REVIEWS



Innovations in ventricular tachycardia ablation

Qussay Marashly¹ · Salim N. Najjar² · Joshua Hahn² · Graham J. Rector² · Muzamil Khawaja² · Mihail G. Chelu^{2,3,4}

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Abstract

Catheter ablation of ventricular arrhythmias (VAs) has evolved significantly over the past decade and is currently a well-established therapeutic option. Technological advances and improved understanding of VA mechanisms have led to tremendous innovations in VA ablation. The purpose of this review article is to provide an overview of current innovations in VA ablation. Mapping techniques, such as ultra-high density mapping, isochronal late activation mapping, and ripple mapping, have provided improved arrhythmogenic substrate delineation and potential procedural success while limiting duration of ablation procedure and potential hemodynamic compromise. Besides, more advanced mapping and ablation techniques such as epicardial and intramyocardial ablation approaches have allowed operators to more precisely target arrhythmogenic substrate. Moreover, advances in alternate energy sources, such as electroporation, as well as stereotactic radiation therapy have been proposed to be effective and safe. New catheters, such as the lattice and the saline-enhanced radiofrequency catheters, have been designed to provide deeper and more durable tissue ablation lesions compared to conventional catheters. Contact force optimization and baseline impedance modulation are important tools to optimize VT radiofrequency ablation and improve procedural success. Furthermore, advances in cardiac imaging, specifically cardiac MRI, have great potential in identifying arrhythmogenic substrate and evaluating ablation success. Overall, VA ablation has undergone significant advances over the past years. Innovations in VA mapping techniques, alternate energy source, new catheters, and utilization of cardiac imaging have great potential to improve overall procedural safety, hemodynamic stability, and procedural success.

Keywords Ventricular tachycardia \cdot Ventricular arrhythmia \cdot Catheter ablation \cdot VT mapping \cdot Stereotactic radiation therapy \cdot Innovation

Abbreviations and acronyms

	•
DEEP	Decrement evoked potential
ECMO	Extracorporeal membrane oxygenation
EGM	Electrogram
ICD	Implantable Cardioverter Defibrillator
IABP	Intra-aortic balloon pump
IES	Intramural Excitable Substrate
ILAM	Isochronal late activation mapping
LGE-MRI	Late gadolinium enhancement on late gado-
	linium enhancement MRI

Mihail G. Chelu Mihail.Chelu@bcm.edu

- ¹ Division of Cardiology, Department of Medicine, Montefiore Medical Center, Albert Einstein College of Medicine, Bronx, New York, USA
- ² Division of Cardiology, Baylor College of Medicine, 7200 Cambridge Suite A6.137, MS: BCM621, Houston, TX 77030, USA
- ³ Baylor St. Luke's Medical Center, Houston, USA
- ⁴ Texas Heart Institute, Houston, USA

MCS	Mechanical circulatory support
MEM	Multielectrode mapping
PFA	Pulse field ablation
PPM	Point-by-point mapping
PVAD	Percutaneous ventricular assist device
SAFE-T	Simultaneous amplitude frequency electro-
	gram transformation
STAR	Stereotactic arrhythmia radioablation
UHD	Ultra-high-density
VA	Ventricular arrhythmia
VT	Ventricular tachycardia
VF	Ventricular fibrillation

1 Introduction

Ventricular arrhythmias (VAs) represent a heterogeneous spectrum ranging from single premature ventricular contractions to sustained ventricular tachycardia (VT) and ventricular fibrillation. VAs are a significant cause of morbidity and mortality [1]. VT ablation is currently an essential therapeutic option [1]. Despite a reduction in ICD shocks with VT ablation, no significant reduction in mortality was demonstrated in multiple randomized controlled studies [2–4]. The VANISH trial has demonstrated that in patients with ischemic cardiomyopathy, VT ablation results in a significant reduction in a composite outcome of death, VT storm, and appropriate ICD shocks compared to escalation of antiarrhythmic drugs [5]. This review article discusses recent innovations in VT ablation, including novel mapping techniques, ablation techniques, alternate energy sources, new catheters, and imaging modalities.

2 Novel VT mapping techniques

Sustained monomorphic ventricular tachycardia is mostly caused by scar-related re-entry [6]. The substrate for re-entry is fibrosis areas interspersed within surviving myocytes leading to regions of conduction blocks and regions of slow conduction [1]. Success of VT ablation depends on accurate mapping and elimination of the VT circuits. In general, the two main approaches for identifying VT circuits are (1) activation/entrainment or pace-mapping and (2) substrate-based mapping. Substrate-based mapping allows for the identification of areas supporting VT re-entry regardless of VT inducibility or hemodynamic intolerance with VT induction [1].

2.1 UHD mapping

UHD mapping involves the use of multielectrode mapping (MEM) catheters with smaller electrodes with tight interelectrode spacing to acquire >1000 mapping points (Fig. 1). Compared to point-by-point mapping (PPM) catheters, MEM catheters have smaller electrodes with tight interelectrode distances, which improves spatial resolution [7, 8]. MEM has a lower sensitivity for far-field signals yet a higher sensitivity for local electrical activity [9, 8]. It has been shown to provide better electroanatomic mapping correlation with contrast-enhanced cardiac magnetic resonance scar assessment [9, 10], shorter radiofrequency ablation time [9], and shorter mapping time [11]. MEM can identify areas of preserved myocardial bundles within heterogeneous scar areas, which would otherwise be recognized as dense scar by mapping with standard ablation catheter (3.5–4 mm electrode) [8].

2.2 RM

Ripple mapping (RM) represents endocardial activation in a dynamic 3-dimensional configuration conveying intracardiac electrograms timing, morphology, and location [12]. It is a feature of CARTO-3[©] electroanatomic mapping system (Biosense Webster Inc, Irvine, CA). RM has been used in humans to identify slow conduction channels within scar areas in ischemic and non-ischemic VT [13-16]. As RM displays all components of an electrogram by dynamic bars corresponding to the electrogram voltage, location, and timing, it allows differentiation between local activation signals originating from within the scar area and far-field signals from nearby myocardial tissue [13, 14]. RM-guided VT ablation has been shown to result in significant short-term and long-term freedom of VT [14, 16]. In addition, Xie et al. showed 97% of RM conduction channels localize within or adjacent to areas of late gadolinium enhancement on late gadolinium enhancement MRI (LGE-MRI) [16].

