

# Predictors of thirty-day readmission in nonagenarians presenting with acute heart failure with preserved ejection fraction: a nationwide analysis

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## ABSTRACT

**BACKGROUND** Acute heart failure with preserved ejection fraction (HFpEF) is a common but poorly studied cause of hospital admissions among nonagenarians. This study aimed to evaluate predictors of thirty-day readmission, in-hospital mortality, length of stay, and hospital charges in nonagenarians hospitalized with acute HFpEF.

**METHODS** Patients hospitalized between January 2016 and December 2018 with a primary diagnosis of diastolic heart failure were identified using ICD-10 within the Nationwide Readmission Database. We excluded patients who died in index admission, and discharged in December each year to allow thirty-day follow-up. Univariate regression was performed on each variable. Variables with  $P$ -value  $< 0.2$  were included in the multivariate regression model.

**RESULTS** From a total of 45,393 index admissions, 43,646 patients (96.2%) survived to discharge. A total of 7,437 patients (15.6%) had a thirty-day readmission. Mean cost of readmission was 43,265 United States dollars (USD) per patient. Significant predictors of thirty-day readmission were chronic kidney disease stage III or higher [adjusted odds ratio (aOR) = 1.20, 95% CI: 1.07–1.34,  $P = 0.002$ ] and diabetes mellitus (aOR = 1.18, 95% CI: 1.07–1.29,  $P = 0.001$ ). Meanwhile, female (aOR = 0.90, 95% CI: 0.82–0.99,  $P = 0.028$ ) and palliative care encounter (aOR = 0.27, 95% CI: 0.21–0.34,  $P < 0.001$ ) were associated with lower odds of readmission. Cardiac arrhythmia (aOR = 1.46, 95% CI: 1.11–1.93,  $P = 0.007$ ) and aortic stenosis (aOR = 1.36, 95% CI: 1.05–1.76,  $P = 0.020$ ) were amongst predictors of in-hospital mortality.

**CONCLUSIONS** In nonagenarians hospitalized with acute HFpEF, thirty-day readmission is common and costly. Chronic comorbidities predict poor outcomes. Further strategies need to be developed to improve the quality of care and prevent the poor outcome in nonagenarians.

By 2030, it is estimated that one every thirty-three patients will have the diagnosis of heart failure (HF). The projected cost estimates of treating HF are 160 billion United States dollars (USD) in direct costs. Because of the aging of the population, greater increase in HF prevalence will be seen in older adults. It is projected that the

number of patients  $> 80$  years with HF will grow by 66% by 2030.<sup>[1]</sup>

HF incidence and prevalence rise dramatically with age due to structural and functional alterations in the cardiovascular system, making HF the most prevalent cardiovascular disease among elderly. HF was reported to be the second leading

cause of hospitalization for patients aged 75 years and above from 2013 to 2018.<sup>[2]</sup>

Most elderly patients with HF have impaired left ventricular diastolic function without significant impairment in left ventricular systolic function, which is called heart failure with preserved ejection fraction (HFpEF).<sup>[3-6]</sup> Increased levels of brain natriuretic peptide, older age, myocardial infarction history, and reduced diastolic function make the prognosis of HFpEF worse.<sup>[7-9]</sup>

Over the years, there have been advances in the treatment of HF, however, the mortality, hospitalization, and readmission rates are still high.

In this study, we aimed to assess the predictors and causes of readmissions with acute HFpEF among nonagenarians in the United States, by using the National Readmission Database (NRD).

## METHODS

### Data Source

This is a retrospective cohort study using the Agency for Healthcare Research and Quality's Healthcare Cost and Utilization Project (HCUP) NRD from January 2016 to December 2018.<sup>[10]</sup> The NRD is the largest publicly available all-payer inpatient health care readmission database in the United States. The NRD is drawn from HCUP State Inpatient Databases containing verified patient linkage numbers that can be used to track a person across hospitals within a State, while adhering to strict privacy guidelines. Unweighted, the NRD contains data from approximately 18 million discharges in the United States each year. Weighted, it estimates roughly 35 million discharges in the United States each year.

The NRD contains both patient and hospital-level information. Up to forty discharge diagnoses and twenty-five procedures are collected for each patient using the International Classification of Diseases, Tenth Revision, Clinical Modification (ICD-10-CM). Patients were tracked during the same year using the variable "nrd\_visitlink", and time between two admissions was calculated using variable "nrd\_daystoevent". National estimates were produced using sampling weights provided by the sponsor. All values presented are weighted estimates.

