

# Characterizing atrial fibrillation symptom improvement following *de novo* catheter ablation

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Aims	Atrial fibrillation (AF) symptom relief is a primary indication for catheter ablation, but AF symptom resolution is not well char- acterized. The study objective was to describe AF symptom documentation in electronic health records (EHRs) pre- and post- ablation and identify correlates of post-ablation symptoms.
Methods and results	We conducted a retrospective cohort study using EHRs of patients with AF ( $n = 1293$ ), undergoing ablation in a large, urban health system from 2010 to 2020. We extracted symptom data from clinical notes using a natural language processing algo- rithm ( <i>F</i> score: 0.81). We used Cochran's <i>Q</i> tests with <i>post-hoc</i> McNemar's tests to determine differences in symptom preva- lence pre- and post-ablation. We used logistic regression models to estimate the adjusted odds of symptom resolution by personal or clinical characteristics at 6 and 12 months post-ablation. In fully adjusted models, at 12 months post-ablation pa- tients, patients with heart failure had significantly lower odds of dyspnoea resolution [odds ratio (OR) 0.38, 95% confidence interval (Cl) 0.25–0.57], oedema resolution (OR 0.37, 95% Cl 0.25–0.56), and fatigue resolution (OR 0.54, 95% Cl 0.34–0.85), but higher odds of palpitations resolution (OR 1.90, 95% Cl 1.25–2.89) compared with those without heart failure. Age 65 and older, female sex, Black or African American race, smoking history, and antiarrhythmic use were also associated with lower odds of resolution of specific symptoms at 6 and 12 months.
Conclusion	The post-ablation symptom patterns are heterogeneous. Findings warrant confirmation with larger, more representative data sets, which may be informative for patients whose primary goal for undergoing an ablation is symptom relief.

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### **Graphical Abstract**



Keywords

Atrial fibrillation • Catheter ablation • Natural language processing • Signs and symptoms

#### **Novelty**

- In this study of nearly 1300 electronic health records of patients with atrial fibrillation undergoing de novo catheter ablation, the majority of patients continue to experience symptoms post-ablation.
- There is significant variability in the specific symptoms that resolve by personal and clinical characteristics with patients who are 65 and older, female sex, Black or African American race, smoking history, antiarrhythmic use, and comorbid heart failure having lower odds of resolution of specific symptoms at 6 and 12 months post-ablation.
- Robust natural language processing methods for extracting the symptom information from electronic health records may add important evidence in cases when patient-reported outcomes data are sparse or only available in small samples.

## Introduction

Atrial fibrillation (AF) is the most common cardiac arrhythmia.<sup>1</sup> Adults living with AF report a variety of physical and psychological symptoms, including dyspnoea, chest pain, fatigue, anxiety, and palpitations, which limit daily functioning and impact health-related quality of life (HRQoL).<sup>2</sup> Catheter ablation is a minimally invasive, percutaneous procedure for symptomatic patients with AF when symptoms and/or heart rhythm are not well controlled by medications, or medications cause intolerable side effects.<sup>3</sup>

Symptom relief is one of the primary indications for performing an ablation<sup>4</sup> and is considered by patients to be more important than rhythm control.<sup>5</sup> However, the impact of ablation on AF symptoms has not been well described. To date, most data about AF symptoms

post-ablation originate from clinical trials whose primary endpoints are medically focused (e.g. mortality, stroke incidence), with HRQoL measured as a secondary endpoint.<sup>6–8</sup> The diverse range of symptoms that patients with AF experience, including mental health symptoms like anxiety, is generally not measured.<sup>9</sup> Moreover, differences in trajectories of symptom relief post-ablation, or in individual symptom burden based on personal and clinical characteristics, have not been well described.<sup>6,7</sup>

Because symptoms are a pre-eminent indication for undergoing ablation, healthcare professionals commonly document AF symptoms before and after ablation in clinical notes in electronic health records (EHRs). Secondary data reuse from EHRs may be an instrument for understanding symptom patterns following catheter ablation. Information stored in encounter notes (e.g. 'Patient presents with fatigue') can be parsed into quantitative data (e.g. 'fatigue = 1'), using techniques such as natural language processing (NLP). Natural language processing is a suite of automated methods used to organize and evaluate the information contained in unstructured clinical notes. Natural language processing is increasingly being used to conduct cardiovascular research and to evaluate patient symptoms.<sup>10,11</sup> To improve understanding of postablation symptom patterns using real-world, routinely collected clinical data, we aimed to extract and describe AF symptom documentation in EHRs before and after *de novo* catheter ablation for AF, and to identify personal and clinical characteristics associated with post-ablation symptom.

## **Methods**

### Study design

We conducted a retrospective cohort study of patients with AF undergoing catheter ablation using data from EHRs at a large, urban academic medical centre in New York City. Data from EHRs exist in structured fields (e.g. patient demographics, comorbidities, encounters, procedures, medications, vital signs, and laboratory results) and unstructured formats (e.g. encounter notes). Both structured and unstructured data were used in this analysis. A description of the steps undertaken to query, clean, and aggregate EHR data in this study is provided in *Figure 1* and described in detail below. This study was approved by the Weill Cornell Medicine Institutional Review Board. The investigation conforms with the principles outlined in the Declaration of Helsinki.

