

# Outcomes of patients with heart failure with preserved ejection fraction undergoing catheter ablation of atrial fibrillation



Amrita Krishnamurthy, MD, Parag Goyal, MD, MSc, Steven M. Markowitz, MD, FHRS, Christopher F. Liu, MD, FHRS, George Thomas, MD, James E. Ip, MD, FHRS, Evelyn M. Horn, MD, Bruce B. Lerman, MD, FHRS, Luke K. Kim, MD, Jim W. Cheung, MD, FHRS

From the Weill Cornell Cardiovascular Outcomes Research Group (CORG), Department of Medicine, Division of Cardiology, Weill Cornell Medicine–New York Presbyterian Hospital, New York, New York.

**BACKGROUND** Limited real-world data exist on early outcomes in patients with heart failure with preserved ejection fraction (HFpEF) undergoing atrial fibrillation (AF) ablation.

**OBJECTIVES** The purpose of this study was to examine and compare rates of index procedural complications and 30-day readmissions after AF ablation in patients with HFpEF, with heart failure with reduced ejection fraction (HFrEF), and without heart failure.

**METHODS** Using the Nationwide Readmissions Database (NRD), we examined 50,299 admissions of adults with heart failure undergoing AF catheter ablation between 2010 and 2014. Using ICD-9-CM codes, we identified procedural complications and causes of readmission after AF ablation.

**RESULTS** From 2010 to 2014, the prevalence of HFpEF among patients undergoing AF ablation increased from 3.05% to 7.35% ( $P$  for trend  $<.001$ ). Compared to patients without heart failure, patients with HFpEF had significantly increased procedural complications and index mortality (8.4% vs 6.2% and 0.30% vs 0.08%, respectively;  $P = .016$  and  $P = .010$ , respectively). Index complication rates between patients with HFpEF and HFrEF were similar. All-

cause 30-day readmissions occurred in 18.3% of patients with HFpEF compared to 9.5% of patients without heart failure ( $P <.001$ ). Compared to no heart failure, the presence of HFpEF was independently associated with all-cause readmissions (adjusted odds ratio 1.52; 95% confidence interval 1.15–1.96;  $P = .002$ ), but not with procedural complications, cardiac readmissions, or early mortality.

**CONCLUSION** Rates of 30-day readmissions after AF ablation are high in patients with HFpEF. However, after adjustment for age and comorbidities, complications and early mortality after AF ablation between patients with HFpEF and those without heart failure are comparable.

**KEYWORDS** Atrial fibrillation; Catheter ablation; Heart failure with preserved ejection fraction; Heart failure with reduced ejection fraction; Outcomes; Readmissions

(Heart Rhythm 0<sup>2</sup> 2022;3:501–508) © 2022 Heart Rhythm Society. Published by Elsevier Inc. This is an open access article under the CC BY-NC-ND license (<http://creativecommons.org/licenses/by-nc-nd/4.0/>).

## Introduction

Heart failure is a leading cause of hospitalizations and readmissions in the United States,<sup>1</sup> and rates of 30- and 90-day heart failure readmissions have increased from 2010 to 2017.<sup>2</sup> Heart failure has been classified as heart failure with preserved ejection fraction (HFpEF) and heart failure with reduced ejection fraction (HFrEF). Importantly, HFpEF is now the most common form of heart failure. Patients with HFpEF tend to be older, be female, and have more comorbidities, including hypertension, renal disease, and atrial fibrillation (AF).<sup>3,4</sup> The combination of heart failure and AF is associated with a worse prognosis.<sup>5</sup> Catheter ablation has been shown to reduce all-

cause mortality, heart failure hospitalizations, and improve left ventricular ejection fraction in those with HFrEF; however, outcomes in HFpEF patients have been less well studied.<sup>6–8</sup> Recent data suggest adoption of an early rhythm control strategy in patients with pre-existing cardiovascular disease, including heart failure, leads to lower risk of adverse cardiovascular outcomes.<sup>9</sup> Compared with antiarrhythmic therapy, catheter ablation has demonstrated improvement in quality of life and survival in both HFpEF as well as HFrEF patients.<sup>10</sup>

Although heart failure has been associated with procedural complications, readmissions, and early mortality after AF ablation,<sup>11</sup> real-world short-term outcomes for the subgroup of patients with HFpEF have not been well described. Both single-center studies and meta-analyses have demonstrated that patients with HFpEF have similar freedom from atrial arrhythmia recurrence and

**Address reprint requests and correspondence:** Dr Jim W. Cheung, Division of Cardiology, Weill Cornell Medical College, 520 East 70th St, Starr 4, New York, NY 10021. E-mail address: [jac9029@med.cornell.edu](mailto:jac9029@med.cornell.edu).

## KEY FINDINGS

- The prevalence of heart failure with preserved ejection fraction (HFpEF) among patients undergoing atrial fibrillation (AF) ablation in the real world increased significantly between 2010 and 2014.
- Patients with HFpEF had more procedural complications, 30-day readmissions, and early mortality after AF ablation compared to patients without heart failure.
- However, after adjustment for age, sex, comorbidities, and hospital factors, the presence of HFpEF was not independently associated with adverse outcomes after AF ablation.

symptomatic improvement as HFrEF patients 12 months after catheter ablation.<sup>12,13</sup> Using a nationally representative administrative database, we sought to identify complications, early mortality, and 30-day readmissions after AF ablation in patients with HFpEF and to compare these outcomes with those of patients with no heart failure and with HFrEF.

