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Procedural Patterns and Safety of Atrial Fibrillation Ablation: Findings from Get With the Guidelines-Atrial Fibrillation

Zak Loring, MD, MHS^{1,2}, DaJuanicia N. Holmes, MS¹, Roland A. Matsouaka, PhD^{1,3}, Anne B. Curtis, MD⁴, John D. Day, MD⁵, Nihar Desai, MD, MPH⁶, Kenneth A. Ellenbogen, MD⁷, Gregory K. Feld, MD⁸, Gregg C. Fonarow, MD⁹, David S. Frankel, MD¹⁰, Jodie L. Hurwitz, MD¹¹, Bradley P. Knight, MD¹², Jose A. Joglar, MD¹³, Andrea M. Russo, MD¹⁴, Mandeep S. Sidhu, MD¹⁵, Mintu P. Turakhia, MD, MAS¹⁶, William R. Lewis, MD¹⁷, Jonathan P. Piccini, MD, MHS^{1,2}

¹Duke Clinical Research Institute, Duke University Medical Center, Durham, NC

²Division of Cardiology, Duke University Medical Center, Durham, NC

³Department of Biostatistics and Bioinformatics, Duke University, Durham, NC

⁴Department of Medicine, University at Buffalo, Buffalo, NY

⁵Intermountain Medical Center Heart Institute, Intermountain Medical Center, Murray, UT

⁶Cardiovascular Medicine Section, Yale University, New Haven, CT

⁷Division of Cardiology, Virginia Commonwealth University, Richmond, VA

⁸Department of Medicine, University of California San Diego, San Diego, CA

⁹Department of Medicine, University of California Los Angeles, Los Angeles, CA

¹⁰Cardiovascular Division, Perelman School of Medicine, University of Pennsylvania, Philadelphia, PA

¹¹North Texas Heart Center, Dallas, TX

¹²Feinberg School of Medicine, Northwestern University, Chicago, IL

¹³Department of Cardiology, University of Texas, Southwestern Medical Center, Dallas, TX

¹⁴Cooper Medical School of Rowan University, Camden, NJ

¹⁵Division of Cardiology, Albany Medical Center, Albany, NY

¹⁶VA Palo Alto Health Care System, Palo Alto, CA; Center for Digital Health, Stanford University, Stanford, CA

¹⁷MetroHealth System Campus, Case Western Reserve University, Cleveland, OH

Abstract

Address for Correspondence: Jonathan P. Piccini, MD, MHS, Electrophysiology Section, Duke University Medical Center, Duke Clinical Research Institute, 200 Morris Street, Durham, NC 27701. Phone: (919) 564-9666; Fax: (919) 668-7057. jonathan.piccini@duke.edu.

Background: Catheter ablation is an increasingly utilized treatment for symptomatic atrial fibrillation (AF). However, there are limited prospective, nationwide data on patient selection and procedural characteristics. This study describes patient characteristics, techniques, treatment patterns, and safety outcomes of patients undergoing AF ablation.

Methods: A total of 3,139 patients undergoing AF ablation between 2016-2018 in the Get With The Guidelines-Atrial Fibrillation registry from 24 US centers were included. Patient demographics, medical history, procedural details and complications were abstracted. Differences between paroxysmal and persistent AF patients were compared using Pearson's Chi-square and Wilcoxon rank-sum tests.

Results: Patients undergoing AF ablation were predominantly male (63.9%) and Caucasian (93.2%) with a median age of 65. Hypertension was the most common comorbidity (67.6%) and persistent AF patients had more comorbidities than paroxysmal AF patients. Drug refractory, paroxysmal AF was the most common ablation indication (Class I, 53.6%) followed by drug refractory, persistent AF (Class I, 41.8%). Radiofrequency (RF) ablation with contact force (CF) sensing was the most common ablation modality (70.5%); 23.7% of patients underwent cryoballoon ablation. Pulmonary vein isolation (PVI) was performed in 94.6% of de novo ablations; the most common adjunctive lesions included left atrial roof or posterior/inferior lines, and cavotricuspid isthmus ablation. Complications were uncommon (5.1%), and were life-threatening in 0.7% of cases.

Conclusions: More than 98% of AF ablations among participating sites are performed for Class I or Class IIA indications. CF-guided RF ablation is the dominant technique and PVI the principal lesion set. In-hospital complications are uncommon and rarely life-threatening.

Introduction

Catheter ablation for atrial fibrillation (AF) has grown significantly as a mainstream therapy with a more than 5-fold increase in utilization between 1999 and 2013.¹ During this time period, professional society guidelines have endorsed catheter ablation as a rhythm control strategy across a range of clinical scenarios.^{2,3} Single-center analyses, European registry studies, and analyses of US administrative claims data have demonstrated an improvement in the safety profile of AF ablation over time.⁴⁻¹¹ However, few data are available describing national practice patterns of procedural techniques and ablation lesion patterns. Additionally, given the major advances and procedural approaches to AF in recent years,¹² analysis of a more contemporary cohort may provide a more accurate assessment of procedural patterns and the safety profile of catheter ablation.

The Get With The Guidelines-Atrial Fibrillation (GWTG-AFIB) registry quality improvement program introduced an ablation module in 2016 that allowed participating sites to report patient characteristics, procedural methods and complication rates of patients undergoing catheter ablation for AF. In this study, we aimed to (1) evaluate the patient characteristics and indications among patients undergoing AF ablation, (2) describe procedural characteristics/techniques and compare them by AF subtype (paroxysmal vs persistent), and (3) assess the frequency of complications.

Methods

Data Collection and Components:

The data used for the analyses presented here were collected as part of the GWTG-AFIB registry. The details of this registry have been described previously.¹³ Briefly, GWTG-AFIB is a national, voluntary quality improvement initiative that began in 2013 in partnership with the AHA and HRS. The goal for the registry is to improve the adoption of guideline-directed therapies and outcomes in patients with AF. The main GWTG-AFIB registry collects data on medical history, hospital care and outcomes of AF patients via an online, interactive case report form. In 2016, a sub-registry module was introduced that allows sites to enroll patients undergoing catheter ablation. Beyond the data collected in the main GWTG-AFIB registry, the ablation module collects data on the indications for ablation, rhythm control history, procedural characteristics, and safety outcomes. The present analysis focuses on patient selection, techniques, and safety outcomes. Participating institutions complied with local regulatory and privacy guidelines, including institutional review board approval where required. Since the GWTG-AFIB registry is a quality improvement program, some institutions consider the program exempt from IRB review. IQVIA served as the registry coordinating center and the Duke Clinical Research Institute served as the analytic core; IRB approval was established for all analytic activities at the analytic core. Data was abstracted by trained personnel using standardized definitions for all data elements.¹³

Indications for Catheter Ablation:

The relative frequency of indications for catheter ablation for AF were reported based on the 2014 AHA/ACC/HRS guidelines, given the study period of 2016-2018 preceded the release of the 2019 Focused Update of these guidelines.^{2,14} Namely, catheter ablation was considered a Class I indication for symptomatic paroxysmal AF refractory to at least one antiarrhythmic drug (AAD), Class IIA for symptomatic persistent AF refractory to at least one AAD, Class IIA as first-line therapy for paroxysmal AF and Class IIB for first-line therapy for persistent AF. The 2019 Focused Update added a Class IIB indication for catheter ablation for treatment of patients with symptomatic AF and heart failure with reduced ejection fraction. This change would not affect the class of recommendations of any of the patients included in the present analysis; however, patients meeting this indication may not be well represented in the study population.