### **Cohort creation**

Electronic health record data at our institution adhere to the Observational Medical Outcomes Partnership (OMOP) Common Data Model, a standardized set of tables and variables that are commonly used for storing and aggregating EHR data across sites.<sup>12,13</sup> The clinical research informatics team at our institution created an OMOP instance, containing structured and unstructured EHR data for patients with a primary diagnosis of paroxysmal AF identified using international classification of diseases (ICD)-9 and ICD-10 codes for at least one visit at our institution between 1 January 2010 and 31 December 2020. From this instance, we created a cohort of patients based on the following eligibility criteria: (i) underwent *de novo* catheter ablation for the treatment of AF, determined using current procedural terminology codes, (ii) age 18 or older, and (iii) at least one encounter note available within 30 days of the date of the ablation.

### Structured data

We extracted demographics, comorbid diagnoses, and medications for patients in the cohort from structured fields. Demographic information included gender, race, ethnicity, and age at the time of the ablation calculated using the date of birth and the date of ablation. We used ICD-9 and ICD-10 billing codes to identify patients diagnosed with comorbid heart failure (systolic or diastolic), hypertension, stroke, transient ischaemic attack. thromboembolism, vascular disease, and diabetes. Using these variables together with age and gender, we calculated the CHA<sub>2</sub>DS<sub>2</sub>-VASC (congestive heart failure, hypertension, age, diabetes mellitus, prior stroke or TIA or thromboembolism, vascular disease, age, sex category) score, a measure of stroke risk in AF, using established scoring criteria.<sup>14</sup> Α CHA<sub>2</sub>DS<sub>2</sub>-VASC score of two or higher is considered a moderate-to-high stroke risk and an indication to start anticoagulation. We summarized CHA<sub>2</sub>DS<sub>2</sub>-VASC as both a continuous variable and categorical (score of  $^{3}2$  vs. <2). We used smoking and alcohol use documentation to create a binary variable characterizing patients as ever having been smokers or ever having admitted to alcohol use (smoker vs. non-smoker, denies vs. admits alcohol use). We did not constrain billing codes, smoking status, or alcohol status to specific dates because of documentation idiosyncrasies based on visit type.<sup>15,10</sup>

We characterized inpatient or outpatient medications according to three categories relevant to catheter ablation for AF: (i) antiarrhythmic agents including amiodarone, dofetilide, dronedarone, flecainide, and propafenone, (ii) rate control medications including atenolol, diltiazem, metoprolol, atenolol, and verapamil, and (iii) anticoagulant medications including apixaban, dabigatran, edoxaban, rivaroxaban, and warfarin. Medication records included prescription refills, single medication administrations (primarily in inpatient visits), and the medication list that is manually maintained by clinicians. Only oral medications were included because they are more likely to reflect a patient's outpatient regimen compared with injections, which are typically administered during inpatient encounters or for short periods of time. We used the start dates and, when available, end dates of each medication record to confirm the patient was taking the medication at specific times of interest during the study period (e.g. pre-ablation, 3, 6, 9, and 12 months post-ablation). Approximately, half of all medication records in our data set had start dates but no end dates. These medication records were included if the start date was within 1 year prior to the ablation year.

As a social determinant of health measure, we determined the social deprivation index (SDI) for each patient using zip codes of home addresses from the demographics table. Social deprivation index is a composite measure of neighbourhood poverty created from seven demographic variables collected from the American Community Survey.<sup>17</sup> The variables include the per cent of individuals in a neighbourhood living in poverty, <12 years of education, single-parent households, living in a rented housing unit, living in an overcrowded housing unit, households without a car, and non-employed adults under 65 years of age. Social deprivation index is calculated at the census tract level and ranges from 1 (least disadvantaged) to 100 (most disadvantaged). In New York City, where this study was conducted and where poverty varies dramatically within relatively small geographical areas, SDI is a more sensitive indicator of socioeconomic status



Figure 1 Diagram of electronic health record (EHR) data extraction for atrial fibrillation patients at our institution.

than county- or state-level measures.<sup>18</sup> For this study, we accessed data sets containing SDI scores for every census tract in the USA, which are made freely available by the creators of the index.<sup>17</sup> We determined the SDI of each patient's community by mapping the zip code with census tracts and subsequently with SDI.

### Unstructured data

We undertook a multistep process to extract symptom information from clinical notes using NLP. We extracted 10 AF symptoms determined through consultation with cardiologists, registered nurses with cardiology expertise, and the literature: anxiety, chest pain, dizziness, dyspnoea, lower extremity oedema, fatigue, malaise, palpitations, syncope, and weakness. We used the NLP software, NimbleMiner,<sup>19</sup> and closely followed the methods outlined in prior NLP-based symptom science research conducted with this tool.<sup>20</sup> The NLP model performance on our corpus of notes was as follows: precision = 0.732, recall = 0.926, *F* score = 0.807. Details regarding our NLP methods are provided in Supplementary material.

After applying the NLP model to extract AF symptoms, we grouped symptoms based on the timeframe that the notes were published (*Figure 2*). This generated a data set indicating whether each symptom was present or absent in the notes at each time point for each patient: preablation, 3, 6, 9, and 12 months post-ablation. If multiple notes were available from a time point, all were analysed, and any mention of a symptom was retained. For example, if two notes were available during the 3-month time point, one describing palpitations and the other not, palpitations were considered endorsed at that time point.