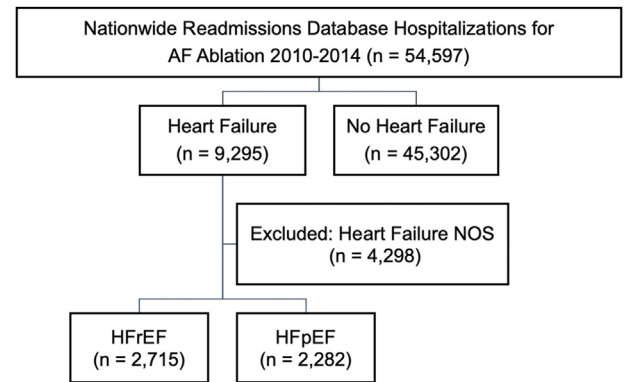
## Methods

### Data source

All data were obtained from the Agency for Healthcare Research and Quality (AHRQ). The National Readmission Database (NRD) is drawn from the Healthcare Cost and Utilization Project (HCUP) State Inpatient Databases. The NRD is an annual, multipayer administrative database that tracks 1 calendar year of discharge data for patients across hospitals within a state.<sup>14–16</sup> The *International Classification of Diseases, Ninth Revision, Clinical Modification (ICD-9-CM)* codes for diagnoses and procedures are available for each patient admission record. We used the NRD using the time period from 2010 to 2014. This study duration was prior to 2015 when the “two midnight” rule for the definition of inpatient admissions was enforced. Because the NRD includes only inpatients, NRD studies of AF catheter ablation after 2014 would exclude outpatient procedures and therefore would significantly undersample the number of AF ablations performed. The Weill Cornell Institutional Review Board deemed ethical approval, and informed consent was not required for the study because all data were derived from a de-identified administrative database. The data that support the findings of this study are available from the corresponding author upon reasonable request. The research reported in this paper adhered to the revised 2013 Helsinki Declaration guidelines.

### Study population

Hospitalizations for catheter ablation of AF were selected by identifying patients with a primary ICD-9-CM diagnosis code for AF (427.31) and a primary



**Figure 1** Flowchart of study population selection. AF = atrial fibrillation; HFpEF = heart failure with preserved ejection fraction; NOS = not otherwise specified; HFrEF = heart failure with reduced ejection fraction.

ICD-9-CM procedure code for catheter ablation (37.34). Patients with a secondary ICD-9-CM code for other arrhythmias or a procedure code for device implantation during the index admission were excluded (Supplemental Table 1). Heart failure subtype was identified via primary ICD-9-CM code for diastolic heart failure (428.30–428.33) and systolic heart failure (428.20–428.23) and were defined in this study as HFpEF and HFrEF, respectively (Figure 1). Patients with combined systolic and diastolic heart failure (428.40–428.43), unspecified heart failure (428.0, 428.9), and left heart failure (428.1) were excluded (n = 4298) (Supplemental Table 2). Because the NRD is reset annually, we excluded patients if they were discharged in December from their index admission in order to ensure that 30-day follow up after discharge was available. We also excluded patients under the age of 18 years and those without mortality data.

### Clinical variables

We included patient- and hospital-level variables extracted from the NRD, including age, sex, median household income, primary payer, hospital region, and hospital size, for baseline characteristic analysis. Patient-level variables and cardiac diagnoses were defined by ICD-9-CM codes, Clinical Classification Software (CCS) codes, and AHRQ comorbidity measures based on the Elixhauser methods as defined in Supplemental Table 3.

### Study endpoints

Our primary endpoints were rates of index procedural complications and 30-day readmissions after AF ablation in patients with HFpEF, HFrEF, and no heart failure, according to the methodology described by HCUP. Procedural complications included cardiac perforation/tamponade, vascular complications, stroke, and other iatrogenic cardiac complications (Supplemental Table 3). ICD-9-CM diagnosis codes were used to identify the causes of 30-day readmissions by organ system (Supplemental Table 4), which included both cardiac

