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## Cardiovascular Outcomes and Trends of Transcatheter vs. Surgical Aortic Valve Replacement Among Octogenarians With Heart Failure: A Propensity Matched National Cohort Analysis

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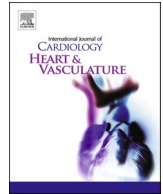
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# Cardiovascular outcomes and trends of Transcatheter vs. Surgical aortic valve replacement among octogenarians with heart failure: A Propensity Matched national cohort analysis

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## ABSTRACT

**Background:** Heart failure (HF) is a complex clinical syndrome with symptoms and signs that result from any structural or functional impairment of ventricular filling or ejection of blood. Limited data is available regarding the in-hospital outcomes of TAVR compared to SAVR in the octogenarian population with HF.

**Methods:** The National Inpatient Sample (NIS) database was used to compare TAVR versus SAVR among octogenarians with HF. The primary outcome was in-hospital mortality. The secondary outcome included acute kidney injury (AKI), cerebrovascular accident (CVA), post-procedural stroke, major bleeding, blood transfusions, sudden cardiac arrest (SCA), cardiogenic shock (CS), and mechanical circulatory support (MCS).

**Results:** A total of 74,995 octogenarian patients with HF (TAVR-HF n = 64,890 (86.5%); SAVR n = 10,105 (13.5%)) were included. The median age of patients in TAVR-HF and SAVR-HF was 86 (83–89) and 82 (81–84) respectively. TAVR-HF had lower percentage in-hospital mortality (1.8% vs. 6.9%;  $p < 0.001$ ), CVA (2.5% vs. 3.6%;  $p = 0.009$ ), SCA (9.9% vs. 20.2%;  $p < 0.001$ ), AKI (17.4% vs. 40.8%;  $p < 0.001$ ), major transfusion (26.4% vs 67.3%;  $p < 0.001$ ), CS (1.8% vs 9.8%;  $p < 0.001$ ), and MCS (0.8% vs 7.3%;  $p < 0.001$ ) when compared to SAVR-HF. Additionally, post-procedural stroke and major bleeding showed no significant difference. The median unmatched total charges for TAVR-HF and SAVR-HF were 194,561\$ and 246,100\$ respectively.

**Conclusion:** In this nationwide observational analysis, TAVR is associated with an improved safety profile for octogenarians with heart failure (both preserved and reduced ejection fraction) compared to SAVR.

**Abbreviations:** TAVR, Transcatheter aortic valve replacement; SAVR, Surgical aortic valve Replacement; HFrEF, Heart Failure with reduced Ejection fraction; HFpEF, Heart Failure with a Preserved ejection fraction; AS, Aortic Stenosis; CVD, cardiovascular disease; RCT, Randomized Controlled Trial; HF, Heart Failure; AKI, Acute Kidney Injury; CVA, Cerebrovascular Accident; LVAD, Left ventricular assist device; LOS, Length of hospital stay; SCA, sudden cardiac arrest; CS, Cardiogenic Shock; MCS, Mechanical Circulatory Support; PSM, Propensity Matched.

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## 1. Introduction

Aortic stenosis (AS) is the most common valvular heart disease in the United States' geriatric population [1]. According to the European and American guidelines, the prevalence of severe AS reaches up to 10% among patients aged 80 and older [2]. Transcatheter aortic valve replacement (TAVR) is a favorable alternative over surgical aortic valve replacement (SAVR) in patients aged 75 and older who; have low, intermediate, or high Society of Thoracic Surgeons (STS) scores or a EuroSCORE II score  $\geq 4\%$ ; and have a history of organ dysfunction according to the European Society of Cardiology (ESC) and the American College of Cardiology (ACC) [3]. According to current guidelines, in octogenarian patients (individuals aged 80–89), surgical risk constitutes the primary factor in determining the therapeutic route as assessed by a multidisciplinary heart team [2,3]. However, the surgical risk may be underestimated in elderly patients due to a lower physiologic reserve, thus providing a rationale for suggesting TAVR over SAVR in this population. Furthermore, TAVR has proven to have similar outcomes compared to surgical repair in severe AS in high and intermediate-risk surgical patients [4]. The less invasiveness of the procedure and faster recovery are likely to be of significant advantage in the octogenarian population.

Cardiovascular Health Study (CHS) criteria can be used to identify higher-risk surgical candidates. Frailty is a geriatric syndrome resulting in decline across multiple physiological systems and serves as a predictor of operative complications and mortality, especially in the context of cardiovascular disease (CVD). Per Kotajarvi et al. and Green et al., despite generally comparable age, disease severity, cardiac function, and comorbid disease burden, re-hospitalizations and death were twice as common in frail compared to non-frail older adults receiving SAVR or TAVR [5,6]. In summary, frailty is prevalent in older adults with severe AS and is associated with increased adverse outcomes and mortality risk following both SAVR and TAVR [5,6].

Heart Failure (HF), as per ACC/AHA heart failure 2022 guidelines, is a complex clinical syndrome with symptoms and signs that result from any structural or functional impairment of ventricular filling or ejection of blood [7]. Guideline-directed medical treatment (GDMT) of HF is directed toward neurohormonal modulation and afterload reduction. However, medical treatment alone does not address the mechanical increase in afterload related to the stenotic valve. Current randomized control trials (RCT) are underway to evaluate the value of unloading the left ventricle through TAVR [8]. In the current study, we used the National Inpatient Sample (NIS) database to evaluate and compare clinical outcomes in octogenarian patients with HF who underwent either TAVR or SAVR. We further investigated whether differences in clinical outcomes exist between patients with a reduced ejection fraction (HFrEF) and those with a preserved ejection fraction (HFpEF) in either group.