### Statistical analysis

Data from structured fields and unstructured fields (extracted using NLP) were merged into a single analytic data set. In addition to evaluating each symptom separately, we created a composite variable reflecting whether the patient had any of the 10 symptoms at each time point. We conducted basic descriptive statistics to describe the demographic and clinical characteristics of the sample as well as symptom prevalence at each time point. We used Cochran's Q tests with post-hoc McNemar's tests to determine whether the prevalence of symptoms differed significantly across time points. Significance was determined using the Benjamini-Hochberg procedure for multiple comparisons correction. Only patients contributing data at each time point (n = 207) were included in the statistical comparisons of prevalence to meet the assumption of the Cochran's Q test that the sample sizes are equal over time. We then used bivariate logistic regression models to estimate the unadjusted odds of symptom prevalence by a range of personal or clinical characteristics at 6 and 12 months post-ablation. We chose these time points because they clear the 3-month post-ablation 'blanking period', and align with endpoints previously reported in clinical trials.<sup>8,21</sup> All patients contributing data at 6 months (n = 651) and 12 months (n = 523) were included in the respective models.

Finally, we conducted an analysis to evaluate the randomness of missing notes among all patients undergoing ablation at our institution. We compared the age, gender, race, ethnicity, and SDI between all patients undergoing ablation, those with pre-ablation notes available (and thus included in our analytic data set), and those with post-ablation notes available, using  $\chi^2$  and *t*-tests for categorical and continuous data, respectively.

## Results

## **Description of the sample**

A total of 1293 patients were included in this analysis. The exclusion cascade is shown in *Figure 3*. The demographic and clinical characteristics of the sample are described in *Table 1*. The mean age of the sample was 65.5 (SD 12.6) years, approximately one-third were female, and two-thirds were White. Approximately 22% lived in a neighbourhood that is considered socially deprived.<sup>17</sup> Nearly half were prescribed an antiarrhythmic medication and three-quarters were prescribed a rate control medication pre-ablation. Nearly half (45%) of the sample had a comorbid diagnosis of heart failure.

# Patterns of symptom prevalence over time

Patterns of symptom prevalence pre-ablation and 3, 6, 9, and 12 months post-ablation are displayed in *Table 2*, and pairwise comparisons are in Supplementary material online, *Table S1*. Almost all (96%) patients had documentation of any AF symptoms pre-ablation, while the remainder were asymptomatic. The most prevalent symptoms documented pre-ablation were dyspnoea (64%), oedema (62%), palpitations (57%), and fatigue (49%). Most patients continued to have documented symptoms at each time point post-ablation (91–95%). There was a significant change in the proportion of patients with documented anxiety from 6 months (41%) to 12 months (36%). There was a significant change in the proportion of patients with documented weakness









from pre-ablation (15%) to all time points post-ablation (20–22%). There were no other significant differences in symptom prevalence across time points.

#### **Table 1** Patient characteristics (n = 1293)

Age (65 and older)	738 (57.1)
Gender (female)	455 (35.2)
Race	
Asian	59 (4.6)
Black or African American	68 (5.3)
Not reported	303 (23.4)
Other race <sup>a</sup>	113 (8.7)
White	750 (58.0)
Ethnicity (Hispanic/Latino)	55 (4.3)
Living in socially deprived neighbourhood <sup>b</sup>	278 (21.5)
Prescribed antiarrhythmic medication	577 (44.6)
Prescribed rate control medication	981 (75.9)
Prescribed anticoagulant medication	921 (71.2)
Alcohol use history	710 (54.9)
Smoking history	324 (25.1)
$CHA_2DS_2$ -VASC score $\geq 2^c$	983 (76.0)
Comorbid heart failure	575 (44.5)

The values are given as n (%).

<sup>a</sup>Includes a small number of American Indian or Alaska Nation (n = 2) and Native Hawaiian or other Pacific Islander (n = 3).

<sup>b</sup>Measured using social deprivation index (SDI) which is a composite measure used to quantify the socioeconomic variation in health outcomes. The patient's zip code was used to correlate to an area-level deprivation calculated based on seven demographic characteristics collected in the American Community Survey. An SDI score above 90 is considered a socially deprived area.<sup>17</sup>

 $^c\text{CHA}_2\text{DS}_2\text{-VASC}$  is a measure of stroke risk in AF. A score  $\geq\!2$  is considered a moderate-to-high stroke risk and an indication to start anticoagulation.  $^{14}$ 