2.3 ILAM

Isochronal late activation mapping (ILAM) is a voltageindependent functional mapping strategy that displays conduction velocity within the scar area visually; with conduction velocity being proportional to isochronal thickness [17]. Areas of slow conduction "deceleration zones" within scar have been recognized to represent critical sites for VT

Fig. 1 Diagram of the grid mapping catheter (the Advisor HD Grid Mapping Catheter). The 4 splines with 4 1-mm equidistanced (3 mm) electrodes allow parallel (along the splines) and perpendicular (across the splines) bipole recordings (reproduced from Okubu et al. [11] with permission)



re-entry (Fig. 2) [18–21]. ILAM has demonstrated that areas of isolated late potentials could be geographically and functionally distant from areas of slow conduction [22]. Anter et al. have shown that areas of slowest conduction during tachycardia exhibited very slow conduction in sinus rhythm and corresponded to VT isthmus site. Compared to voltage and EGM criteria for identification of VT isthmus, areas of significant conduction slowing (>75%) during sinus rhythm had a significantly higher positive predictive value for identification of VT critical zones [23]. ILAM allows for identification of VT critical sites during sinus rhythm with a higher sensitivity and specificity compared of voltage and EGM mapping.

2.4 DEEP mapping

Decrement evoked potential (DEEP) mapping aims at identifying near-field potentials that demonstrate delayed or "decremental" conduction after a single "evoked" extrastimulus [24]. DEEP mapping exhibits improved sensitivity and significantly higher specificity in detecting areas of VT diastolic pathways compared to late potentials [24, 25]. A multicenter study showed that DEEP mapping-guided VT radiofrequency ablation (RFA)–reduced VT inducibility after ablation by 80% and resulted in a significant reduction in number of shocks and VT burden within 6 months postablation as well as 75% freedom of VT at 6 months of follow-up [25].

2.5 Omnipolar mapping

Bipolar electrograms are relatively spared from far-field signal artifacts compared to unipolar electrograms. However, other factors such as angle and contact force of electrode, activation wavefront orientation relative to bipole pair, myocardial fiber orientation, electrode size, and interelectrode spacing can influence bipolar electrograms morphology [26, 27]. Omnipolar mapping aims at deriving electrograms that reflect wavefront direction and velocity independent of electrode orientation [28]. Omnipolar electrograms are obtained using the method described by Deno et al. [28]. An omnipolar electrogram represents a unifying electrogram created from simultaneous adjacent bipoles from an electric field caused by a passing activation wavefront [29]. Compared to unipole- and bipole-derived data, omnipole-derived conduction velocity and activation direction in swine models exhibited more consistent measurements as well as higher sensitivity for shifts in activation direction [30]. Omnipolar mapping has been shown to provide an enhanced delineation of VT substrate, increased identification of conduction



Fig.2 Area of slow conduction within scar indicated by long fractionated electrogram (red circle) corresponds to area of isochronal crowding — "deceleration zone." The critical isthmus site/termi-

nation site courses through the deceleration zone during VT (reproduced from Aziz et al. [22] with permission)

channels, as well as a higher and more consistent peakto-peak voltages compared to bipolar mapping [31, 32]. Besides, it provided a better correlation with evidence of scar on LGE-MRI [32]. No randomized clinical trials have compared omnipolar mapping to other forms of EGM recording [33]. Moreover, omnipolar mapping requires dedicated electro-anatomical mapping systems as well as more sophisticated computational efforts, which might be a limiting factor for its widespread use [33].

2.6 SAFE-T

The simultaneous amplitude frequency electrogram transformation (SAFE-T) mapping strategy is an automated process that recognizes high-frequency potentials during sinus rhythm using the Hilbert-Huang transform (HHT) processing tool for nonlinear and nonstationary signals [34, 35]. The SAFE-T value is a product of the instantaneous amplitude and instantaneous frequency of an electrogram. It is derived from the HHT processing tool and is expressed as (Hz mV). Louise et al. have shown that a SAFE-T cutoff value of >3Hz mV provides 92% sensitivity and 78% specificity for predicting VT origin [34]. It has been shown to provide improved identification of VT isthmus site compared to unipolar and bipolar low-voltage zones and late potential mapping [35]. Besides, areas of scar identified via abnormal SAFE-T values were significantly smaller than area of unipolar low-voltage zones, bipolar low-voltage zones, and bipolar scar, which can potentially lead to a reduction in VT burden with a more limited area of ablation [35], which can limit procedural complications associated with larger areas of ablation as in full scar homogenization techniques.

2.7 Future directions

Targeting late potentials during sinus rhythm has been shown to reduce VT recurrence after ablation [36–38]. Novel VT mapping techniques, as highlighted above, allow for mapping and targeting sinus rhythm surrogates of VT critical isthmus sites, such as identification of slow conduction channels with RM, areas of slow conduction "deceleration zones" with ILAM, and near-field potentials with delayed or "decremental" conduction with DEEP mapping. This provides a significant advantage compared to conventional mapping techniques by evading hemodynamic instability associated with inducing VT. Novel mapping techniques also allow for identification of potential VT circuits in cases of inconsistent inducibility. Those mapping techniques can be utilized solely or complementary to conventional techniques. In addition to more effective and efficient mapping and VT ablation, we believe such novel techniques can also improve VT ablation procedural safety profile as it would limit peri-procedural hemodynamic compromise and would also allow for reducing the duration of the procedure.

Many trials (Table 1) are focused on evaluating new mapping technologies, as more precise identification of VT circuits could lead to improved ablation outcomes.