### Study Population

Our study population was patients aged 90 years and above admitted between January 2016 and December 2018 with a primary diagnosis of diastolic HF (ICD-10 codes: I50.30, I50.31, and I50.33). Unfortunately, no ICD codes exclusively existed for HFpEF for our patient population. We excluded patients with systolic failure or combined systolic/diastolic HF, and patients who were discharged in December each year to allow thirty-day follow-up. Patients who died in index admission were excluded from evaluating readmission outcomes but included in other secondary outcomes pertaining to index admission only.

NRD variables were used to identify patients and hospital characteristics. Patient characteristics included age, gender, median household income, and primary insurance. Hospital characteristics included hospital bed size and teaching status. ICD-10 codes used in our analysis are included in Table 1.

In accordance with the HCUP data use agreement, we excluded any variable containing a small number of observations ( $\leq 10$ ) that could pose risk of person identification or data privacy violation.

### Missing Data

Data on median household income and primary insurance were missing in 1.09% and 0.06% of hospitalizations, respectively. Other key variables had no missing values. In-hospital mortality and total charges outcomes were missing in 0.06% and 1.54% of hospitalizations, respectively. All hospitalizations with missing values were excluded from our analysis.

### Study Outcomes

Primary outcome was thirty-day readmission. Secondary outcomes were in-hospital mortality, length of stay (LOS), and total hospital charges in index admission. In-hospital mortality, LOS, and total hospital charges were directly coded in NRD.

Index admission was defined as the first admission with the primary diagnosis of diastolic HF without prior admission in the thirty-day period. A readmission was defined as any readmission within thirty days of the index admission. For patients who



**Table 1** The ICD-10-CM used to identify key variables.

Diseases	ICD-10 codes
Acute or acute on chronic diastolic heart failure	I50.30, I5031, I5033
Non-diastolic heart failure (exclusion)	I50.40, I50.41, I50.42, I50.43, I50.20, I50.21, I50.22, I50.23, I50.81, I50.810, I50.811, I50.813, I50.814
Myocardial infarction	I21.x, I22.x, I25.2
Cardiac arrhythmias	I44.1-I44.3, I45.6, I45.9, I47.x-I49.x, R00.0, R00.1, R00.8, Z95.0
Pulmonary circulation disorders	I26.x, I27.x, I28.0, I28.8, I28.9
Peripheral vascular disorders	I70.x, I71.x, I72.x, I73.1, I73.8, I73.9, I77.1, I77.7, I79.0, I79.1, I79.8, I79.2, K55.1, K55.8, K55.9, Z95.8, Z95.9
Chronic pulmonary disease	I27.8, I27.9, J40.x-J47.x, J60.x-J67.x, J68.4, J70.1, J70.3, J84, J96.1
Diabetes mellitus	E08.9, E09.9, E10.9, E11.9, E13.9, E08.2-E08.8, E09.x, E10.2-E10.8, E11.2-E11.8, E12.2-E12.8, E13.2-E13.8
Hypothyroidism	E00.x-E03.x
Obesity	E66.x, Z68.3, Z68.4, Z68.5
Liver disease	B18.x, I85.x, K70.x, K71.1, K71.3-K71.5, K71.7, K72.x-K74.x, K75.4, K75.8, K76.0, K76.2-K76.9, Z94.4
Peptic ulcer disease, no bleeding	K25.5, K25.7, K25.9, K26.5, K26.7, K26.9, K27.5, K27.7, K27.9, K28.5, K28.7, K28.9
Lymphoma	C81.x-C86.x, C88.x, C90.0, C90.2, C90.3, C96.x, D47.Z9
Metastatic cancer	C77.x-C80.x, R18.0
Rheumatoid arthritis/collagen, vascular disease	L94.0, M32.x, L94.1, M35.x, L94.3, M45.x, M05.x, M46.5, M06.x, M46.1, M08.x, M46.8, M12.0, M46.9, M12.3, M48.8, M49.x, M30.x, M31.0 8-M31.3
Fluid and electrolytes disorders	E22.2
Coagulopathy	D66.x-D68.x, D69.1, D69.3-D69.6
Obesity	E66.x, Z68.3, Z68.4, Z68.5
Alcohol abuse	F10, E52, G62.1, I42.6, K29.2, K70.0, K70.3, K70.9, T51.x, Z71.4
Drug abuse	F11.x-F16.x, F18.x, F19.x, Z71.5
Psychosis	F20.x, F22.x-F25.x, F28.x, F29.x, F30.1, F30.2, F31.2, F31.6, F44.8
Depression	F20.4, F31.3-F31.5, F32.x, F33.x, F34.1, F41.2, F43.2
Prior myocardial infarction	I25.2
Prior percutaneous coronary intervention	Z98.61, Z95.5
Prior coronary artery bypass grafting	Z95.1
Chronic kidney disease stage III or higher	N18.3, N18.30, N18.31, N18.32, N18.4, N18.5, N18.6, N18.9
Atrial fibrillation	I48, I48.0, I48.1, I48.11, I48.19, I48.2, I48.20, I48.21, I48.91
Palliative care encounter	Z51.5
Blood transfusion	30230N0, 30230N1, 30233N0, 30233N1, 30240N0, 30240N1, 30243N0, 30243N1, 30233P0, 30233P1, 30230P0, 30230P1, 30240P0, 30240P1, 30243P0, 30243P1
Dyslipidemia	E78.5

