ORIGINAL RESEARCH

Differences in Healthcare Use Between Patients With Persistent and Paroxysmal Atrial Fibrillation Undergoing Catheter-Based Atrial Fibrillation Ablation: A Population-Based Cohort Study From Ontario, Canada

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BACKGROUND: Patients with persistent atrial fibrillation (AF) undergoing catheter-based AF ablation have lower success rates than those with paroxysmal AF. We compared healthcare use and clinical outcomes between patients according to their AF subtypes.

METHODS AND RESULTS: Consecutive patients undergoing AF ablation were prospectively identified from a population-based registry in Ontario, Canada. Via linkage with administrative databases, we performed a retrospective analysis comparing the following outcomes between patients with persistent and paroxysmal AF: healthcare use (defined as AF-related hospitalizations/emergency room visits), periprocedural complications, and mortality. Multivariable Poisson modeling was performed to compare the rates of AF-related and all-cause hospitalizations/emergency room visits in the year before versus after ablation. Between April 2012 and March 2016, there were 3768 consecutive patients who underwent first-time AF ablation, of whom 1040 (27.6%) had persistent AF. The mean follow-up was 1329 days. Patients with persistent AF had higher risk of AF-related hospitalization/emergency room visits (hazard ratio [HR], 1.21; 95% CI, 1.09–1.34), mortality (HR, 1.74; 95% CI, 1.15–2.63), and periprocedural complications (odds ratio, 1.36; 95% CI, 1.02–1.75) than those with paroxysmal AF. In the overall cohort, there was a 48% reduction in the rate of AF-related hospitalization/emergency room visits in the year after versus before ablation (rate ratio [RR], 0.52; 95% CI, 0.48–0.56). This reduction was observed for patients with paroxysmal (RR, 0.45; 95% CI, 0.41–0.50) and persistent (RR, 0.74; 95% CI, 0.63–0.87) AF.

CONCLUSIONS: Although patients with persistent AF had higher risk of adverse outcomes than those with paroxysmal AF, ablation was associated with a favorable reduction in downstream AF-related healthcare use, irrespective of AF type.

Key Words: ablation atrial fibrillation outcomes research registry

atheter-based atrial fibrillation (AF) ablation is an effective therapy for patients with symptomatic AF. Current AF guidelines strongly endorse ablation, particularly for symptomatic patients with AF in whom medical therapy is not effective.¹⁻³ These recommendations are largely based on studies that

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CLINICAL PERSPECTIVE

What Is New?

- From a prospectively identified, populationbased cohort of consecutive catheter-based atrial fibrillation (AF) ablation in Ontario, Canada, the risk of death, periprocedural complications, and time to first AF-related hospitalization or emergency room visit was higher among patients with persistent AF relative to those with paroxysmal AF.
- Irrespective of AF type (persistent or paroxysmal AF), patients who underwent catheter-based AF ablation had lower rates of AF-related hospitalization or emergency room visit in the year after ablation when compared with the year before ablation.

What Are the Clinical Implications?

- Among patients undergoing catheter-based AF ablation, clinicians should be cognizant that the patient's AF type (persistent or paroxysmal) is an important marker of adverse outcomes after ablation.
- Although the rate of AF-related hospitalization or emergency room visit was high within the first 30 days after ablation, catheter-based AF ablation was associated with an overall reduction in the rates of AF-related healthcare use within the first year.
- Additional research is needed to delineate whether AF ablation is associated with downstream cost savings for healthcare systems, and future research on the efficacy of catheterbased AF ablation should focus on healthcare use as a key outcome.

Nonstandard Abbreviations and Acronyms

- ER emergency room
- HCU healthcare use
- PY person-years

enrolled patients with paroxysmal AF with single procedural success rates of up to 70% to 80%.^{3,4} The success rate of ablation for patients with persistent AF is typically lower, often with the need for repeat ablation.^{4,5}

To date, efficacy measures of AF ablation in clinical trials have focused on freedom from AF recurrence and/or improvement in patients' quality of life.^{6–9} There is a paucity of data on how reductions in AF recurrence may translate to impact healthcare use (HCU), such as downstream hospitalization or emergency room (ER) visits, particularly on a population-based level.^{10–14} Understanding these resource implications is particularly important given the current climate of fiscal constraint. Accordingly, we sought to address this knowledge gap by using data from a population-based registry in Ontario, Canada, to evaluate HCU between patients with persistent and paroxysmal AF who underwent catheter-based ablation. We hypothesize that AF-related HCU would be reduced among patients who undergo ablation. Given that patients with persistent AF are likely to have a greater burden of medical comorbidities, we hypothesize that their postablation outcomes would be less favorable than those with paroxysmal AF.

METHODS

Analytic methods and/or study materials can be made available to researchers to reproduce the results or to replicate this study in other data sets. Patient-level data will not be available because of privacy regulations in Ontario, Canada. Qualified researchers trained in human subject confidentiality protocols may contact Dr Wijeysundera, Dr Verma, and Dr Ha to request access to the analytic methods and study materials.

Design

We conducted a retrospective cohort study from a prospectively collected, population-based registry of patients who underwent AF ablation in Ontario. Ontario is Canada's most populous province, with \approx 14.6 million inhabitants, constituting \approx 38% of the national population. All residents in Ontario receive universal health coverage from a single payer, the Ministry of Health and Long Term Care of Ontario. We adhered to the Strengthening the Reporting of Observational Studies in Epidemiology (STROBE) guidelines for reporting of a cohort study.¹⁵

Data Sources

The study population consisted of consecutive patients undergoing AF ablation in the province of Ontario between April 1, 2012, and March 30, 2016. Patients were prospectively identified, and their demographics were collected by an ablation database within the CorHealth Cardiac Registry. CorHealth is a nonprofit agency funded by the Ministry of Health and Long Term Care, which tracks and collects information on all invasive cardiac procedures performed in Ontario. For each AF ablation procedure, institutions must complete and electronically submit a registry form to CorHealth. This electronic form includes information pertaining to the patient's demographics, cardiac history, AF type (paroxysmal or persistent), and intraprocedural details. Each data field collected by the CorHealth Cardiac Registry is predefined in a data dictionary. The CorHealth registry also uses populated entries to calculate the CHA₂DS₂-VASc score (Congestive heart failure [1 point], Hypertension [1 point], Age \geq 65-74 years [1 point], Age \geq 75 years [2 points], Diabetes mellitus [1 point], Prior stroke, transient ischemic attack, or arterial thromboembolism [2 point], Vascular disease [1 point], or Female sex [1 point]). The registry also calculated the HATCH score (Hypertension [1 point], Age \geq 75 years [1 point], Chronic obstructive pulmonary disease [1 point], or Heart failure [1 point]) to predict AF progression from paroxysmal to persistent.

We then linked patients identified in the AF ablation cohort to health administrative databases to supplement baseline medical comorbidities and to ascertain outcomes based on the International Classification of Diseases, Tenth Revision (ICD-10), coding system. These data sets were linked using unique encoded identifiers and were analyzed at ICES (Toronto, Canada). The following administrative databases were used: Registered Persons Database, Canadian Institute for Health Information Discharge Abstract Database, Canadian Institute for Health Information Same Day Surgery, National Ambulatory Care Reporting System database, Ontario diabetes mellitus database, Ontario hypertension database, and the Ontario chronic obstructive pulmonary disease database. Socioeconomic status was determined on the basis of neighborhood income from 2006 or 2014 Canadian Census data for patients who underwent ablation before or after January 1, 2014, respectively. Prescription data of patients who were aged ≥65 years were obtained from the Ontario Drug Benefits database.

Study Population

The study cohort consisted of patients who underwent catheter-based AF ablation in Ontario, Canada, between April 1, 2012, and March 31, 2016, constituting 4 fiscal years. Patients were excluded if they did not have a valid Ontario Health Insurance Plan number or who were <18 years of age at the time of ablation. For patients who underwent >1 AF ablation during the study period, the first AF ablation was identified as the index event. The patient's AF type (paroxysmal or persistent) was defined according to criteria set forth by current AF clinical guidelines.^{1–3} The end of the study follow-up period was December 31, 2017, or the day of the patient's death, whichever occurred first.

Outcomes

The primary outcome of this study was AF-related HCU, defined as a composite of AF-related

hospitalization or ER visit. AF must be listed as the most responsible diagnosis (main diagnosis code 148) to gualify as an AF-related hospitalization or ER visit. This code had a specificity of 93.0% (95% Cl. 91.6%-94.2%) and a sensitivity of 96.6% (95% CI, 94.1%-98.2%) in the Canadian Institute for Health Information National Ambulatory Care Reporting System database when compared with chart abstraction.¹⁶ Secondary outcomes were (1) overall HCU based on all-cause hospitalization or ER visit, (2) mortality, and (3) periprocedural complications. A periprocedural complication was defined as having any of the following outcomes which resulted in hospitalization or ER visit within 30 days of ablation: bleeding, transfusion, cardiac, respiratory, vascular, neurologic, non-central nervous system thromboembolic, lower extremity, renal, infectious, venous thrombotic, or a medical event requiring surgical or interventional management. These outcomes were identified from the Canadian Institute for Health Information Discharge Abstract Database or National Ambulatory Care Reporting System database. A list of codes used in the identification of periprocedural complications is listed in Tables S1 and S2. All study outcomes were defined using ICD-10 codes.