## 2. Methods

### 2.1. Data source

We analyzed data from the NIS database from 2015 to 2018. NIS is part of the healthcare cost and utilization project (HCUP) databases. The Agency for Healthcare Research and Quality (AHRQ) sponsors these databases [9]. The NIS database represents nearly 95% of the US population and includes 20% of discharge patient data from nearly 1000 hospitals. The NIS undergoes annual quality assessments confirming its internal validity. Additionally, the NIS is a publicly available database with de-identified data; therefore, Institutional Review Board approval was not required for our study.

### 2.2. Study population

We selected a HF cohort of TAVR and SAVR using the International Classification of Disease, Tenth Edition, Clinical Modification (ICD-10-

CM) codes for demographics, baseline comorbidities, matching variables, and outcomes. The codes to generate cohorts are summarized in **Supplemental S1**. Further, TAVR-HF and SAVR HF were created using baseline TAVR and SAVR index cases in NIS samples. HFrEF and HFpEF are defined per 2022 ACC/AHA/Heart failure guidelines as left ventricular ejection fraction (LVEF)  $\leq 40\%$  and LVEF  $\geq 50\%$  respectively. Furthermore, the guideline also further sub-classify HF into HF with mildly reduced EF (HFmrEF) and HF with improved EF (HFimpEF) as LVEF 41%–49% and previous LVEF  $< 40\%$  and a follow up measurement of LVEF  $> 40\%$  [7].

The cohort selection flow diagram is shown in **Fig. 1**. Both STS and Euroscore II cannot be calculated from the NIS. As a result, we used the elixhauser index, AHRQ risk severity, and mortality index to calculate the combined estimate of the patients' risk profiles [10].

The inclusion criteria for our study consisted of patients aged 80–90 with a history of HF who underwent TAVR or SAVR. Patients were then sub-grouped into HFrEF and HFpEF groups. Exclusion criteria consisted of all patients under 80 or greater than 90 years of age. Octogenarians who received both SAVR and TAVR or had AS in the setting of congenital rheumatic heart disease were also excluded.

### 2.3. Outcomes measured

The primary outcome was in-hospital mortality. Secondary outcomes included acute kidney injury (AKI), cerebrovascular accident (CVA), post-procedural stroke, major bleeding-bleeding as defined by the Valve Academic Research Consortium (VARC), blood transfusions, sudden cardiac arrest (SCA), cardiogenic shock (CS), mechanical circulatory support (MCS: including left ventricular assist device, pVAD, and ECMO). Tertiary outcomes included quality measures such as length of hospital stay (LOS) and cost of hospitalization. The common variable definitions are shown in **Supplementary S2**.

### 2.4. Statistical analysis

Categorical variables were reported as frequencies with percentages using Pearson's chi-square test and compared using logistic regression for accurate documentation. In contrast, continuous variables were reported as weighted means with standard deviation (normal distribution) or median with interquartile ranges (IQR) for skewed distribution. Outcome's frequency and percentages of the unmatched cohort were reported using Pearson's chi-square test and logistic regression. Propensity matching (PSM) was done using *Entropy near matching balance* for mean, median, and skewness weighted using the STATA *ebalance* module (**Supplemental S3**). PSM Entropy balance is superior to another propensity matching of any kind including nearest neighbor matching, pruning, or inverse probability treatment weighting [11]. PSM was done for cohorts TAVR-HF and SAVR-HF. The matching was done to eliminate confounding effects secondary to baseline demographics, comorbidities, and STS score components (**Supplemental S3**). Matched cohort data including characteristics and outcomes were as percentages, frequencies, and p-values using Pearson's chi-square and logistic regression. Further subgroup analysis was performed for HF with reduced ejection fraction (HFrEF) and preserved ejection fraction (HFpEF) in terms of TAVR and SAVR. Trend analysis was also performed for all outcomes for TAVR-HF and SAVR-HF using Pearson's chi-square. All analyses were conducted using appropriate stratifying, clustering, and weighting samples provided by Healthcare Cost and Utilization Project regulations [12,13]. Discharge weights provided by NIS were applied for all analyses to develop national representative procedures for this study. Statistical analysis was performed using *STATA Version 16.1, College Station, TX: StataCorp LLC* [14].