# Associations between patient characteristics and symptom resolution

At 6 months post-ablation, in fully adjusted models, patients 65 and older had significantly lower odds of palpitations resolution [odds ratio (OR) 0.58, 95% confidence interval (CI) 0.40-0.85] compared with patients under 65 (Table 3). Females had significantly lower odds of anxiety resolution (OR 0.46, 95% CI 0.31-0.69), dizziness resolution (OR 0.65, 95% CI 0.45–0.96), and palpitations resolution (OR 0.42, 95% CI 0.29–0.61) compared with males. Asian patients had significantly higher odds of anxiety resolution (OR 4.70, 95% Cl 1.36-29.60) compared with White patients. Black or African American patients had significantly lower odds of malaise resolution (OR 0.17, 95% CI 0.04-0.91) compared with White patients. Patients living in socially deprived neighbourhoods had significantly higher odds of chest pain resolution (OR 1.73, 95% CI 1.06-2.92) compared with those not living in socialdeprived neighbourhoods. Patients with an active antiarrhythmic prescription at 6 months had significantly lower odds of dizziness resolution (OR 0.61, 95% CI 0.41–0.93) and syncope resolution (OR 0.55, 95% CI 0.34–0.91) compared with those with no prescription. Patients with a history of alcohol use had significantly higher odds of weakness resolution (OR 1.87, 95% Cl 1.01–3.46) compared with those with no history. Patients with a smoking history had significantly lower odds of oedema resolution (OR 0.60, 95% CI 0.42–0.87) compared with those with no history. Patients with comorbid heart failure had significantly lower odds of chest pain resolution (OR 0.51, 95% CI 0.33-0.78), fatigue resolution (OR 0.46, 95% CI 0.30-0.69), dyspnoea resolution (OR 0.41, 95% CI 0.28-0.59), and oedema resolution (OR 0.33, 95% Cl 0.23–0.48), but significantly higher odds of palpitations resolution (OR 1.90, 95% CI 1.30-2.81), compared with those without heart failure.

At 12 months post-ablation, in fully adjusted models, patients 65 and older had significantly lower odds of palpitations resolution (OR 0.60, 95% CI 0.39–0.91) compared with those under 65 (*Table 4*). Females had significantly lower odds of anxiety resolution (OR 0.53, 95% CI 0.32–0.86) and palpitations resolution (OR 0.42, 95% CI 0.28–0.63) compared with males. Black or African American patients had significantly lower odds of syncope resolution (OR 0.30, 95% CI 0.12–0.84) compared with White patients. Patients with an active anti-arrhythmic prescription at 12 months had significantly lower odds of

#### Table 2 Symptom prevalence by time point among patients undergoing catheter ablation<sup>a</sup>

	Pre-ablation, <i>n</i> = 1293, <i>n</i> (%)	3 months, <i>n</i> = 822, <i>n</i> (%)	6 months, <i>n</i> = 651, <i>n</i> (%)	9 months, <i>n</i> = 471, <i>n</i> (%)	12 months, <i>n</i> = 523, <i>n</i> (%)	P-value <sup>b</sup>
Any symptoms	1246 (96)	769 (94)	592 (91)	449 (95)	495 (95)	0.47
Anxiety	444 (34)	312 (38)	264 (41)	204 (43)	190 (36)	0.009
Chest pain	551 (43)	351 (43)	278 (43)	220 (47)	218 (42)	0.29
Dizziness	569 (44)	346 (42)	276 (42)	198 (42)	211 (40)	0.75
Fatigue	637 (49)	386 (47)	291 (45)	223 (47)	243 (46)	0.36
Malaise	104 (8)	77 (9)	63 (10)	46 (10)	55 (11)	0.12
Palpitations	736 (57)	453 (55)	358 (55)	245 (52)	294 (56)	0.89
Dyspnoea	823 (64)	474 (58)	379 (58)	301 (64)	300 (57)	0.74
Oedema	796 (62)	473 (58)	355 (55)	290 (62)	299 (57)	0.99
Syncope	332 (26)	226 (28)	202 (31)	145 (31)	146 (28)	0.06
Weakness	194 (15)	182 (22)	132 (20)	103 (22)	111 (21)	0.006

Bold values indicate statistical significance at P < 0.05 and correspond to footnote 'b'.

<sup>a</sup>The prevalence is calculated as the proportion of patients with symptoms among those contributing data at each time point.

<sup>b</sup>Significant at P < 0.05. Significance is determined using Cochran's Q tests.