Procedural Characteristics, Safety Endpoints, and Quality Measures:

The characteristics of catheter ablation procedures were reported, including pre-procedural management/imaging, sedation strategy, intra-procedural imaging, ablation technique/lesion sets, provocative testing and procedure times. These variables were compared between AF subtypes (paroxysmal vs persistent). Paroxysmal AF was defined as AF self-terminating within seven days of recognized onset; whereas, persistent AF was defined as AF self-terminating beyond seven days or was terminated electrically or pharmacologically. Frequency of intra-procedural and post-procedural complications were reported for all patients as well as by AF subtype. As an exploratory analysis, the baseline characteristics and procedural management of patients with and without post-procedural complications were compared.

Statistical Analysis

Baseline patient characteristics were described overall and by AF subtype using percentages for categorical variables and medians with interquartile range (IQR, 25th - 75th percentiles) for continuous variables. Differences in these characteristics were compared using Pearson's Chi-square tests for categorical variables and Wilcoxon rank-sum tests for continuous variables. Comparisons were made using non-missing data only. All tests were 2-sided and statistical significance was declared when $p < 0.05$. All analyses were performed with SAS software version 9.4 (SAS Institute, Cary, NC).

Results

Study Population and Baseline Characteristics

Between January 4, 2013 and November 22, 2018 there were 65,565 patients who were registered from 141 sites participating in GWTG-AFIB. We excluded patients who were missing gender information (n=131), received comfort care only on hospital day 0 or 1 (n=822), from hospitals without the ablation module (n=61,102), were missing all data on ablation technique (n=187), received cavotricuspid isthmus (CTI) ablation only (n=115), had long-standing persistent AF (n=32), had other atrial arrhythmias such as left atrial (LA) flutter or LA tachycardia (n=35), or were missing data on their rhythm (n=2). The final cohort contained 3,139 patients from 24 hospitals with a median of 84 patients per site (IQR 10-236). Of participating sites, 85.1% were academic teaching hospitals with more than half (54.2%) having 500 or more beds (site characteristics in Supplemental Table).

Baseline characteristics are presented in Table 1. The median age was 65 years (58-71), 36.1% of patients were female, 93.2% were Caucasian and 2.6% of patients were African American. Most patients had private insurance (64.6%), although a substantial minority (29.9%) had Medicare with a higher prevalence among patients with persistent AF compared with paroxysmal AF (33.7% vs 27.1%, $p=0.001$). The most common comorbidities were hypertension (67.6%), obstructive sleep apnea (OSA, 31.9%), stage 3 chronic kidney disease (CKD, 24.3%), coronary artery disease (18.9%), diabetes mellitus (18.8%), and heart failure (18.2%), with 8.2% of included patients having a left ventricular ejection fraction $< 40\%$. Each of these conditions were more prevalent among patients with persistent AF ($p < 0.001$ for all). Median CHA₂DS₂VASc score¹⁵ was 2 (1-4) and median HAS-BLED score¹⁶ was 2 (1-2); both were higher among persistent AF patients ($p < 0.001$). The median LA diameter was 4.2cm (3.7-4.8) for all patients, with persistent AF patients having larger diameters than paroxysmal AF patients (4.4 [4.0-5.0] vs 4.0 [3.5-4.5], $p < 0.0001$).

Catheter Ablation Frequency by Indication

Most patients (78.1%) underwent first-time (*de novo*) ablation. A total of 18.0% had had one prior ablation, 3.0% had had two prior ablations and less than 1% had had 3 prior ablations (Table 1). Patients with persistent AF were more likely to have undergone prior ablation ($p=0.0002$). The most common indication for catheter ablation was paroxysmal AF refractory to at least one AAD (53.6%) followed by drug refractory persistent AF (41.8%) (Figure 1). Less than 1 in 20 procedures were performed as first line therapy for paroxysmal AF (3.0%) or persistent AF (1.6%).

Procedural Techniques and Characteristics

Patients were treated with AADs at the time of admission for their ablation hospitalization 59.6% of the time (Figure 2). The most commonly used AAD was flecainide (14.6%). Patients with paroxysmal AF were more likely to receive AADs overall (63.4% vs 54.6%) and more likely to receive flecainide, sotalol, dronedarone or propafenone ($p<0.0001$ for all). Persistent AF patients were more likely to receive amiodarone or dofetilide. Peri-procedural management strategies are described in Table 2. The most common peri-procedural anticoagulation strategy was uninterrupted anticoagulation (defined as no interruption to the patient's anticoagulant medication regimen prior to the ablation procedure, 73.6%); bridging (temporary interruption of a patient's chronic anticoagulant medication with use of a short-acting parenteral anticoagulant) was uncommon (4.3%). Pre-procedure transesophageal echocardiography (TEE) was performed in 49.0% of patients, more commonly among persistent AF patients ($p<0.0001$). Pre-procedure computed tomography (CT) was used in most cases (60.3%), while magnetic resonance imaging (MRI) was uncommon (2.9%).

The most common sedation strategy was general anesthesia with endotracheal intubation (87.7%), although 5.9% of patients had conscious sedation and 5.8% of patients had general anesthesia with use of a laryngeal mask airway. Intracardiac echocardiography and 3D electroanatomic mapping were used in more than 90% of cases; whereas, intraoperative TEE and rotational angiography use were uncommon.

The most common energy modality utilized was irrigated radiofrequency (RF) ablation with contact force (CF) sensing (70.5%) followed by cryoballoon (23.7%). Patients with paroxysmal AF were more likely to undergo cryoballoon ablation than persistent AF patients. Provocative testing was common and done most frequently with burst pacing (68.5%). Isoproterenol and adenosine were also commonly used, with isoproterenol more frequently used in persistent AF patients and adenosine more commonly used in paroxysmal AF patients. Median procedure times were 180 minutes (IQR 140-230), with 37 minutes of ablation (IQR 17-59) and 16 minutes of fluoroscopy (IQR 8-28). Persistent AF patients had longer ablation, fluoroscopy and total procedure times ($p<0.0001$ for all).

