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**ORIGINAL RESEARCH** 

# Machine Learning Identification of Modifiable Predictors of Patient Outcomes After Transcatheter Aortic Valve Replacement



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#### ABSTRACT

**BACKGROUND** Transcatheter aortic valve replacement (TAVR) is an important treatment option for patients with severe symptomatic aortic stenosis. It is important to identify predictors of excellent outcomes (good clinical outcomes, more time spent at home) after TAVR that are potentially amenable to improvement.

**OBJECTIVES** The purpose of the study was to use machine learning to identify potentially modifiable predictors of clinically relevant patient-centered outcomes after TAVR.

**METHODS** We used data from 8,332 TAVR cases (January 2016-December 2021) from 21 hospitals to train random forest models with 57 patient characteristics (demographics, comorbidities, surgical risk score, lab values, health status scores) and care process parameters to predict the end point, a composite of parameters that designated an excellent outcome and included no major complications (in-hospital or at 30 days), post-TAVR length of stay of 1 day or less, discharge to home, no readmission, and alive at 30 days. We used recursive feature elimination with cross-validation and Shapley Additive Explanation feature importance to identify parameters with the highest predictive values.

**RESULTS** The final random forest model retained 29 predictors (15 patient characteristics and 14 care process components); the area under the curve, sensitivity, and specificity were 0.77, 0.67, and 0.73, respectively. Four potentially modifiable predictors with relatively high Shapley Additive Explanation values were identified: type of anesthesia, direct movement to stepdown unit post-TAVR, time between catheterization and TAVR, and preprocedural length of stay.

**CONCLUSIONS** This study identified four potentially modifiable predictors of excellent outcome after TAVR, suggesting that machine learning combined with hospital-level data can inform modifiable components of care, which could support better delivery of care for patients undergoing TAVR. (JACC Adv 2024;3:101116) © 2024 The Authors. Published by Elsevier on behalf of the American College of Cardiology Foundation. This is an open access article under the CC BY-NC-ND license (http://creativecommons.org/licenses/by-nc-nd/4.0/).

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The authors attest they are in compliance with human studies committees and animal welfare regulations of the authors' institutions and Food and Drug Administration guidelines, including patient consent where appropriate. For more information, visit the Author Center.

#### ABBREVIATIONS AND ACRONYMS

AS = aortic stenosis

2

AUC = area under the curve

- LOS = length of stay
- ML = machine learning

**RF** = random forest

SHAP = Shapley Additive Explanation

STS = Society of Thoracic Surgeons

**TAVR** = transcatheter aortic valve replacement

ortic stenosis (AS) is the most common valve disease in the United States,<sup>1</sup> affecting 4.4% of adults aged  $\geq$ 65 years.<sup>2</sup> Severe AS occurs when there is a hemodynamically significant narrowing of the aortic valve. Untreated AS is correlated with high morbidity and mortality.<sup>3,4</sup>

Transcatheter aortic valve replacement (TAVR) is an alternative to surgical aortic valve replacement for patients with severe AS.<sup>3</sup> Results of the recent PARTNER 3 trial suggest that TAVR is becoming the preferred choice for candidates of all risk levels in pa-

tients aged  $\geq$ 65 years.<sup>5</sup> Despite recent advances in technology and techniques, complications leading to extended lengths of stay or readmission are common post-TAVR.<sup>6</sup> As mortality rates for most cardiovascular procedures have declined over the past 2 to 3 decades, there has been greater emphasis on understanding quality of care among patients who survive these interventions.

Existing evidence has examined factors impacting outcomes post-TAVR but has not identified a combination of pre- and peri-procedural levers that can help achieve outcomes that matter most to patients. Based on a recent study, patients care about being able to do specific day-to-day activities, maintaining independence, reducing pain, symptoms and suffering, and staying alive.<sup>7</sup> These goals can be translated to concrete outcomes, and achieving these patient goals can be termed a "Tier-1" outcome. With the increasing volume of TAVR procedures, more data are available that can be leveraged to improve patient outcomes.<sup>8</sup>

Machine learning (ML) techniques use algorithms to detect patterns in large, complex datasets to make predictions. ML has the potential to make novel or more accurate predictions than traditional statistical models.<sup>9</sup> ML holds promise for cardiovascular medicine, as an increasing volume of high-quality data can be strategically combined with the drive for greater efficiencies in health care systems and the growing demand for personalized care.<sup>8</sup>

Previous ML models have identified parameters that are predictive of patient risk levels for transcatheter aortic valve implantation/TAVR<sup>5,10,11</sup> and percutaneous coronary intervention.<sup>12</sup> However, these studies have predominantly identified nonmodifiable patient characteristics as predictors of patient outcomes. Using ML to identify predictors that are modifiable at the clinical point of care would create opportunities for health care providers and systems to improve their delivery of care to patients. In this study, we used a large multicenter dataset to develop a random forest (RF) ML model to identify modifiable care process predictors of Tier-1 outcomes 30-days post-TAVR. We present the accuracy of the RF model and identify the most important modifiable care process predictors, including how they influenced patient outcomes in our dataset.