3 Innovations in ablation techniques

3.1 VT RFA

3.1.1 Bipolar RFA

Conventionally, VT RFA is performed in a unipolar configuration, in which an irrigated-tip catheter is used, and energy is applied between the catheter tip a reference electrode on the patient's skin (Fig. 3A). However, despite the effectiveness of this approach in arrhythmia ablation, VT recurrence is still significant, which is in part related to the failure of transmural lesion creation [39]. Bipolar RFA ablation involves energy application between two closely placed catheter tips (Fig. 3B), which leads to a focused high-density energy delivery to the myocardial tissue between the two catheter tips, which improves lesion transmurality [40-42]. It has also been shown that bipolar ablation can reduce the time needed to create transmural lesions compared to unipolar ablation [40]. Bipolar RFA is particularly useful when targeting VT circuits deep within the tissue such as interventricular septum and intramyocardial tissue as unipolar RFA may not achieve sufficient energy delivery to create a transmural lesion [42-44]. Bipolar ablation is shown to be safe and effective when conventional unipolar ablation strategies fail to eradicate VT, even when performed in critical structures such as VA arising close to the His-bundle [45, 46].

3.1.2 Sequential and simultaneous unipolar RFA (Fig. 3C)

Sequential unipolar RFA involves the application of unipolar RFA energy in sequential order from endocardial then epicardial sites or vice versa, whereas simultaneous unipolar RFA involves the delivery of unipolar RFA simultaneously from endocardial and epicardial sites. Simultaneous RFA has been shown to be highly effective and safe in cases of failed sequential RFA [47, 48]. Compared to sequential ablation, in simultaneous ablation, radiofrequency energy is delivered from both sites of the myocardium leading to decreased temperature loss via convective cooling from the opposite side of the unipolar energy source as well as increased current density and heat load and delivery to intramyocardial tissue creating deeper lesions [47, 48]. It has been previously shown that unipolar RFA with an output of 30 W creates a lesion with a depth of 5 to 8 mm [49]. Thus, in cases where the distance between the endocardial and

Trial title NCT number Primary outcome Ablation of Ventricular Tachycardia Guided by Multi-site Pacing Using NCT04477499 VT recurrence at once year f Parallel Mapping Ripple Mapping Guided Ablation of Ischaemic Ventricular Tachycar- NCT03997201 VT or ICD events within one dia: A Multi-centre Prospective Clinical Trial. NCT03997201 VT or ICD events within one	NCT number Drimory outcome	H
Ablation of Ventricular Tachycardia Guided by Multi-site Pacing Using NCT04477499 VT recurrence at once year f Parallel Mapping Ripple Mapping Guided Ablation of Ischaemic Ventricular Tachycar- NCT03997201 VT or ICD events within one dia: A Multi-centre Prospective Clinical Trial.		1 Ime perspective
Ripple Mapping Guided Ablation of Ischaemic Ventricular Tachycar- NCT03997201 VT or ICD events within one dia: A Multi-centre Prospective Clinical Trial.	NCT04477499 VT recurrence at once year following ablation with a parallel mapping technique	Prospective
	NCT03997201 VT or ICD events within one year of ablation with a parallel mapping technique	Prospective
Paced Electrogram Feature Analysis (PEFA) for Ablation Targeting in NCT03870854 All-cause mortality, appropr Ischaemic Ventricular Tachycardia PEFA mapping.	NCT03870854 All-cause mortality, appropriate ICD shocks, VT storm, non-ICD shocked treated VT, and incessant VT after 3 years of ablation using PEFA mapping.	Prospective
UltraSOUND-based Characterization of Ventricular Tachycardia SCAR NCT03862339 Accuracy of Intracardiac Ech and Arrhythmogenic Substrate; The SOUNDSCAR Study arrhythmogenic substrate d	NCT03862339 Accuracy of Intracardiac Echocardiography (ICE) to specifically define arrhythmogenic substrate during VT ablation	Prospective
Non-Invasive 3D Mapping for Identifying Origin and Preferential NCT03831516 Accuracy of the Medtronic C Propagation of Ventricular Arrhythmias	NCT03831516 Accuracy of the Medtronic CardioInsight vest in identifying ablation targets	Prospective, observational
Peri-Procedural Transmural Electrophysiological Imaging of Scar- NCT03713866 VT termination and recurren Related Ventricular Tachycardia	NCT03713866 VT termination and recurrence rate within 6 months of procedure	Interventional (clinical trial)
Ablation at Virtual-hEart pRedicted Targets for VT NCT03536052 VT inducibility after virtual	NCT03536052 VT inducibility after virtual heart-guided VT ablation	Interventional (clinical trial)
Real-time Automated Program for IDentification of VT Origin NCT03523286 VT non-inducibility at end o electroanatomical mapping and ECGs acquired during	NCT03523286 VT non-inducibility at end of ablation after RAPID-VT software (3D electroanatomical mapping system derived from cardiac CT imaging and ECGs accurited during VT induction) mapping.	Interventional (clinical trial)

epicardial sites is >5 to 8 mm, simultaneous RFA should be considered over sequential RFA [48]. Compared to bipolar ablation, in simultaneous ablation, the two ablation catheters are independent of each other, which allows for individual control of power delivery at each side of the myocardial tissue, which theoretically can decrease the risk of surrounding tissue thermal damage (e.g., coronary arteries) and steam pop formation [44, 47, 48].