**Table 1** Baseline characteristics of hospitalizations stratified by heart failure subtype

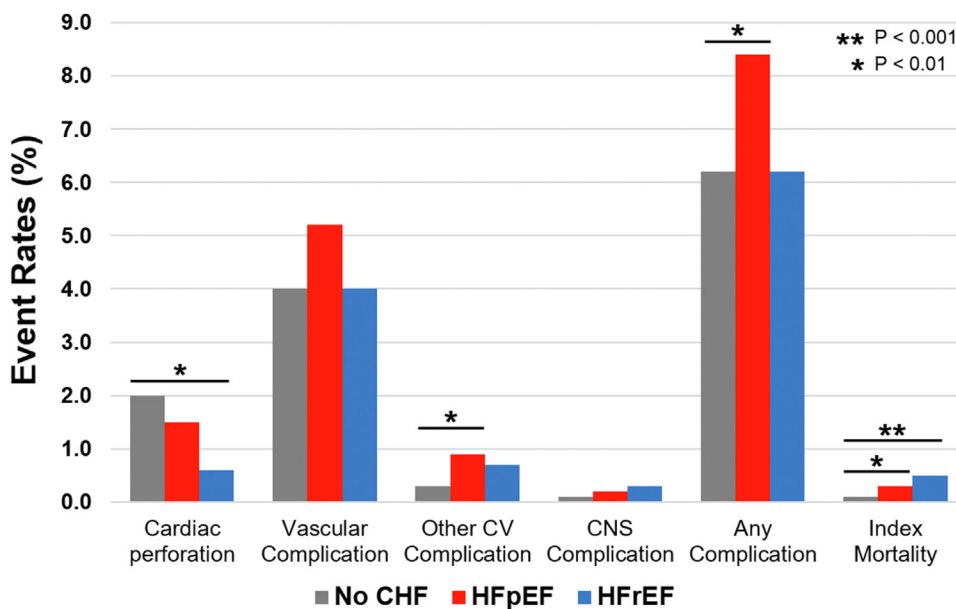
Characteristics	No CHF	HFrEF	HFpEF
No. of AF ablations	45,302	2715	2282
Age (y) [mean (SE)]	63.4 (0.15)	68.2 (0.39)	72.3 (0.39)
Age group (y)			
<65	22,541 (49.8)	927 (34.1)	504 (22.1)
65–74	16,263 (35.9)	910 (33.5)	777 (34.0)
≥75	6498 (14.3)	878 (32.4)	1002 (43.9)
Female	16,663 (36.8)	870 (32.0)	1403 (61.5)
CAD	10,257 (22.6)	1411 (52.0)	927 (40.6)
Previous PCI	3250 (7.1)	395 (14.6)	252 (11.0)
Previous PPM	4495 (9.9)	457 (16.8)	726 (31.8)
Previous ICD	1092 (2.4)	957 (35.2)	95 (4.2)
Hypertension	26,821 (59.2)	1242 (45.8)	1279 (56.03)
Diabetes mellitus	8178 (18.1)	891 (32.8)	727 (31.9)
Hyperlipidemia	7505 (46.1)	606 (49.5)	629 (56.9)
Obesity	6727 (14.8)	478 (17.6)	564 (24.7)
History of stroke	2515 (5.6)	205 (7.6)	220 (9.7)
Valvular disease	4997 (11.0)	664 (24.4)	594 (26.0)
Peripheral vascular disease	1257 (2.8)	205 (7.6)	128 (5.6)
Pulmonary hypertension	781 (1.7)	231 (8.5)	281 (12.3)
Chronic lung disease	5593 (12.3)	690 (25.4)	712 (31.2)
Smoking	2154 (4.8)	190 (7.0)	148 (6.5)
Renal disease	1945 (4.3)	748 (27.5)	452 (19.8)
Alcohol abuse	477 (1.1)	56 (2.1)	27 (1.2)
Substance abuse	164 (0.4)	24 (0.9)	8 (0.3)
Cancer	388 (0.9)	47 (1.7)	63 (2.8)
Anemia	1983 (4.4)	320 (11.8)	359 (15.7)
Coagulopathy	677 (1.5)	113 (4.2)	91 (4.0)
Elixhauser co-morbidity score >4	3248 (7.2)	1261 (46.4)	1160 (50.8)
Hospital AF ablation volume			
First tertile (lowest)	15,096 (33.3)	1387 (51.1)	1110 (48.6)
Second tertile	15,363 (33.9)	872 (32.1)	723 (31.7)
Third tertile (highest)	14,843 (32.8)	455 (16.8)	449 (19.7)
Teaching hospital	33,733 (74.5)	2024 (74.6)	1627 (71.3)
Median household income			
First quartile (lowest)	8173 (18.3)	697 (26.0)	531 (23.6)
Second quartile	10,274 (23.0)	718 (26.8)	595 (26.5)
Third quartile	12,098 (27.2)	614 (22.9)	510 (22.7)
Fourth quartile (highest)	13,997 (31.4)	653 (24.3)	610 (27.2)
Primary payer			
Medicare	22,431 (49.5)	1929 (71.1)	1768 (77.5)
Medicaid	1255 (2.8)	92 (3.4)	47.7 (2.1)
Private including HMO	20,183 (44.6)	592 (21.8)	406 (17.8)
Self-pay/no charge/other	1417 (3.1)	98 (3.6)	60 (2.6)
Hospital region			
Urban	44,945 (99.2)	2678 (98.6)	2199 (96.4)
Hospital bed size			
Small	1494 (3.3)	94 (3.5)	114 (5.0)
Medium	7773 (17.2)	446 (16.4)	458 (20.0)
Large	36,035 (79.5)	2175 (80.1)	1710 (74.9)
Length of stay ≥3 days	10,670 (23.6)	1664 (61.3)	1421 (62.3)
LOS, days	0.89 [0.40–1.92]	2.83 [1.12–5.63]	2.88 [1.13–5.84]
Cost (\$)	23,199 [16,940–29,773]	18,685 [10,014–29,284]	22,728 [13,179–31,355]

Values are given as n (%) or mean [interquartile range] unless otherwise indicated.