Characteristic	Any symptoms	Anxiety	Chest pain	Dizziness	Dyspnoea	Oedema
Age (ref: under 65, <i>n</i> = 255)	1.16 (0.63–2.09)	0.95 (0.61–1.47)	0.96 (0.63–1.46)	1.16 (0.78–1.75)	0.86 (0.60–1.25)	1.00 (0.69–1.47)
Gender (ref: Male, $n = 408$ )	0.78 (0.41–1.42)	0.46 (0.31–0.69)	0.71 (0.48–1.06)	0.65 (0.45–0.96)	0.82 (0.57–1.17)	1.00 (0.69–1.45)
Race (ref: White, $n = 393$ )						
Asian $(n = 34)$	0.92 (0.21–2.90)	4.70 (1.36–29.60)	0.75 (0.33–1.87)	0.65 (0.30–1.55)	0.87 (0.41–1.91)	0.52 (0.23–1.16)
Black/African American ( $n = 40$ )	0.71 (0.16–2.24)	1.05 (0.49–2.41)	0.74 (0.36–1.61)	0.60 (0.30–1.24)	1.10 (0.53–2.33)	0.48 (0.23–1.00)
Ethnicity (ref: not Hispanic/Latino, $n = 616$ )	1.48 (0.32–5.16)	0.42 (0.19–1.00)	0.47 (0.21–1.12)	0.71 (0.31–1.75)	0.55 (0.24–1.24)	0.58 (0.24–1.37)
Neighbourhood SDI <sup>a</sup> (ref: not deprived, $n = 477$ )	1.10 (0.56–2.07)	1.07 (0.68–1.75)	1.73 (1.06–2.92)	1.46 (0.93–2.37)	1.09 (0.73–1.65)	1.02 (0.68–1.54)
Antiarrhythmic rx <sup>b</sup> (ref: no rx, $n = 503$ )	1.65 (0.89–3.00)	0.94 (0.59–1.52)	1.15 (0.73–1.86)	0.61 (0.41–0.93)	0.87 (0.59–1.30)	1.17 (0.78–1.76)
Rate control $rx^{b}$ (ref: no $rx$ , $n = 347$ )	1.13 (0.64–1.98)	0.68 (0.45–1.02)	0.75 (0.51–1.12)	0.79 (0.54–1.16)	0.91 (0.64–1.29)	0.80 (0.56–1.13)
Alcohol history (ref: no history, $n = 224$ )	0.91 (0.51–1.67)	1.07 (0.70–1.62)	0.91 (0.61–1.37)	0.80 (0.54–1.18)	1.28 (0.89–1.83)	1.19 (0.83–1.71)
Smoking history (ref: no history, $n = 407$ )	1.06 (0.58–1.89)	0.74 (0.49–1.13)	0.88 (0.58–1.34)	0.72 (0.48–1.07)	0.71 (0.50–1.02)	0.60 (0.42–0.87)
Heart failure (ref: no heart failure, $n = 251$ )	0.96 (0.52–1.76)	0.68 (0.44–1.04)	0.51 (0.33–0.78)	0.76 (0.51–1.13)	0.41 (0.28–0.59)	0.33 (0.23–0.48)
		Fatigue	Malaise	Palpitations	Syncope	Weakness
Age (ref: under 65, <i>n</i> = 255)		0.94 (0.63–1.41)	0.94 (0.32–2.90)	0.58 (0.40–0.85)	0.75 (0.47–1.22)	0.70 (0.37–1.34)
Gender (ref: Male, $n = 408$ )		0.94 (0.63–1.39)	2.42 (0.73–11.10)	0.42 (0.29–0.61)	0.65 (0.41–1.03)	0.55 (0.30–1.00)
Race (ref: White, $n = 393$ )						
Asian $(n = 34)$		1.16 (0.50–3.03)	0.43 (0.07–8.34)	1.12 (0.51–2.61)	0.98 (0.36–3.46)	1.59 (0.43–10.30)
Black/African American ( $n = 40$ )		0.94 (0.46–1.98)	0.17 (0.04–0.91)	0.72 (0.35–1.51)	0.59 (0.27–1.40)	0.73 (0.28–2.15)
Ethnicity (ref: not Hispanic/Latino, $n = 616$ )		1.82 (0.70–5.71)	NA	0.89 (0.39–2.14)	1.17 (0.42–4.20)	NA
Neighbourhood SDI <sup>a</sup> (ref: not deprived, $n = 477$ )		1.23 (0.79, 1.97)	1.13 (0.35, 5.05)	1.13 (0.74, 1.75)	0.89 (0.54, 1.51)	1.23 (0.60, 2.80)
Antiarrhythmic rx <sup>b</sup> (ref: no rx, $n = 503$ )		0.66 (0.44–1.02)	0.87 (0.29–3.23)	0.92 (0.61–1.42)	0.55 (0.34–0.91)	0.59 (0.32–1.13)
Rate control rx <sup>b</sup> (ref: no rx, $n = 347$ )		1.05 (0.72–1.53)	0.50 (0.17–1.41)	0.73 (0.50–1.04)	0.64 (0.41–1.01)	0.62 (0.33–1.14)
Alcohol history (ref. no history, $n = 224$ )		1.19 (0.81–1.76)	0.72 (0.22–2.11)	1.41 (0.97–2.05)	0.63 (0.38–1.02)	1.87 (1.01–3.46)
Smoking history (ref: no history, $n = 407$ )		0.84 (0.56–1.26)	0.46 (0.16–1.30)	1.05 (0.71–1.55)	1.03 (0.64–1.68)	0.59 (0.32–1.11)
Heart failure (ref: no heart failure, $n = 251$ )		0.46 (0.30–0.69)	1.18 (0.39–3.66)	1.90 (1.30–2.81)	0.84 (0.51–1.35)	0.53 (0.27–1.02)
Bold values indicate statistical significance at $P < 0.05$ . <sup>a</sup> Social deprivation index (SDI) measured at the census tract lew <sup>b</sup> Rx = active prescription at 6 months post-ablation, per electro	el. SDI <90 is considered no nic health records.	ıt deprived.				

associations between patient characteristics and resolution of any symptoms and specific symptoms 12 months post-ablation among com pre-ablation, <i>n</i> = 523, odds ratios (ORs), and 95% confidence intervals (95% CIs)	Any symptoms Anxiety Chest pain Dizziness Dyspnoea Oedema	1.16 (0.57–2.32) 1.48 (0.87–2.61) 0.93 (0.58–1.49) 1.38 (0.85–2.28) 0.69 (0.46–1.04) 1.12 (0.74–1.69)
<b>Table 4</b> Fully adjusted associations between patient ch patients with the symptom pre-ablation, $n = 523$ , odds r	Characteristic Any symptom	Age (ref: under 65, <i>n</i> = 207) 1.16 (0.57–2.32)