Figure 3 illustrates the lesion sets employed in patients undergoing ablation for the first time ($N=2,418$, 77% of patients). Overall, pulmonary vein isolation (PVI) was performed in 94.6% of cases. The most common adjunctive lesion set was right sided CTI ablation for atrial flutter (32.9%), LA roof line (30.8%), and LA posterior/inferior line (24.0%). Persistent AF patients were more likely to receive a LA roof line (42.1% vs 21.7%, $p<0.0001$), LA posterior/inferior line (36.0% vs 14.3%, $p<0.0001$), posterior wall (PW) isolation (24.0% vs 8.4%, $p<0.0001$), lateral mitral isthmus line (5.4% vs 2.7%, $p=0.0020$), or septal mitral isthmus line (5.4% vs 3.1%, $p=0.0062$). Complex fractionated electrogram ablation (CFAE) and focal impulse and rotor modulation (FIRM) were performed in 9.2% and 10.7% of patients respectively, with higher rates in persistent AF (11.4% vs 7.4% $p=0.0015$ for CFAE, 14.0% vs 8.1% $p<0.0001$ for FIRM). Among patients with right sided CTI ablation for atrial flutter, 8.4% of patients had a documented baseline rhythm of atrial flutter (typical or atypical).

Figure 4 illustrates the lesion sets employed stratified by first time vs repeat ablation (N=721). Patients undergoing repeat ablation were less likely to receive PVI (94.6% vs 81.4%, $p<0.0001$) and were more likely to receive adjunctive lesions including LA posterior/inferior line (33.4% vs 24.0%, $p<0.0001$), lateral mitral isthmus line (13.0% vs 3.9%, $p<0.0001$), septal mitral isthmus line (11.1% vs 4.1%), LA appendage isolation (6.9% vs 3.6%, $p=0.0002$), SVC isolation (8.6% vs 3.8%, $p<0.0001$), and CFAE (13.6% vs 9.2%, $p=0.0001$).

Safety Outcomes

In-hospital complications were noted in 160 patients (5.1% of the total population, Table 3). Complications were more common in patients with persistent AF compared with paroxysmal AF (91 cases [6.7%] vs 69 cases [3.9%], $p=0.0004$). The most common complications reported were volume overload (39 cases, 1.2%), hematoma (33 cases, 1.1%), hemopericardium (14 cases, 0.4%), need for pericardiocentesis (11 cases, 0.4%), and urinary tract infection (11 cases, 0.4%). Life-threatening complications were uncommon, with nine cases of tamponade, seven cases of hemorrhage requiring transfusion, four strokes or transient ischemic attacks (TIAs), and one death. There were no reports of atrioesophageal fistula.

The demographics and medical history of patients who experienced a procedural complication and those who did not are compared in Table 4. Patients who had procedural complications were more likely to be older, female, and have hypertension, heart failure stage 3 CKD or COPD. They were also more likely to have larger left atria and have higher thromboembolic and/or bleeding risk scores. Procedural management patterns by complication presence are described in Table 5. Patients who had complications were less likely to have interrupted anticoagulation. They were more likely to have a preoperative TEE, intraoperative intra-cardiac echocardiography (ICE) and less likely to have had rotational angiography. Use of irrigated RF with CF was more frequent among those with complications, whereas cryoballoon use was less frequent. Frequency of specific complication types was similar across energy and catheter types with the exception of volume overload/pulmonary edema which was seen exclusively among patients with RF and contact force (36/2206 patients [1.6%]) and RF without contact force (3/271 patients, [1.1%]) and was not seen in the 626 patients receiving cryoballoon ablation (N=626) or in whom “other” was selected as their energy and catheter type (N=27). Patients with complications more frequently had provocative testing with isoproterenol as well as longer fluoroscopy and total procedure times. Patients with complications had higher rates of ablation on the LA roof, LA posterior/inferior wall, LA appendage, right sided CTI and more targeted ganglion ablation. Complication rates were similar across regions. While 94.4% of patients with complications were treated at hospitals identified as academic or teaching hospitals; it should be noted that, 85.1% of patients were treated at this hospital type (Supplemental Table). Patients with complications were treated at sites with higher case volumes than those without complications (245 vs 244 cases over the study period, $p=0.01$).

Discussion

In this analysis of greater than 3,000 patients undergoing catheter ablation across 24 US centers, catheter ablation for AF was highly consistent with guideline recommendations. The vast majority (>95%) of cases were performed for Class I or Class IIA indications, most patients received antiarrhythmic drug therapy prior to ablation, and anticoagulation was typically not interrupted for the procedure. Open-irrigated RF ablation with CF sensing was the most common ablation modality used, although cryoballoon ablation accounted for nearly a quarter of all cases. Beyond PVI, the most common adjunctive ablation included ablation of the CTI, the LA roof/posterior wall, and rotor ablation. In-hospital complications were uncommon and mortality was rare. These findings confirm and extend prior observations that catheter ablation at experienced centers is safe.

The demographics of the current study of patients undergoing AF ablation are similar to those reported in the World Wide Survey (WWS), National Inpatient Sample (NIS), and broader analyses of AF patients in the GWTG-AFIB registry.^{11,17,18} Compared with the GWTG-AFIB patients with elevated CHA₂DS₂-VASc scores, this catheter ablation population had a lower prevalence of women, as well as African American, and Hispanic patients. Sex and racial disparities have been observed in the GWTG-Heart Failure registry as well as other studies demonstrating that minorities are less likely to receive invasive cardiovascular therapies.^{19,20} These disparities may have contributed to their under-representation in the present study.

The majority of patients were referred for ablation for a Class I indication (drug refractory, paroxysmal AF), with a large proportion referred for Class IIA indications (drug refractory, persistent AF). Less than 1 in 20 patients were referred for first line therapy for paroxysmal or persistent AF. The fact that more than 95% of ablations were performed for Class IA or IIA indications suggests that AF ablation practice is very consistent with guideline recommendations.

The 2017 HRS/EHRA/ECAS/APHRS/SOLAECE expert consensus statement on catheter ablation for AF recommends performance of the ablation procedure without interruption of warfarin (Class I), dabigatran (Class I), rivaroxaban (Class I), or another direct acting oral anticoagulant (Class IIA).²¹ Alternatively, the consensus recommendations counsel that holding 1-2 doses with resumption after the procedure is reasonable (Class IIA).²¹ These data from GWTG-AFIB suggest that anticoagulation in clinical practice is highly consistent with these recommendations. The COMPARE trial randomized patients to continuous warfarin vs a low molecular weight heparin bridge and showed that thromboembolic events were higher with the bridging strategy.²² Thus, it is not surprising that only 4.3% of patients in our study had bridging of anticoagulation in the peri-procedural period.

Open-irrigated radiofrequency ablation was the dominant energy source in the registry with a substantial minority of patients receiving cryoballoon-based ablation. The FIRE AND ICE trial demonstrating non-inferiority of cryoballoon ablation to RF ablation was published in 2016 (near the beginning of this study period).²³ While most cryoballoon ablations were done in patients with paroxysmal AF, 37% of the patients receiving cryoballoon ablation had

persistent AF despite the absence of randomized data in these patients. While observational data has been promising,²⁴ ongoing randomized trials of cryoballoon ablation for the treatment of persistent AF, such as IRON-ICE trial will provide valuable data to guide future practice ([ClinicalTrials.gov NCT03365700](https://ClinicalTrials.gov/NCT03365700)).