AF = atrial fibrillation; CAD = coronary artery disease; CHF = congestive heart failure; HFpEF = heart failure with preserved ejection fraction; HFrEF = heart failure with reduced ejection fraction; HMO = health maintenance organization; ICD = implantable cardioverter-defibrillator; LOS = length of stay; PCI = percutaneous coronary intervention; PPM = permanent pacemaker.

and noncardiac causes. Subgroups of cardiac causes of readmissions included atrial fibrillation/tachycardia and congestive heart failure ([Supplemental Table 5](#)). In our analysis, we included only the first readmission within

30 days after discharge from the index admission for AF ablation. Index and early mortality included mortality occurring at the index admission or at 30-day readmission, respectively, after AF ablation.



**Figure 2** Index procedural complications after atrial fibrillation ablation among patients with no heart failure, with heart failure with preserved ejection fraction (HFpEF), and with heart failure with reduced ejection fraction (HFrEF). CHF = congestive heart failure; CNS = central nervous system; CV = cardiovascular.

### Statistical analysis

All analyses were performed using SAS software, Version 9.4 (SAS Institute, Cary, NC). Discharge weight provided by the NRD was used for all analyses to obtain national estimates. All analyses accounted for hospital-level clustering of patients and complex survey sampling design. Both patient- and hospital-level variables were used for baseline characteristic analysis. For descriptive analyses, we compared baseline patient- and hospital-level variables of AF ablation patients stratified by heart failure phenotype. Categorical variables are given as frequencies, and continuous variables are given as mean (standard error) or median [interquartile range]. Baseline characteristics and outcomes were compared by Rao-Scott  $\chi^2$  test for categorical variables. Survey-specific linear regression or Mann-Whitney–Wilcoxon nonparametric test was used for continuous variables. To identify predictors of complications, readmissions, and mortality, we created multivariable logistic regression models for the outcome of interest by including covariates that had univariate significance for the outcome ( $P < .10$ ) (Supplemental Table 6). All  $P$  values were corrected for multiple comparisons using the Bonferroni method. All tests were 2-sided, with  $P < .05$  considered significant.

## Results

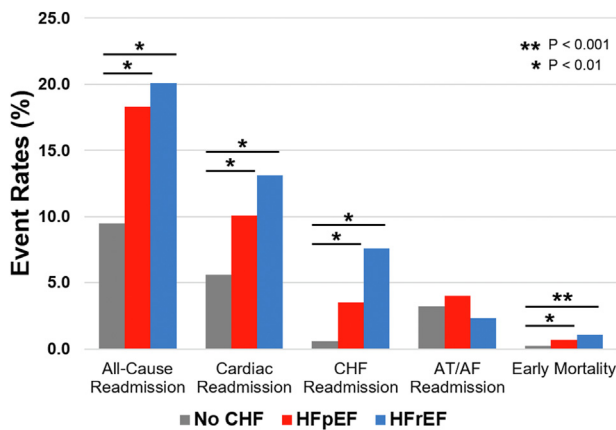
### Study population and trends in prevalence of HFpEF among patients undergoing AF ablation

A total of 50,299 admission records from the NRD of patients undergoing catheter ablation of AF from 2010 to 2014 were included in the study analysis. Median length of hospital stay was 1 day [0.4–1.9], suggesting that the majority of AF catheter ablation procedures were overnight stays (Table 1). In this group, 2282 (4.5%) had a diagnosis of HFpEF, 2715

(5.4%) had a diagnosis of HFrEF, and 45,302 (90.0%) had no heart failure (Figure 1). There was a significant increase in the prevalence of heart failure among those undergoing AF catheter ablation during the study period. From 2010 to 2014, the prevalence of HFrEF increased from 3.82% to 7.63% ( $P$  for trend  $< .0001$ ), while the prevalence of HFpEF increased from 3.05% to 7.35% ( $P$  for trend  $< .0001$ ) (Supplemental Figure 1). Compared to patients without heart failure, patients with HFpEF were older, were more likely to be female, and were more likely to have coronary artery disease, previous percutaneous coronary intervention, previous pacemaker implantation, diabetes, obesity, previous stroke, valvular disease, peripheral vascular disease, pulmonary hypertension, and lung disease (Table 1). Patients with HFrEF were less likely to have kidney disease. Compared to patients with HFrEF, patients with HFpEF were older, more likely to be female, and more likely to have previous pacemaker implantation, hypertension, obesity, previous stroke, pulmonary hypertension, and lung disease.