Characteristic	Any symptoms	Anxiety	Chest pain	Dizziness	Dyspnoea	Oedema
Age (ref: under 65, <i>n</i> = 207)	1.16 (0.57–2.32)	1.48 (0.87–2.61)	0.93 (0.58–1.49)	1.38 (0.85–2.28)	0.69 (0.46–1.04)	1.12 (0.74–1.69)
Gender (ref: Male, $n = 330$ )	0.70 (0.33–1.40)	0.53 (0.32–0.86)	0.78 (0.50–1.24)	0.65 (0.41–1.02)	0.71 (0.48–1.06)	1.03 (0.70–1.52)
Race (ref: White, $n = 323$ )						
Asian $(n = 26)$	NA	1.39 (0.46–6.06)	1.30 (0.47–4.60)	0.97 (0.35–3.45)	1.35 (0.57–3.48)	0.39 (0.17–0.91)
Black/African American ( $n = 32$ )	0.44 (0.07–1.62)	1.15 (0.45–3.59)	0.61 (0.28–1.41)	0.56 (0.25–1.36)	0.88 (0.40–1.94)	0.90 (0.42–1.96)
Ethnicity (ref: not Hispanic/Latino, $n = 495$ )	1.28 (0.18–5.77)	0.63 (0.24–1.90)	0.57 (0.23–1.50)	1.12 (0.44–3.32)	0.66 (0.27–1.62)	0.49 (0.20–1.22)
Neighbourhood SDI <sup>a</sup> (ref: not deprived, $n = 370$ )	1.52 (0.73–3.04)	1.19 (0.68–2.19)	1.37 (0.82–2.40)	0.98 (0.60–1.65)	1.16 (0.75–1.83)	1.04 (0.68–1.61)
Antiarrhythmic rx <sup>b</sup> (ref. no rx, $n = 400$ )	0.96 (0.41–2.08)	0.89 (0.49–1.69)	1.04 (0.59–1.92)	0.58 (0.35–1.00)	1.08 (0.67–1.77)	1.06 (0.66–1.72)
Rate control $rx^{b}$ (ref: no $rx$ , $n = 246$ )	1.63 (0.85–3.18)	0.73 (0.44–1.19)	0.65 (0.42–1.02)	0.83 (0.53–1.32)	0.97 (0.66–1.44)	0.71 (0.48–1.04)
Alcohol history (ref. no history, $n = 183$ )	1.33 (0.67–2.79)	0.85 (0.50–1.40)	1.23 (0.78–1.93)	0.77 (0.48–1.23)	1.39 (0.94–2.06)	1.10 (0.75–1.63)
Smoking history (ref: no history, $n = 316$ )	0.79 (0.39–1.55)	0.65 (0.40–1.08)	0.80 (0.50–1.28)	0.72 (0.45–1.15)	0.62 (0.42–0.93)	0.70 (0.47–1.03)
Heart failure (ref: no heart failure, $n = 192$ )	1.09 (0.55–2.21)	0.73 (0.43–1.20)	0.75 (0.46–1.19)	0.97 (0.60–1.54)	0.38 (0.25–0.57)	0.37 (0.25–0.56)
		Fatigue	Malaise	Palpitations	Syncope	Weakness
Age (ref: under $65$ , $n = 207$ )		1.15 (0.73–1.84)	1.51 (0.42–6.22)	0.60 (0.39–0.91)	1.07 (0.58–2.04)	1.35 (0.61–3.19)
Gender (ref: Male, $n = 330$ )		0.76 (0.49–1.17)	2.60 (0.62–18.00)	0.42 (0.28–0.63)	0.77 (0.42–1.41)	0.85 (0.40–1.83)
Race (ref: White, $n = 323$ )						
Asian ( $n = 26$ )		1.72 (0.63–6.02)	NA	1.31 (0.52–3.82)	1.56 (0.30–28.6)	NA
Black/African American ( $n = 32$ )		0.98 (0.47–2.20)	NA	0.56 (0.26–1.23)	0.30 (0.12–0.84)	0.36 (0.12–1.21)
Ethnicity (ref: not Hispanic/Latino, $n = 495$ )		1.13 (0.41–3.65)	0.26 (0.03–5.81)	0.62 (0.26–1.50)	0.71 (0.25–2.37)	3.20 (0.57–60.90)
Neighbourhood SDI <sup>a</sup> (ref: not deprived, $n = 370$ )		0.82 (0.51–1.34)	0.78 (0.21–3.67)	0.68 (0.44–1.07)	0.74 (0.40–1.42)	0.77 (0.35–1.82)
Antiarrhythmic rx <sup>b</sup> (ref. no rx, $n = 400$ )		1.02 (0.60–1.78)	0.41 (0.11–1.96)	1.17 (0.69–2.02)	0.75 (0.38–1.59)	0.42 (0.19–0.98)
Rate control $rx^{b}$ (ref: no $rx$ , $n = 246$ )		0.83 (0.54–1.28)	0.71 (0.20–2.90)	0.69 (0.46–1.05)	0.53 (0.29–0.95)	0.93 (0.44–2.05)
Alcohol history (ref: no history, $n = 183$ )		1.31 (0.84–2.03)	1.00 (0.25–3.51)	1.28 (0.85–1.93)	0.51 (0.26–0.96)	3.10 (1.46–6.86)
Smoking history (ref: no history, $n = 316$ )		0.54 (0.35–0.83)	1.45 (0.39–6.97)	0.82 (0.54–1.26)	0.70 (0.39–1.27)	0.54 (0.25–1.16)
Heart failure (ref: no heart failure, $n = 192$ )		0.54 (0.34–0.85)	1.30 (0.36–4.91)	1.90 (1.25–2.89)	1.13 (0.60–2.10)	0.71 (0.31–1.57)
Bold values indicate statistical significance at $P < 0.05$ . <sup>35</sup> ocial deprivation index (SDI) measured at the census tract le <sup>b</sup> Sx = active prescription at 6 months post-ablation, per electr	vel. SDI <90 is considered no: onic health records.	t deprived.				