ICD-10-CM: International Classification of Diseases, Tenth Edition, Clinical Modification.

were readmitted multiple times during the thirty-day post admission, only the first readmission was included.

### Statistical Analysis

Data analysis was performed using STATA 17.0 (StataCorp, College Station, Texas, USA). Data were

expressed as a percentage for categorical variables and mean  $\pm$  SD for continuous variables. Univariate regression analysis was used to calculate unadjusted odds ratio for the primary and secondary outcomes. Multivariate regression analysis was used to adjust for the potential confounders and calculate adjusted odds ratio (aOR). A logistic regres-



sion model was used for binary outcome and linear regression for continuous outcome. The models were built by including the variables that were associated with the outcome of interest on univariable regression analysis with a cut-off *P*-value of 0.20. Continuous variables were compared using the independent Student's *t*-test and categorical variables were compared using the Pearson's chi-squared test. All statistical tests were two-sided, and *P*-value < 0.05 was considered statistically significant.

## RESULTS

### Patient Characteristics

From 107 million discharges included in NRD from January 2016 to December 2018, our cohort included 45,393 index admissions of whom 43,646 patients (96.2%) survived to discharge. A total of 7,437 patients were readmitted in thirty-day period post discharge from index hospitalization. Baseline characteristics were stratified according to readmission status.

Female constituted 70.3% of readmitted patients. Medicare was the primary insurance in both groups (*P* = 0.042). Patients who were readmitted were more likely to have chronic ischemic heart disease (40.8% vs. 36.8%, *P* < 0.001), chronic kidney disease (CKD) stage III or higher (21.8% vs. 16.8%, *P* < 0.001), chronic pulmonary disease (33.2% vs. 29.8%, *P* < 0.001), diabetes mellitus (25.6% vs. 21.6%, *P* < 0.001), and hypertension (77.7% vs. 75.2%, *P* = 0.002). Readmitted patients had less palliative care encounter (2.3% vs. 7.8%, *P* < 0.001). Other patient and hospital characteristics are included in Table 2.

### Thirty-day Readmission

Of 43,646 patients who survived to discharge from index admission, 7,437 patients (17.0%) were readmitted within thirty days. Of those readmitted, 24 patients (0.32%) were discharged to a nursing facility. The mean cost of readmission was 43,265 USD per patient. Mean LOS of readmission was 5.46 days. Readmission due to cardiovascular etiologies constituted 49% of all readmissions followed by respiratory etiologies (13%) and infectious

etiologies (9%). The most common specific causes of readmission were HF (37%) followed by sepsis (8%) and pneumonia (6%). Etiologies of readmission are presented in Figure 1.

Independent predictors of readmission were admission to teaching hospital (aOR = 1.09, 95% CI: 1.01–1.18, *P* = 0.021), chronic ischemic heart disease (aOR = 1.11, 95% CI: 1.02–1.22, *P* = 0.022), CKD stage III or higher (aOR = 1.20, 95% CI: 1.07–1.34, *P* = 0.002), chronic pulmonary disease (aOR = 1.14, 95% CI: 1.05–1.23, *P* = 0.001), diabetes mellitus (aOR = 1.18, 95% CI: 1.07–1.29, *P* = 0.001), fluid and electrolyte disorders (aOR = 1.13, 95% CI: 1.05–1.22, *P* = 0.002), and LOS greater than two days (aOR = 1.20, 95% CI: 1.09–1.32, *P* < 0.001). Female (aOR = 0.90, 95% CI: 0.82–0.99, *P* = 0.028), and palliative care encounter (aOR = 0.27, 95% CI: 0.21–0.34, *P* < 0.001), were independently associated with decreased odds of readmission (Table 3).