### Rates of AF ablation procedural complications among patients with HFpEF

Procedural complications associated with AF ablation, stratified by heart failure type, are shown in Figure 2. Compared to patients without heart failure, patients with HFpEF had higher rates of any procedural complication (8.4% vs 6.2%;  $P = .016$ ) and index mortality (0.3% vs 0.08%;  $P = .010$ ). Rates of cardiac perforation (1.5% vs 2.0%;  $P = .403$ ), vascular complications (5.2% vs 4.0%;  $P = .099$ ), and central nervous system complications (0.2% vs 0.3%;  $P = 0.504$ ) were similar between patients with HFpEF and those without heart failure. Compared to patients with HFrEF, patients with HFpEF had a trend toward higher rates of procedural complications (8.4% vs 6.2%;  $P = .09$ ) but



**Figure 3** Thirty-day readmissions after atrial fibrillation (AF) ablation among patients with no heart failure, with HFpEF, and with HFrEF. AT = atrial tachycardia. Other abbreviations as in Figure 2.

similar rates of index mortality after AF ablation (0.3% vs 0.49%;  $P = .44$ ).

**Thirty-day readmission rates among patients with HFpEF after AF ablation**

Compared to patients without heart failure, patients with HFpEF had higher rates of all-cause 30-day readmissions (18.3% vs 9.5%;  $P < .001$ ), cardiac readmissions (10.1% vs 5.6%;  $P < .001$ ), heart failure readmission (3.5% vs 0.61%;  $P < .001$ ), noncardiac readmissions (8.2% vs 3.8%;  $P < .001$ ), and early mortality (0.67% vs 0.25%;  $P = .007$ ) (Figure 3). Compared to patients with HFrEF, patients with HFpEF had similar rates of 30-day, all-cause readmissions,

cardiac readmissions, noncardiac readmissions, and early mortality but fewer heart failure readmissions (3.5% vs 7.6%;  $P < .001$ ). Readmissions for atrial arrhythmias were similar among patients with HFpEF, with HFrEF, and without heart failure.

**HFpEF and HPrEF as predictors of outcomes**

We used logistic regression analysis to identify the relationship between HFpEF, HFrEF, and index procedural complications as well as 30-day readmissions. After adjusting for age, medical comorbidities, and hospital factors, HFpEF was not independently associated with procedural complications (Table 2) or index mortality (Table 3). With respect to 30-day readmissions, compared to patients without heart failure, HFpEF was independently associated with all-cause readmissions (adjusted odds ratio [aOR] 1.52; 95% confidence interval [CI] 1.17–1.99;  $P < .01$ ) and heart failure readmissions (aOR 2.42; 95% CI 1.51–3.86;  $P < .001$ ) but not cardiac readmission, noncardiac readmissions, or early mortality.

**Discussion**

Using a nationally representative, all-payer administrative database, we identified several important findings in this real-world analysis of early outcomes after AF catheter ablation in HFpEF patients. First, the prevalence of HFpEF among patients undergoing AF ablation increased significantly between 2010 and 2014. Second, compared to patients without heart failure, patients with HFpEF had more procedural complications, all-cause 30-day readmissions, cardiac readmissions, noncardiac readmissions, and early mortality

**Table 2** Association between heart failure subtype and index procedural complications (absence of CHF as reference)

Outcome	Unadjusted OR (95% CI)	Unadjusted P value	Adjusted OR (95% CI)	Adjusted P value
<b>Cardiac perforation</b>				
No HF	Reference		Reference	
HFrEF	0.28 (0.14–0.57)	<.001	0.31 (0.15–0.65)	.002
HFpEF	0.75 (0.39–1.46)	.404	0.68 (0.35–1.29)	.236
<b>Vascular complication</b>				
No HF	Reference		Reference	
HFrEF	1.00 (0.72–1.39)	.988	0.89 (0.59–1.35)	.579
HFpEF	1.30 (0.95–1.78)	.099	0.82 (0.58–1.17)	.269
<b>Other CV complication</b>				
No HF	Reference		Reference	
HFrEF	1.88 (1.21–2.92)	.005	1.55 (0.99–2.45)	.058
HFpEF	2.13 (1.03–4.41)	.041	1.77 (0.85–3.68)	.128
<b>CNS complication</b>				
No HF	Reference		Reference	
HFrEF	0.99 (0.34–2.82)	.978	1.01 (0.35–2.90)	.992
HFpEF	0.61 (0.14–2.61)	.508	0.59 (0.14–2.53)	.479
<b>Any complication</b>				
No HF	Reference		Reference	
HFrEF	1.00 (0.77–1.31)	.986	0.91 (0.67–1.24)	.554
HFpEF	1.39 (1.06–1.81)	.017	0.95 (0.71–1.28)	.758
<b>Index mortality</b>				
No HF	Reference		Reference	
HFrEF	6.09 (2.33–15.89)	<.001	1.98 (0.60–6.53)	.264
HFpEF	3.76 (1.27–11.14)	.0168	1.12 (0.40–3.55)	.758

CI = confidence interval; CNS = central nervous system; CV = cardiovascular; HF = heart failure; OR = odds ratio; other abbreviations as in Table 1.