### METHODS

**DATA SOURCE.** This retrospective study utilized deidentified data from the Biome multicenter data repository (Biome Analytics), which aggregates clinical and financial data from 21 hospitals in the United States with TAVR programs. This repository included 11 teaching and 10 nonteaching hospitals, comprising public and private institutions. Thirteen of the hospitals were located on the West Coast. Twenty of the hospitals conducted more than 50 TAVR procedures per year, and 13 of the hospitals conducted over 100 TAVRs annually.

Clinical data fields were those defined by the National Cardiovascular Data Registry and Transcatheter Valve Therapy Registry.<sup>13</sup> Other data fields were derived from line-item cost accounting data provided by member hospitals.

**STUDY POPULATION.** The study population included all TAVR cases from January 2016 through December 2021 at member hospitals. Cases with missing data in  $\geq 1$  model predictors were excluded. To be included, cases were required to satisfy the following criteria: "Elective" as the admission status, transfemoral access, not valve-in-valve TAVR, procedure was not aborted, length of stay (LOS) was not an outlier (LOS outlier defined as case with admit or discharge dates that was null, discharge date before the admit date, or post-TAVR LOS >100 days), the discharge location was not null, and there was matched administrative/hospital cost data. LOS was defined as the time from admission to discharge. Discharge status was defined as the discharge status recorded during the index event.

**STUDY OUTCOMES.** The study's end point was a composite outcome designed to indicate whether a case had an excellent outcome at 30 days. The components of the composite outcome were defined based on parameters that are clinically relevant and that matter to patients: ability to do a specific activity (proxy measure: discharged  $\leq$ 1 day postprocedure), maintaining independence (proxy measure: discharged home, no readmission within 30 days of procedure), reducing/eliminating pain or symptoms (proxy measure: no major complications), and staying alive.<sup>7</sup>



To qualify as an excellent outcome, termed "Tier-1," a patient must have met *all* of the following criteria: 1) No major complications in-hospital or within 30 days of procedure, including acute kidney injury, stroke, hemorrhage, or vascular complications. Occurrence of any of these complications disqualified the patient from a Tier-1 outcome; 2) No pacemaker insertion in-hospital or within 30 days of procedure; 3) A postprocedural LOS of  $\leq 1$  day; 4) Discharged to home; 5) No readmission within 30 days of procedure; and 6) Alive at 30 days.

The outcome parameters were binary, and the outcome was defined to ensure that components can be assessed using registry data. All other cases were designated as Not Tier-1.

**POTENTIAL PREDICTORS.** Potential predictor variables were identified based on expert clinical opinion and the experience of member hospitals. A total of 57 potential predictors were identified (Supplemental Table 1). Each parameter was classified as a patient characteristic or a process characteristic potentially amenable to improvement. Some process characteristics were potentially modifiable, such as day of the week on which the procedure occurs in the case of an elective procedure, and some were not modifiable (eg, procedure duration).

**STUDY DESIGN AND MODEL DEVELOPMENT**. Cases were randomly divided (70:30) into a development set and a test set. To address high rates of missing observations for two potentially important features (potential predictors), imputation was performed. For the Kansas City Cardiomyopathy Questionnaire-12 score, the median value was imputed for missing values (5.8%). For the five-meter walk test, missing values (10.1%) were randomly imputed to either the maximum value (60 seconds) or the median, with probability based on the proportion of the maximum value in observed cases. Categorical predictors were coded ordinally, with a value of 1 corresponding to the lowest likelihood of a Tier-1 outcome.