3.2 Epicardial VT ablation

Epicardial VT ablation has been clinically utilized in certain VT scenarios where a concealed VT circuit is present, such as arrhythmogenic right ventricular cardiomyopathy (ARVC), inferolateral non-ischemic cardiomyopathy VT, post-infarction transmural substrate, and in cases where endocardial ablation fails to control VT [50-53]. In patients with scar-related VT, a combined endocardial-epicardial ablation approach is associated with a decreased risk of VT recurrence and subsequent mortality compared to the endocardial-only ablation approach [52]. Moreover, using bipolar substrate mapping for scar identification, endocardial-epicardial VT ablation via scar homogenization is associated with a significantly higher success rate at 5 years of follow-up and reduced need for antiarrhythmic drugs compared to endocardial-only ablation [54]. Interestingly, epicardial adjunctive ablation was performed on all patients assigned to the endocardial-epicardial VT ablation treatment group despite VT being non-inducible following endocardial ablation [54]. Better outcomes with adjunctive epicardial ablations seem to be related to superior access to the epicardium and deep intramural tissue [54]. Recently, Matos et al. showed data from a multicentric observational registry demonstrating that endocardial-epicardial VT ablation whether done in a combined or sequential strategy to be associated with lower VT recurrence and lower all-cause death in patients with ischemic or nonischemic cardiomyopathy who underwent repeat VT ablations [55]. Despite reports of increased complications with epicardial access [52], complications can be significantly reduced with improved skill and experience [53, 54].

Besides the conventional subxiphoid epicardial access technique introduced by Sosa et al. [56], new techniques have been developed for improved epicardial access safety and success. Kumar et al. described the needle-in-needle technique in which subxiphoid access to the cardiac silhouette is performed via an 18-G Cook needle followed by the insertion of a 21-G micropuncture needle into the 18-G Cook needle [57], which allows pericardial access via a 21-G needle rather than a 17-G or 18-G needle. Epicardial access via a micropuncture needle is associated with decreased risk of large pericardial effusion when compared to epicardial access using a large-bore needle [57, 58]. Di

Fig. 3 Radiofrequency ablation. (A) Unipolar radiofrequency ablation. (B) Bipolar radiofrequency ablation. (C) Simultaneous unipolar radiofrequency ablation



Biase et al. described an epicardial access technique using the EpiAccess system (EpiEP, Inc., New Haven, CT) in which a fiber optic–equipped needle provides real-time confirmation of epicardial access via pressure waveform monitoring [59]. Moreover, real-time pressure waveform monitoring allows recognition of needle contact with the RV wall, which improves procedural safety [59]. Carbon dioxide (CO_2) pericardial space insufflation has been utilized to aid with subxiphoid epicardial access, which can be performed via a right atrial appendage exit procedure (Fig. 4) [60] or via coronary venous system exit procedure [61]. Furthermore, advanced imaging modalities such as CT and CMR have been reported to guide epicardial access and improve procedural safety [62, 63].

3.3 Intramyocardial VT mapping and ablation

Endocardial and epicardial ablation and mapping techniques are limited to myocardial tissue close to the catheter and often fail to identify intramural excitable substrates [64, 65]. Sapp et al. demonstrated that intramural catheter ablation was feasible and resulted in 12 months free of VT in four out of eight patients with VT refractory to antiarrhythmic drugs and ablations [66]. In a larger multicenter study with 31 refractory VT patients, intramural catheter ablation resulted in 48% free of VT at 6 months of follow-up [65]. Stevenson et al. used an ablation catheter or a multi-electrode catheter for initial mapping, then a needle electrode was inserted into the myocardium for unipolar pacing to assess tissue excitability at potential VT target sites based on low-voltage areas [65]. Areas of IES had higher endocardial and intramural needle unipolar voltage as well as higher endocardial and intramural needle electrogram frequency compared to scar areas [67]. Coronary venous system (CVS) mapping and ablation have been proposed as an effective strategy for targeting VA originating from intramural or epicardial foci [68] and is particularly important when arrhythmogenic substrate delineating via endocardial and epicardial mapping techniques is suboptimal such as NICM VT arising from the septum [69–71]. Briceño et al. demonstrated that in patients with suspected intramural septal VA, CVS mapping can improve arrhythmogenic substrate identification and guide endocardial ablation [69]. In all cases, intramural activation and pace maps were better than endocardial and epicardial mapping [69]. CVS mapping is particularly important when epicardial approach is limited as in patients with prior history of CABG due to epicardial adhesions and scarring. Karimianpour et al. have demonstrated that CVS mapping is safe, feasible, and effective in patients with prior history of CABG [72]. Accurate delineation of CVS anatomy via advanced imaging, such as CT, is important when performing CVS mapping as identification of CVS branches within



Fig. 4 Right atrial appendage exit procedure. An example of transatrial guidewire exit: Anteroposterior (A) and lateral (B) contrast angiograms of the right atrial appendage from a catheter positioned at its crest. (C) The back end of a 0.014-inch guidewire crosses the right atrial appendage. (D) A microcatheter is advanced over the guidewire into the pericardial space (reproduced from Greenbaum et al. [60] with permission)

proximity of the arrhythmogenic substrate [72]. Romero et al. have described a novel technique for intramyocardial ablation using intracoronary wire for a patient with prior failed ablation attempts for VA arising from the LV summit [73].

3.4 Impedance modulation

RFA using irrigation catheters is a power-controlled procedure; thus, lesion formation is affected by power output and baseline impedance [74, 75]. Barkagan et al. have demonstrated a strong inverse correlation between baseline impedance and radiofrequency current output and lesion volume [74]. Shapira-Daniels et al. demonstrated that adding or adjusting return patches in patients with failed VT RFA reduces baseline impedance, increases in current output, and improves procedural success [75]. RFA with lower ionic concentration solutions, such as half-normal saline (HNS) and 5% dextrose in water (5DW) has been shown to result in increased energy delivery to tissue and larger lesion formation [76–78]. Nguyen et al. demonstrated that the use of HNS as the irrigant during VT ablation in patients who had failed standard ablation procedures with normal saline being the cooling RFA irrigant is safe and effective [79]. Half normal saline irrigation, compared to normal saline, exhibits a higher electrical impedance and thus is thought to result in less electrical current dispersion and more focused delivery of current to the cardiac tissue [79]. Steam pop incidence has been observed to be higher using HNS and D5W [76, 79]. Baseline impedance modulation is an important tool to maximize lesion formation and improve procedural success. However, more data regarding optimal baseline impedance values and safety profile is essential for better procedural optimization.