**Table 3** Association between heart failure subtype and 30-day readmissions (absence of CHF as reference)

Outcome	Unadjusted OR (95% CI)	Unadjusted <i>P</i> value	Adjusted OR (95% CI)	Adjusted <i>P</i> value
All-cause readmission				
No HF	Reference		Reference	
HFrEF	2.39 (1.97–2.90)	<.001	1.56 (1.19–2.05)	.001
HFpEF	2.14 (1.79–2.55)	<.001	1.52 (1.17–1.98)	.002
Cardiac readmission				
No HF	Reference		Reference	
HFrEF	2.52 (2.03–3.12)	<.001	1.98 (1.44–2.73)	<.001
HFpEF	1.88 (1.52–2.34)	<.001	1.36 (0.98–1.93)	.064
HF readmission				
No HF	Reference		Reference	
HFrEF	13.25 (9.52–18.46)	<.001	5.99 (4.21–8.53)	<.001
HFpEF	6.04 (4.10–8.88)	<.001	2.42 (1.51–3.86)	<.001
AT/AF readmission				
No HF	Reference		Reference	
HFrEF	0.71 (0.49–1.05)	.085	1.07 (0.59–1.94)	.882
HFpEF	1.24 (0.86–1.77)	.246	0.93 (0.55–1.56)	.772
Noncardiac readmission				
No HF	Reference		Reference	
HFrEF	1.86 (1.39–2.48)	<.001	1.12 (0.80–1.58)	.520
HFpEF	2.23 (1.70–2.90)	<.001	1.25 (0.94–1.67)	.124
Early mortality				
No HF	Reference		Reference	
HFrEF	4.39 (2.34–8.23)	<.001	1.79 (0.66–4.88)	.252
HFpEF	2.66 (1.26–5.60)	<.001	0.95 (0.42–2.16)	.904

AT = atrial tachycardia; other abbreviations as in [Tables 1 and 2](#).

after AF ablation compared to patients without heart failure. After adjustment for age, sex, comorbidities, and hospital factors, the presence of HFpEF was independently associated with all-cause and heart failure readmissions but not independently associated with procedural complications, cardiac readmissions, or early mortality. Our study suggests that many adverse outcomes seen in patients with HFpEF undergoing AF ablation are driven by age, sex, and comorbidities rather than the presence of HFpEF itself.

In patients with left ventricular systolic dysfunction, AF ablation can improve left ventricular ejection fraction, exercise capacity, and quality of life, as well as reduce hospitalizations and mortality.<sup>6–8</sup> Studies of AF ablation outcomes are more limited in patients with HFpEF. There are some phenotypic differences between patients with AF who have HFpEF vs those with HFrEF. In a study by Melenovsky et al<sup>17</sup> that evaluated echocardiographic and catheterization parameters of the left atrium in 198 heart failure patients and 40 heart failure-free controls, both HFrEF and HFpEF patients had more dilated atria and greater degrees of left atrial dysfunction. Compared to patients with HFrEF, patients with HFpEF can have a higher burden of AF due to higher left atrial peak pressures and higher left atrial stiffness. Among patients with HFpEF, left atrial dysfunction is associated with increased mortality. Several smaller studies have shown that catheter ablation of AF in patients with HFpEF can be safe and is associated with reduced heart failure events. In an observational, single-center, retrospective cohort study, Aldaas et al<sup>18</sup> examined outcomes of 547 patients, including 51 (9%) with HFpEF and 40 (7%) with HFrEF, who underwent radiofrequency catheter ablation

for AF. There were no significant differences in recurrence of atrial arrhythmias, procedural complications, or 5-year survival in patients with HFpEF vs patients with HFrEF vs patients without heart failure. All-cause hospitalizations were more common in HFpEF and HFrEF patients than in those without heart failure.<sup>18</sup>

Larger-scale, prospective, randomized clinical trials of outcomes after AF catheter ablation in patients with HFpEF are needed. Single-center studies to date, such as STALL AF-HFpEF (A Prospective Study Using Invasive Haemodynamic Measurements Following Catheter Ablation for AF and Early HFpEF), which examined 54 patients with AF, 35 of whom also had HFpEF, have demonstrated that AF catheter ablation can lead to improvements in quality of life and hemodynamic parameters such as postexercise pulmonary capillary wedge pressure in patients with HFpEF.<sup>19</sup> Additional prospective trials seek to examine various outcomes in HFpEF patients undergoing AF catheter ablation, including exercise capacity and quality of life<sup>20</sup> and arrhythmia recurrence,<sup>21</sup> are currently under recruitment. Once these data are available, there may be further expansion in the patient population offered early AF ablation.

Our study provides real-world insight into short-term outcomes after AF catheter ablation in patients with HFpEF, and its findings suggest that HFpEF does not seem to independently confer increased rates of procedural complications, index procedural mortality, or early mortality. However, as with HFrEF, HFpEF is independently associated with increased risk of all-cause 30-day readmissions. Although there was a trend toward HFpEF as an independent predictor of cardiac readmissions, this was not seen with regard to

readmissions for atrial tachycardia/AF recurrence. Taken together, these findings underscore the importance of patient optimization by electrophysiologists and heart failure specialists in the periprocedural period.