The RF model was implemented in Python version 3.8 using the scikit-learn library version 1.01.<sup>14</sup> Each forest contained 100 trees, and the threshold for classification of a TAVR case to Tier-1 was a predicted Tier-1 outcome by  $\geq$ 50 trees. RF model hyperparameters were selected using grid search with bootstrap samples and 3-fold cross-validation. Hyperparameters searched were max\_depth (maximum = 15), max\_features (sqrt or log2), min samples\_leaf (maximum = 5), and min\_samples\_split (maximum = 10). Other hyperparameters were at default values. Predictors were introduced into the

TABLE 1 Patient Characteristics, Process Components, and Outcomes Across Development and Test Sets						
	Overall Cohort (N = 8,332)	Development Set (n = 5,832)	Test Set (n = 2,500)	P Value		
Patient characteristics						
Age (y)	$\textbf{79.53} \pm \textbf{8.52}$	$\textbf{79.5} \pm \textbf{8.54}$	$\textbf{79.61} \pm \textbf{8.48}$	0.60		
Female	3,662 (44)	2,575 (44.2)	1,087 (43.5)	0.57		
STS risk score	$4.04\pm3.65$	$4.02\pm3.6$	$\textbf{4.07} \pm \textbf{3.75}$	0.56		
Preprocedural KCCQ-12 Overall Score	$51.99 \pm 24.56$	$51.95 \pm 24.62$	$\textbf{52.11} \pm \textbf{24.44}$	0.79		
Glomerular filtration rate (mL/min)	$\textbf{66.02} \pm \textbf{26.29}$	$\textbf{65.94} \pm \textbf{26.14}$	$\textbf{66.23} \pm \textbf{26.64}$	0.64		
Body mass index (kg/m <sup>2</sup> )	$\textbf{27.81} \pm \textbf{6.52}$	$\textbf{27.85} \pm \textbf{6.48}$	$\textbf{27.71} \pm \textbf{6.59}$	0.36		
Platelets (per µL)	$208,\!602\pm71,\!218$	$\textbf{208,882} \pm \textbf{72,506}$	$\textbf{207,949} \pm \textbf{68,128}$	0.58		
Body surface area (m <sup>2</sup> )	$1.32\pm0.47$	$1.32\pm0.47$	$1.32\pm0.47$	0.66		
Five-meter walk test (s)	$14.41 \pm 18.36$	$14.22\pm18.18$	$14.85\pm18.75$	0.15		
Ejection fraction (%)	$\textbf{58.19} \pm \textbf{12.54}$	$58.18 \pm 12.51$	$\textbf{58.21} \pm \textbf{12.6}$	0.93		
Conduction defect	2,804 (33.7)	1,942 (33.3)	862 (34.5)	0.30		
Atrial fibrillation/flutter	2,651 (31.8)	1,859 (31.9)	792 (31.7)	0.86		
Diabetes mellitus	2,819 (33.8)	1,975 (33.9)	844 (33.8)	0.93		
Prior stroke	790 (9.5)	548 (9.4)	242 (9.7)	0.69		
Prior pacemaker	939 (11.3)	662 (11.4)	277 (11.1)	0.72		
Acute coronary syndrome	181 (2.2)	122 (2.1)	59 (2.4)	0.44		
Prior cardiogenic shock	23 (0.3)	15 (0.3)	8 (0.3)	0.62		
Hostile chest	393 (4.7)	266 (4.6)	127 (5.1)	0.31		
Chronic lung disease	1,914 (23)	1,326 (22.7)	588 (23.5)	0.44		
Prior coronary artery bypass graft	1,048 (12.6)	706 (12.1)	342 (13.7)	0.05		
Prior percutaneous coronary intervention	2,146 (25.8)	1,484 (25.5)	662 (26.5)	0.32		
Prior cardiac arrest	16 (0.2)	10 (0.2)	6 (0.2)	0.51		
Infective endocarditis	38 (0.5)	31 (0.5)	7 (0.3)	0.12		
Current or recent smoker within 1 y	362 (4.3)	247 (4.2)	115 (4.6)	0.45		
Currently on dialysis	309 (3.7)	206 (3.5)	103 (4.1)	0.19		
Immunocompromised	575 (6.9)	406 (7)	169 (6.8)	0.74		
Transient ischemic attack	599 (7.2)	426 (7.3)	173 (6.9)	0.53		
Hypertension	7,372 (88.5)	5,158 (88.4)	2,214 (88.6)	0.88		
Peripheral arterial disease	1,409 (16.9)	982 (16.8)	427 (17.1)	0.79		
Number of previous cardiac surgeries	$\textbf{0.17}\pm\textbf{0.42}$	$\textbf{0.16}\pm\textbf{0.42}$	$\textbf{0.18} \pm \textbf{0.44}$	0.06		
Heart failure hospitalization in the past 2 weeks	6,525 (78.3)	4,558 (78.2)	1,967 (78.7)	0.59		
Previous implantable cardioverter-defibrillator	183 (2.2)	132 (2.3)	51 (2)	0.52		
Prior aortic valve procedure	151 (1.8)	96 (1.7)	55 (2.2)	0.08		

Continued on the next page

RF model in stages. In the baseline stage, 33 patient characteristics were included, and the recursive feature elimination with cross-validation procedure was used to eliminate parameters with very low predictive power. At the next stage, 24 process characteristics were introduced, and 29 total predictors were retained after the recursive feature elimination with cross-validation elimination in the final model (Supplemental Table 2).