Exposure

The exposure of interest was AF type (paroxysmal or persistent AF).

Statistical Analysis

Non-normally distributed continuous variables were reported as medians with 25th and 75th percentiles and compared with the Kruskal-Wallis test. Normally distributed continuous variables were reported as means with standard deviations and compared with the Student's t-test. Categorical variables were reported as proportions and were compared using the χ^2 statistic. For the primary outcome analysis of AFrelated HCU, the rate of AF-related hospitalizations and ER visits was reported as cases per 100 person-years (PY) during the year before index ablation and after discharge from ablation. In addition, rates were reported during the following phases of the periablation period: 1 to 30, 31 to 90, and 91 to 365 days before ablation and 1 to 30, 31 to 90, and 91 to 365 days after discharge from ablation. These phases were chosen to examine for potential temporal effects. We used a multivariable generalized estimating equation for Poisson regression modeling to examine the association between AF type (persistent versus paroxysmal) on rates of AF-related hospitalizations or ER visits before and after ablation, using time as an offset term and incorporating an interaction term for AF type×time. The generalized estimating equation model allowed us to account for clustering of outcomes in the preablation and postablation period as well as overdispersion. A similar analytic approach was performed to evaluate all-cause HCU.

The analysis of all other secondary outcomes, except for periprocedural complications, was performed by time-to-event analyses, with the date of index ablation as time 0. Multivariable Cox regression analyses were performed to assess the association of AF type (paroxysmal versus persistent) with study outcomes. We used cause-specific hazard competing risk models to analyze nonfatal outcomes and to account for the competing risk of death. Statistical measures of association were reported as hazard ratios (HRs) with 95% Cls. Periprocedural complications occurring within 30 days after index ablation were classified as binary outcomes (yes or no). Multivariable logistic regression analysis was performed to assess the association between AF type and occurrence of periprocedural complications. For this analysis, statistical measures of association were reported as odds ratios with 95% Cls.

All models were adjusted for the following factors: AF type (paroxysmal versus persistent), age, sex, rural residence, Charlson comorbidity score, CHA₂DS₂-VASc score, diabetes mellitus, hypertension, chronic obstructive pulmonary disease, stroke/ transient ischemic attack/non-central nervous system thromboembolism, history of myocardial infarction, history of cardiovascular disease, history of any vascular disease, history of percutaneous coronary intervention or coronary artery bypass grafting surgery, sleep apnea, income (divided in guintiles), and left ventricular ejection function. For analyses examining the rates of hospitalization or ER visits, statistical measures of significance were reported as rate ratios (RRs) with corresponding 95% CIs. For all outcomes, a 2-sided P<0.05 was considered to be statistically significant. Statistical analyses were performed with SAS 9.4 (SAS Institute, Cary, NC).

Ethics

This study was approved by the institutional review board at Sunnybrook Health Sciences Centre (Toronto, Ontario, Canada). All data pertaining to the project are housed and analyzed by CorHealth and ICES, which are prescribed entities under Ontario's health information privacy legislation (section 45 of

Table.Baseline Characteristics of Patients UndergoingParoxysmal and Persistent AF Ablation

Characteristic	Paroxysmal AF (N=2728)	Persistent AF (N=1040)	P Value
Age, y	59.64±10.39	60.75±10.04	0.003
Age >65 y	846 (31.0)	351 (33.8)	0.107
Men	1821 (66.8)	756 (72.7)	<0.001
Heart failure	492 (18.0)	313 (30.1)	<0.001
Diabetes mellitus	472 (17.3)	186 (17.9)	0.674
Hypertension	1699 (62.3)	681 (65.5)	0.069
COPD	358 (13.1)	172 (16.5)	0.007
Stroke/TIA	59 (2.2)	25 (2.4)	0.654
Non-CNS thromboembolism	17 (0.6)	8 (0.8)	0.622
PVD	20 (0.7)	11 (1.1)	0.324
MI	104 (3.8)	37 (3.6)	0.713
CABG	56 (2.1)	35 (3.4)	0.019
PCI	186 (6.8)	75 (7.2)	0.671
LVEF, %			<0.001
≥50	1571 (57.6)	529 (50.9)	
35–49	116 (4.3)	102 (9.8)	
<35	38 (1.4)	53 (5.1)	
Not recorded	1003 (36.8)	356 (34.2)	
CHA ₂ DS ₂ -VASc score, mean±SD	1.94±1.49	2.12±1.58	0.001
CHA ₂ DS ₂ -VASc score			0.006
0	477 (17.5)	173 (16.6)	
1	725 (26.6)	239 (23.0)	
2	655 (24.0)	230 (22.1)	
3	457 (16.8)	205 (19.7)	
≥4	414 (15.2)	193 (18.6)	
HATCH score, mean±SD	1.28±1.26	1.63±1.46	<0.001
HATCH score			<0.001
0	803 (29.4)	258 (24.8)	
1	1098 (40.2)	344 (33.1)	
2	328 (12.0)	135 (13.0)	
≥3	499 (18.3)	303 (29.1)	
Charlson score, mean±SD	0.31±0.77	0.40±0.88	0.001
Charlson score category		-	
0	2215 (81.2)	786 (75.6)	<0.001
1	292 (10.7)	157 (15.1)	
≥2	221 (8.1)	97 (9.3)	
Rural residence	414 (15.2)	194 (18.7)	0.009
Income quintile			
1 (Lowest)	314 (11.5)	118 (11.3)	0.268
2	419 (15.4)	153 (14.7)	
3	498 (18.3)	200 (19.2)	

Table 1. Continued

Characteristic	Paroxysmal AF (N=2728)	Persistent AF (N=1040)	P Value
4	601 (22.0)	258 (24.8)	
5 (Highest)	896 (32.8)	311 (29.9)	

Values are mean±SD or number (percentage). AF indicates atrial fibrillation; CABG, coronary artery bypass grafting; CHA₂DS₂-VASc, Congestive heart failure (1 point), Hypertension (1 point), Age \geq 65-74 years (1 point), Age \geq 75 years (2 points), Diabetes mellitus (1 point), Prior stroke, TIA, or arterial thromboembolism (2 points), Vascular disease (1 point), or Female sex (1 point); CNS, central nervous system; COPD, chronic obstructive pulmonary disease; HATCH, Hypertension (1 point), Age \geq 75 years (1 point), TIA or stroke (1 point), Chronic obstructive pulmonary disease (1 point), or Heart failure (1 point); LVEF, left ventricular ejection fraction; MI, myocardial infarction; PCI, percutaneous coronary intervention; PVD, peripheral vascular disease; and TIA, transient ischemic attack.

Ontario's Personal Health Information Privacy Act, regulation 329/04, section 18). This permits ICES to receive and use health information without the need for patient-level written consent for the purposes of analysis and compilation of statistical information

about the healthcare system of Ontario, Canada. This information is strictly intended for the purpose of analysis and/or compiling statistical information with respect to the management of, evaluation, or monitoring of the allocation of healthcare resources. No personal identification information is identified in the reporting of results. ICES is fully compliant with the Ontario Health Information Privacy law, with privacy practice approved by the Ontario Privacy Commissioner.

RESULTS

Study Cohort

Between April 1, 2012, and March 30, 2016, there were 4513 patients who underwent AF ablation in 10 Ontario centers. Fifteen (0.3%) patients were excluded because of age <18 years or missing age/sex/income quintile. In addition, 687 (15.2%) patients were excluded as they had previous AF ablation before the start date

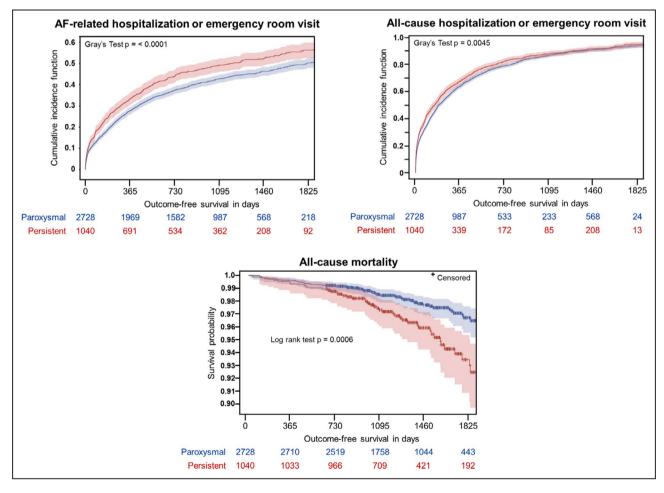


Figure 1. Rates of atrial fibrillation (AF)-related hospitalization or emergency room (ER) visit, all-cause hospitalization or ER visit, and all-cause mortality among patients who underwent catheter-based AF ablation, stratified by AF type (persistent or paroxysmal).