### Study limitations

This was a retrospective study based on administrative data from the NRD, which relies on entered ICD-9-CM codes. This introduces potential error from miscoding, undercoding or overcoding, and missing data. Notably, the ICD-9-CM does not include specific codes for HFrEF or HFpEF, so codes for systolic and diastolic heart failure, respectively, were used as surrogate identifiers of these conditions. In recent years, efforts have been made to standardize the definition of HFpEF using scoring systems. Reddy et al<sup>22</sup> created the H<sub>2</sub>FPEF score, which utilizes clinical and echocardiographic criteria to discern HFpEF from noncardiac causes of dyspnea. In addition, the European Society of Cardiology created the HFA-PEFF score, which uses biomarker as well as functional and morphologic echocardiographic criteria, in an algorithm used to diagnose HFpEF.<sup>23</sup> Because echocardiographic parameters were not available in the NRD, these scoring systems could not be used to further ascertain the diagnosis of HFpEF in our study population. Given the likely underestimation of HFpEF prevalence in our study, complication and readmission rates may be overestimated. With publication of studies showing improved outcomes with AF ablation for patients with HFrEF,<sup>6</sup> a more contemporary analysis of patients with heart failure undergoing AF ablation may reveal different patient characteristics. The NRD does not include data on clinical variables such as New York Heart Association (NYHA) functional class, atrial fibrillation burden, and medications; therefore, we were unable to explore the impact of these patient factors on outcomes. Procedural variables such as type of catheter ablation, procedural length, and operator expertise, also are not included in the NRD. There have been numerous advances in the practice of AF catheter ablation since this study period. Improvements in catheter-based technologies have led to greater procedural safety, shorter procedural durations, and more durable pulmonary vein isolation.<sup>24</sup> In addition, emerging technologies such as irreversible electroporation have shown promise in early trials, with larger-scale data on safety and efficacy forthcoming.<sup>25</sup> Given this, the readmissions and procedural complications rates seen in our study are expected to be higher than those seen with a contemporary cohort. Although the NRD is designed to approximate the national distribution of hospital characteristics, it does not include data from all 50 states in the United States; therefore, certain hospital types may be overrepresented or underrepresented, thus limiting the generalizability of this study. Moreover, given that the NRD is an inpatient database, variability in coding of inpatient vs outpatient procedures can lead to heterogeneity in patient inclusion across institutions. Specifically, the exclusion of patients undergoing AF catheter ablation as outpatients may have biased our study population. Finally, estimates of early mortality

may be underrepresented because out-of-hospital deaths before readmission would not be included in the analysis.

### Conclusion

In this nationally representative real-world cohort, rates of 30-day readmissions after AF catheter ablation are higher in patients with HFpEF than in patients without heart failure. After adjustment, HFpEF was not an independent predictor of overall index procedural complications, cardiac readmissions, index mortality, or early mortality after AF catheter ablation. Further studies examining the role of patient and procedural factors are warranted to determine the clinical predictors of readmissions in heart failure patients after AF catheter ablation. A multidisciplinary approach to optimize periprocedural heart failure management may reduce readmissions in patients with HFpEF who undergo catheter ablation of AF.

**Funding Sources:** This work was supported by grants from the Michael Wolk Heart Foundation, the New York Cardiac Center, Inc., and the New York Weill Cornell Medical Center Alumni Council. The Michael Wolk Heart Foundation, the New York Cardiac Center, Inc., and the New York Weill Cornell Medical Center Alumni Council had no role in the design and conduct of the study, the collection, analysis, and interpretation of the data, or the preparation, review, or approval of the manuscript.

**Disclosures:** Dr Cheung has received consulting fees from Abbott, Biotronik, Boston Scientific, and Zoll Medical; research grant support from Boston Scientific; and fellowship grant support from Abbott, Biosense Webster, Biotronik, Boston Scientific, and Medtronic. Dr Markowitz has received consulting fees from Preventice Medical and fees from Boston Scientific for serving on a data safety monitoring board. Dr Goyal is supported by American Heart Association Grant 20CDA35310455, National Institute on Aging Grant K76AG064428, and Loan Repayment Program award L30AG060521; receives personal fees for medicolegal consulting related to heart failure; and has received honoraria from Akcea Therapeutics Inc and Bionest Inc. Dr Horn has received fees from V-wave Ltd and SoniVie for serving on a data safety monitoring board and clinical events committee. All other authors report no relevant disclosures.

**Authorship:** All authors attest they meet the current ICMJE criteria for authorship.

**Patient Consent:** Informed consent was not required for the study because all data were derived from a de-identified administrative database.

**Ethics Statement:** The Weill Cornell Institutional Review Board deemed ethical approval, and informed consent was not required for the study because all data were derived from a de-identified administrative database. The research reported in this paper adhered to the revised 2013 Helsinki Declaration guidelines.

### Supplementary data

Supplementary data associated with this article can be found in the online version at <https://doi.org/10.1016/j.hroo.2022.06.012>.

### References

1. Ambrosy AP, Fonarow GC, Butler J, et al. The global health and economic burden of hospitalizations for heart failure: lessons learned from hospitalized heart failure registries. *J Am Coll Cardiol* 2014;63:1123–1133.
2. Khan MS, Sreenivasan J, Lateef N, et al. Trends in 30- and 90-day readmission rates for heart failure. *Circ Heart Fail* 2021;14:e008335.