The 2017 HRS/EHRA/ECAS/APHRS/SOLAECE expert consensus statement affirms that PVI should be the cornerstone of AF ablation. The most frequent lesion sets employed in adjunctive ablation were CTI ablation, LA roof/posterior wall ablation, and catheter ablation of focal sources. Randomized comparisons of adjunctive substrate ablation have been neutral. In the STAR AF-II trial, neither linear ablation nor CFAE improved the rate of recurrence of AF for persistent AF.^{25,26} Similarly, FIRM ablation has been shown to have no additional benefit over PVI in persistent AF in the recently presented REAFFIRM trial ([ClinicalTrials.gov NCT02274857](https://ClinicalTrials.gov/NCT02274857)); however, the results of this trial were not available during the study period. There has been enthusiasm regarding ablation of the posterior wall, particularly given its shared embryologic origin with the pulmonary veins.²⁷ Some randomized data suggests that addition of posterior wall isolation to PVI improves maintenance of sinus rhythm.²⁸ This study demonstrates that, while PVI continues to be the cornerstone of AF ablation, adjunctive ablation is commonly performed. The uncertainty over the benefit of adjunctive ablation is supported by the significant heterogeneity in its use in clinical practice, including in GWTG-AFIB.

In-hospital complications are uncommon following AF ablation, but they remain an important concern for electrophysiologists, referring physicians, and patients. These data confirm that AF ablation is a safe procedure with a low in-hospital complication rate in specialized centers. This is particularly true with respect to potentially life-threatening complications. A total of 21 life-threatening complications were reported for the 3,139 patients (nine cases of tamponade, seven hemorrhages requiring transfusion, three strokes, one TIA, and one death) representing less than 1% of all patients. These overall complication rates and in-hospital mortality are in line with those previously reported from prior studies, including the WWS, as well as the NIS and provides insight into the real world experience of modern AF ablation.^{5,17} Rates of postoperative hemorrhage, pericardial complications, and neurological complications were lower in the GWTG-AFIB cohort compared to the NIS (2.86% vs 0.2%, 1.99% vs 0.1%, and 1.05% vs 0.1% respectively). Higher ablation volume has been shown to be associated with a lower odds of mortality and complications.¹¹ Thus, it is possible that the lower observed rates of these complications is a reflection of the experience of the enrolling centers which were predominantly (85.1%) academic teaching hospitals.

While the low number of overall in-hospital complications limits the ability to identify predictors or drivers of complications, we did describe the characteristics and procedural management of patients who had or did not have procedural complications. Not surprisingly, patients with higher baseline risk profiles had higher rates of complications as has been demonstrated in analysis of the NIS.¹¹ The higher use of preoperative TEE among patients with complications may be reflective of clinicians using this pre-procedure imaging more frequently in high risk patients. The higher rates of ICE use among patients with complications may follow a similar trend; alternatively, the larger bore venous access that

ICE requires may carry additional risk or ICE may be introduced as a reaction to a complication (e.g. to assess for effusion). Use of contact force-guided RF was more common in patients with complications; whereas cryoballoon ablation was less common. Additionally, patients with complications had more non-PVI ablation. This is consistent with other studies demonstrating higher complication rates with RF ablation compared to cryoballoon and more complications with additional ablation beyond PVI. RF ablation is often performed using continuously irrigated catheters which may increase the risk of pulmonary edema, one of the more frequently reported complications. At present, there is no class I indication for non-PVI ablation in the LA in the 2017 HRS/EHRA/ECAS/APHRS/SOLAECE expert consensus statement.^{21,29} However, clinicians and investigators continue to pursue ablation beyond PVI in an attempt to improve AF-free survival. Ongoing randomized trials will help clarify the role of non-PVI ablation moving forward. Given the association between ablation of non-PV triggers and improved outcomes,³⁰ provocation testing with isoproterenol is often used to identify non-pulmonary vein triggers of AF, and thus may be closely related to non-PVI ablation, and also lacks a Class I indication.^{21,31} Patients with complications were seen at sites that had higher median ablation case volumes. While this difference was statistically significant, the relatively similar absolute values of case volumes suggest that this is likely a reflection of high volume centers amassing more complications as a result of doing more procedures. This analysis compared patients with any complication to those with no complications and therefore equally weighted minor complications (e.g. urinary tract infections) and serious events (e.g. tamponade, death). Thus, the overall results are meant to be descriptive, and hypothesis generating; as with any observational analysis, no causal relationship should be inferred.

The study has several limitations. Data were collected over a three year span (2016-2018) during which time several large randomized controlled trials were published and thus may not fully reflect current practice patterns.^{32,33} The 3,139 patients in this study came from only 24 sites with a median of 84 patients per site and thus may not be representative of patterns in smaller, low-volume practices. Additionally, sites participating in the GWTG-AFIB registry may be more likely to deliver guideline-indicated therapy, which may have introduced selection bias. Sites enrolled consecutive patients to the registry to get as complete an assessment of care patterns as possible without selection bias within sites. Data entry and medical chart review was performed by providers without external audit which raises the potential for inaccurate reporting; however, providers receive training on data entry and a recent analysis of the similarly designed GWTG-Stroke initiative demonstrated an overall composite accuracy of data entry of 96.1% suggesting that the data abstraction in GWTG registries is reliable.³⁴ The present analysis evaluated in-hospital complications and does not capture complications that may present after hospital discharge such as atriopharyngeal fistulae. Future analyses are planned to evaluate medium to long term outcomes including complications, mortality, readmission and resource utilization.

Conclusions

Catheter ablation of AF in the United States among 24 sites participating in the GWTG-AFIB quality improvement registry is performed for Class I or Class IIA indications in more than 98% of cases, consistent with current guideline recommendations. CF-guided RF is the

most common ablation modality and PVI remains the dominant ablation lesion set. Major in-hospital complications including tamponade, stroke and death are infrequent.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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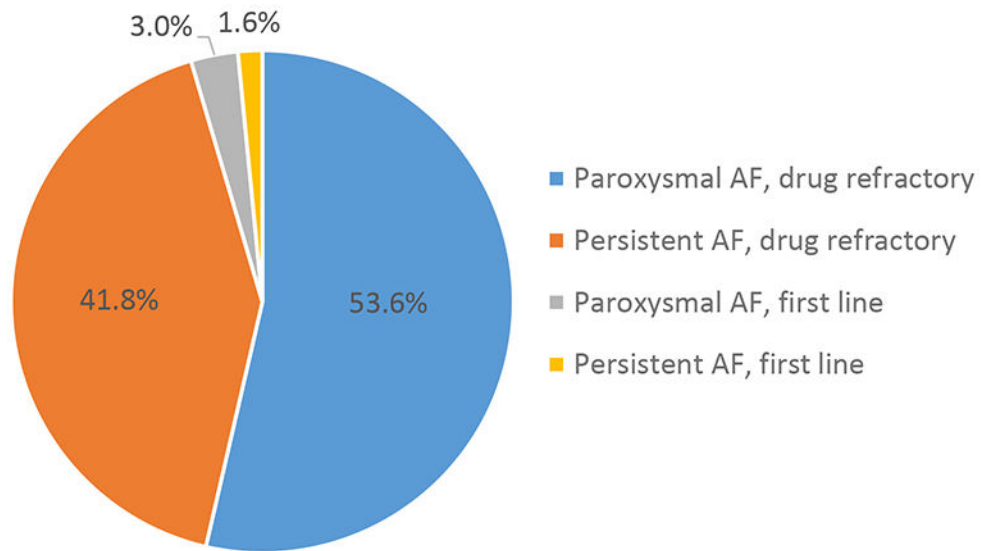