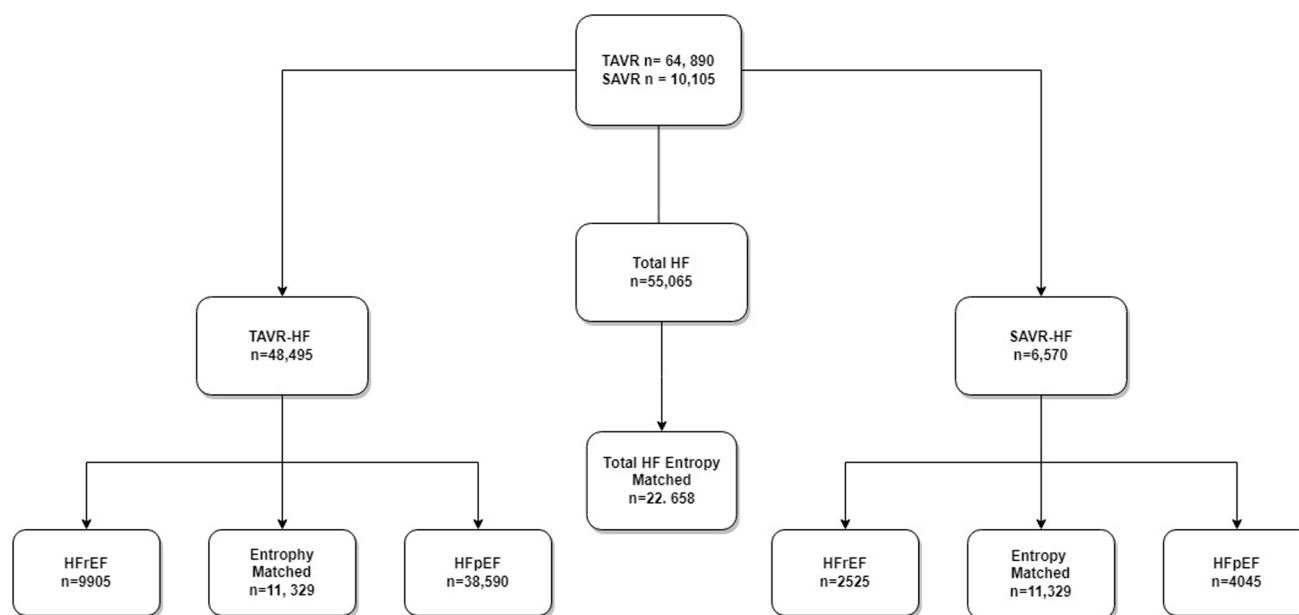


Fig. 1. Selection of cohort for crude and matched cohort.

### 3. Results

#### 3.1. Demographic and baseline comorbidities

A total of 74,995 octogenarian HF patients (TAVR-HF  $n = 64,890$  (86.5%); SAVR-HF  $n = 10,105$  (13.5%)) were included in our study. Patients' median age for TAVR-HF and SAVR-HF was 86 (IQR: 83–89) and 82 (IQR: 81–84); 48.8% and 36.8% were females, respectively. The most common procedure setting in TAVR and SAVR was elective and accounted for 79.7% and 63.4%, respectively. The baseline demographics, hospital characteristics, and comorbidities are shown in (Table 1). Among the population stratified based on ejection fraction of the left ventricle in the HF octogenarian cohort, a total of 38,590 (79.6%) HFpEF and 9905 (20.4%) HFrEF patients underwent TAVR. In contrast, in the SAVR cohort, 4045 (61.6%) and 2525 (38.4%) patients had HFpEF and HFrEF, respectively (Table 1). After PSM, we included 11,329 patients in each study cohort (TAVR-HF and SAVR-HF) (Table 1).

#### 3.2. Comparison of primary and secondary outcomes between TAVR and SAVR HF

Patients undergoing TAVR-HF as compared to SAVR-HF have lower percentage in-hospital mortality (1.8% vs. 6.9%;  $p = 0.000$ ), CVA (2.5% vs. 3.6%;  $p = 0.009$ ), SCA (9.9% vs. 20.2%;  $p < 0.001$ ), AKI (17.4% vs. 40.8%;  $p < 0.001$ ), major transfusion (26.4% vs. 67.3%;  $p < 0.000$ ), CS (1.8% vs. 9.8%;  $p < 0.001$ ), and MCS (0.8% vs. 7.3%;  $p < 0.001$ ). Additionally, post-procedural stroke (0.9% vs. 1.0%;  $p = 0.08$ ), and major bleeding (1.8% vs. 1.8%;  $p = 0.89$ ) showed no significant difference (Table 2).

PSM results for TAVR-HF in comparison to SAVR-HF were consistent in terms of in-hospital mortality (1.8% vs. 8.2%;  $p < 0.001$ ), SCA (9.6% vs. 13.2%;  $p = 0.02$ ), AKI (17.2% vs. 27.4%;  $p < 0.001$ ), transfusion (25.9% vs. 31.4%;  $p = 0.03$ ), CS (1.8% vs. 3.5%;  $p < 0.001$ ) and MCS (0.8% vs. 4.4%;  $p < 0.001$ ). However, CVA (2.5% vs. 1.7%;  $p = 0.17$ ), post-procedural stroke (0.9% vs. 0.5%;  $p = 0.25$ ) and major bleeding (1.7% vs. 1.0%;  $p = 0.1$ ) showed no significant difference (Table 2). The overall frequency and percentages of weighted unmatched and PSM outcomes are shown in Table 1; Table 2.

Further subgrouping to compare frequency and percentage of complications for HFrEF and HFpEF group showed that in TAVR-HFrEF had

lower in-hospital mortality (1.9% vs. 7.7%;  $p < 0.001$ ), SCA (10% vs. 19.4%;  $p < 0.001$ ), AKI (21% vs. 45.1%;  $p < 0.001$ ), transfusion (30.2% vs. 71.7%;  $p < 0.001$ ), CS (3.1% vs. 14.9%;  $p < 0.001$ ), and MCS (1.7% vs. 12.5%;  $p < 0.001$ ); while there was no significant difference in CVA (2.6% vs. 3.4%;  $p = 0.33$ ), post-procedural CVA (0.8% vs. 1.2%;  $p = 0.35$ ), and major bleeding (2.2% vs. 2.4%;  $p = 0.83$ ) when compared to SAVR-HFrEF (Table 2).

Similarly, TAVR-HFpEF also had a lower percentage of in-hospital mortality, SCA, AKI, transfusion, CS, and MCS, where there was no significant difference in CVA, post-procedural stroke, and major bleeding when compared to SAVR-HFpEF (Table 2).

#### 3.3. The trend of complications between TAVR-HF and SAVR-HF

TAVR-HF had lower trends from 2015 to 2018 regarding in-hospital mortality, post-procedural stroke, SCA, AKI, major bleeding, transfusion, CS, and MCS (Fig. 2). On the other hand, SAVR-HF had high trends of in-hospital mortality, AKI, CS, and a lower trend of CVA, post-procedural CVA, and major bleeding (Fig. 2). There was no significant difference in trend in SAVR-HF among SCA, transfusion, and MCS (see Fig. 3).