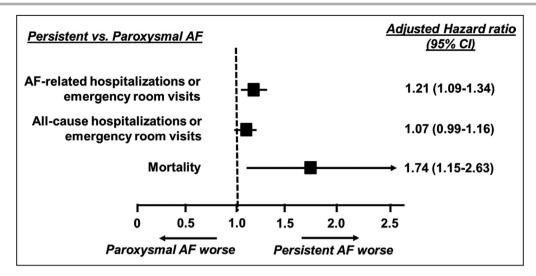


Figure 2. Multivariable regression analysis of outcomes between paroxysmal and persistent atrial fibrillation (AF) patients undergoing catheter-based AF ablation.

of the study cohort. After excluding another 43 (1.1%) patients with missing data on AF type, the final cohort consisted of 3768 patients who underwent de novo AF ablation in Ontario (Figure S1). The mean follow-up period was 1329 days, and 2467 (65.5%) patients had at least 3 years of follow-up.

There were 2728 (72.4%) patients with paroxysmal and 1040 (27.6%) patients with persistent AF in the final ablation cohort. The baseline characteristics of the study cohort are shown in the Table. Compared with patients with paroxysmal AF who underwent ablation, patients with persistent AF were slightly older (60.8±10.0 versus

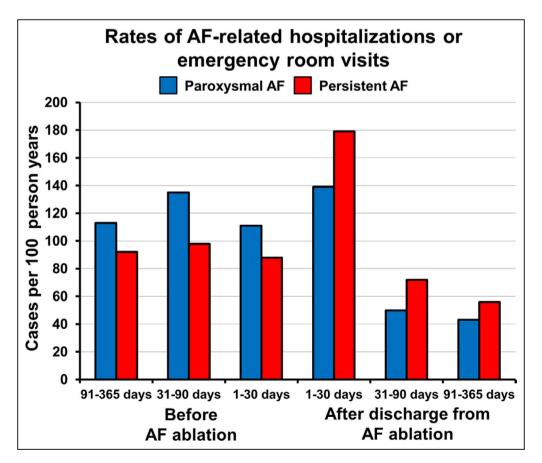


Figure 3. Rates of atrial fibrillation (AF)-related hospitalizations or emergency room visits in the year before AF ablation and in the year after discharge from ablation, according to AF type.

Healthcare Use Outcomes of AF Ablation

59.6±10.4 years; P=0.003), more likely to be men (72.7% versus 66.8%; P<0.001), more likely to reside in rural areas (18.7% versus 15.2%; P=0.009), more likely to have a history of heart failure (30.1% versus 18.0%; P<0.001), and more likely to have chronic obstructive pulmonary disease (16.5% versus 13.1%; P=0.007). In addition, patients with persistent AF had higher CHA₂DS₂-VASc scores (2.1±1.6 versus 1.9±1.5 points; P=0.002), HATCH scores (1.6±1.5 versus 1.3±1.3 points; P<0.001), and Charlson scores (0.40±0.88 versus 0.31±0.77 point; P=0.001) when compared with those with paroxysmal AF. The prevalence of previous stroke or transient ischemic attack was similar between the 2 groups (persistent versus paroxysmal AF, 2.4% versus 2.2%; P=0.65). Among the 1197 patients who were >65 years old, 947 (79.1%) were prescribed with an oral anticoagulant within 90 days before ablation. Of these 947 patients, 702 (74.1%) were treated with a direct oral anticoagulant (apixaban, dabigatran, or rivaroxaban). At 300 to 365 days after ablation, 661 of these patients (69.7%) continued with the same oral anticoagulant agent. Rates of oral anticoagulant continuation at 300 to 365 days after ablation ranged from 66% to 78% (Table S3).

AF-Related HCU

In the overall cohort, 29.1% of patients had at least one hospitalization or ER visit within 1 year after discharge from ablation (Figure 1). Patients with persistent AF had a 21% higher risk of AF-related hospitalization or ER visit relative to those with paroxysmal AF (adjusted HR, 1.21; 95% Cl, 1.09–1.34) (Figure 2). In the year before ablation, the rate of AF-related hospitalizations or ER visits was 109 cases per 100 PY in the overall cohort. During the first year after discharge from ablation, the rate of AFrelated hospitalizations or ER visits was 57 cases per 100 PY in the overall cohort, 68 cases per 100 PY for patients with persistent AF, and 53 cases per 100 PY for patients with paroxysmal AF. The temporal distribution of the rate of AF-related hospitalizations and ER visits in the year before and after index ablation for patients with persistent and paroxysmal AF is shown in Figure 3. In the first year after discharge from ablation, the rate of AF-related hospitalization or ER visits was highest within the first 30 days (148 cases per 100 PY). Subsequently, there was a 64% and 70% reduction in AF-related HCU between 31 to 90 days and 91 to 365 days after ablation (57 and 47 cases per 100 PY) when compared with the

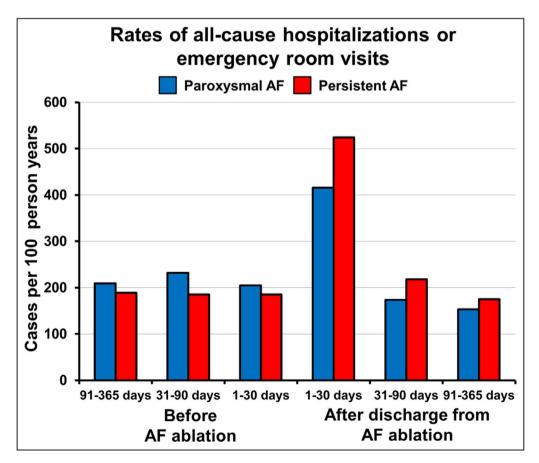


Figure 4. Rates of all-cause hospitalizations or emergency room visits in the year before atrial fibrillation (AF) ablation and in the year after discharge from ablation, according to AF type.

first 30 days after ablation. This pattern was observed for both AF types (Figure 3).

Using multivariable Poisson modeling, patients undergoing AF ablation had a 48% reduction in the rate of AF-related hospitalizations or ER visits within the first year after discharge from ablation when compared with the year before (adjusted RR, 0.52; 95% CI, 0.48–0.56; P<0.0001). The adjusted RR for AF-related hospitalization or ER visit was 0.74 (95% CI, 0.63–0.87; P=0.002) and 0.45 (95% CI, 0.41–0.50; P<0.0001) for patients with persistent and paroxysmal AF, respectively. This suggested that both groups of patients had fewer AF-related hospitalizations or ER visits in the year after ablation relative to the previous year. The magnitude of AF-related HCU reduction was more pronounced among patients with persistent AF ($P_{interaction} < 0.0001$).

All-Cause HCU

The rate of all-cause hospitalization or ER visit within 1 year after discharge from ablation was 54.5% in the overall cohort (Figure 1). The risk of all-cause-related hospitalization or ER visit was similar between patients undergoing persistent and paroxysmal AF ablation (adjusted HR, 1.07; 95% CI, 0.99–1.16; P=0.10) (Figure 2). After adjustment with Poisson modeling, our study cohort had a 9% reduction in the rate of all-cause hospitalizations or ER visits in the year after discharge from ablation when compared with the year before (RR, 0.91; 95% CI, 0.87–0.96; P<0.001). When stratified by AF type, a 16% reduction in the rate of all-cause hospitalizations or ER visits was observed among patients with paroxysmal AF in the year after discharge from ablation when compared with the year before (adjusted RR, 0.84; 95% CI, 0.79-0.89; P<0.001). On the other hand, patients with persistent AF had a 12% increase in the rate of all-cause hospitalizations or ER visits in the year after discharge from ablation when compared with the year prior (adjusted RR, 1.12; 95% Cl, 1.02-1.24; P=0.02). A statistically significant interaction was observed in the adjusted RRs between patients with paroxysmal and persistent AF (P_{interaction}<0.0001). The time course of all-cause HCU after ablation before and after index ablation, stratified by patients with persistent and paroxysmal AF, is shown in Figure 4.

Mortality

The mortality rates of the entire study cohort at 1 year and at the end of study follow-up were 0.45% and 2.60%, respectively (Figure 1). The median age of patients who died was 66 years. Patients undergoing persistent AF ablation were at higher risk of death when compared with those with paroxysmal AF (adjusted HR, 1.74; 95% CI, 1.15–2.63; P=0.008).