Figure 1. Indications for Catheter Ablation:

Percentage of patients referred for catheter ablation for paroxysmal atrial fibrillation (AF) refractory to at least one antiarrhythmic drug (AAD), persistent AF refractory to at least one AAD, first line therapy for paroxysmal AF, and first line therapy for persistent AF.

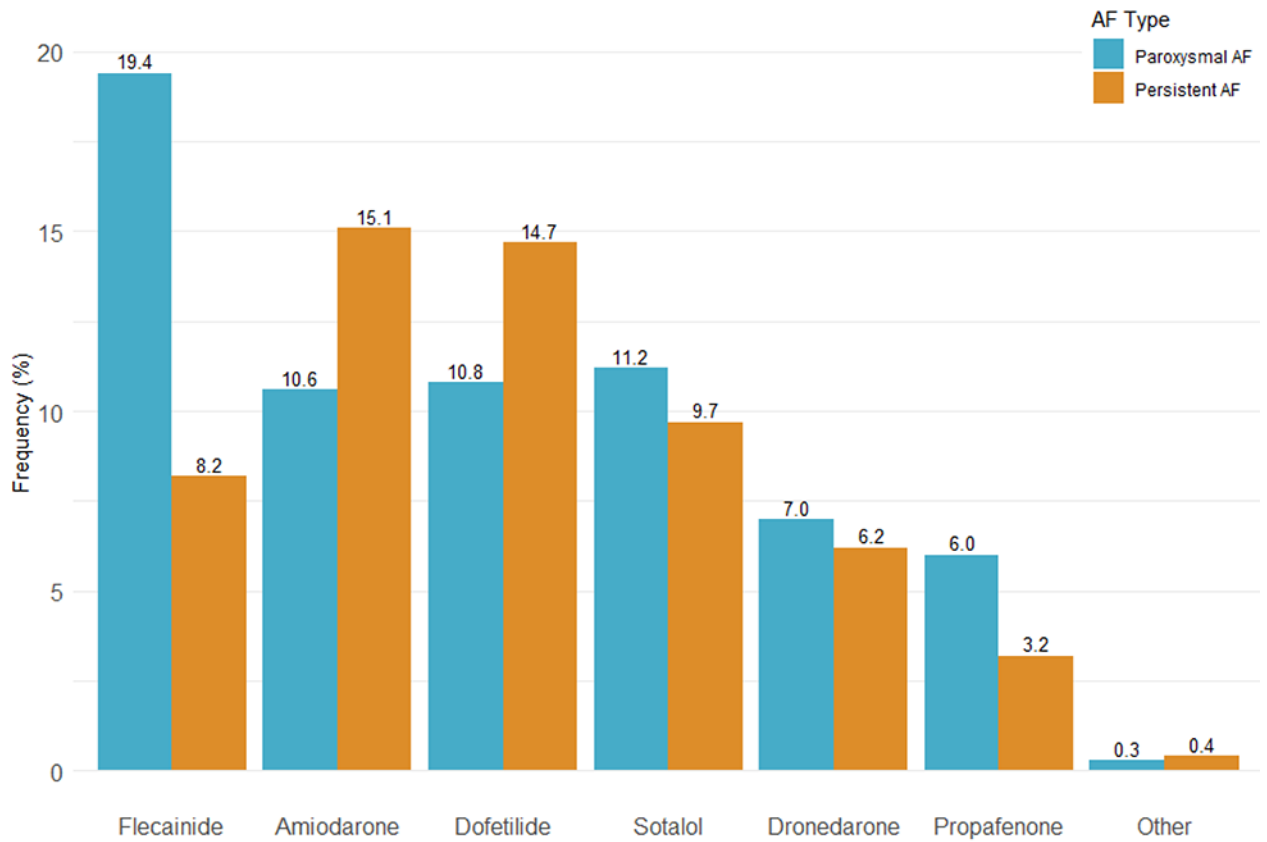


Figure 2. Antiarrhythmic Use:

Frequency of antiarrhythmic use prior to catheter ablation subgrouped by atrial fibrillation (AF) subtype. $P < 0.0001$ for all comparisons between paroxysmal and persistent AF.

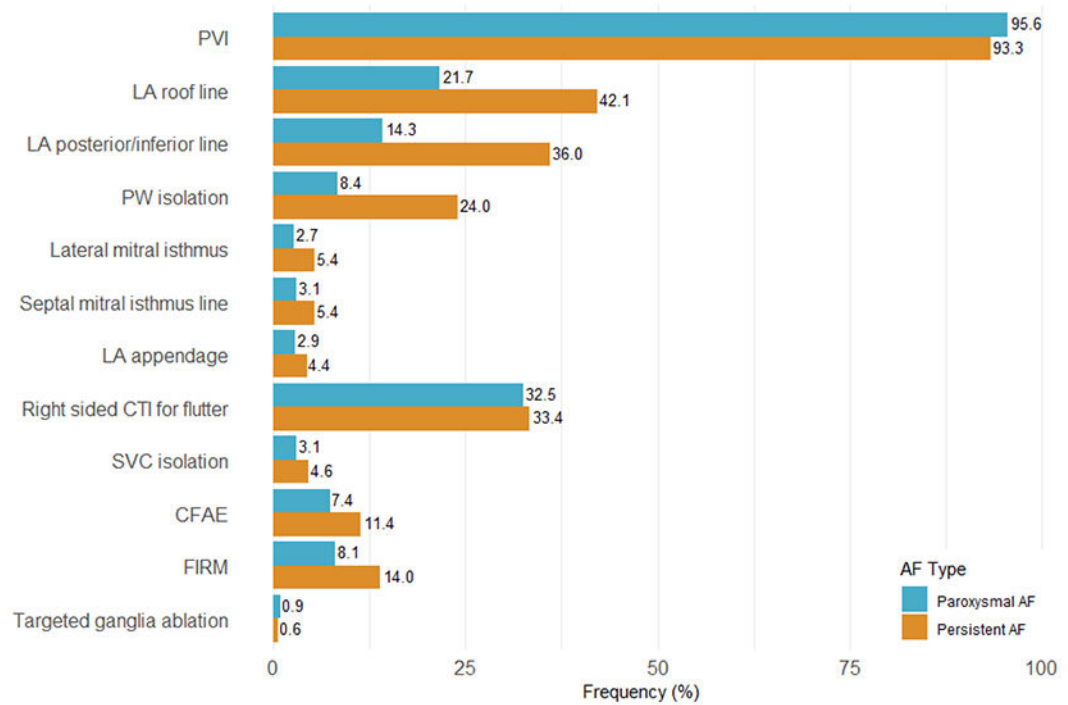


Figure 3. Ablation Techniques for First Ablations:

Frequency of different ablation lesion sets for patients undergoing first ablations (N=2418) subgrouped by atrial fibrillation (AF) subtype. Abbreviations: PVI = pulmonary vein isolation; LA = left atrial; PW = posterior wall; CTI = cavotricuspid isthmus; SVC = superior vena cava; CFAE = complex fractionated atrial electrogram; FIRM = focal impulse and rotor modulation

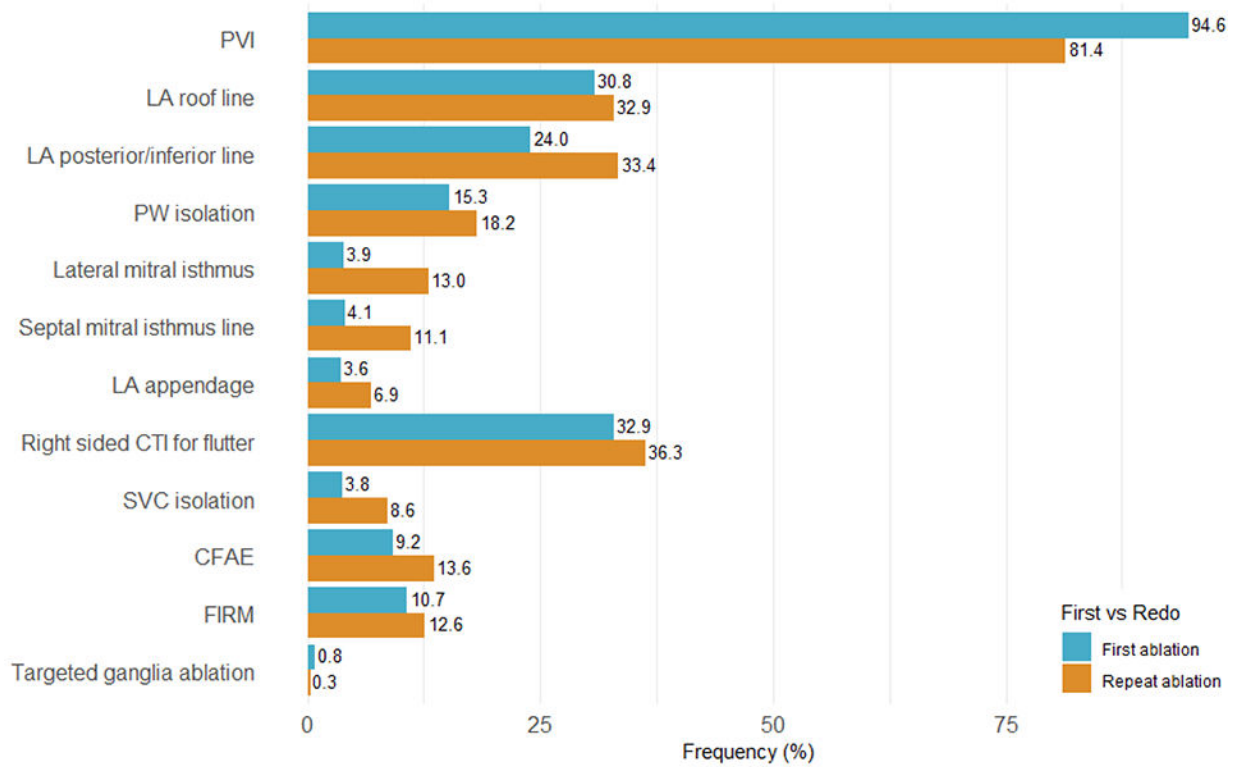


Figure 4. Ablation Techniques for First Time vs Repeat Ablations:

Frequency of different ablation lesion sets for patients undergoing first time ablation (N=2418) vs repeat ablations (N=721). Abbreviations: PVI = pulmonary vein isolation; LA = left atrial; PW = posterior wall; CTI = cavotricuspid isthmus; SVC = superior vena cava; CFAE = complex fractionated atrial electrogram; FIRM = focal impulse and rotor modulation

Table 1:**Baseline Characteristics:**

Demographics and medical history for all included patients and by AF subtype. Data presented as median (IQR) or N (%) as appropriate.

	Overall	Paroxysmal AF	Persistent AF	P-value
N	3139	1778 (56.6%)	1361 (43.4%)	
Age, years	65 (58-71)	64 (57-70)	66 (60-72)	<0.0001
Female	1132 (36.1%)	711 (40%)	421 (30.9%)	<0.0001
Race/ethnicity				0.6949
White	2924 (93.2%)	1653 (93.0%)	1271 (93.4%)	
Black/African American	80 (2.6%)	43 (2.4%)	37 (2.7%)	
Hispanic	28 (0.9%)	19 (1.1%)	9 (0.7%)	
Other	107 (3.4%)	63 (3.5%)	44 (3.2%)	
Insurance				0.0010
Private	2021 (64.6%)	1200 (67.7%)	821 (60.4%)	
Medicare	483 (15.4%)	251 (14.2%)	232 (17.1%)	
Private + Medicare	453 (14.5%)	228 (12.9%)	225 (16.6%)	
Medicaid	152 (4.8%)	81 (4.6%)	71 (5.2%)	
None	22 (0.7%)	12 (0.7%)	10 (0.74%)	
Hypertension	2122 (67.6%)	1102 (62.0%)	1020 (74.9%)	<0.0001
OSA	1000 (31.9%)	505 (28.4%)	495 (36.4%)	<0.0001
Coronary artery disease	593 (18.9%)	295 (16.6%)	298 (21.9%)	0.0002
Diabetes Mellitus	589 (18.8%)	284 (16%)	305 (22.4%)	<0.0001
Heart Failure	571 (18.2%)	201 (11.3%)	370 (27.2%)	<0.0001
LVEF <40%	243 (8.2%)	80 (4.8%)	163 (12.5%)	<0.0001
Thyroid disease	505 (16.1%)	284 (16.0%)	221 (16.2%)	0.8413
On dialysis	6 (0.2%)	3 (0.2%)	3 (0.2%)	0.7425
Stage 3 CKD	623 (24.3%)	308 (22.0%)	315 (27.2%)	0.0023
Cancer	365 (11.6%)	201 (11.3%)	164 (12.1%)	0.5187
COPD	256 (8.2%)	129 (7.3%)	127 (9.3%)	0.0352
CVA/TIA	245 (7.8%)	129 (7.3%)	116 (8.5%)	0.1895
Peripheral vascular disease	95 (3.0%)	37 (2.1%)	58 (4.3%)	0.0004
Family history of AF	82 (2.6%)	39 (2.2%)	43 (3.2%)	0.0927
Left atrial diameter, cm	4.2 (3.7-4.8)	4.0 (3.5-4.5)	4.4 (4.0-5.0)	<0.0001
CHA₂DS₂VASc score	2 (1-4)	2 (1-3)	3 (2-4)	<0.0001
ORBIT bleeding score	1 (0-2)	1 (0-2)	1 (0-2)	0.0003
HAS-BLED Score	2 (1-2)	1 (1-2)	2 (1-2)	<0.0001
Prior major bleeding	94 (3.0%)	44 (2.5%)	50 (3.7%)	0.0151
Prior ablations for AF				0.0002
0	2447 (78.1%)	1432 (80.6%)	1015 (74.8%)	
1	563 (18.0%)	293 (16.5%)	270 (19.9%)	
2	95 (3.0%)	42 (2.4%)	53 (3.9%)	