### In-hospital Mortality

A total of 1,727 patients died in index hospitalization. Independent predictors of in-hospital mortality were private insurance (aOR = 2.07, 95% CI: 1.25–3.44, *P* = 0.005), acute myocardial infarction (aOR = 1.40, 95% CI: 1.11–1.75, *P* = 0.004), CKD stage III or higher (aOR = 1.26, 95% CI: 1.01–1.58, *P* = 0.042), cardiac arrhythmias (aOR = 1.46, 95% CI: 1.11–1.93, *P* = 0.007), pulmonary circulation disorder (aOR = 1.27, 95% CI: 1.07–1.51, *P* = 0.006), paralysis (aOR = 3.81, 95% CI: 1.61–9.03, *P* = 0.002), liver disease (aOR = 2.06, 95% CI: 1.27–3.35, *P* = 0.003), weight loss (aOR = 2.01, 95% CI: 1.63–2.49, *P* < 0.001), fluid and electrolyte disorders (aOR = 2.05, 95% CI: 1.77–2.37, *P* < 0.001), and aortic stenosis (aOR = 1.36, 95% CI: 1.05–1.76, *P* = 0.020). Paradoxically, history of percutaneous coronary intervention (aOR = 0.61, 95% CI: 0.43–0.87, *P* = 0.007), and dyslipidemia (aOR = 0.75, 95% CI: 0.64–0.88, *P* < 0.001) were associated with lower odds of in-hospital mortality (Table 4).

### Length of Stay

Mean LOS in our cohort was 4.72 days in index admission. Teaching hospital and large hospital size were associated with mean increased LOS of 0.29 days, and 0.44 days, respectively. Palliative care encounter was similarly associated with mean in-

Table 2 Baseline characteristics according to readmission status.

Characteristics	Thirty-day readmission		P-value
	No readmission (n = 36,209)	Readmission (n = 7,437)	
Female	26,470 (73.1%)	5,226 (70.3%)	0.001
Median household income quartile for zip code in percentile			0.869
< 25 <sup>th</sup>	7,158 (19.8%)	1,485 (20.0%)	
25 <sup>th</sup> -50 <sup>th</sup>	9,732 (26.9%)	1,950 (26.2%)	
50 <sup>th</sup> -75 <sup>th</sup>	9,980 (27.6%)	2,069 (27.8%)	
> 75 <sup>th</sup>	8,948 (24.7%)	1,868 (25.1%)	
Insurance			0.042
Medicare	34,750 (96.0%)	7,203 (96.9%)	
Medicaid	159 (0.4%)	29 (0.4%)	
Private	919 (2.5%)	159 (2.1%)	
Self-pay/other/no charge	360 (1.0%)	47 (0.6%)	
Hospital bed size			0.680
Small	8,241 (22.7%)	1,644 (22.1%)	
Medium	10,716 (29.6%)	2,195 (29.5%)	
Large	17,253 (47.6%)	3,598 (48.4%)	
Teaching hospital	20,668 (57.1%)	4,402 (59.2%)	0.028
Admission on weekend	8,953 (24.7%)	1,920 (25.8%)	0.205
Comorbidities			
Cardiac arrhythmias	25,909 (71.6%)	5,414 (72.8%)	0.175
Peripheral vascular disease	4,177 (11.5%)	881 (11.8%)	0.607
Atrial fibrillation	22,629 (62.5%)	4,772 (64.2%)	0.084
Aortic stenosis	4,962 (13.7%)	1,113 (15.0%)	0.058
Hypertension	27,221 (75.2%)	5,781 (77.7%)	0.002
Diabetes mellitus	7,826 (21.6%)	1,904 (25.6%)	< 0.001
Chronic kidney disease stage III-V	6,088 (16.8%)	1,624 (21.8%)	< 0.001
Pulmonary circulation disorder	9,724 (26.9%)	2,019 (27.1%)	0.731
Chronic ischemic heart	13,314 (36.8%)	3,038 (40.8%)	< 0.001
History of percutaneous coronary intervention	2,646 (7.3%)	613 (8.2%)	0.053
History of myocardial infarction	3,135 (8.7%)	661 (8.9%)	0.661
History of coronary artery bypass graft	3,199 (8.8%)	696 (9.4%)	0.313
Chronic lung disease	10,792 (29.8%)	2,469 (33.2%)	< 0.001
Obesity	1,788 (4.7%)	383 (4.8%)	0.632
Dyslipidemia	14,442 (39.9%)	2,950 (39.7%)	0.822
Weight loss	2,576 (7.1%)	568 (7.6%)	0.298
Fluid and electrolyte disorder	12,349 (34.1%)	2,723 (36.6%)	0.004
Hypothyroidism	10,554 (29.1%)	2,217 (29.8%)	0.422
Coagulopathy	2,376 (6.6%)	469 (6.3%)	0.591
Chronic blood loss anemia	287 (0.8%)	70 (0.9%)	0.378
Solid tumor without metastasis	764 (2.1%)	136 (1.8%)	0.284
Rheumatoid arthritis/collagen vascular disorders	1,062 (2.9%)	215 (2.8%)	0.905
Depression	3,467 (9.6%)	699 (9.4%)	0.759
Blood transfusion	753 (2.1%)	36 (0.5%)	0.730
Palliative care encounter	2,840 (7.8%)	170 (2.3%)	0.000
Length of stay > 2 d	27,827 (76.9%)	5,978 (80.4%)	0.000