Complications

The incidence rates of any periprocedural complication occurring within 7 and 30 days after AF ablation were 6.6% and 7.4%, respectively. Patients with persistent AF had higher odds of periprocedural complications within 30 days after ablation compared with those with paroxysmal AF (odds ratio, 1.36; 95% CI, 1.02–1.75).

The full results of multivariable regression models evaluating AF-related hospitalization/ER visit, mortality, and periprocedural complications were shown in Tables S4 to S6.

DISCUSSION

There are 3 important findings in this contemporary, population-based study of patients undergoing catheter-based AF ablation in Ontario, Canada. First, patients undergoing persistent AF ablation had a greater comorbidity burden along with higher mortality and complication rates when compared with patients with paroxysmal AF. This was reflected by trends toward greater risk of all-cause hospitalization or ER visits among patients with persistent AF. Second, we observed that ablation was associated with a reduction in the rate of AF-related hospitalizations or ER visits in the first year after ablation when compared with the year before ablation. Third, although patients with persistent AF were more likely to have adverse outcomes, our study showed that ablation, irrespective of AF type, was associated with a significant reduction in downstream AF-related HCU.

Healthcare Use

Few studies have addressed whether AF ablation reduces the rates of hospitalizations before and after the procedure. There is a need to define whether this invasive intervention may yield downstream benefits because patients experience high rates of hospitalization or ER visits during the early (<30 days) postablation phase.¹⁰ This is an important consideration because randomized trials have reported mixed results on the impact of AF ablation on death and stroke.^{17,18} Our study showed that AF ablation was associated with a 48% reduction in the rate of AFrelated hospitalizations or ER visits in the year after ablation when compared with the year before. Our findings, derived from a population-based cohort of consecutive patients undergoing AF ablation, were consistent with a recent analysis from the Truven Health MarketScan database in the United States by Guo et al, which reported a 56% reduction of AFrelated hospitalizations after ablation.¹¹

Interestingly, our data on the time course of AFrelated HCU showed the highest use during the first 3 months, followed by a substantial reduction

between 3 and 12 months after ablation. The first 3 months after ablation is considered to be a period during which AF-related hospitalization or ER visits are high because of arrhythmia recurrences, presumably related to the postablation "healing phase." In fact, most clinical trials censor arrhythmic events occurring during the first 3 months (the so-called "blanking period").³ Previous studies reported hospitalization rates of up to 8% in the first 90 days, which were similarly observed in our patient cohort.^{11–14} On the other hand, whether HCU remains elevated at beyond 3 months after ablation is not well defined. Our study showed that hospitalization from 91 to 365 days was 14.5%, which was higher than those reported by other studies.¹¹ However, unlike other study cohorts, which examined subsets of the overall population undergoing AF ablation, our results were based on an entire population-based cohort of patients undergoing AF ablation who were prospectively and consecutively identified. This approach likely yielded less biased estimates when compared with studies that examined only portions of the entire ablation population or restricted their study cohort according to patients' insurance status.

The temporal distribution of AF-related hospitalizations or ER visits in our study showed that the high rate during the first 30 days after ablation was balanced by subsequent reduction of events from 91 to 365 days after ablation, resulting in an overall lowering of event rates in the year after ablation when compared with the year before. This phenomenon was observed regardless of AF type, as evidenced by a reduction of 55% and 26% in the rate of AF-related hospitalizations or ER visits for patients with paroxysmal and persistent AF, respectively. This finding supports the notion that catheter-based ablation for patients with paroxysmal AF is effective in reducing downstream AF-related HCU. Despite the higher risk of adverse outcomes experienced by patients with persistent AF after ablation, their downstream AF-related HCU profile was also favorable.

Among patients with persistent AF, we observed a 12% increase in their rate of all-cause hospitalization or ER visit in the year after ablation when compared with the year before ablation. On the other hand, the proportion of these visits due to AF-related hospitalization or ER visits decreased from 49% to 32% over the same timeframe. Therefore, the increase in HCU after ablation for patients with persistent AF was related to non-AF causes. In this patient subset, the decrease in the rate of AF-related HCU after ablation was offset by a small increase in HCU attributable to non-AF diagnoses. This might be explained by the fact that patients with persistent AF exhibited greater medical comorbidity burden than those with paroxysmal AF, as suggested in the Table.

Clinical Implications

Our findings are pertinent to physicians, patients, and health policy makers because they may assist in planning of HCU around the time of ablation and calculating the cost-benefit ratio for various types of patients with AF. Furthermore, by understanding the time course of HCU after ablation, an important quality improvement focus should consist of interventions to mitigate early hospital readmission or ER use.

Limitations

Our study has several limitations. Although patients were prospectively and consecutively identified, the possibility of missing and/or erroneous categorization could still have occurred in our study. Although a distinguishing aspect of our study was our ability to identify the AF type (persistent versus paroxysmal) of each patient, 2 caveats need to be considered. First, categorization of AF type was dependent on the electrophysiologist who completed the CorHealth AF ablation registry form. Although the delineation of persistent versus paroxysmal AF was based on accepted definitions in consensus documents/guidelines, misclassification could occur, which would introduce bias to our results. Second, among patients with persistent AF, we did not further characterize which of them had long-standing persistent AF. This information might be of interest in terms of providing additional insight on the prognostic significance of patients with this advanced form of persistent AF. Third, our study outcomes were determined by linkage with health administrative databases, making it prone to bias related to diagnostic misclassification and/or inaccuracy because the specificity and sensitivity of ICD-10 codes are not 100%. However, these sources of bias should not preferentially affect one exposure group over another. This limitation is universal in all studies that use population-based administrative databases to ascertain outcomes. Given the use of administrative codes, we could not provide further information on several pertinent aspects, such as details of the type of AF-related admissions or ER visits that occurred (eg, tachycardia related to AF, cardioversion for AF, or heart failure attributable to a primary diagnosis of AF), types of AF ablation strategy used, periprocedural/intraprocedural anticoagulation regimen, information on AF duration before and after ablation, and causes of death. Finally, this study only assessed 1-year outcomes. A longer time horizon may have shown an even greater reduction in HCU, as suggested by previous studies.¹⁹

CONCLUSIONS

Patients with persistent AF undergoing catheter-based AF ablation experienced higher rates of adverse outcomes

and greater HCU relative to those with paroxysmal AF. Regardless of AF type, ablation importantly reduced the rate of AF-related hospitalizations or ER visits in the year after ablation when compared with the year before.