	Overall	Paroxysmal AF	Persistent AF	P-value
3	29 (0.9%)	10 (0.6%)	19 (1.4%)	

Abbreviations: AF = atrial fibrillation; OSA = obstructive sleep apnea; LVEF = left ventricular ejection fraction; CKD = chronic kidney disease; COPD = chronic obstructive pulmonary disease; CVA/TIA = cerebrovascular accident/transient ischemic attack

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Table 2:
Procedural Characteristics:

Use of peri-procedural anticoagulation (AC), imaging, catheter types and provocation testing as well as procedural times for overall population and by AF subtype. Data presented as N (%) or median (IQR) as appropriate.

	Overall	Paroxysmal AF	Persistent AF	p-value
N	3139	1778 (56.6%)	1361 (43.4%)	
Peri-procedure AC strategy				0.0134
Uninterrupted	2280 (73.6%)	1299 (74.4%)	981 (72.4%)	
Interrupted	688 (22.2%)	389 (22.3%)	299 (22.1%)	
Bridging	132 (4.3%)	58 (3.3%)	74 (5.5%)	
Pre-procedure Imaging				
Preoperative TEE	1535 (49.0%)	762 (43.0%)	773 (56.9%)	<0.0001
CT	1889 (60.3%)	1106 (62.3%)	783 (57.7%)	0.0079
MRI	91 (2.9%)	66 (3.7%)	25 (1.8%)	0.0019
Intra-procedure Imaging				
Intra-cardiac echocardiography	2867 (91.5%)	1628 (91.8%)	1239 (91.2%)	0.5954
3D electroanatomic mapping	2843 (90.8%)	1602 (90.3%)	1241 (91.4%)	0.3008
Intra-operative TEE	436 (13.9%)	248 (14.0%)	188 (13.8%)	0.9134
Rotational angiography	176 (5.6%)	103 (5.8%)	73 (5.4%)	0.6042
Energy and catheter type				
Irrigated RF with contact force	2206 (70.5%)	1222 (69.0%)	984 (72.5%)	0.0335
Irrigated RF without contact force	271 (8.7%)	135 (7.6%)	136 (10.0%)	0.0182
Cryoballoon	743 (23.7%)	468 (26.4%)	275 (20.2%)	<0.0001
Other	53 (1.7%)	29 (1.6%)	24 (1.8%)	0.7788
Provocation testing*				
Burst pacing	1652 (68.5%)	942 (67.9%)	710 (69.4%)	0.4222
Isoproterenol	1148 (47.6%)	626 (45.1%)	552 (51.0%)	0.0040
Adenosine	602 (25.0%)	394 (28.4%)	208 (20.3%)	<0.0001
Other	224 (9.3%)	149 (10.7%)	75 (7.3%)	0.0044
Procedure time (minutes)				
Ablation time	37 (17-59)	35 (17.5-53)	41 (17-67)	<0.0001
Fluoroscopy time	16 (8-28)	15 (7-26)	18 (10-30)	<0.0001
Total procedure time	180 (140-230)	174 (137-218)	190 (146-240)	<0.0001

Abbreviations: AF= atrial fibrillation; AC = anticoagulation; CT = computed tomography; MRI = magnetic resonance imaging; TEE = transesophageal echocardiography; RF = radiofrequency.

* All patients in this cohort received provocation testing of some type.

Table 3:**Complications:**

Number of intra-procedural or immediate post-procedure complications for all patients and by AF subtype.

	Overall	Paroxysmal AF	Persistent AF	p-value
N	3139	1778 (56.6%)	1361 (43.4%)	N
Total cases with complications	160 (5.1%)	69 (3.9%)	91 (6.7%)	0.0004
Volume overload	39 (1.2%)	14 (0.8%)	25 (1.8%)	0.0086
Hematoma	33 (1.1%)	14 (0.8%)	19 (1.4%)	0.0979
Hemopericardium	14 (0.4%)	10 (0.6%)	4 (0.3%)	0.2628
Pericardiocentesis	11 (0.4%)	7 (0.4%)	4 (0.3%)	0.6762
Urinary tract infection	11 (0.4%)	8 (0.4%)	3 (0.2%)	0.2805
Anesthesia complication	9 (0.3%)	4 (0.2%)	5 (0.4%)	0.4602
Tamponade	9 (0.3%)	8 (0.4%)	1 (0.08%)	0.0559
Pseudoaneurysm	8 (0.2%)	3 (0.2%)	5 (0.4%)	0.2744
Hemorrhage requiring transfusion	7 (0.2%)	3 (0.2%)	4 (0.3%)	0.4618
Phrenic nerve injury	6 (0.2%)	1 (0.1%)	5 (0.4%)	0.0481
Retroperitoneal bleed	3 (0.1%)	1 (0.1%)	2 (0.2%)	0.4154
Arteriovenous fistula	3 (0.1%)	0 (0.0%)	3 (0.2%)	0.0477
Aspiration	3 (0.1%)	2 (0.1%)	1 (0.07%)	0.7256
Stroke	3 (0.1%)	2 (0.1%)	1 (0.07%)	0.7256
Transient ischemic attack	1 (0.03%)	1 (0.1%)	0 (0.0%)	0.3815
Perforation or tamponade requiring surgery	1 (0.03%)	1 (0.1%)	0 (0.0%)	0.3815
Death	1 (0.03%)	0 (0.0%)	1 (0.07%)	0.2532
Other	45 (1.4%)	18 (1.0%)	27 (2.0%)	0.0234

Table 4:
Baseline characteristics by complication occurrence:

Demographics and medical history for all included patients and by complication occurrence. Data presented as median (IQR) or N (%) as appropriate