Data are presented as n (%).



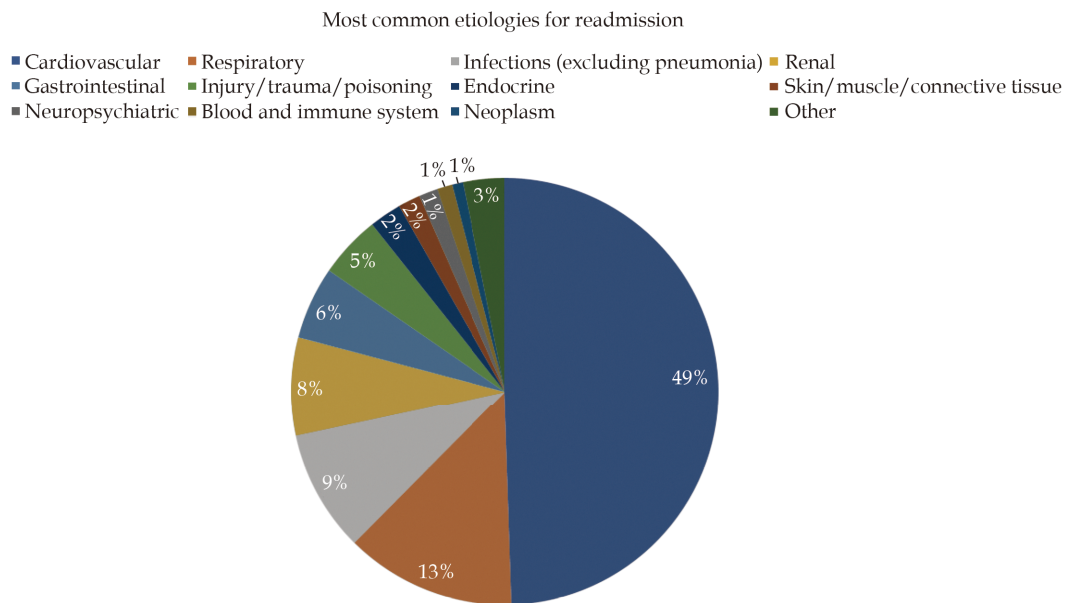


Figure 1 Etiologies of readmission.

Table 3 Predictors of thirty-day readmission.

Predictor	Adjusted OR	95% CI		P-value
		Lower limit	Upper limit	
Female	0.90	0.82	0.99	0.028
Teaching hospital	1.09	1.01	1.18	0.021
Chronic ischemic heart disease	1.11	1.02	1.22	0.022
Chronic kidney disease stage III or higher	1.20	1.07	1.34	0.002
Chronic pulmonary disease	1.14	1.05	1.23	0.001
Diabetes mellitus	1.18	1.07	1.29	0.001
Fluid and electrolyte disorders	1.13	1.05	1.22	0.002
Length of stay > 2 d	1.20	1.09	1.32	< 0.001
Palliative care encounter	0.27	0.21	0.34	< 0.001

CI: confidence interval; OR: odds ratio.

creased LOS of 0.79 days. Other patient characteristics such as chronic ischemic heart disease, CKD stage III or higher, pulmonary hypertension, chronic pulmonary disease, weight loss, were independently associated with mean increased LOS. All differences mentioned above were statistically significant ( $P < 0.05$ ) (Table 5).