ARTICLE INFORMATION

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Supplementary Material

Tables S1–S6 Figure S1

REFERENCES

- January CT, Wann LS, Alpert JS, Calkins H, Cigarroa JE, Cleveland JC Jr, Conti JB, Ellinor PT, Ezekowitz MD, Field ME, et al. 2014 AHA/ ACC/HRS guideline for the management of patients with atrial fibrillation: a report of the American College of Cardiology/American Heart Association Task Force on practice guidelines and the Heart Rhythm Society. *Circulation*. 2014;130:e199–e267.
- Kirchhof P, Benussi S, Kotecha D, Ahlsson A, Atar D, Casadei B, Castella M, Diener HC, Heidbuchel H, Hendriks J, et al. 2016 ESC guidelines for the management of atrial fibrillation developed in collaboration with EACTS. *Eur Heart J*. 2016;37:2893–2962.
- Calkins H, Hindricks G, Cappato R, Kim YH, Saad EB, Aguinaga L, Akar JG, Badhwar V, Brugada J, Camm J, et al. 2017 HRS/EHRA/ECAS/ APHRS/SOLAECE expert consensus statement on catheter and surgical ablation of atrial fibrillation. *Heart Rhythm.* 2017;14:e275–e444.
- Verma A, Macle L. Persistent atrial fibrillation ablation: where do we go from here? Can J Cardiol. 2018;34:1471–1481.
- 5. Kirchhof P, Calkins H. Catheter ablation in patients with persistent atrial fibrillation. *Eur Heart J.* 2017;38:20–26.
- Piccini JP, Lopes RD, Kong MH, Hasselblad V, Jackson K, Al-Khatib SM. Pulmonary vein isolation for the maintenance of sinus rhythm in patients with atrial fibrillation: a meta-analysis of randomized, controlled trials. *Circ Arrhythm Electrophysiol.* 2009;2:626–633.
- Wynn GJ, Das M, Bonnett LJ, Panikker S, Wong T, Gupta D. Efficacy of catheter ablation for persistent atrial fibrillation: a systematic review and meta-analysis of evidence from randomized and nonrandomized controlled trials. *Circ Arrhythm Electrophysiol*. 2014;7:841–852.
- Mark DB, Anstrom KJ, Sheng S, Piccini JP, Baloch KN, Monahan KH, Daniels MR, Bahnson TD, Poole JE, Rosenberg Y, et al. Effect of catheter ablation vs medical therapy on quality of life among patients with atrial fibrillation: the CABANA randomized clinical trial. *JAMA*. 2019;321:1275–1285.
- Blomström-Lundqvist C, Gizurarson S, Schwieler J, Jensen SM, Bergfeldt L, Kennebäck G, Rubulis A, Malmborg H, Raatikainen P, Lönnerholm S, et al. Effect of catheter ablation vs antiarrhythmic medication on quality of life in patients with atrial fibrillation: the CAPTAF randomized clinical trial. JAMA. 2019;321:1059–1068.
- Shah RU, Freeman JV, Shilane D, Wang PJ, Go AS, Hlatky MA. Procedural complications, rehospitalizations, and repeat procedures after catheter ablation for atrial fibrillation. *J Am Coll Cardiol.* 2012;59:143–149.
- Guo J, Nayak HM, Besser SA, Beaser A, Aziz Z, Broman M, Ozcan C, Tung R, Upadhyay GA. Impact of atrial fibrillation ablation on recurrent hospitalization: a nationwide cohort study. *JACC Clin Electrophysiol*. 2019;5:330–339.
- Noseworthy PA, Kapa S, Haas LR, Van Houten H, Deshmuk AJ, Mulpuru SK, McLeod CJ, Asirvatham SJ, Friedman PA, Shah ND, et al. Trends and predictors of readmission after catheter ablation for atrial fibrillation, 2009–2013. *Am Heart J*. 2015;170:483–489.
- Arora S, Lahewala S, Tripathi B, Mehta V, Kumar V, Chandramohan D, Lemor A, Dave M, Patel N, Patel NV, et al. Causes and predictors of readmission in patients with atrial fibrillation undergoing catheter ablation: a national population-based cohort study. J Am Heart Assoc. 2018;7:e009294. DOI: 10.1161/JAHA.118.009294.
- Freeman JV, Tabada GH, Reynolds K, Sung SH, Liu TI, Gupta N, Go AS. Contemporary procedural complications, hospitalizations, and emergency visits after catheter ablation for atrial fibrillation. *Am J Cardiol.* 2018;121:602–608.
- STROBE statement: STrengthening the Reporting of OBservational studies in Epidemiology. Available at: https://www.strobe-statement. org. Accessed December 31, 2019.
- Atzema CL, Austin PC, Chong AS, Dorian P. Factors associated with 90-day death after emergency department discharge for atrial fibrillation. *Ann Emerg Med.* 2013;61:539–548.
- Marrouche NF, Brachmann J, Andresen D, Siebels J, Boersma L, Jordaens L, Merkely B, Pokushalov E, Sanders P, Proff J, et al. Catheter ablation for atrial fibrillation with heart failure. *N Engl J Med.* 2018;378:417–427.
- Packer DL, Mark DB, Robb RA, Monahan KH, Bahnson TD, Poole JE, Noseworthy PA, Rosenberg YD, Jeffries N, Mitchell LB, et al. Effect of catheter ablation vs antiarrhythmic drug therapy on mortality, stroke, bleeding, and cardiac arrest among patients with atrial fibrillation: the CABANA randomized clinical trial. JAMA. 2019;321:1261–1274.
- Gao L, Moodie M. Modelling the lifetime cost-effectiveness of catheter ablation for atrial fibrillation with heart failure. *BMJ Open*. 2019;9:e031033.

SUPPLEMENTAL MATERIAL

 Table S1. Administrative Codes Employed in the Identification of Periprocedural Complications.

Grouping	ICD-10 or CCI code or OHIP billing code
Bleeding complications	D78.02; D78.22; E36.02; E89.811; G97.32; G97.52; H59.121; H59.122; H59.123; H59.129;
	H59.321; H59.322; H59.323; H59.329; H95.22; H95.42; I97.410; I97.411; I97.418; I97.42;
	I97.610; I97.611; I97.618; I97.620; J95.62; J95.831; K91.62; K91.841; L76.02; L76.22;
	M96.811; M96.831; N99.62; N99.821; D78.12; E36.12; G97.49; H59.229; H95.32; I97.51;
	197.52; J95.72; K91.72; L76.12; M96.821; N99.72; T88.8XXA; K66.1
	I60; I61; I62.0; I62.1; I62.9; K92.0; K92.1; K92.2; I85.0; I98.20; I98.3; K22.10; K22.12;
	K22.14; K22.16; K25.0; K25.2; K25.4; K25.6; K26.0; K26.2; K26.4; K26.6; K27.0; K27.2;
	K27.4; K27.6; K28.0; K28.2; K28.4; K28.6; K29.0; K63.80; K31.80; K55.20; K62.5; K92.2;
	K66.1; R04.0; R04.1; R04.2; R04.8; R04.9; H35.6; H43.1; H45.0; M25.0; M79.81; I32.0;
	N02.0; N02.1; N02.2; N02.3; N02.4; N02.5; N02.6; N02.7; N02.8; N02.9; R31.0; R31.1; R31.9;
	N93.8; N93.9; N95.0; R58; D68.3
Transfusion	1.LZ.19.^^
Vascular complications	T81.718A; T81.72XA; T80.1XXA; S09.0XXS; S15.9XXS; S45.909S; S55.909S; S65.909S;
vascular complications	S75.909S; S85.909S; S95.909S; S25.90XS; S35.90XS; T88.9XXS; I77.0; I77.2; S15.009A;
	S15.309A; S15.209A; S09.0XXA; S15.8XXA; S15.9XXA; S25.00XA; S25.109A; S25.20XA;
	S15.507A; S15.207A; S15.07AAA; S15.07AAA; S15.07AAA; S15.07AAA; S25.007A; S25.107A; S25.207A; S25.207A; S25.309A; S25.409A; S25.509A; S25.90XA; S45.809A; S45.009A; S45.109A; S45.209A;
	S55.109A; S65.109A; S55.009A; S65.009A; S65.209A; S65.309A; S65.409A; S65.509A;
	S45.809A; S75.009A; S75.209A; S85.309A; S85.409A; S85.009A; S85.509A; S85.009A;
	S85.509A; S85.109A; S85.139A; S85.809A; S85.169A; S95.109A; S75.809A; S95.809A;
	1.IA.80; 1.IB.80; 1.IC.80; 1.ID.80; 1.IJ.80; 1.IK.80; 1.IM.80; 1.IN.80; 1.IS.80; 1.JE.80;
	1.JJ.80; 1.JK.80; 1.JL.80; 1.JM.80; 1.JQ.80; 1.JT.80; 1.JU.80; 1.JW.80; 1.JX.80; 1.JY.80;
	1.KG.80;
	1.KR.80; 1.KT.80; 1.KV.80; 1.KX.80; 1.KY.80; 1.KZ.80
Vascular complications	S75.909S; S85.909S; S95.909S; S75.009A; S75.209A; S85.309A; S85.409A; S85.009A;
involving the lower	S85.509A; S85.009A; S85.509A; S85.109A; S85.139A; S85.809A;
extremities	S85.169A; S95.109A; S75.809A; S95.809A

Respiratory complications	J95.851; J95.89; J95.859; J95.88; J95.89; J95.811; J90; J94.2; J94.8; 1.GV.52.^^
Cardiac complications	I97.710; I97.790; I97.88; I97.89; I31.2; I31.4; T82.519A; T82.529A; T82.539A; T82.599A;
	T82.528A; T82.538A; T82.598A; 1.HA.52; I21.x; I22.x; I25.2; I50.x; 1.HM.80; 1.HM.78;
	1.HP.80; 1.HP.78; 1.HZ.80
Neurologic complications	I97.811; I97.821; I60.0; I60.1; I60.2; I60.3; I60.4; I60.5; I60.6; I60.7; I60.9; I61.x; I63.0; I63.1;
	I63.2; I63.4; I63.5; I63.8; I63.9; I64.x; H34.1; G45.0; G45.1; G45.2; G45.3; G45.8; G45.9; I63;
	I65; I66; G97.32; G97.52; G97.49
Genitourinary complications	N99.89
Complications involving	Z341; E650; E682; M134; M137; Z401; R790; R795; R862; R781; R708; R783; R784; R785;
interventional/surgical	R855; R856; R808; R873; R864; R825; R826; R818; R819; R820; Z226; 1.VZ.70; 1.IA.80;
therapy	1.IB.80; 1.IC.80; 1.ID.80; 1.IJ.80; 1.IK.80; 1.IM.80; 1.IN.80; 1.IS.80; 1.JE.80; 1.JJ.80;
	1.JK.80; 1.JL.80; 1.JM.80; 1.JQ.80; 1.JT.80; 1.JU.80; 1.JW.80; 1.JX.80; 1.JY.80; 1.KG.80;
	1.KR.80; 1.KT.80; 1.KV.80; 1.KX.80; 1.KY.80; 1.KZ.80; 1.HA.52; 1.HM.80; 1.HM.78;
	1.HP.80; 1.HP.78; 1.HZ.80; 1.GV.52.^^

CCI indicated Canadian Classification of Health Interventions; OHIP, Ontario Health Insurance Plan; ICD-10 = International Statistical Classification of Diseases and Related Health Problems, 10^{th} revision.