	Overall	No complication	Complication	P-value
N	3138*	2978 (94.9%)	160 (5.1%)	
Age, years	65 (58-71)	65 (58-71)	68 (62-73)	<0.0001
Female	1131 (36.0%)	1053 (35.4%)	78 (48.8%)	0.0006
Hypertension	2121 (67.6%)	1999 (67.1%)	122 (76.2%)	0.0163
OSA	999 (31.8%)	949 (31.9%)	50 (31.3%)	0.8704
Coronary artery disease	593 (18.9%)	555 (18.6%)	38 (23.8%)	0.1076
Diabetes Mellitus	589 (18.8%)	563 (18.9%)	26 (16.2%)	0.4021
Heart Failure	570 (18.2%)	525 (17.6%)	45 (28.1%)	0.0008
LVEF <40%	243 (8.2%)	229 (8.1%)	14 (9.2%)	0.6488
Thyroid disease	505 (16.1%)	473 (15.8%)	32 (20.0%)	0.1675
On dialysis	6 (0.2%)	6 (0.2%)	0 (0.0%)	0.5699
Stage 3 CKD	622 (24.3%)	576 (23.9%)	46 (31.3%)	0.0414
Cancer	364 (11.6%)	343 (11.5%)	21 (13.1%)	0.5363
COPD	256 (8.2%)	233 (7.8%)	23 (14.4%)	0.0032
CVA/TIA	245 (7.8%)	231 (7.8%)	14 (8.8%)	0.6483
Peripheral vascular disease	95 (3.0%)	89 (3.0%)	6 (3.8%)	0.5840
Family history of AF	82 (2.6%)	77 (2.6%)	5 (3.1%)	0.6770
Left atrial diameter, cm	4.2 (3.7-4.8)	4.2 (3.7-4.8)	4.4 (3.9-5.0)	0.0490
CHA₂DS₂VASc score	2 (1-4)	2 (1-4)	3 (2-4)	<0.0001
ORBIT bleeding score	1 (0-2)	1 (0-2)	1 (0-3)	<0.0001
HAS-BLED Score	2 (1-2)	1 (1-2)	2 (1.5-3)	<0.0001
Prior major bleeding	94 (3.0%)	86 (2.9%)	8 (5.0%)	0.1241

* One patient had missing information regarding complications so these analysis are out of 3,138 patients. Abbreviations: OSA = obstructive sleep apnea; LVEF = left ventricular ejection fraction; CKD = chronic kidney disease; COPD = chronic obstructive pulmonary disease; CVA/TIA = cerebrovascular accident/transient ischemic attack; AF = atrial fibrillation

Table 5:
Procedural characteristics by complication occurrence:

Procedural characteristics for patients by complication occurrence. Data presented as median (IQR) or N (%) as appropriate

	Overall	No complication	Complication	P-value
N	3138*	2978 (94.9%)	160 (5.1%)	
Peri-procedure AC strategy				<.0001
Uninterrupted	2279 (73.5%)	2150 (73.1%)	129 (81.1%)	
Interrupted	688 (22.2%)	673 (22.9%)	15 (9.4%)	
Bridging	132 (4.3%)	117 (4.0%)	15 (9.4%)	
Pre-procedure Imaging				
Preoperative TEE	1534 (49.0%)	1438 (48.4%)	96 (60.0%)	0.0043
CT	1888 (60.3%)	1785 (60.1%)	103 (64.4%)	0.2796
MRI	91 (2.9%)	89 (3.0%)	2 (1.2%)	0.2005
Intra-procedure Imaging				
Intra-cardiac echocardiography	2866 (91.5%)	2712 (91.3%)	154 (96.2%)	0.0279
3D electroanatomic mapping	2842 (90.8%)	2690 (90.5%)	152 (95.0%)	0.0578
Intra-operative TEE	435 (13.9%)	407 (13.7%)	28 (17.5%)	0.1758
Rotational angiography	176 (5.6%)	173 (5.8%)	3 (1.9%)	0.0347
Energy and catheter type				
Irrigated RF with contact force	2205 (70.5%)	2076 (69.9%)	129 (80.6%)	0.0038
Irrigated RF without contact force	271 (8.7%)	257 (8.7%)	14 (8.8%)	0.9672
Cryoballoon	743 (23.7%)	725 (24.4%)	18 (11.2%)	0.0001
Other	53 (1.7%)	51 (1.7%)	2 (1.2%)	0.6552
Provocation testing				
Burst pacing	1652 (68.5%)	1563 (68.5%)	89 (69.0%)	0.9110
Isoproterenol	1147 (47.6%)	1063 (46.6%)	84 (65.1%)	<0.0001
Adenosine	602 (25.0%)	573 (25.1%)	29 (22.5%)	0.5005
Other	224 (9.3%)	221 (9.7%)	3 (2.3%)	0.0051
Procedure time (minutes)				
Ablation time	37 (17-59)	38 (18-59)	36 (0-88)	0.9531
Fluoroscopy time	16 (8-28)	16 (8-28)	20 (11-32)	0.0081
Total procedure time	180 (140-230)	179 (140-226)	215 (164-273)	<0.0001
Lesion sets				
PVI	2875 (91.6%)	2736 (91.9%)	139 (86.9%)	0.0262
LA roof line	872 (31.3%)	8510 (30.8%)	62 (40.8%)	0.0097
LA posterior/inferior line	731 (24.3%)	679 (25.8%)	52 (34.2%)	0.0219
Lateral mitral isthmus	170 (6.1%)	156 (5.9%)	14 (9.2%)	0.1003
Septal mitral isthmus	162 (5.8%)	150 (5.7%)	12 (7.9%)	0.2610
LA appendage	122 (4.4%)	110 (4.2%)	12 (7.9%)	0.0296
R sided CTI for flutter	938 (33.7%)	875 (33.2%)	63 (41.4%)	0.0375

	Overall	No complication	Complication	P-value
SVC isolation	137 (5.0%)	126 (4.8%)	11 (7.2%)	0.1747
CFAE	285 (10.2%)	269 (10.2%)	16 (10.5%)	0.9037
FIRM	311 (11.2%)	296 (11.2%)	15 (9.9%)	0.6001
Targeted ganglion ablation	19 (0.7%)	15 (0.6%)	4 (2.6%)	0.0027
Hospital characteristics				
Region				0.0893
West	400 (12.8%)	379 (12.7%)	21 (13.1%)	
South	1273 (40.6%)	1194 (40.1%)	79 (49.4%)	
Midwest	359 (11.4%)	346 (11.6%)	13 (8.1%)	
Northeast	1106 (35.2%)	1059 (35.6%)	47 (29.4%)	
Academic/Teaching Hospital	2648 (85.1%)	2497 (84.6%)	151 (94.4%)	0.0007
Ablation Case Volume	244 (192-404)	244 (192-404)	245 (229-487)	0.0109

* One patient had missing information regarding complications so these analysis are out of 3,138 patients. Abbreviations: AF= atrial fibrillation; AC = anticoagulation; CT = computed tomography; MRI = magnetic resonance imaging; TEE = transesophageal echocardiography; RF = radiofrequency