3. Goyal P, Almarzooq ZI, Horn EM, et al. Characteristics of hospitalizations for heart failure with preserved ejection fraction. *Am J Med* 2016;129:635. e15–e26.
4. Steinberg BA, Zhao X, Heidenreich PA, et al. Trends in patients hospitalized with heart failure and preserved left ventricular ejection fraction: prevalence, therapies, and outcomes. *Circulation* 2012;126:65–75.
5. Goyal P, Almarzooq ZI, Cheung J, et al. Atrial fibrillation and heart failure with preserved ejection fraction: Insights on a unique clinical phenotype from a nationally-representative United States cohort. *Int J Cardiol* 2018;266:112–118.
6. Marrouche NF, Kheirkhahan M, Brachmann J. Catheter ablation for atrial fibrillation with heart failure. *N Engl J Med* 2018;379:492.
7. Geng J, Zhang Y, Wang Y, et al. Catheter ablation versus rate control in patients with atrial fibrillation and heart failure: a multicenter study. *Medicine (Baltimore)* 2017;96:e9179.
8. Prabhu S, Taylor AJ, Costello BT, et al. Catheter ablation versus medical rate control in atrial fibrillation and systolic dysfunction: the CAMERA-MRI study. *J Am Coll Cardiol* 2017;70:1949–1961.
9. Kirchhof P, Camm AJ, Goette A, et al. Early rhythm-control therapy in patients with atrial fibrillation. *N Engl J Med* 2020;383:1305–1316.
10. Packer DL, Piccini JP, Monahan KH, et al. Ablation versus drug therapy for atrial fibrillation in heart failure: results from the CABANA trial. *Circulation* 2021; 143:1377–1390.
11. Cheng EP, Liu CF, Yeo I, et al. Risk of mortality following catheter ablation of atrial fibrillation. *J Am Coll Cardiol* 2019;74:2254–2264.
12. Black-Maier E, Ren X, Steinberg BA, et al. Catheter ablation of atrial fibrillation in patients with heart failure and preserved ejection fraction. *Heart Rhythm* 2018; 15:651–657.
13. Aldaas OM, Lupercio F, Darden D, et al. Meta-analysis of the usefulness of catheter ablation of atrial fibrillation in patients with heart failure with preserved ejection fraction. *Am J Cardiol* 2021;142:66–73.
14. Agency for Healthcare Research and Quality. Introduction to the HCUP Nationwide Readmissions Database (NRD). 2010–2015. Available from, [https://www.hcup-us.ahrq.gov/db/nation/nrd/Introduction\\_NRD\\_2010-2015.pdf](https://www.hcup-us.ahrq.gov/db/nation/nrd/Introduction_NRD_2010-2015.pdf).
15. Healthcare Cost and Utilization Project (HCUP). Overview of the Nationwide Readmissions Database (NRD). Available from, <http://www.hcup-us.ahrq.gov/nrdoverview.jsp>.
16. HCUP Quality Control Procedures Deliverable #1707.05. Available from, <https://www.hcup-us.ahrq.gov/db/quality.pdf>.
17. Melenovskiy V, Hwang SJ, Redfield MM, Zakeri R, Lin G, Borlaug BA. Left atrial remodeling and function in advanced heart failure with preserved or reduced ejection fraction. *Circ Heart Fail* 2015;8:295–303.
18. Aldaas OM, Malladi CL, Mylavarapu PS, et al. Comparison of outcomes after ablation of atrial fibrillation in patients with heart failure with preserved versus reduced ejection fraction. *Am J Cardiol* 2020;136:62–70.
19. Sugumar H, Nanayakkara S, Vizi D, et al. A prospective STudy using invAsive haemodynamic measurements foLLowing catheter ablation for AF and early HFpEF: STALL AF-HFpEF. *Eur J Heart Fail* 2021; 23:785–796.
20. Ablation Versus Medical Management of Atrial Fibrillation in HFpEF (AMPERE). Available from, <https://clinicaltrials.gov/ct2/show/NCT04282850>.
21. Catheter Ablation for Atrial Fibrillation in Preserved Ejection Fraction (AFFECT). Available from, <https://clinicaltrials.gov/ct2/show/NCT04317911>.
22. Reddy YNV, Carter RE, Obokata M, Redfield MM, Borlaug BA. A simple, evidence-based approach to help guide diagnosis of heart failure with preserved ejection fraction. *Circulation* 2018;138:861–870.
23. Pieske B, Tschöpe C, de Boer RA, et al. How to diagnose heart failure with preserved ejection fraction: the HFA-PEFF diagnostic algorithm: a consensus recommendation from the Heart Failure Association (HFA) of the European Society of Cardiology (ESC). *Eur Heart J* 2019;40:3297–3317.
24. Parameswaran R, Al-Kaisey AM, Kalman JM. Catheter ablation for atrial fibrillation: current indications and evolving technologies. *Nat Rev Cardiol* 2021; 18:210–225.
25. Loh P, van Es R, Groen MHA, et al. Pulmonary vein isolation with single pulse irreversible electroporation: a first in human study in 10 patients with atrial fibrillation. *Circ Arrhythm Electrophysiol* 2020;13:e008192.