3.5 Contact force

Contact force has been utilized as an indicator of atrial fibrillation (AF) ablation efficacy. Using irrigated contact force-sensing catheters, Elsokkari et al. examined the contact force needed to achieve electrical unexcitability postablation [80]. Although contact force did not predict postablation electrical unexcitability, a contact force of 10 g or more within the scar region was associated with a significant association with post-ablation electrical unexcitability [80]. In addition, they have found that the retrograde aortic approach to the left ventricle resulted in better contact force with the anterior wall, whereas the transseptal approach had a better contact force with the lateral wall [80]. However, Hendriks et al. showed no difference in VT ablation procedural outcomes or safety profile between contact-force and non-contact force-sensing catheters [81]. Limited data exist regarding measures of VT ablation efficacy. More data is certainly needed to assess the role of contact force sensing catheters and the required contact force for effective and successful VT ablation.

3.6 ICE

Integration of intracardiac echocardiography (ICE) with VT ablation procedure has been shown to improve procedural overall safety and success by improving visualization of intracavitary structures, direct visualization of the ablation catheter, monitoring of the contact between the catheter and intracavitary structures, and better understanding of catheter positioning in relation to landmark structures [82, 83]. ICE-guided mapping and ablation is especially important in cases of papillary muscle VT/PVCs as anatomic visualization of PM and catheter is essential [82]. The use of ICE during VT ablation has been shown to significantly lower risk for 12-month VT-related admissions and repeat VT ablation compared to VT ablations without ICE [84].

Given the mixed procedural success of VT RFA as well as the risk of adverse effects, alternate modalities for cardiac ablation have become an area of increasing interest. Besides, many clinical trials are evaluating original strategies of VT ablation beyond the standard catheter ablation technique (Table 2).

3.7 Electroporation

Electroporation, also known as pulsed-field ablation (PFA), is achieved simply by applying a voltage between electrodes that contact tissue through various energy modalities including direct, alternating, or pulsed direct current [85]. Although the mechanism for electroporation is not completely understood, the general principle is that electric fields created contact cell membranes and cause pore formation and cell damage which can be reversible or permanent depending on the magnitude and duration of energy applied [86]. PFA is safe, effective, and results in durable ablation lesions in AF [87, 88]. In animal models, PFA has been shown to significantly reduce the window of vulnerability to ventricular fibrillation from baseline values with no evidence of underlying myocardial damage [89]. Ventricular endocardial PFA has been shown to result in homogenous focal fibrosis with preservation of nerves and vessels on histologic assessment [90, 91]. In addition, PFA has been shown to create ablative lesions rapidly (<10 s of PFA energy) [91]. Im et al. have shown that PFA produces greater histologic lesion depths in infarcted myocardial regions, which makes it an important choice for energy delivery in cases of post-infarction cardiomyopathy VT [91]. Recently, Tan et al. described a novel approach to PFA of ventricular myocardium using screw-in pacemaker leads [92]. Active fixation of ablation catheter improves stability and myocardial contact, which results in improved reliability [92]. Besides, it has also been shown that lesion durability improves with higher energy deliveries [92].

3.8 Ethanol ablation

Ethanol ablation is utilized as a last-resort option when radiofrequency ablation fails, especially with recent advances in mapping techniques and epicardial ablation [93–97]. Transarterial coronary ethanol ablation (TCEA) can be particularly useful in cases of septal VT with no endocardial origin and poor epicardial approach candidacy. In addition, TCEA can be considered in patients with a history of cardiac surgery as the epicardial approach may be challenging due to scarring [98]. Due to technical challenges involved with TCEA, retrograde coronary venous ethanol ablation (RCVEA) has become increasingly popular [93, 99]. This approach is thought to have less risk of canulating coronary veins as well as avoiding issues with coronary stenosis which can complicate the procedure significantly. Although ethanol ablation has been shown to improve VT control in the appropriate setting, it is associated with significant morbidity and mortality. Procedural complications occur in around 14% of cases, most commonly in the form of permanent or transient complete heart block, and risk of periprocedural mortality has been reported to be around 1% [100]. In addition, several anatomic variations can limit long-term durability of ethanol ablation such as the absence of a target vessel to deliver ethanol to the myocardial region of interest and the presence or development of vein-to-vein collaterals leading to ethanol bypassing the arrhythmogenic substrate [100].

3.9 STAR

Stereotactic arrhythmia radioablation (STAR) for VT was first performed in 2012 for a patient with sustained VT refractory to antiarrhythmic medications [101]. It involves the use of highly focused external beam radiotherapy for local concentrated ablative radiation doses with minimal injury to nearby tissues. The use of STAR for VT treatment has been shown to reduce initial VT recurrence, ICD therapies, and VT burden, in small studies [102-104]. In addition, STAR can reduce antiarrhythmic medication use and improve quality of life [103]. However, recurrences were observed in long-term follow-up [102]. STAR has been shown to be effective after one therapeutic session [102, 105, 106]. Although no significant adverse reactions were reported, low-grade pneumonitis has been reported [107]. Preliminary results from the ENCORE-VT study have shown two probable or definite related adverse events occurred 2 years after the procedure, including a pericardial effusion at 2.2 years and a gastropericardial fistula at 2.4 years [103]. The ideal location and size of the target volume for STAR remain uncertain [108]. On a molecular level, Zhang et al. have shown that STAR-treated tissue exhibited an increase in sodium channels expression (Nav1.5) and gap junctions (Cx43), which increase signal propagation leading to the prevention of re-entry [106]. Clinically, a significant reduction in QRS complex interval and a significant increase in conduction velocity were observed [106].

It is important to note that STAR depends on precise identification of arrhythmogenic substrate such as ischemic scar; thus, its application to other causes of VT such as NICM-related VT or infiltrative disease-related VT might be limited as the arrhythmogenic substrate is diffusely scattered throughout the myocardium. Early evidence of STAR therapy for refractory ventricular arrhythmias has been promising with regard to safety and efficacy outcomes; however, evidence is still limited. Larger randomized clinical trials are needed to make significant claims regarding the utility of this novel non-invasive cardiac ablation strategy [109].