#### 3.4. Comparison of quality measures between TAVR and SAVR with HF

The median length of stay in TAVR-HF and SAVR-HF was three days and ten days, respectively. The median unmatched total charges for TAVR-HF and SAVR-HF were 194,561\$ and 246,100\$, respectively. The median Elixhauser index was 6, and the median AHRQ risk mortality index was 3 for both cohorts (Table 1). TAVR and SAVR trends of quality measure showed a high trend of increasing hospital stay cost of SAVR-HF compared to TAVR-HF (Table 3). LOS trend comparison between TAVR-HF and SAVR-HF showed high LOS among SAVR cohorts, but the trend of the SAVR intragroup from 2015 to 2018 was stable (Table 3).

### 4. Discussion

This national cohort included 74,995 octogenarian patients with HF. Major findings of the study include: 1) TAVR-HF, compared to SAVR-HF, had a significantly lower percentage of in-hospital mortality, SCA, CVA, AKI, major transfusion, CS, and MCS. 2) There was no significant difference in the percentage of post-procedural stroke and major bleeding

**Table 1**

Showing Baseline demographics, comorbidities, and descriptive complications among TAVR and SAVR with Heart Failure Octogenarian groups for both unmatched and propensity matched cohorts.

Analyte	Unmatched Cohort			Propensity Matched Cohort		
	TAVR-HF (n = 64,890)	SAVR-HF (n = 10,105)	p-value	TAVR-HF (n = 11,329)	SAVR-HF (n = 11,329)	p-value
<b>Age (median; IQRs) years</b>	86 (83–89)	82 (81–84)		86(23–89)	82(81–84)	
<b>Year n (%)</b>						
2015	3,480 (5.4)	1,270 (12.6)	0.000	–		–
2016	17,230 (26.6)	3,580 (35.4)	0.000			0.000
2017	21,510 (33.1)	2,825 (28)	0.000			0.000
2018	22,675 (34.9)	2,430 (24)	0.000			0.000
<b>Sex n (%)</b>						
Male	33,475 (51.6)	6,455 (63.9)	0.000	5,816 (51.3)	5,816 (51.3)	1.000
Female	31,415 (48.4)	3,650 (36.1)	0.000	5,513 (48.7)	5,513 (48.7)	1.000
<b>Race n (%)</b>						
White	55,190 (90.5)	8,160 (87.6)	0.000	10,252 (90.5)	10,272 (90.7)	0.449
Black	2,175 (3.6)	315 (3.4)	0.000	402 (3.5)	321.9 (2.8)	0.449
Hispanic	2,700 (4.4)	640 (6.9)	0.000	502 (4.4)	585.8 (5.2)	0.449
Asian/PI	815 (1.3)	185 (2)	0.000	152 (1.3)	143.8 (1.3)	0.449
<b>Transfers (%)</b>						
Not Transferred	59,770 (92.3)	8,665 (86.1)	0.000	10,476 (92.5)	10,531 (93)	0.391
Transferred	3,990 (6.2)	1,235 (12.3)	0.000	678 (6)	567.9 (5)	0.391
<b>Elective (%)</b>						
Non-elective	13,110 (20.3)	3,690 (36.6)	0.000	2,273 (20.1)	2,273 (20.1)	1.000
Elective	51,380 (79.7)	6,385 (63.4)	0.000	9,056 (79.9)	9,056 (79.9)	1.000
<b>Hospital Bed Size n (%) [Values vary by Region &amp; Control]</b>						
Small	4,275 (7)	850 (9.6)	0.003	771 (6.8)	788 (7)	0.973
Medium	12,525 (20.4)	1,940 (22)	0.003	2,303 (20.3)	2,269 (20)	0.973
Large	61,415 (72.6)	8,835 (68.4)	0.003	8,255 (72.9)	8,272 (73)	0.973
<b>Hospital Location &amp; Teaching Status n (%)</b>						
Rural	645 (1.1)	175 (2)	0.000	115 (1)	100.8 (0.9)	0.861
Urban Non-Teaching	6,915 (10.1)	1,320 (14.9)	0.000	1,071 (9.5)	1,099.4 (9.7)	0.861
Urban Teaching	54,575 (88.9)	7,340 (83.1)	0.000	10,143 (89.5)	10,128.8 (89.4)	0.861