Codes to define peri-procedural complications were derived from a compendium of diagnostic codes from ICD-10, procedure codes from CCI, and physician billing codes from OHIP.

Canadian Classification of Health Interventions, 4th edition, Canadian Institute of Health Information (2015). https://www.cihi.ca/sites/default/files/cci_volume_four_2015_en_0.pdf. Accessed April 25 2018.

Schedule of benefits. Physician services under the Health Insurance Act. Ministry of health and Long Term Care, Ontario (2015). http://www.health.gov.on.ca/en/pro/programs/ohip/sob/physserv/sob_master11062015.pdf. Accessed April 25 2018.

Table S2. Administrative Codes with their Associated Definitions Employed in the Identification of Periprocedural Complications.

Grouping	ICD-10, CCI, or OHIP code	Description
Bleeding complications	D78.02	Intraoperative hemorrhage and hematoma of the spleen complicating other procedure
	D78.22	Postprocedural hemorrhage of the spleen following other procedure
	E36.02	Intraoperative hemorrhage and hematoma of an endocrine system organ or structure complicating other procedure
	E89.811	Postprocedural hemorrhage of an endocrine system organ or structure following other procedure
	G97.32	Intraoperative hemorrhage and hematoma of a nervous system organ or structure complicating other procedure
	G97.52	Postprocedural hemorrhage of a nervous system organ or structure following other procedure
	H59.121	Intraoperative hemorrhage and hematoma of right eye and adnexa complicating other procedure
	H59.122	Intraoperative hemorrhage and hematoma of left eye and adnexa complicating other procedure
	H59.123	Intraoperative hemorrhage and hematoma of eye and adnexa complicating other procedure, bilateral
	H59.129	Intraoperative hemorrhage and hematoma of unspecified eye and adnexa complicating other procedure
	H59.321	Postprocedural hemorrhage of right eye and adnexa following other procedure
	H59.322	Postprocedural hemorrhage of left eye and adnexa following other procedure
	H59.323	Postprocedural hemorrhage of eye and adnexa following other procedure, bilateral
	H59.329	Postprocedural hemorrhage of unspecified eye and adnexa following other procedure
	H95.22	Intraoperative hemorrhage and hematoma of ear and mastoid process complicating other procedure
	H95.42	Postprocedural hemorrhage of ear and mastoid process following other procedure
	I97.410	Intraoperative hemorrhage and hematoma of a circulatory system organ or structure complicating a cardiac catheterization
	I97.411	Intraoperative hemorrhage and hematoma of a circulatory system organ or structure complicating a cardiac bypass

I97.418	Intraoperative hemorrhage and hematoma of a circulatory system organ or structure
I97.42	complicating other circulatory system procedure Intraoperative hemorrhage and hematoma of a circulatory system organ or structure
I97.610	 complicating other procedure Postprocedural hemorrhage of a circulatory system organ or structure following a cardiac catheterization
 I97.611	Postprocedural hemorrhage of a circulatory system organ or structure following cardiac bypass
<u> </u>	Postprocedural hemorrhage of a circulatory system organ or structure following earlie by pass Postprocedural hemorrhage of a circulatory system organ or structure following other
	circulatory system procedure
I97.620	Postprocedural hemorrhage of a circulatory system organ or structure following other procedure
J95.62	Intraoperative hemorrhage and hematoma of a respiratory system organ or structure complicating other procedure
 J95.831	Postprocedural hemorrhage of a respiratory system organ or structure following other procedure
K91.62	Intraoperative hemorrhage and hematoma of a digestive system organ or structure complicating other procedure
K91.841	Postprocedural hemorrhage of a digestive system organ or structure following other procedure
L76.02	Intraoperative hemorrhage and hematoma of skin and subcutaneous tissue complicating other procedure
L76.22	Postprocedural hemorrhage of skin and subcutaneous tissue following other procedure
M96.811	Intraoperative hemorrhage and hematoma of a musculoskeletal structure complicating other procedure
 M96.831	Postprocedural hemorrhage of a musculoskeletal structure following other procedure
N99.62	Intraoperative hemorrhage and hematoma of a genitourinary system organ or structure complicating other procedure
N99.821	Postprocedural hemorrhage of a genitourinary system organ or structure following other procedure
D78.12	Accidental puncture and laceration of the spleen during other procedure
E36.12	Accidental puncture and laceration of an endocrine system organ or structure during other procedure
G97.49	Accidental puncture and laceration of other nervous system organ or structure during other procedure
 H59.229	Accidental puncture and laceration of unspecified eye and adnexa during other procedure
H95.32	Accidental puncture and laceration of the ear and mastoid process during other procedure

	I97.5 1	Accidental puncture and laceration of a circulatory system organ or structure during a
		circulatory system procedure
	I97.52	Accidental puncture and laceration of a circulatory system organ or structure during other
		procedure
	J95.72	Accidental puncture and laceration of a respiratory system organ or structure during other
		procedure
	K91.72	Accidental puncture and laceration of a digestive system organ or structure during other
		procedure
	L76.12	Accidental puncture and laceration of skin and subcutaneous tissue during other procedure
	M96.821	Accidental puncture and laceration of a musculoskeletal structure during other procedure
	N99.72	Accidental puncture and laceration of a genitourinary system organ or structure during other
		procedure
	T88.8XXA	Other specified complications of surgical and medical care, not elsewhere classified, initial
		encounter
	K66.1	Hemoperitoneum
Transfusion	1.LZ.19.^^	Transfusion
Vascular	T81.718A	Complication of other artery following a procedure, not elsewhere classified, initial encounter
complications		
	T81.72XA	Complication of vein following a procedure, not elsewhere classified, initial encounter
	T80.1XXA	Vascular complications following infusion, transfusion and therapeutic injection, initial
		encounter
	S09.0XXS	Injury of blood vessels of head, not elsewhere classified, sequela
	S15.9XXS	Injury of unspecified blood vessel at neck level, sequela
	S45.909S	Unspecified injury of unspecified blood vessel at shoulder and upper arm level, unspecified
		arm, sequela
	S55.909S	Unspecified injury of unspecified blood vessel at forearm level, unspecified arm, sequela
	S65.909S	Unspecified injury of unspecified blood vessel at wrist and hand level of unspecified arm,
		sequela
	S75.909S	Unspecified injury of unspecified blood vessel at hip and thigh level, unspecified leg, sequela
	S85.909S	Unspecified injury of unspecified blood vessel at lower leg level, unspecified leg, sequela
	S95.909S	Unspecified injury of unspecified blood vessel at ankle and foot level, unspecified leg, sequela
	S25.90XS	Unspecified injury of unspecified blood vessel of thorax, sequela
	S35.90XS	Unspecified injury of unspecified blood vessel at abdomen, lower back and pelvis level, sequela

T88.9XXS	Complication of surgical and medical care, unspecified, sequela
177.0	Arteriovenous fistula, acquired
I77.2	Rupture of artery
S15.009A	Unspecified injury of unspecified carotid artery, initial encounter
S15.309A	Unspecified injury of unspecified internal jugular vein, initial encounter
S15.209A	Unspecified injury of unspecified external jugular vein, initial encounter
S09.0XXA	Injury of blood vessels of head, not elsewhere classified, initial encounter
15.8XXA	Injury of other specified blood vessels at neck level, initial encounter
S15.9XXA	Injury of unspecified blood vessel at neck level, initial encounter
S25.00XA	Unspecified injury of thoracic aorta, initial encounter
S25.109A	Unspecified injury of unspecified innominate or subclavian artery, initial encounter
S25.20XA	Unspecified injury of superior vena cava, initial encounter
25.309A	Unspecified injury of unspecified innominate or subclavian vein, initial encounter
S25.409A	Unspecified injury of unspecified pulmonary blood vessels, initial encounter
S25.509A	Unspecified injury of intercostal blood vessels, unspecified side, initial encounter
S25.90XA	Unspecified injury of unspecified blood vessel of thorax, initial encounter
S45.809A	Unspecified injury of other specified blood vessels at shoulder and upper arm level, unspecified
	arm, initial encounter
S45.009A	Unspecified injury of axillary artery, unspecified side, initial encounter
S45.109A	Unspecified injury of brachial artery, unspecified side, initial encounter
S45.209A	Unspecified injury of axillary or brachial vein, unspecified side, initial encounter
S55.109A	Unspecified injury of radial artery at forearm level, unspecified arm, initial encounter
S65.109A	Unspecified injury of radial artery at wrist and hand level of unspecified arm, initial encounter
S55.009A	Unspecified injury of ulnar artery at forearm level, unspecified arm, initial encounter
S65.009A	Unspecified injury of ulnar artery at wrist and hand level of unspecified arm, initial encounter
S65.209A	Unspecified injury of superficial palmar arch of unspecified hand, initial encounter
S65.309A	Unspecified injury of deep palmar arch of unspecified hand, initial encounter
S65.409A	Unspecified injury of blood vessel of unspecified thumb, initial encounter
S65.509A	Unspecified injury of blood vessel of unspecified finger, initial encounter
S45.809A	Unspecified injury of other specified blood vessels at shoulder and upper arm level, unspecified
	arm, initial encounter
S75.009A	Unspecified injury of femoral artery, unspecified leg, initial encounter