Trial title	NCT number	Primary outcome	Time perspective
A Prospective, Multi-center, Post Market Randomized Controlled Trial Comparing VT Ablation Outcomes Using Remote MAGNETIC Navigation Guided Substrate Mapping and Ablation Versus Manual Approach in a Low LVEF Population	NCT02637947	Freedom from VT within 12 months	Interventional (clinical trial)
Phase I/II Study of 4-D Navigated Non-invasive Radiosurgical Ablation of Ventricular Tachycardia	NCT03601832	Composite of death, arrhythmic storm, or appropriate ICD therapy within 90 days, as well as radiation side effects at 18 months	Interventional (clinical trial)
Stereotactic Arrhythmia Radioablation for Ventricular Tachycardia Management	NCT04065802	Reduction in VT burden at 6 months and a composite safety endpoint of radiation-induced side effects at 90 days	Interventional (clinical trial)
Stereotactic Ablative Radiotherapy (SAbR) for Treatment of Ventricular Tachycardia (VT) Refractory Standard Invasive Ablation Techniques	NCT04334811	Comparative analysis of appropriate ICD therapies at 6 months before and after SAbR treatment	Interventional (clinical trial)
Radiosurgery for the Treatment of Refractory Ventricular Extrasystoles and Tachycardias (RAVENTA)	NCT03867747	Safety: acute toxicity and serious adverse events following radiosurgery at 30 days	Interventional (clinical trial)
Noninvasive Cardiac Radioablation for Ventricular Tachycardia	NCT04162171	Comparative analysis of VT events 6 months before and after and volume of spared healthy tissue	Interventional (clinical trial)
STereotactic RadioAblation by Multimodal Imaging for VT (STRA-MI-VT)	NCT04066517	Safety: serious adverse events at 12 months Efficacy: VT burden at 12 months	Interventional (clinical trial)
Proton Particle Therapy for Cardiac Arrhythmia Extracorporeal Energy Source Ablation of Cardiac Tissue: A First Stage Early Feasibility Study	NCT04392193	Mortality, ICD shocks, recurrent VT, and cardiac arrest (all at 2-year follow-up)	Interventional (clinical trial)

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4 Novel VT ablation catheters

Several new catheters and ablation approaches have been developed to provide more durable and ablation lesions and better identification and ablation of the arrhythmogenic substrate within the epicardium and deep intramural tissue (Table 3).

4.1 Lattice tip focal ablation catheter

While the lattice tip ablation catheter was not specifically designed for VT ablation and much of the supportive data are limited to atrial ablation studies, there is growing interest in this catheter's utility for VT. The lattice catheter, Sphere-9 (Affera, Inc., MA, USA), has dual capacity for both radiofrequency (RF) as well as pulsed-field ablation (PF). Its design involves an expandable 8F irrigated spherical catheter with a lattice electrode surface that allows for higher current delivery at lower density compared to conventional catheters, thereby limiting the risk of tissue overheating and steampop formation [110–112]. In addition, its 9 mini-electrodes create a large area of conductive heating which provides deeper ablation lesions (Fig. 5) [110, 112, 113]. Additionally, it provides improved stability due to the compressibility of the lattice mesh and the complex lattice topography which attenuates sliding across the endocardial surface. The lattice catheter exhibits a similar safety profile in vivo studies compared to the conventional irrigated catheter in terms of steam-pop formation, char formation, and thrombus formation on the catheter [110]. Overall, although further studies on VT ablation in humans with the Sphere-9 lattice catheter are warranted, preliminary data on its use appear promising for the future of arrhythmia management [110].

4.2 SERF catheter ablation

The saline-enhanced radiofrequency (SERF) ablation catheter uses a 25-G needle to deliver radiofrequency energy and degassed warmed saline injection into myocardial tissue [114]. The saline stabilizes the electrode temperature and acts as a medium to deliver energy deep into the tissue [114]. The SERF ablation catheter utilizes convective heating and reduces the chances of steam pops [114]. Animal models have shown that SERF ablation produces large, near-transmural lesions, without evidence of myocardial perforation, wall thinning, or aneurysm formation [114]. SERF catheters offer reproducible diameters, controllable volumes, less thrombolytic material, no steam pops, and can be reliably transmural when needed [114, 115]. High-power needle/saline lesions have been demonstrated to result in significantly deeper and larger lesions when compared to

4				
Type of ablation technique/catheter	· Energy	Design/approach	Advantages	Disadvantages
Lattice Tip	RF/PFA	Expandable 8F irrigated spherical catheter with a lattice electrode surface.	Higher current density, less risk for steam pops, large area for heating; toggle between PF and RF, stability of catheter	Uncertainty regarding depth of lesions for ventricular arrhythmias
SERF	RF	25-G needle to deliver RF energy and degassed warmed saline injection into myocardial tissue.	Convective heating and reduces the chances of steam pops; reproducible diameters, con- trollable volumes, less thrombolytic mate- rial, minimal steam pops and deep lesions	Theoretical risk for reducing ventricular func- tion or promoting wall aneurysm or rupture
Epicardial ablation	RF	Epicardial access	Higher success rate compared to endocardial only ablation.	Epicardial access complications.
Intramyocardial ablation	RF	Intramural needle catheter. Coronary venous system approach. Intracoronary approach.	Improved intramyocardial substrate deline- ation.	Technically complex with increased theoretical risk of mechanical complications.
STAR	High doses of radia- tion	Noninvasive radiation therapy that delivers precise high dose of radiation.	Noninvasive, precise	The ideal location and size of the target vol- ume for STAR remains uncertain; radiation side effects (fibrosis, effusions); longer time to efficacy, may not prevent recurrence
RF radiofrequency, PFA pulsed fiel	d ablation, SEK	<i>AF</i> saline-enhanced radiofrequency, <i>STAR</i> stered	otactic arrhythmia radioablation	

Fig. 5 The lattice catheter provides a larger effective surface area compared to standard linear irrigated electrodes, allowing higher current delivery at lower density, which reduces the risk of tissue overheating (reproduced from Shapira-Daniels et al. [112] with permission)



4-mm-tip and irrigated-tip lesions [115]. Clinical data is limited but there are many promising active trials (Table 4) studying the SERF catheter among other novel ablation catheters and devices.