<b>Hospital Region n (%)</b>						
Northeast	14,875 (24.2)	2,130 (24.1)	0.266	2,773 (24.5)	2,797.3 (24.7)	0.003
Midwest	14,255 (23.2)	2,135 (24.2)	0.266	2,549 (22.5)	2,916.9 (25.7)	0.003
South	19,825 (32.3)	2,600 (29.4)	0.266	3,748 (33.1)	2,939.2 (25.9)	0.003
West	12,460 (20.3)	1,970 (22.3)	0.266	2,259 (19.9)	2,675 (23.6)	0.003
<b>Weekend Admission n (%)</b>						
Monday-Friday	62,365 (96.1)	9,320 (92.2)	0.000	10,908 (96.3)	10,908 (96.3)	1.000
Saturday-Sunday	2,530 (3.9)	785 (7.8)	0.000	421(3.7)	421(3.7)	1.000
<b>Comorbidities</b>						
Pulmonary Circulation Disorders	13,000 (20)	2190 (21.7)	0.117	2234 (19.7)	2234 (19.7)	1.000
Chronic Pulmonary Disease	16,825 (25.9)	2,200 (21.8)	0.000	2932 (25.9)	2932 (25.9)	1.000
Diabetes Uncomplicated	9,325 (14.4)	1,215 (12)	0.006	1573 (13.9)	1573 (13.9)	1.000
Diabetes Complicated	11,190 (17.2)	1,665 (16.5)	0.423	2001 (17.7)	2001(17.7)	1.000
Hypothyroidism	14,670 (22.6)	1,665 (16.5)	0.000	2590 (22.9)	2590 (22.9)	1.000
Renal Failure	26,125 (40.3)	3,190 (31.6)	0.000	4513 (39.8)	4513 (39.8)	1.000
Peptic Ulcer Disease (excluding bleeding)	410 (0.6)	100 (1)	0.066	71 (0.6)	71 (0.6)	1.000
Lymphoma	475 (0.7)	50 (0.5)	0.233	89 (0.8)	89 (0.8)	1.000
Metastatic Cancer	325 (0.5)	20 (0.2)	0.058	52 (0.5)	52 (0.5)	1.000
Solid Tumor Without Metastasis	1,475 (2.3)	180 (1.8)	0.190	251 (2.2)	251 (2.2)	1.000
Rheumatoid Arthritis/Collagen Vascular	2,910 (4.5)	305 (3)	0.003	513 (4.5)	513 (4.5)	1.000
Coagulopathy	8,420 (13)	4,455 (44.1)	0.000	1421 (12.5)	1421 (12.5)	1.000
Obesity	7,605 (11.7)	1,360 (13.5)	0.028	1345 (11.9)	1345 (11.9)	1.000
Weight Loss	2,495 (3.8)	965 (9.5)	0.000	442 (3.9)	442 (3.9)	1.000
Fluid and Electrolyte Disorders	10,170 (15.7)	4,500 (44.5)	0.000	1758 (15.5)	1758 (15.5)	1.000
Blood Loss Anemia	805 (1.2)	120 (1.2)	0.843	137 (1.2)	137 (1.2)	1.000
Deficiency Anemia	2,765 (4.3)	390 (3.9)	0.416	479 (4.2)	479 (4.2)	1.000
Alcohol Abuse	395 (0.6)	135 (1.3)	0.000	69 (0.6)	69 (0.6)	1.000
Drug Abuse	75 (0.1)	15 (0.1)	0.691	15 (0.1)	15 (0.1)	1.000
HD	545 (0.8)	135 (1.3)	0.024	105 (0.9)	105 (0.9)	1.000
HTN	10,990 (16.9)	2,670 (26.4)	0.000	1,694 (15)	1,694 (15)	1.000
PAD	7,990 (12.3)	860 (8.5)	0.000	1,384 (12.2)	1,384 (12.2)	1.000
Family History of CAD	4,605 (7.1)	730 (7.2)	0.834	790 (7)	790 (7)	1.000
OSA	6005	965	0.600	1,032 (9.1)	1,032 (9.1)	1.000
Liver Disease	795 (1.2)	155 (1.5)	0.279	139 (1.2)	139 (1.2)	1.000
Alcohol	355 (0.5)	135 (1.3)	0.000	62 (0.5)	62 (0.5)	1.000
Smoking	23,215 (35.8)	3,485 (34.5)	0.268	4,103 (36.2)	4,103 (36.2)	1.000
Pneumonia	995 (1.5)	575 (5.7)	0.000	169 (1.5)	169 (1.5)	1.000
Hx of PCI	1,790 (2.8)	135 (1.3)	0.000	302 (2.7)	302 (2.7)	1.000
Hx of CABG	10,665 (16.4)	465 (4.6)	0.000	1,837 (16.2)	1,837 (16.2)	1.000
Previous MI	8,700 (13.4)	760 (7.5)	0.000	1,530 (13.5)	1,530 (13.5)	1.000
CAD	46,430 (71.5)	7,105 (70.3)	0.279	8,106 (71.6)	8,106 (71.6)	1.000
Syncope	715 (1.1)	115 (1.1)	0.889	126 (1.1)	126 (1.1)	1.000
Endocarditis	215 (0.3)	290 (2.9)	0.000	42 (0.4)	42 (0.4)	1.000