To evaluate the RF model's predictive power, sensitivity, specificity, positive predictive value, and negative predictive value were calculated using the test set. As an additional validation check, a logistic regression model was estimated in Python using the same outcome and predictor variables as the final RF model. Receiver-operator characteristic curves and the total area under the curve (AUC) were calculated for RF and logistic regression models. A calibration curve was constructed for the RF model to compare the predicted vs actual proportion of Tier-1 cases in the test set for each of the 20 bins. Shapley Additive Explanation (SHAP) values were calculated for each TAVR case for each parameter in the RF model and summed across test set cases to show both the magnitude and direction of each parameter to the model's overall predictions. As a confirming diagnostic, the Gini feature importance was calculated for each parameter in the RF model.

Model development and reporting were conducted in accordance with the critical questions posed by van Smeden et al<sup>15</sup> and the standards outlined by the Transparent Reporting of a multivariable prediction model for Individual Prognosis or Diagnosis Initiative.<sup>16</sup>

TABLE 1 Continued				
	Overall Cohort (N = 8,332)	Development Set (n = 5,832)	Test Set (n = 2,500)	P Value
Process characteristics				
Anesthesia type: moderate sedation	4,593 (55.1)	3,215 (55.1)	1,378 (55.1)	1.00
Cardiopulmonary bypass	11 (0.1)	8 (0.1)	3 (0.1)	0.84
Carotid ultrasound	352 (4.2)	254 (4.4)	98 (3.9)	0.37
Time elapsed between diagnostic catheterization and procedure (days)	$\textbf{46.7} \pm \textbf{65.23}$	$\textbf{46.13} \pm \textbf{64.22}$	$\textbf{47.95} \pm \textbf{67.51}$	0.24
Closure device used	7,166 (86.0)	5,013 (86.0)	2,153 (86.12)	0.84
Contrast volume (mL)	$\textbf{97.9} \pm \textbf{53.6}$	$\textbf{97.5} \pm \textbf{53.4}$	$\textbf{98.9} \pm \textbf{54.1}$	0.30
Direct to step-down (bypass ICU)	2,237 (26.9)	1,555 (26.7)	682 (27.3)	0.56
5-m walk documentation compliance	7,487 (89.9)	5,249 (90)	2,238 (89.5)	0.5
Foley catheter	575 (6.9)	409 (7)	166 (6.6)	0.54
Intraprocedure inotrope positive	2,151 (25.8)	1,506 (25.8)	645 (25.8)	0.98
KCCQ-12 documentation compliance	7,854 (94.3)	5,497 (94.3)	2,357 (94.3)	0.97
Preprocedural length of stay (d)	$\textbf{0.49} \pm \textbf{7.22}$	$\textbf{0.55} \pm \textbf{8.56}$	$\textbf{0.35}\pm\textbf{1.65}$	0.24
Mechanical assist device placed at start of procedure	36 (0.4)	23 (0.4)	13 (0.5)	0.42
Pulmonary function testing	1,318 (15.8)	915 (15.7)	403 (16.1)	0.62
Preprocedure testing	718 (8.6)	509 (8.7)	209 (8.4)	0.58
Procedure day of the week				0.90
Monday	1,799 (21.6)	1,258 (21.6)	541 (21.6)	
Tuesday	1,315 (15.8)	920 (15.8)	395 (15.8)	
Wednesday	2,703 (32.4)	1,906 (32.7)	797 (31.9)	
Thursday	1,714 (20.6)	1,186 (20.3)	528 (21.1)	
Friday	785 (9.4)	549 (9.4)	236 (9.4)	
Weekend	16 (0.2)	13 (0.2)	3 (0.1)	
Procedure duration (min)	$\textbf{0.91} \pm \textbf{0.85}$	$\textbf{0.9}\pm\textbf{0.82}$	$\textbf{0.94} \pm \textbf{0.9}$	0.04
Procedural start time (h)	$\textbf{10.69} \pm \textbf{2.8}$	$10.66 \pm 2.79$	$10.75 \pm 2.83$	0.18
Sentinel Protect System	1,671 (20.1)	1,161 (19.9)	510 (20.4)	0.61
Swan-Ganz catheterization	540 (6.5)	379 (6.5)	161 (6.4)	0.92
TAVR and percutaneous coronary intervention during admission	192 (2.3)	157 (2.7)	35 (1.4)	<0.01
Valve sheath access method (percutaneous)	8,249 (99)	5,777 (99.1)	2,472 (98.9)	0.46
Procedure location				0.21
Catheter lab	2,608 (31.3)	1,832 (31.4)	776 (31)	
Hybrid cath lab suite	2,499 (30)	1,780 (30.5)	719 (28.8)	
Hybrid OR suite	3,225 (38.7)	2,220 (38.1)	1,005 (40.2)	
Valve type: balloon expandable	6,965 (83.6)	4,860 (83.3)	2,105 (84.2)	0.33

Values are mean  $\pm$  SD or n (%). *P* values < 0.05 are indicated in **bold**.