S75.209A	Unspecified injury of greater saphenous vein at hip and thigh level, unspecified leg, initial
	encounter
S85.309A	Unspecified injury of greater saphenous vein at lower leg level, unspecified leg, initial
	encounter
S85.409A	Unspecified injury of lesser saphenous vein at lower leg level, unspecified leg, initial encounter
S85.009A	Unspecified injury of popliteal artery, unspecified leg, initial encounter
S85.509A	Unspecified injury of popliteal vein, unspecified leg, initial encounter
S85.009A	Unspecified injury of popliteal artery, unspecified leg, initial encounter
S85.509A	Unspecified injury of popliteal vein, unspecified leg, initial encounter
S85.109A	Unspecified injury of unspecified tibial artery, unspecified leg, initial encounter
S85.139A	Unspecified injury of anterior tibial artery, unspecified leg, initial encounter
S85.809A	Unspecified injury of other blood vessels at lower leg level, unspecified leg, initial encounter
S85.169A	Unspecified injury of posterior tibial artery, unspecified leg, initial encounter
S95.109A	Unspecified injury of plantar artery of unspecified foot, initial encounter
S75.809A	Unspecified injury of other blood vessels at hip and thigh level, unspecified leg, initial
	encounter
S95.809A	Unspecified injury of other blood vessels at ankle and foot level, unspecified leg, initial
	encounter
1.IA.80	Ascending aorta repair
1.IB.80	Aortic arch repair
1.IC.80	Descending aorta repair
1.ID.80	Aortic NEC repair
1.IJ.80	Coronary artery repair
1.IK.80	Coronary vein repair
1.IM.80	Pulmonary artery repair
1.IN.80	Pulmonary vein repair
1.IS.80	Vena caval repair
1.JE.80	Carotid artery repair
1.JJ.80	Brachiocephalic artery repair
1.JK.80	Subclavian artery repair
1.JL.80	Internal mammary artery repair
1.JM.80	Arteries of arm NEC repair

1.JQ.80	Jugular vein repair
1.JT.80	Subclavian vein repair
1.JU.80	Veins of arm NEC repair
1.JW.80	Intracranial vessel repair
1.JX.80	Other vessels of head, neck, or spine NEC, repair
1.JY.80	Thoracic vessels NEC, repair
1.KG.80	Arteries of the leg NEC, repair
1.KR.80	Vein of leg NEC, repair
1.KT.80	Vessels of the pelvis, perineum and gluteal region repair
1.KV.80	Artery NEC, repair
1.KX.80	Vein NEC, repair
1.KY.80	Artery with vein, repair
1.KZ.80	Blood vessels NEC, repair
I60; I61;	Bleeding
I62.0; I62.1;	
I62.9; K92.0;	
K92.1; K92.2;	
185.0; 198.20;	
I98.3; K22.10;	
K22.12;	
K22.14;	
K22.16;	
K25.0; K25.2;	
K25.4; K25.6;	
K26.0; K26.2;	
K26.4; K26.6;	
K27.0; K27.2;	
K27.4; K27.6;	
K28.0; K28.2;	
K28.4; K28.6;	
K29.0;	
K63.80;	
K31.80;	

	IZEE OD	
	K55.20;	
	K62.5; K92.2;	
	K66.1; R04.0;	
	R04.1; R04.2;	
	R04.8; R04.9;	
	H35.6; H43.1;	
	H45.0;	
	M25.0;	
	M79.81;	
	I32.0; N02.0;	
	N02.1; N02.2;	
	N02.3; N02.4;	
	N02.5; N02.6;	
	N02.7; N02.8;	
	N02.9; R31.0;	
	R31.1; R31.9;	
	N93.8; N93.9;	
	N95.0; R58;	
	D68.3	
Vascular	S75.909S	Unspecified injury of unspecified blood vessel at hip and thigh level, unspecified leg, sequela
complications		
involving the		
lower		
extremities		
	S85.909S	Unspecified injury of unspecified blood vessel at lower leg level, unspecified leg, sequela
	S95.909S	Unspecified injury of unspecified blood vessel at ankle and foot level, unspecified leg, sequela
	S75.009A	Unspecified injury of femoral artery, unspecified leg, initial encounter
	S75.209A	Unspecified injury of greater saphenous vein at hip and thigh level, unspecified leg, initial
	0.000	encounter
	S85.309A	Unspecified injury of greater saphenous vein at lower leg level, unspecified leg, initial
	500100711	encounter
	S85.409A	Unspecified injury of lesser saphenous vein at lower leg level, unspecified leg, initial encounter
	S85.009A	Unspecified injury of popliteal artery, unspecified leg, initial encounter
	505.007A	onspective injury of populear attery, unspective leg, initial encounter

	S85.509A	Il second find in instructionalities and an anticipation of the second terms
		Unspecified injury of popliteal vein, unspecified leg, initial encounter
	S85.009A	Unspecified injury of popliteal artery, unspecified leg, initial encounter
	S85.509A	Unspecified injury of popliteal vein, unspecified leg, initial encounter
	S85.109A	Unspecified injury of unspecified tibial artery, unspecified leg, initial encounter
	S85.139A	Unspecified injury of anterior tibial artery, unspecified leg, initial encounter
	S85.809A	Unspecified injury of other blood vessels at lower leg level, unspecified leg, initial encounter
	S85.169A	Unspecified injury of posterior tibial artery, unspecified leg, initial encounter
	S95.109A	Unspecified injury of plantar artery of unspecified foot, initial encounter
	S75.809A	Unspecified injury of other blood vessels at hip and thigh level, unspecified leg, initial encounter
	S95.809A	Unspecified injury of other blood vessels at ankle and foot level, unspecified leg, initial encounter
Respiratory complications	J95.851	Ventilator associated pneumonia
	J95.89	Other postprocedural complications and disorders of respiratory system, not elsewhere classified
	J95.859	Other complication of respirator [ventilator]
	J95.88	Other intraoperative complications of respiratory system, not elsewhere classified
	J95.89	Other postprocedural complications and disorders of respiratory system, not elsewhere classified
	J95.811	Postprocedural pneumothorax
	J90	Pleural effusion, not elsewhere classified
	J94.2	Hemothorax
	J94.8	Other specified pleural conditions
	1.GV.52.^^	Chest tube insertion
Cardiac complications	I97.710	Intraoperative cardiac arrest during cardiac surgery
	I97.790	Other intraoperative cardiac functional disturbances during cardiac surgery
	I97.88	Other intraoperative complications of the circulatory system, not elsewhere classified
	197.89	Other postprocedural complications and disorders of the circulatory system, not elsewhere classified

	I31.2	Hemopericardium, not elsewhere classified		
	I31.4	Cardiac tamponade		
	T82.519A	Breakdown (mechanical) of unspecified cardiac and vascular devices and implants, initial		
		encounter		
	T82.529A	Displacement of unspecified cardiac and vascular devices and implants, initial encounter		
	T82.539A	Leakage of unspecified cardiac and vascular devices and implants, initial encounter		
	T82.599A	Other mechanical complication of unspecified cardiac and vascular devices and implants, initial encounter		
	T82.528A	Displacement of other cardiac and vascular devices and implants, initial encounter		
	T82.538A	Leakage of other cardiac and vascular devices and implants, initial encounter		
	T82.598A	Other mechanical complication of other cardiac and vascular devices and implants, initial encounter		
	1.HA.52	Drainage of the pericardium		
	1.HM.80;1.H	Repair of atrium		
	M.78			
	1.HP.80; 1.HP.78	Repair of ventricle		
	1.HZ.80	Repair of heart NEC		
	I21.x; I22.x;	Myocardial infarction		
	125.2			
	I50.x	Heart failure		
Genitourinary complications	N99.89	Other postprocedural complications and disorders of genitourinary system		
Neurologic complications	197.811	Intraoperative cerebrovascular infarction during other surgery		
•	I97.821	Postoperative cerebrovascular infarction during other surgery		
	G97.32	Intraoperative hemorrhage and hematoma of a nervous system organ or structure complicating other procedure		
	G97.52	Postprocedural hemorrhage of a nervous system organ or structure following other procedure		
	G97.49	Accidental puncture and laceration of other nervous system organ or structure during other procedure		
	I60.0; I60.1;	Stroke, TIA, or intracranial bleeding		
	1			