5 Mechanical circulatory support during VT ablation

Periprocedural hemodynamic decompensation occurs in 11% of patients [116]. IABPs provide relatively low cardiac output augmentation (~0.5 L/min) in comparison to other MCS devices, and their use is limited when used during VT ablation, as their mechanism of counterpulsation relies on a regular, stable, non-tachycardic rhythm to provide support [117, 118]. The TandemHeart (CardiacAssist Inc., Pittsburgh, PA) is a percutaneous left-atrial-to-femoral-artery bypass system and requires two large venous and arterial access sites. Its use is associated with vascular

complications [117]. In addition, its trans-septal approach to the left atrium can interfere with transseptal mapping [117, 119]. TandemHert-assisted VT ablation has demonstrated its safety and feasibility [120, 121]. The Impella (2.5/ CP/5.0) (Abiomed, Danvers, MA) is the most commonly used MCS device employed during VT ablation. Its axial flow pump unloads the LV and provides around 2.5–5 L/ min of cardiac output, but its use can cause electromagnetic interference in magnetic-based mapping systems [117, 122]. Short-term MCS use has not been shown to decrease VT recurrence or increase VT-free survival long term [120–125].

Veno-arterial ECMO uses an external centrifugal pump and membrane oxygenator to extract blood from the venous system and shunt it back into the arterial system, which can provide biventricular support and augment cardiac output upwards of 4.5 L/min [119, 126]. Despite these advantages, outcomes associated with the use of ECMO for hemodynamic support during VT ablation are mixed. In a

 Table 4
 Innovations in catheters and ablation approaches

Trial title	NCT number	Primary outcome	Time perspective
Saline Enhanced Radiofrequency (SERF) VT Ablation Early Feasibility Study (EFS)	NCT03628534	Efficacy: VT non-inducibility following abla- tion via SERF catheter Safety: number of serious adverse events	Interventional (clinical trial)
Saline-Enhanced Radiofrequency (SERF) Cath- eter Ablation for the Treatment of Ventricular Tachycardia	NCT02994446	Serious adverse events, MACE, and VT non- inducibility following SERF catheter ablation	Interventional (clinical trial)
FLExAbility Sensor Enabled Substrate Tar- geted Ablation for the Reduction of Ventricu- lar Tachycardia (LESS-VT) Study	NCT03490201	Rate of complications at 7 days and VT burden at 6 months	Interventional (clinical trial)
Outcomes of Intramural Needle Ablation for Recurrent Ventricular Tachycardia That Has Failed Conventional Radiofrequency Ablation	NCT02799693	Hospitalization or recurrence of VT at 6 months and serious adverse events within 30 days of ablation	Observational
Endocardial-Epicardial Radiofrequency Abla- tion of Ventricular Tachycardia in Patients With Ischemic Cardiomyopathy	NCT04512911	Freedom from VT after 1st and 2nd interven- tions at 12 months, respectively	Interventional (clinical trial)

study of 64 patients undergoing VT ablation, pre-emptive use of EMCO was associated with improved survival and lower rates of VT recurrence, while another case series of 21 patients who required ECMO as "rescue therapy" following the development of hemodynamic compromise during VT ablation showed a higher rate of complications and decreased VT-free survival [127, 128].

Santangeli et al. have identified eight clinical factors associated with a high risk for acute hemodynamic decompensation during VT ablation: advanced age (>60 years), ischemic cardiomyopathy, advanced heart failure status with EF <25%, NYHA class III/IV, COPD, DM, VT storm at presentation, and use of general anesthesia [116]. Santangeli et al. have developed the PAINESD risk score to identify patients with a high risk for acute hemodynamic decompensation [129]. In a propensity score–matched analysis, Muser et al. have shown that the prophylactic use of MCS during VT ablation in high-risk patients, identified using the PAINESD score, is associated with a reduction in mortality [130, 131].

Despite mixed data regarding the use of generalized prophylactic use of MCS devices during VT ablation, we believe that selective prophylactic use of MCS during VT ablation for patients with a high risk of acute hemodynamic decompensation is critical for patients' safety and procedural success. Larger randomized controlled clinical trials are needed for more accurate and precise identification of patients who would benefit from MCS during VT ablation.

6 Innovations in imaging

6.1 MRI

LGE-MRI has been shown to correlate well with the location and character of myocardial scarring [132]. Augmentation of electro-anatomical mapping (EAM) for the identification of arrhythmogenic foci with LGE-MRI has been an ongoing area of advancement, with well-established pre-clinical and preliminary evidence of feasibility (Fig. 6) [133, 134]. In small studies, MRI-guided ablation was shown to reduce VT recurrence [135] and radiofrequency applications [136]. Moreover, T1-weighted imaging have been studied for use in assessing necrosis at ablation sites [137, 138]. Lesion dimensions on T1-weighted imaging has been shown to correlate strongly with necrosis in pathology [139]. As inadequate ablation lesion formation is believed to be responsible for post-ablation VT recurrence, the acquisition of inversion-weighted T1 MRI during ablation may improve the durability of VT ablation.



Fig. 6 (A) An example of ventricular tachycardia (VT) ablation performed using endocardial electroanatomical mapping without magnetic resonance imaging (MRI). (B) An example of VT ablation performed with electroanatomical mapping with MRI-derived scar integration. The scar detected on late-gadolinium enhancement MRI corresponded to areas of low-voltage, fractionated, and isolated potentials found on the electroanatomical mapping. Ao, aorta; PA, pulmonary artery (reproduced from Cuculich et al. [135] with permission)

6.2 Real-time MRI

Real-time MRI during ablation offers the benefit of increasing target precision as compared to traditional methods which are limited by errors in mapping registration [137]. Registration errors up to 3.5 mm are introduced by the integration of previously obtained cardiac imaging (CT or MRI) with EAM [140]. Real-time MR-guided procedures are hypothesized to be able to reduce targeting errors such that arrhythmogenic foci of 1–2 mm can be accurately targeted [141]. In porcine models, voltage mapping guided by real-time MRI was 57% sensitive for a scar on LGE-MRI and 96% specific [142].