### Total Hospital Charges

Mean total hospital charges in our cohort was 32,554 USD in index admission. Size of the hospital (moderate and large) and teaching hospitals were independent predictors of increased cost of hospitalization. Medicaid insurance and palliative care en-

counter were similarly associated with increased cost. Other patient characteristics such as CKD stage III or higher, cardiac arrhythmias, pulmonary circulation disorders, peripheral vascular disorders, liver disease, fluid and electrolyte disorders, blood loss anemia, and aortic stenosis were amongst predictors of increased hospitalization cost (Table 6).

### DISCUSSION

Patients above the age of 85 years constitute more than 9% of patients admitted to hospitals in the United States.<sup>[11]</sup> Hospitalizations and health care spending for elderly are expected to rise as the pop-

**Table 4 Predictors of in-hospital mortality outcome.**

Predictor	Adjusted OR	95% CI		P-value
		Lower limit	Upper limit	
Private insurance	2.07	1.25	3.44	0.005
Acute myocardial infarction	1.40	1.11	1.75	0.004
Chronic kidney disease stage III or higher	1.26	1.01	1.58	0.044
Cardiac arrhythmias	1.46	1.11	1.93	0.007
History of percutaneous coronary intervention	0.61	0.43	0.87	0.007
Dyslipidemia	0.75	0.64	0.88	< 0.001
Aortic stenosis	1.36	1.05	1.76	0.020
Pulmonary circulation disorder	1.27	1.07	1.51	0.006
Liver disease	2.06	1.27	3.35	0.003
Weight loss	2.01	1.63	2.49	< 0.001
Fluid and electrolyte disorders	2.05	1.77	2.37	< 0.001

CI: confidence interval; OR: odds ratio.

**Table 5 Predictors of increased length of stay outcome.**

Predictor	Coefficient	95% CI		P-value
		Lower limit	Upper limit	
Admission in weekend	-0.1815489	-0.2946481	-0.0684497	0.002
Hospital bed size				
Large	0.4448497	0.2913729	0.5983266	< 0.001
Median household income quartile for zip code in percentile				
< 25 <sup>th</sup>	Reference	Reference	Reference	
50 <sup>th</sup> -75 <sup>th</sup>	-0.1998943	-0.361739	-0.0380496	0.015
Teaching hospital	0.2929085	0.1721228	0.4136942	< 0.001
Chronic ischemic heart disease	0.1744351	0.0285726	0.3202976	0.019
Chronic kidney disease stage III or higher	0.3922964	0.2345545	0.5500383	< 0.001
Pulmonary hypertension	1.599025	1.108897	2.089153	< 0.001
Cardiac arrhythmias	0.2393916	0.0443117	0.4344714	0.016
Pulmonary circulation disorder	0.3753799	0.2564289	0.4943308	< 0.001
Chronic pulmonary disease	0.5235521	0.4129308	0.6341734	< 0.001
Diabetes mellitus	0.2975333	0.104529	0.4905376	0.003
Coagulopathy	0.6498321	0.4151741	0.8844902	< 0.001
Obesity	0.5458434	0.3242828	0.7674041	< 0.001
Weight loss	1.288857	1.012733	1.564981	< 0.001
Fluid and electrolyte disorders	1.260683	1.129377	1.391989	< 0.001
Depression	0.2089386	0.0207869	0.3970904	0.030
History of percutaneous coronary intervention	-0.3453232	-0.5227807	-0.1678657	< 0.001
History of coronary artery bypass graft	-0.205737	-0.3938539	-0.0176201	0.032
Blood transfusion	1.896131	1.42946	2.362801	< 0.001
Palliative care encounter	0.7950352	0.5829619	1.007109	< 0.001

CI: confidence interval.

ulation continues to age. Disease-specific interventions are not well studied in elderly population.<sup>[12]</sup> Several studies have documented predictors of read-

mission of HF in the general population.<sup>[13-17]</sup> However, few small studies evaluated HF in general or HFpEF in elderly population.<sup>[18,19]</sup> Our study is the largest



Table 6 Predictors of total hospital charges outcome.