**\*Abbreviations:** PCI: Percutaneous Coronary Intervention; CABG: Coronary Artery Bypass Graft; CAD: Coronary Artery Disease; MI: Myocardial infarction; AIDS: Acquired Immunodeficiency Syndrome; HIV: Human Immunodeficiency Virus; Peripheral Artery Disease (PAD); Hypertension (HTN); Hemodialysis (HD); CVA (cerebral vascular accident); SCA (sudden cardiac arrest); MCS (mechanical circulatory support); AKI (acute kidney injury); OSA (obstructive sleep apnea).

**Table 2**

Unmatched Cohort and Propensity Matched Outcomes of SAVR vs TAVR Among Heart Failure, Heart Failure with reduced and preserved ejection fraction in Octogenarian Population.

Outcomes	Unmatched Cohort			Propensity Matched Cohort		
	TAVR-HF n (%)	SAVR-HF n (%)	p-value	TAVR-HF n (%)	SAVR-HF n (%)	p-value
<b>In-Hospital Mortality</b>	1145 (1.8)	695 (6.9)	0.000	206 (1.8)	926.6 (8.2)	0.000
HFpEF	185 (1.9)	195 (7.7)	0.000	33 (2)	146 (8.7)	0.001
HFpEF	580 (1.5)	210 (5.2)	0.000	107 (1.6)	325.9 (4.8)	0.006
<b>Post-procedural stroke</b>	605 (0.9)	100 (1)	0.080	104 (0.9)	54.6 (0.5)	0.251
SCA	6450 (9.9)	2045 (20.2)	0.000	1087 (9.6)	1495.5 (13.2)	0.018
HFpEF	990 (10)	490 (19.4)	0.000	161 (9.5)	238.2 (14.1)	0.362
HFpEF	3795 (9.8)	800 (19.8)	0.000	651 (9.5)	798.6 (11.7)	0.346
<b>AKI</b>	11,265 (17.4)	4125 (40.8)	0.000	1951 (17.2)	3099.2 (27.4)	0.000
HFpEF	2080 (21)	1140 (45.1)	0.000	357 (21.2)	681.7 (40.4)	0.002
HFpEF	6040 (15.7)	1495 (37)	0.000	1053 (15.4)	2134.8 (31.3)	0.000
<b>Major Bleeding</b>	1185 (1.8)	180 (1.8)	0.891	195 (1.7)	111.7 (1)	0.104
HFpEF	220 (2.2)	60 (2.4)	0.831	34 (2)	17.1 (1)	0.418
HFpEF	675 (1.7)	65 (1.6)	0.781	111 (1.6)	60.3 (0.9)	0.379
<b>Transfusion</b>	17,110 (26.4)	6805 (67.3)	0.000	2931 (25.9)	3559.6 (31.4)	0.026
HFpEF	2995 (30.2)	1810 (71.7)	0.000	513 (30.4)	561 (33.3)	0.651
HFpEF	9430 (24.4)	2570 (63.5)	0.000	1632 (23.9)	1943.2 (28.5)	0.232
<b>Cardiogenic Shock</b>	1180 (1.8)	995 (9.8)	0.000	202 (1.8)	391.4 (3.5)	0.000
HFpEF	310 (3.1)	375 (14.9)	0.000	53 (3.1)	105.9 (6.3)	0.078
HFpEF	410 (1.1)	270 (6.7)	0.000	75 (1.1)	224.8 (3.3)	0.001
<b>Mechanical Circulatory Support (LVAD or pVAD or ECMO)</b>	535 (0.8)	735 (7.3)	0.000	90 (0.8)	495.5 (4.4)	0.000
HFpEF	165 (1.7)	315 (12.5)	0.000	28 (1.7)	201.4 (11.9)	0.000
HFpEF	175 (0.5)	130 (3.2)	0.000	29 (0.4)	209.4 (3.1)	0.000

**Abbreviations:** CVA: Cerebral Vascular Accident; TAVR: Transaortic Valve Replacement; SAVR: Surgical Aortic Valve Replacement; SCA: Sudden Cardiac Arrest; AKI: Acute Kidney Injury; LVAD: Left Ventricular Assist Device; pVAD: Percutaneous Ventricular Assist Device; ECMO: Extracorporeal Membrane Oxygenation; HF: Heart Failure; HFpEF: Heart Failure with reduced ejection fraction; HFpEF: Heart Failure with preserved ejection fraction.

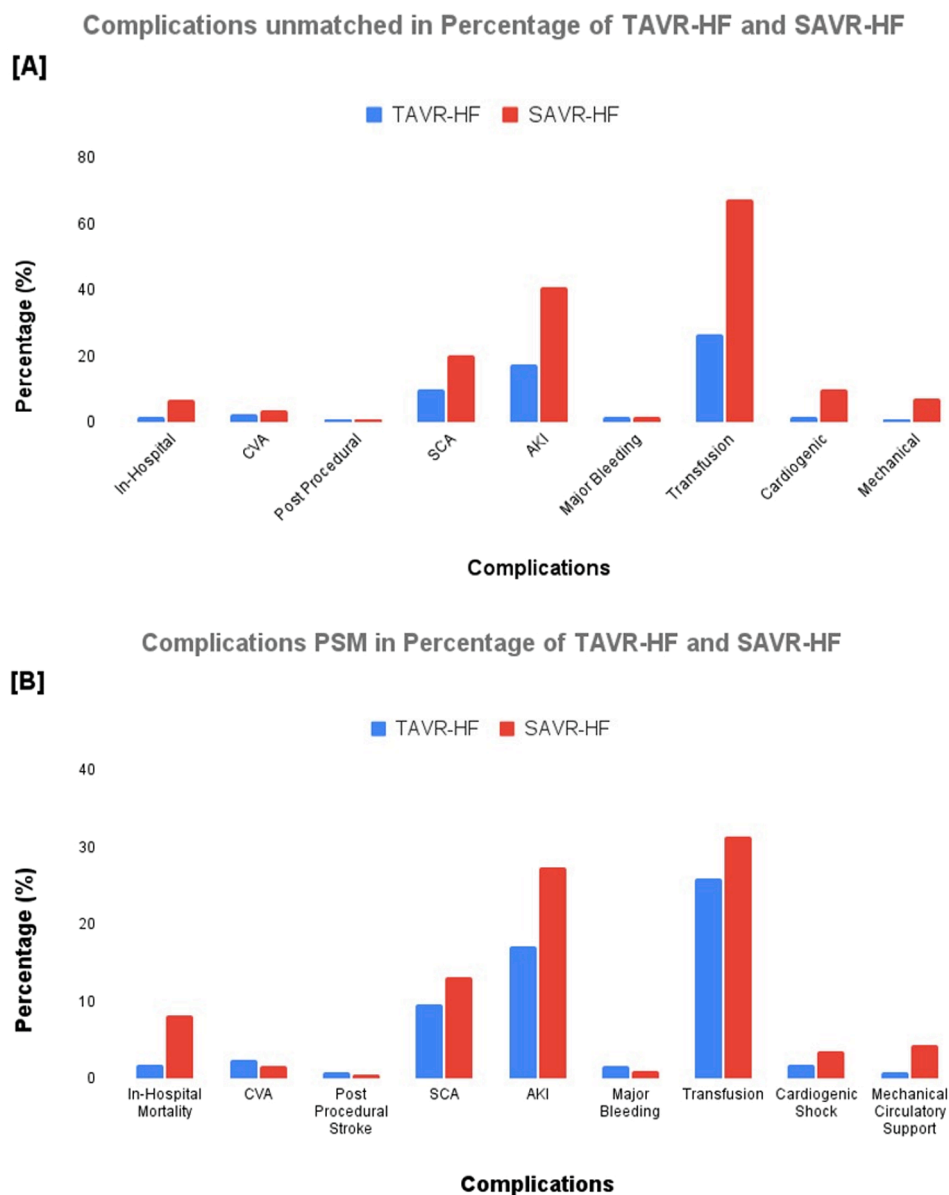
in TAVR-HF and SAVR-HF. 3) In the subgroups, TAVR-HFpEF and TAVR-HFpEF had a lower risk of in-hospital mortality, SCA, AKI, transfusion, CS, and MCS compared to SAVR-HFpEF and SAVR-HFpEF. 4) There was no significant difference in CVA incidence, post-procedural stroke, and major bleeding between the subgroups. 5) Lower trends were observed for in-hospital mortality, post-procedural stroke, SCA, AKI, major