7 ECGi

Electrocardiographic imaging (ECGi) is the use of a dense array of body-surface electrocardiograms to non-invasively reconstruct a digital 3D model of a patient's heart [143]. A 252-electrode vest is used to collect the electrographic potentials which are then integrated with anatomical data from a high-resolution non-contrast cardiac CT [144]. Activation maps, voltage maps, isopotential maps, and phase maps can be created through post-processing algorithms (Fig. 7A) [144]. ECGi can accurately identify the origin of PVCs [145]. Simultaneous EAM and ECGi showed moderate concordance in activation-repolarization maps. The average spatial resolution of ECGi for sites of focal activation was 13.2 mm, which may not be sufficient for the guidance of VT ablation [146]. ECGi could be utilized to identify areas of low-voltage fractionated EGMs during sinus rhythm, which can aid in identifying the VT critical sites [147, 148]. However, current evidence is very limited and ECGi clinical validation in large patient cohorts has been a challenge [143].

An example of ECGi is VIVO "View into Ventricular Onset" (Catheter Precision, Mt. Olive, NJ), which integrates standard 12-lead ECG, a 3D image of ECG lead placement, and myocardial model from pre-procedural imaging (CT/ MRI) to localize ventricular arrhythmias (Fig. 7B) [149], which allows ORS vectors to be accurately calculated and localized by the anatomical region of the heart [150]. In a prospective observational study of VT ablation with standard EAM, VIVO ECGi predicted VT focus with 88% accuracy [149]. Correlation between ECGi electrograms and invasive epicardial recordings is overall well but exhibits significant variability [151]. However, even in cases with suboptimal correlation, improved electrogram correlation is found within 20 mm in the majority cases [151]. Thus, ECGi can potentially decrease mapping time during VT ablation by reducing the mapping area to a 2-cm radius area [151]. In addition, ECGi can be used to plan the strategy of VT ablation by distinguishing epicardial or endocardial VT origin based on EGM morphology [151].



Fig.7 (**A**) Electrocardiographic imaging (ECGi): Body surface electrical potentials from 250 electrodes are integrated with geometric data from computed tomography via ECGi software algorithms (reproduced from Rudy et al. [154] with permission). (**B**) View into ventricular onset (VIVO; Catheter Precision, Mt. Olive, NJ): Cardiac CT or MRI is integrated with a 3D image of the patient torso with ECG electrodes applied to register the position of the 12-lead ECG and produce an activation map (reproduced from Misra et al.

[149] with permission). (C) Virtual heart arrhythmia ablation targeting (VAAT): Algorithmic reconstruction of the myocardium via MRI (a) allows prediction of ablation targets (c) (reproduced from Prakosa et al., [152] with permission). (D) InHEART: Preprocedural CT images are processed such that endocardium is segments, myocardial wall thinning is identified, a channelness filter is applied, and the Eikonal simulation predicts ablation targets (reproduced from Cedilnik et al. [153] with permission)

8 Computational models

8.1 Virtual heart

Virtual heart arrhythmia ablation targeting (VAAT) involves an algorithmic reconstruction model of an individual patient's myocardium from LGE-MRI data to identify optimal ablation targets for inducible reentrant VT pathways by simulation (Fig. 7C). As VAAT targeting predicts sites of arrhythmia due to underlying substrate, not solely the currently active culprit, it may decrease VT recurrence and reduce the need for repeat procedures [152].

VAAT-guided ablation matched EAM-based targets with smaller lesion sizes in both porcine models and retrospective human studies. In a five-patient prospective study, VAATguided ablation achieved acute lesion success, demonstrating clinical feasibility [152]. VAAT has been developed for scar-related VT and its utility outside of this setting is uncertain [152].

8.2 InHEART

A novel framework for the integration of computational modeling with pre-procedural CT has been advanced. A computational model applying an Eikonal model of wave-front propagation to thickness maps of the myocardial wall can create an individualized 3D simulated model of myocardial electrophysiology. Application of an image-based algorithm to the resultant model allows automatic prediction of VT isthmuses (Fig. 7D). Concordance with invasive EAM was seen in only 5 of 9 human subjects undergoing VT ablation. Such a model can be rapidly applied pre-procedurally to aid intervention planning and identify targets; however, data on clinical outcomes are lacking [153].

9 Conclusion

Approaches to VT ablations have developed rapidly to enhance its success, safety, and accuracy. Innovation in VT mapping techniques has lessened the requirement for mapping of poorly hemodynamically tolerated VTs to achieve satisfactory ablation efficacy. We believe that ILAM provides not only a novel approach to functional VT ablation during sinus rhythm but also a better understanding of VT by identifying area of slow conduction. In addition, we believe that epicardial access techniques as well as ICE are essential skills in the electrophysiologist's toolbox when performing VT ablation. Mastering epicardial access allows for more efficient and effective ablation by targeting epicardial arrhythmogenic substrate as well as intramyocardial arrhythmogenic substrate via simultaneous unipolar RFA and bipolar RFA. New energy sources, such as electroporation and stereotactic radiation therapy, remain in experimental phases but are promising if current limitations can be overcome. New ablation catheters, such as the lattice-tip ablation catheter, are promising modalities that will potentially improve lesion creation and limit tissue overheating. Moreover, the introduction of various imaging modalities to assess arrhythmogenic substrate identification and ablation success evaluation will potentially enhance the overall success, safety, and accuracy of VT ablation. Despite existing small feasibility studies on electrocardiographic imaging and computational models, they are still not validated in large studies.

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Declarations

Competing interests The authors declare no competing interests.

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