weakness resolution (OR 0.42, 95% CI 0.19–0.98) compared with those with no prescription. Patients with an alcohol use history had significantly higher odds of weakness resolution (OR 3.10, 95% CI 1.46–6.86) compared with those with no history. Patients with a smoking history had significantly lower odds of dyspnoea resolution (OR 0.62, 95% CI 0.42–0.93) and fatigue resolution (OR 0.54, 95% CI 0.35–0.83) compared with those with no history. Patients with heart failure had significantly lower odds of dyspnoea resolution (OR 0.38, 95% CI 0.25–0.57), oedema resolution (OR 0.37, 95% CI 0.25–0.56), and fatigue resolution (OR 0.54, 95% CI 0.34–0.85), but higher odds of palpitations resolution (OR 1.90, 95% CI 1.25–2.89) compared with those without heart failure.

### **Evaluation of data missingness**

Among the 2531 patients with a documented ablation at the institution, 1293 (51.1%) had pre-ablation notes and were included in our analytic data set, and 1053 (41.6%) had any post-ablation notes. Compared with all patients undergoing ablations, patients with pre-ablation notes were significantly older, fewer were males, and fewer were non-White (see Supplementary material online, *Table S2*). Compared with all patients undergoing ablations, patients with any post-ablation notes were significantly older, fewer were females, and fewer were non-White. There were no differences by ethnicity or SDI in either comparison.

## Discussion

In this analysis of EHRs of nearly 1300 patients with AF undergoing catheter ablation, we found that the majority of patients continue to experience AF symptoms per EHRs, but that there is significant variability in the specific symptoms that resolve by personal and clinical characteristics. Importantly, age 65 and older, female sex, Black or African American race, smoking history, antiarrhythmic use, and comorbid heart failure are associated with lower odds of resolution of specific symptoms at 6 and 12 months. Importantly, nuances regarding the data sources, including the lack of characterization of subtle changes in symptom burden and potential differences in clinician perception of patient symptoms, exist which may have influenced these results. Therefore, these findings warrant confirmation in future investigations with larger, more representative data sets. Nonetheless, this study raises questions about whether specific subgroups of patients with AF symptoms will experience symptom relief post-ablation. Providing patients with realistic expectations of symptom relief post-ablation is an important component of building trust and conducting high-quality shared decision-making around catheter ablation.

A 2016 meta-analysis<sup>22</sup> and multiple recent clinical trials<sup>8,21</sup> and registry studies<sup>23</sup> have confirmed that catheter ablation produces lasting improvements in quality of life at 12 months, and even 2 years,<sup>2</sup> post-ablation. At the same time, multiple studies have also acknowledged differences in symptom outcomes post-ablation. Qualitative and survey studies alike show that many patients report fatigue<sup>25</sup> and increased anxiety<sup>26,27</sup> persisting for several months post-ablation, which they report is upsetting for patients who hoped for faster recoveries after the procedure. Similarly, we found that most patients continued to report at least one symptom post-ablation and that the prevalence of patients with documented anxiety symptoms increased incrementally throughout the post-ablation period, before falling back to baseline at 12 months. This is consistent with prior reports of anxiety about AF recurrence post-ablation.<sup>26,28</sup> Given that anxiety and negative affect can exacerbate other AF symptoms and lead to recurrence,<sup>7,29</sup> addressing post-ablation anxiety should be a priority.

Our findings regarding differences in symptoms by gender and antiarrhythmic use are corroborated by prior studies. A 2-year observational cohort of over 10 000 patients with AF from the ORBIT-AF registry reported significant differences in the types and severity of AF symptoms by gender, but this was not focused on the ablation period.<sup>23</sup> A separate observational cohort following patients with AF 2 years post-ablation reported that patients on continued anticoagulant and/or antiarrhythmic medication had higher odds of AF symptoms.<sup>24</sup> We also uncovered associations between Black/African American race and post-ablation symptoms that have not been previously reported to our knowledge, but our finding was limited by the small proportion of patients identified as this race and high amount of missingness of this variable in EHRs. A previous study suggests patients with missing race data are more likely to represent non-White individuals.<sup>30</sup> Finally, in our study, heart failure was strongly associated with differences in symptom prevalence. The American Heart Association, American College of Cardiology, and Heart Rhythm Society (AHA/ACC/HRS) guidelines for AF support catheter ablation for symptomatic patients with comorbid heart failure because of new evidence, suggesting it is associated with lower hospitalization and mortality rates.<sup>31</sup> Our study suggests heart failure patients may experience relief of some symptoms but not all, and that symptoms that are more directly correlated with heart failure itself (dyspnoea, oedema) may persist post-ablation.