bleeding, transfusion, CS, and MCS in the TAVR-HF. 6) Higher trends were observed for in-hospital mortality, AKI, and CS in SAVR-HF. As a result, our findings suggest that TAVR may be a more promising approach to treating AS in this vulnerable population.

After the onset of symptomatic valvular HF, the average survival in untreated AS is approximately two years [1]. Currently, recommended evidence-based operative treatments for AS are TAVR and SAVR [15]. It has been demonstrated that for high surgical risk patients, utilization of TAVR is associated with better outcomes [16]. Octogenarians are a unique subset of patients with high surgical risk due to depleting physiological reserves, age, and pathologic processes, including associated comorbidities [17,18]. In this population, a recent study comparing the outcome of in-hospital mortality in patients undergoing TAVR and SAVR found no significant difference [19]. However, no studies have investigated whether TAVR has lower rates of in-hospital mortality in patients who are both octogenarians and at high surgical risk due to coexisting HF. This is important since approximately 10% of octogenarians may have HF, placing them at very high surgical risk [20].

Our study found that in-hospital mortality in octogenarians with HF was 1.8% in patients undergoing TAVR. The in-hospital mortality rate in our analysis is lower than previous studies comparing octogenarians undergoing TAVR reporting in-hospital mortality ranging between 3 and 4.2% [21–24]. Our lower in-hospital mortality rate in the TAVR group likely reflects improved design and delivery of the prosthesis [16,23–25]. Additionally, lower in-hospital mortality in our study may be due to the positive effects of aortic valve replacement on decreasing afterload in HF patients with significant AS [16]. In our study, the in-hospital mortality rate for SAVR was 6.9%. This is consistent with the previous studies suggesting an in-hospital mortality rate ranging between 2 and 7.1% [21–24]. Higher rates of in-hospital mortality of SAVR in our study are likely secondary to our sample consisting of HF patients only, compared to previous studies including healthier octogenarians at lower surgical risk, particularly for open-heart surgery. Finally, we also found significant differences in in-hospital mortality in octogenarians with either HFpEF or HFpEF undergoing TAVR compared to SAVR. The in-hospital mortality of octogenarians in TAVR-HFpEF and TAVR-HFpEF was 1.9% and 7.7% compared to SAVR-HFpEF, and SAVR-HFpEF 7.7% and 5.2%, respectively. Our findings on in-hospital mortality are in parallel with the recent findings of Sheng et al. that TAVR may be of benefit over SAVR in all octogenarians regardless of comorbidity burden [24].

Our study also found an association between the use of TAVR in octogenarians with HF and lower in-hospital complications such as CVA, post-procedural stroke, AKI, transfusion, CS, and MCS compared to SAVR. Compared to the lower incidence of stroke in HF patients undergoing TAVR in our analysis, Hijri et al. and Brennan et al. demonstrated no change in stroke between TAVR and SAVR groups [4,19]. Leon et al. demonstrated a lower risk of stroke in the transfemoral access cohort undergoing TAVR when compared with SAVR, whereas in the transthoracic access cohort, no significant difference was observed between the two groups [23]. According to Hijri et al., TAVR demonstrated a lower incidence of AKI, similar to our analysis [19]. Our findings of lower incidence of AKI in HF patients undergoing TAVR are consistent with Reardon et al. and Leon et al., who demonstrated similar findings in patients at intermediate risk undergoing TAVR vs. SAVR [23,26]. Increased requirement of blood transfusions in HF patients undergoing SAVR as compared to those undergoing TAVR according to our analysis is also consistent with higher transfusion requirement in intermediate-risk patients undergoing SAVR as compared to TAVR as illustrated by Reardon et al. and Leon et al. [23,26]. Decreased LOS demonstrated by Hijri et al. is also consistent with our analysis in patients with HF undergoing TAVR [19]. Our findings suggesting that TAVR is associated



**Fig. 2.** (A) Complications of the unmatched percentage of TAVR-HF and SAVR-HF. (B) Complications of the propensity-matched percentage of TAVR HF and SAVR HF.

with better in-hospital outcomes in octogenarians with HF are likely related to the recent improved design and delivery of the prosthesis [16,23–25]. A recent study suggests that older patients and patients with significant comorbidities were more likely to undergo transapical than transfemoral access for TAVR [4]. However, emerging evidence shows that transfemoral access may be associated with lower in-hospital mortality, LOS, AKI, and CS [27]. An increase in transfemoral access may, therefore, potentially explain lower complications compared to the past. A comparison of quality measures between TAVR-HF and SAVR-HF groups demonstrated decreased LOS (3 vs. ten days) and median unmatched total charges (194,561\$ vs. 246,100\$) for the TAVR-HF group in our analysis. Our analysis also demonstrated lower trends of in-hospital mortality, post-procedural CVA, SCA, AKI, major bleeding, transfusion, CS, and MCS in the TAVR-HF group compared to SAVR-HF.