Predictor	Coefficient	95% CI		P-value
		Lower limit	Upper limit	
Medicare insurance	15,215.58	6,895.664	23,535.5	< 0.001
Hospital bed size				
Small	Reference	Reference	Reference	
Medium	4,446.771	2,346.773	6,546.77	< 0.001
Large	11,161.2	8,971.176	13,351.23	< 0.001
Median household income quartile for zip code in percentile				
> 75 <sup>th</sup>	5,876.8	3,440.12	8,313.48	< 0.001
Teaching hospital	4,286.967	2,416.891	6,157.044	< 0.001
Chronic kidney disease stage III or higher	3,319.49	1,715.971	4,923.009	< 0.001
Cardiac arrhythmias	3,729.229	1,962.791	5,495.667	< 0.001
Pulmonary circulation disorder	2,534.161	1,380.285	3,688.036	< 0.001
Peripheral vascular disorder	3,540.677	1,819.354	5,262.000	< 0.001
Complicated hypertension	2,131.445	795.0957	3,467.794	0.002
Atrial fibrillation	-2,397.392	-4,154.082	-640.7016	0.007
Aortic stenosis	2,386.994	865.6449	3,908.343	0.002
Chronic pulmonary disease	7,383.334	6,307.357	8,459.312	< 0.001
Paralysis	34,052.55	9,375.435	58,729.65	0.007
Uncomplicated diabetes mellitus	2,092.701	631.921	3,553.481	0.005
Complicated diabetes mellitus	3,564.021	1,787.301	5,340.741	< 0.001
Liver disease	6,318.949	1,322.822	11,315.08	0.013
Coagulopathy	6,340.614	3,975.907	8,705.321	< 0.001
Obesity	3,353.469	1,309.475	5,397.464	0.001
Weight loss	11,092.86	8,198.294	13,987.42	< 0.001
Fluid and electrolyte disorders	9,141.854	8,005.137	10,278.57	< 0.001
Palliative care encounter	4,673.144	2,784.688	6,561.599	< 0.001
Blood transfusion	19,899.15	14,073.98	25,724.31	< 0.001

CI: confidence interval.

and first to report data exclusively in nonagenarians presenting with HFpEF.

In our analysis, we identified several independent predictors of readmission, in-hospital mortality, increased LOS, and total hospital costs in nonagenarians presenting with acute or acute on chronic HFpEF.

We observed a 17% thirty-day readmission rate in HFpEF nonagenarian population, which was comparable to other previous studies that documented thirty-day readmission rates from 18% to 25%.<sup>[15,20-22]</sup> Cardiovascular etiologies were responsible for 49% of readmissions, particularly HF (37%), followed by

pulmonary etiologies (17%), pneumonia (6%), infectious etiologies (9%), and renal etiologies (7%). General etiologies of readmissions were similar to a study done by Arora, *et al.*<sup>[20]</sup> However, a higher percentage of HF readmissions was observed in our analysis which was done exclusively in nonagenarians. Our population had a high burden of chronic comorbidities, which likely have impacted readmission outcomes. We found chronic ischemic heart disease, CKD stage III or higher, chronic pulmonary disease, and diabetes mellitus to be independent predictors of readmission in nonagenarians. Although females constituted the majority of our



cohort (72.6%), female was associated with less readmission odds, which was observed by Stolfo, *et al.*<sup>[23]</sup> in a prior study. In contrast to a prior study done using NRD,<sup>[20]</sup> blood loss anemia, packed red blood cells transfusion, and discharge to a nursing facility were not found to be independent predictors of readmission in nonagenarians. LOS greater than two days in index admission predicted readmission. This could be explained by the higher comorbidity burden in this age group. Our study demonstrated the strong impact of palliative care encounter on prevention of future readmission although it was poorly utilized (only 6.9% of our cohort received palliative care service). This finding could open avenues for palliative care utilization in this age group with emphasis on quality of life rather than quantity.

