outcome

Reported ventricular tachycardia recurrence rates after postinfarct ventricular tachycardia ablation procedures vary widely, but typically range between 30 and 50% in the larger studies. The most common origin of recurrent ventricular tachycardia is from areas adjacent to prior ablation lesions suggesting that the ventricular tachycardia exit points may have been altered by ablation [48].

The only endpoint criterion endorsed by current guidelines is noninducibility of ventricular tachycardia by programmed stimulation [5]. Absence of inducible ventricular tachycardia has been associated with better outcome [49], but has the limitation of the probabilistic nature of ventricular tachycardia inducibility. The predictive accuracy of programmed stimulation has also been disappointing [50]. Ventricular tachycardia recurrence has been observed in 29% of patients who were rendered noninducible acutely [51]; this may reflect healing and contraction of initial radiofrequency lesions [52]. The various techniques of substrate-based ablation described above have introduced additional acute endpoints, such as elimination of all late or abnormal electrograms [27,31,53]. Development of more reliable endpoints for ventricular tachycardia ablation procedures remains a high priority.

The impact of ventricular tachycardia ablation on mortality is not well defined. A recent meta-analysis found a lower mortality in patients in whom ventricular tachycardia was rendered noninducible by programmed stimulation [49]. The analysis, however, did not adjust for differences in other risk predictors. A retrospective analysis performed by Yokokawa et al. [54*] involving 1064 patients from seven centers suggested noninducibility as an independent predictor of survival. The study could not, however, evaluate the independent impact of ventricular tachycardia recurrence on mortality. A strong association between freedom from ventricular tachycardia and transplant-free survival during follow-up was demonstrated in a retrospective multicenter study involving 2061 patients [55*], and patients with early ventricular tachycardia recurrence, within seven days after ablation, had a greater than two-fold increase in mortality in one series [56].

Complications of ventricular tachycardia ablation

The most common complications are related to vascular access and usually resolve spontaneously [57]. Cardiac tamponade occurs in about 1% of patients [58]. From a nationwide inpatient sample database, Palaniswamy et al. [59] reported major complication rates of 11.2% (including vascular: 6.9%, cardiac: 4.3%, and neurologic: 0.5%) and in-hospital mortality rates (1.6%) in the United States during 2002–2011. Most larger trials, however, reported lower complication rates [4,6,7], and at our center, complications related to the ablation are seen in about 5%. The majority of ablation-related mortality is because of uncontrolled ventricular arrhythmia and associated heart failure.

Future directions

Incorporating pathophysiological considerations beyond anatomic scar assessment in mapping of ventricular tachycardia substrate may prove useful. In a recent pilot study, Klein et al. [60*] described a
novel mapping approach for postinfarction ventricular tachycardia ablation by integrating iodine-123-labeled-metaiodobenzylguanidine scintigraphy with voltage mapping, enabling identification of viable but denervated myocardial tissue.

The quality and durability of ablation lesions will continue to be a major determinant of clinical outcome. Technology allowing the monitoring of lesion size will continue to play an important role in improving outcomes. Electroporation is a nonthermocool ablation modality capable of creating deep myocardial lesions. Animal studies have suggested its potential application in epicardial ablation [61].

CONCLUSION
Catheter ablation for ventricular tachycardia associated with ischemic cardiomyopathy can significantly reduce ventricular tachycardia recurrences, and this should translate into an improved quality of life for patients who are having recurrent ventricular tachycardia terminated by ICD shocks and should be considered early in the management of these patients. Complication rates are acceptable, but recurrence rates remain significant. Technological advances in mapping and ablation are expected to further improve success rates.

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R.M.J. receives consulting fees from St Jude Medical, Boston Scientific, and Biosense Webster, Inc.; W.G.S. is coholder of a patent for needle ablation that is co-owned by Boston Scientific, and Biosense Webster, Inc.; W.G.S. is an associate editor of the Journal of Interventional Cardiac Electrophysiology and the Journal of Intervventional Cardiology.

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« of special interest
£ of outstanding interest

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14. Wijnmaalen AP, van der Geest RJ, van Huls van Taxis CF, et al. Improved outcomes. Electroporation is a nonthermal ablation modality capable of creating deep myocardial lesions. Animal studies have suggested its potential application in epicardial ablation [61].

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55. The study showed that noninducibility after ventricular tachycardia ablation in patients with postinfarction ventricular tachycardia – the only end point criteria endorsed by current guidelines is independently associated with lower mortality.


57. The study showed that noninducibility after ventricular tachycardia ablation in patients with postinfarction ventricular tachycardia – the only end point criteria endorsed by current guidelines is independently associated with lower mortality.


63. The study describes a novel mapping approach by integrating molecular imaging of sympathetic denervation in ventricular tachycardia ablation procedures. Functional imaging beyond anatomic scar assessment in mapping of ventricular tachycardia substrate may prove useful.