Subgroup analysis of our patients revealed that TAVR-HFrEF had lower in-hospital mortality, SCA, AKI, transfusion, CS, and MCS when compared with SAVR- HFrEF. In contrast, there was no significant difference in CVA, post-procedural CVA, and major bleeding between TAVR-HFrEF and SAVR-HFrEF. Whereas TAVR-HFrEF also had a lower

percentage of in-hospital mortality, SCA, AKI, transfusion, CS, and MCS compared to SAVR-HFrEF. There was no significant difference in CVA, post-procedural CVA, and major bleeding in patients with HFpEF undergoing TAVR or SAVR. As a result, TAVR use in octogenarians with HF (both in HFrEF and HFpEF subgroups) compared to SAVR is a promising option.

## 5. Limitations

Our study's main limitation was that given data was selected from NIS, the observational and retrospective data had some inherent selection bias. We attempted to decrease the selection bias by entropy matching to match the mean, median, and variance of nearby matching. Secondly, NIS outcomes are in-hospital, and it does not capture the procedural details, the frailty of the patients included, and long-term outcomes or medication-induced changes in outcomes. Furthermore, given the cross-sectional snapshot data nature, we could not do a long-term follow-up to report improvement in ejection fraction after TAVR.



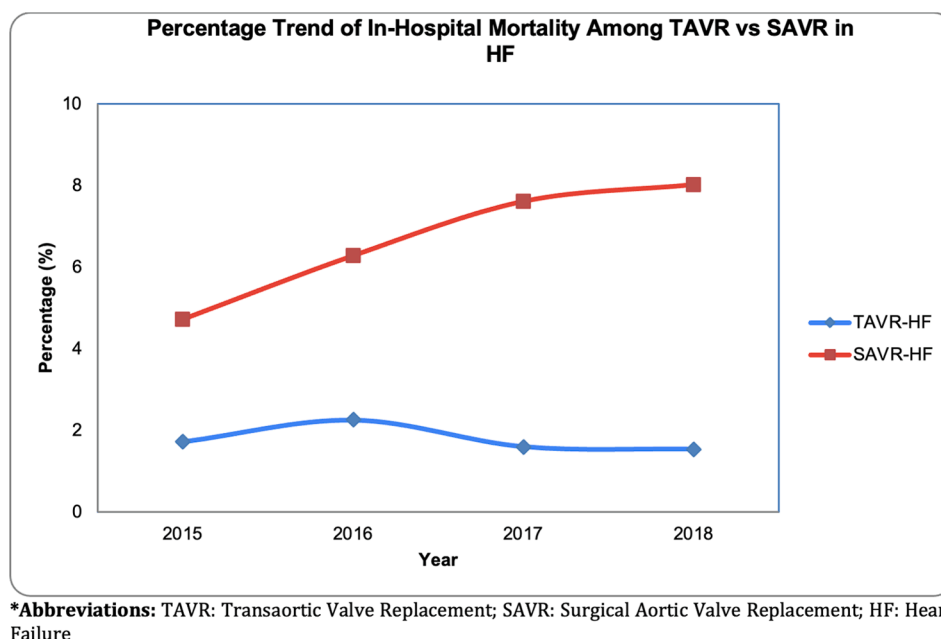


Fig. 3. Trend Percentage of In-Hospital Mortality in unmatched TAVR-HF and SAVR-HF.

Table 3

Trend of hospital cost and length of stay between TAVR and SAVR with HF in Octogenarian Population.

variable	TAVR-HF Median Cost (IQR)	SAVR-HF Median Cost (IQR)	TAVR-HF LOS [Median (IQR)] days	SAVR-LOS [Median (IQR)] days
2015	195,788\$ (141 282)	217,983\$ (184 796)	4 (4)	10 (8)
2016	198,450\$ (139 528)	233,911\$ (186 586)	3 (4)	10 (9)
2017	187,849\$ (137 886)	245,692\$ (207 935)	3 (3)	10 (8)
2018	196,157\$ (141 501)	286,817\$ (256 135)	2 (3)	10 (9)
Pooled	193,819 \$ (142,201\$- 282,323\$)	244,369\$ (168,669 \$-386,322\$)	3 (2-5)	10 (7-16)

Abbreviations: TAVR: transcatheter aortic valve replacement; HF: Heart Failure; LOS: Length of Stay; IQR: Interquartile Range; SAVR: surgical aortic valve replacement.

## 6. Conclusion

In our study of octogenarians, TAVR-HF had lower odds of in-hospital mortality, CVA, AKI, major transfusion, CS, and MCS than SAVR-HF. In the subgroup analysis of HFrEF and HFpEF for TAVR and SAVR, TAVR in both subgroups had a lower risk of in-hospital mortality, SCA, AKI, transfusion, CS, and MCS. Therefore, our study suggests that octogenarians with HF may benefit from TAVR in both HFrEF and HFpEF as the first-line treatment of their AS.

## Declaration of Competing Interest

The authors declare that they have no known competing financial interests or personal relationships that could have appeared to influence the work reported in this paper.

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## Registration number of clinical studies

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## Appendix A. Supplementary data

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