Findings from this study and other investigations of post-ablation symptoms may be informative to patients whose primary goal for undergoing ablation is symptom reduction or resolution. The AHA/ ACC/HRS guidelines for AF support catheter ablations for the treatment of symptomatic paroxysmal AF in the setting of refractory AF or intolerance to antiarrhythmic medications.<sup>4</sup> Symptom control is also a primary goal for patients who opt for ablation.<sup>5</sup> In practice, the decision to undergo ablation requires a nuanced balance of benefits, including the potential to relieve symptoms, and complications, including the high frequency of recurrence and need for repeat ablation.  $^{32,33}$ Shared decision-making, supported by high-quality decision aids, could improve patient guidance as they weigh risks and benefits.<sup>34,35</sup> However, a recent study demonstrated that only one in five patients with AF engaged in shared decision-making around rhythm control strategies, and over half did not understand what different rhythm control options were available to them.<sup>36</sup> Thus, while shared decisionmaking interventions and decision aids supporting anticoagulation decisions for AF have proliferated in recent years,<sup>37–45</sup> there remains a lack of attention on AF symptom and rhythm management.<sup>35</sup> Once generated, high-quality evidence about post-ablation symptom outcomes could be presented to patients to facilitate shared decision-making.

As symptoms drive poor quality of life among patients with cardiovascular disease, comparisons of different methods for collecting high-quality evidence about symptoms should be prioritized. Symptom mining from EHRs to conduct symptom science work is still in its nascent stages, and the accuracy and usefulness of the information retrieved could be improved.<sup>11</sup> For example, in this study, we were only able to characterize symptoms by their presence or absence; in reality, symptom burden is a continuum. More subtle changes in symptom burden (e.g. improvement but not altogether resolved) could potentially be captured with more sophisticated, machine-learning-based NLP techniques, but nonetheless relies on clinicians to document these changes. Symptom documentation in EHRs is conducted by clinicians, who may de-prioritize certain symptoms, misunderstand patient reports, or conduct incomplete symptom assessments.

Nonetheless, real-world evidence from EHRs may offer benefits over patient-reported outcomes in clinical trials and disease registries, which are typically time and resource intensive to conduct,<sup>46</sup> may include younger, healthier, predominantly White populations due to trial eligibility criteria and patient mistrust of researchers,<sup>47</sup> and often face significant challenges with missing longitudinal patient-reported outcomes data.<sup>8</sup> In this single-site study, we were able to access over 33 000 EHRs and ultimately study the symptoms of ~1300 patients across 10 years, with significantly less time and resource utilization compared with a clinical trial of the same size. With use of common data models such as OMOP, aggregation of records across multiple

institutions is possible. Given these potential benefits, EHRs may provide a complementary mechanism for studying symptoms when clinical trials are infeasible. Future studies comparing patient self-reported symptoms to those documented in EHRs by clinicians will illuminate the degree to which symptom documentation in EHRs reflects patients' true symptom status.

Additional limitations of this study include that it was a single-arm study without a comparator cohort (e.g. patients with AF on medications alone). Additionally, we observed missing data across the periand post-ablation periods, which may have been biased by age, gender, and race. In particular, patients in this sample were significantly older than the broader population of patients undergoing ablation at our institution. This missing datum probably reflects a mixture of both true missingness (inconsistent attendance at follow-up appointments and/ or follow-up outside the institution) and documentation bias (some symptoms not documented and some notes not stored in the institution's OMOP instance). Similarly, data for certain variables may have been underreported and dates associated with the data may have been incorrect in some cases; this was particularly true for medication administrations (which lacked end dates) and smoking and alcohol use, which were historical variables and not representative of active use at each time point.

## Implications for practice

This analysis of symptom information documented in institutional EHRs suggests that symptom resolution post-ablation varies by specific symptoms and specific patient characteristics. Specifically, we noted trends in characteristics such that patients with comorbid heart failure, age 65 and older, female sex, Black or African American race, smoking history, and antiarrhythmic use may be less likely to experience resolution of specific symptoms 6 and 12 months post-ablation. While warranting confirmation in future research, this information may be informative for practice, as it could facilitate high-quality shared decision-making among patients and clinicians considering catheter ablation to treat AF symptoms.

## Conclusion

This study demonstrated that post-ablation symptom patterns are heterogeneous. Though confirmation with larger, more representative data sets is needed, these findings may be informative for patients whose primary goal for undergoing an ablation is symptom relief.

## **Author contributions**

M.R.T. conceptualized the study, acquired funding, collected and aggregated the data, completed the data analysis, and drafted the original manuscript. M.R.T., A.V., W.G., B.T., and M.H. completed the data curation (annotating the gold standard corpus). J.P. provided methodological guidance and supervision. D.S. provided clinical domain-specific guidance and supervision. A.V., W.G., B.T., M.H., J.P., and D.S. provided critical feedback on the data analysis and reviewed and edited the manuscript.

## Supplementary material

Supplementary material is available at European Journal of Cardiovascular Nursing online.

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**Conflict of interest:** M.R.T. and J.P. are affiliated with Iris OB Health LLC (equity). M.R.T. is affiliated with Boston Scientific Corp (consulting). The remaining authors have no disclosures.

## Data availability

The data underlying this article cannot be shared publicly due to the privacy of individuals who participated in the study, as determined by the Weill Cornell Medicine Institutional Review Board. The data will be shared on reasonable request with the corresponding author.

## **Ethics** approval

This study was approved by the Weill Cornell Medicine Institutional Review Board.

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