	160.2; 160.3;	
	I60.4; I60.5;	
	I60.6; I60.7;	
	I60.9; I61.x;	
	I63.0; I63.1;	
	I63.2; I63.4;	
	I63.5; I63.8;	
	I63.9; I64.x;	
	H34.1; G45.0;	
	G45.1; G45.2;	
	G45.3; G45.8;	
	G45.9; I63;	
	165; 166	
Non-CNS	I74; K55.0;	Non-CNS thromboembolic events
thromboembolic	N28.0; G95.1;	
complications	I23.6; I51.3	
-		
Complications	Z341	Closed drainage effusion or pneumothorax
involving		
interventional/s		
urgical therapy		
	E650; E682	Cardiac pump bypass
	M134; M137	Thoracotomy
	Z401	Aspiration of pericardium
	R790	Suture of lacerated major artery
	R795	Repair of lacerated major artery or microscopic repair of digital artery
	R862	Repair of lacerated major artery or microscopic repair of digital artery by bypass or
		interposition graft
	R781	Ligation of artery
	R708	Ligation of internal iliac artery
	R783; R784;	Aorto-iliac repair
	R785	
	R855	Common femoral/profunda femoris repair - as sole procedure
	R862 R781 R708 R783; R784; R785	Repair of lacerated major artery or microscopic repair of digital artery by bypass or interposition graft Ligation of artery Ligation of internal iliac artery Aorto-iliac repair

R856	Extended profundoplasty
R808	Femoral aneurysm - reconstruction or excision with graft
R873	Thrombin injection of femoral artery pseudoaneurysm
R864	Repair of false aneurysm at groin anastomosis
R825	Resection of AV aneurysm or fistula with major graft
R826	Resection of AV aneurysm or fistula with minor graft
R818; R819;	Repair of a lacerated major vein
R820	
Z226	Soft tissue incision and drainage (surgical)
1.VZ.70	Leg incision, Not other specified
1.IA.80	Ascending aorta repair
1.IB.80	Aortic arch repair
1.IC.80	Descending aorta repair
1.ID.80	Aortic NEC repair
1.IJ.80	Coronary artery repair
1.IK.80	Coronary vein repair
1.IM.80	Pulmonary artery repair
1.IN.80	Pulmonary vein repair
1.IS.80	Vena caval repair
1.JE.80	Carotid artery repair
1.JJ.80	Brachiocephalic artery repair
1.JK.80	Subclavian artery repair
1.JL.80	Internal mammary artery repair
1.JM.80	Arteries of arm NEC repair
1.JQ.80	Jugular vein repair
1.JT.80	Subclavian vein repair
1.JU.80	Veins of arm NEC repair
1.JW.80	Intracranial vessel repair
1.JX.80	Other vessels of head, neck, or spine NEC, repair
1.JY.80	Thoracic vessels NEC, repair

1.KG.80	Arteries of the leg NEC, repair
1.KR.80	Vein of leg NEC, repair
1.KT.80	Vessels of the pelvis, perineum and gluteal region repair
1.KV.80	Artery NEC, repair
1.KX.80	Vein NEC, repair
1.KY.80	Artery with vein, repair
1.KZ.80	Blood vessels NEC, repair
1.HA.52	Drainage of the pericardium
1.HM.80;	Repair of atrium
1.HM.78	
1.HP.80;	Repair of ventricle
1.HP.78	
1.HZ.80	Repair of heart NEC
1.GV.52.^^	Chest tube insertion

CCI indicated Canadian Classification of Health Interventions; OHIP, Ontario Health Insurance Plan; ICD-10 = InternationalStatistical Classification of Diseases and Related Health Problems, 10^{th} revision.

Codes to define peri-procedural complications were derived from a compendium of diagnostic codes from ICD-10, procedure codes from CCI, and physician billing codes from OHIP.

Canadian Classification of Health Interventions, 4th edition, Canadian Institute of Health Information (2015). https://www.cihi.ca/sites/default/files/cci_volume_four_2015_en_0.pdf. Accessed April 25 2018.

Schedule of benefits. Physician services under the Health Insurance Act. Ministry of health and Long Term Care, Ontario (2015). http://www.health.gov.on.ca/en/pro/programs/ohip/sob/physserv/sob_master11062015.pdf. Accessed April 25 2018.

	Percentage of patients who filled a prescription for the same oral anticoagulant at 300-365 days after discharge from ablation (%)
Apixaban prescribed within 90 days from ablation (n=134)	78.3%
Dabigatran prescribed within 90 days from ablation (n=274)	66.4%
Rivaroxaban prescribed within 90 days from ablation (n=294)	69.7%
Warfarin prescribed within 90 days from ablation (n=245)	66.1%

Table S3. Oral anticoagulation use after AF ablation for Ontario patients who were >65 years old (2012-16).

Covariate	Hazard ratio	95% Confidence interval	P-value
Persistent AF	1.74	1.15-2.63	0.01
(vs. paroxysmal AF)			
Age (per year increase)	1.03	1.00-1.06	0.046
Charlson score	1.40	1.20-1.64	< 0.001
(per point increase)			
CHA2DS2-VaSc score	1.12	0.87-1.46	0.38
(per point increase)			
Income quintile			
1 (lowest)	Referent	-	-
2	0.50	0.24-1.05	0.07
3	0.84	0.44-1.60	0.60
4	0.85	0.45-1.61	0.62
5 (highest)	0.51	0.27-0.98	0.04
Female	0.69	0.41-1.17	0.17
Rural residence	1.19	0.73-1.95	0.49
Diabetes	1.45	0.85-2.48	0.17
Hypertension	1.21	0.65-2.23	0.55
COPD	1.72	1.08-2.74	0.02
Stroke/TIA/Non-CNS thromboembolism	1.08	0.40-2.93	0.87
MI	1.40	0.60-3.27	0.44
PCI or CABG	0.86	0.36-2.06	0.73
LVEF			
≥50%	Referent	-	-
35-49%	1.42	0.68-2.93	0.35
<35%	0.60	0.18-2.05	0.42
Not reported	0.94	0.60-1.48	0.80

 Table S4. Multivariable Analysis of Factors Associated with All-Cause Mortality after AF Ablation.

Covariate	Hazard ratio	95% Confidence interval	P-value
Persistent AF	1.21	1.09-1.34	< 0.001
(vs. paroxysmal AF)			
Age (per year increase)	1.00	0.99-1.01	0.67
Charlson score	1.03	0.96-1.10	0.48
(per point increase)			
CHA2DS2-VaSc score	1.08	1.01-1.16	0.02
(per point increase)			
Income quintile			
1 (lowest)	Referent	-	
2	0.92	0.77-1.01	0.36
3	0.91	0.76-1.08	0.26
4	0.88	0.74-1.04	0.14
5 (highest)	1.00	0.85-1.17	0.96
Female	1.03	0.91-1.16	0.68
Rural residence	1.00	0.88-1.14	0.97
Diabetes	0.81	0.70-0.95	0.007
Hypertension	0.98	0.86-1.11	0.75
COPD	1.09	0.96-1.25	0.20
Stroke/TIA/Non-CNS thromboembolism	0.71	0.50-1.01	0.06
MI	0.99	0.74-1.34	0.96
PCI or CABG	0.78	0.56-1.05	0.11
LVEF			
≥50%	Referent	-	
35-49%	1.00	0.81-1.23	0.99
<35%	0.78	0.56-1.09	0.15
Not reported	1.03	0.93-1.14	0.53

 Table S5. Multivariable Analysis of Factors Associated with AF-Related Hospitalization or ER Visit.

Covariate	Odds ratio	95% Confidence interval	P-value
Persistent AF	1.34	1.02-1.75	0.04
(vs. paroxysmal AF)			
Age (per year increase)	1.00	0.98-1.02	0.99
Charlson score	1.05	0.96-1.24	0.53
(per point increase)			
CHA2DS2-VaSc score	1.31	1.05-1.63	0.02
(per point increase)			
Income quintile			
1 (lowest)	1.66	1.11-2.48	0.03
2	1.35	0.92-1.98	0.42
3	1.06	0.73-1.54	0.31
4	1.10	0.77-1.57	0.44
5 (highest)	Referent	-	-
Female	1.18	0.84-1.67	0.34
Rural residence	0.80	0.56-1.14	0.22
Diabetes	1.67	0.45-1.00	0.05
Hypertension	1.08	0.74-1.58	0.68
COPD	0.90	0.63-1.28	0.55
Stroke/TIA/Non-CNS thromboembolism	0.63	0.33-1.20	0.16
MI	1.36	0.68-2.70	0.38
PCI or CABG	1.59	0.72-3.52	0.25
LVEF			
≥50%	1.05	0.80-1.38	0.35
35-49%	1.35	0.82-2.22	0.49
<35%	1.37	0.68-2.77	0.56
Not reported	Referent	-	

 Table S6. Multivariable Analysis of Factors Associated with Peri-Procedural Complications within 30 Days of Ablation.

Figure S1. Flow Diagram of Patients Included in the Study.