In-hospital mortality rate in index admission was 3.8% in our cohort, which is close to average mortality in hospitalized patients aged 75 years and higher (4.3%–4.6%).<sup>[24]</sup>

Compared to readmission predictors, chronic comorbidities such as cardiac arrhythmias, aortic stenosis, liver disease, pulmonary circulatory disorders, and CKD stage III or higher were independently associated with increased odds of in-hospital mortality. Interestingly, dyslipidemia and history of percutaneous coronary intervention were associated with lower odds of in-hospital mortality, which was thought to be due to prescribed statins and other goal-directed medical therapy for coronary artery disease. However, Organized Program to Initiate Lifesaving Treatment in Hospitalized Patients with Heart Failure (The OPTIMIZE-HF) study on 48,612 hospitalized patients with acute HF showed similar results regarding hyperlipidemia despite that only 66% of patients diagnosed with hyperlipidemia were on statins or other lipid lowering therapy.<sup>[16]</sup>

Chronic comorbidities were also identified as predictors of increased LOS and total hospital charges such as hypertension, diabetes mellitus, and chronic pulmonary disease. Blood transfusion was associated with increased LOS and increased total hospital charges, but it did not affect readmission nor in-hospital mortality outcomes. Readmission within thirty days was more costly on average compared to index admission (mean cost: 43,265 USD vs. 32,554

USD), which is likely due to increased LOS in the readmitted cohort (mean LOS: 5.46 days vs. 4.72 days). It is worth mentioning that discharge to nursing facilities was higher in the second admission compared to the index admission (0.32% vs. 0.15%,  $P = 0.29$ ), which probably added to the overall health care cost.

## LIMITATIONS

Our study has certain limitations. Firstly, NRD uses ICD codes for diagnosis, which is subject to coding errors. Secondly, the differentiation between volume overload due to HFpEF and advanced CKD can be challenging. Both conditions often co-exist, and we are unable to differentiate between the primary disease processes driving the hospitalization. The primary outcome of our study is the rate of thirty-day readmission post-discharge and in-hospital mortality may be a competing risk endpoint, particularly in this age group, thus assessing the composite endpoint of thirty-day readmission or death would be an area of future research. We cannot identify patients who may have expired without being re-hospitalized in our database. The information pertaining to the longitudinal follow-up of patients, information related to race, ethnicity, individual operator, and procedure level is also not available in the NRD. Moreover, factors influencing patient prognosis such as medications and echocardiography findings such as diastolic grading are absent. Last but not least, the study was retrospective, which is subject to confounding bias not typically seen in prospective trials.

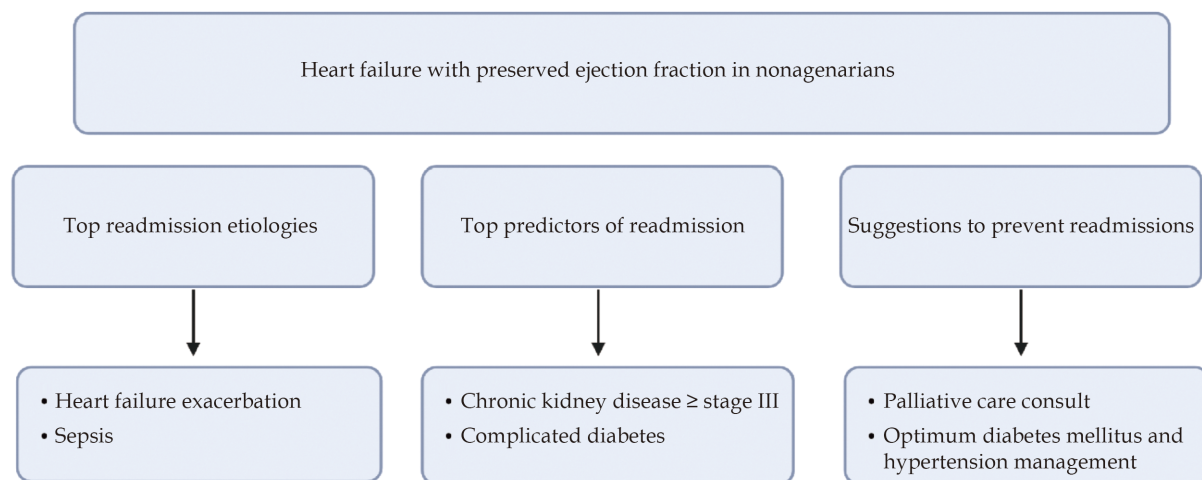
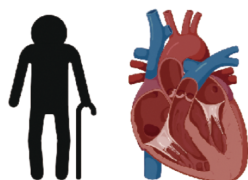
## CONCLUSIONS

We identified several predictors of thirty-day readmissions, in-hospital mortality, increased LOS, and hospitalization cost amongst nonagenarians admitted with HFpEF. Having knowledge of these predictors should help guide further strategies targeting reduction of readmissions, decreasing healthcare costs, and improving the quality of care patients receive (Figure 2).

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**Figure 2 Central illustration.**

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