ICU = intensive care unit; KCCQ12 = Kansas City Cardiomyopathy Questionnaire 12; OR = operating room; STS = Society of Thoracic Surgeons; TAVR = transcatheter aortic valve replacement.

### RESULTS

**DATASET CHARACTERISTICS.** A total of 14,562 TAVR cases were identified from the Biome multicenter data repository (Biome Analytics) (Figure 1). After exclusion, the total number of cases included in the analysis was 8,332. Patient characteristics and process components were similar across development and test sets (Table 1). The mean age of patients in the overall cohort was 79.5  $\pm$  8.5 years, and 56% were male. The mean Society of Thoracic Surgeons (STS) risk score was 4.0  $\pm$  3.7.

Approximately half of the overall cohort had a Tier-1 outcome (n = 3,793, 45.5%) (Supplemental Table 3). The most common reason a patient was not classified with a Tier-1 outcome was LOS >1 day (n = 3,943, 86.9% of all Non-Tier-1 cases). Other common reasons were  $\geq$ 1 complications (N = 1,618, 35.7% of all Non-Tier-1 cases) and readmission within 30 days (n = 525, 11.6% of all Non-Tier-1 cases).

**RANDOM FOREST ALGORITHM PERFORMANCE EVALUATION.** The final RF model with 29 predictors had good sensitivity (0.67) and specificity (0.73) for outcome differentiation, with an AUC of 0.77 (**Figure 2**). Positive predictive value and negative predictive value were 0.67 and 0.73, respectively. Of the 29 predictors, 15 were patient characteristics and 14 were care process components, of which four were



potentially modifiable at the point of care and contributed heavily toward achieving the Tier-1 outcome (Central Illustration).

When evaluated on the test set of 2,500 patients, the false positive rate and false negative rate were each 15% (**Figure 3A**), and good calibration was achieved between predicted and actual outcomes (**Figure 3B**). The logistic regression model constructed with the same parameters had AUC of 0.77 (Supplemental Table 4, Supplemental Figure 1).

**ASSESSMENT OF PREDICTOR IMPORTANCE.** From the SHAP summary plot, we identified four highly impactful modifiable predictors: receipt of moderate sedation, direct movement of the patient to a stepdown unit, lower preprocedural LOS, and fewer days between diagnostic catheterization and TAVR that increased the likelihood of a Tier-1 outcome (**Figure 4**). These four predictors also had the highest Gini feature importance among modifiable parameters in the final RF model.

#### DISCUSSION

In this study, we describe a ML algorithm that uses patient-level data to predict excellent (Tier 1) outcomes following TAVR with high sensitivity and specificity. Modifiable parameters with the highest predictive importance for excellent outcomes were the type of anesthesia used (with moderate sedation being predictive of good outcomes after TAVR), patient movement directly to a stepdown unit postprocedure, minimal time between diagnostic catheterization and the procedure, and shorter preprocedural LOS. These results demonstrate that patient-level data can help predict excellent outcomes following TAVR. The goal of our model was to leverage these patient-level data to inform heart team and hospital efforts to improve modifiable parameters of care, given the observed variation in those parameters-and TAVR outcomes-across hospitals.

The current study is unique in that the predefined outcome (Tier-1/Not Tier-1) is a composite of parameters of clinical and patient-centered importance. Existing algorithms like the STS risk score calculate the risk of mortality and morbidity but do not measure other outcomes that matter to patients. The Tier-1 composite outcome accounts for traditional outcomes like mortality but also includes additional outcomes important to patients including survival, short LOS, no major complications (those associated with worse survival or quality of life at 1 year<sup>17</sup>), and no readmission.

In defining Tier-1, the focus was to ensure the inclusion of components that are clinically relevant and matter to patients while focusing on variables that can be captured in administrative or registry data. Patient-centered care has been identified as a key component of health care quality in the cardiovascular disease space.<sup>18,19</sup> Strategies such as shared decision-making that elevate patient priorities have been growing in importance to health care providers, facilities, payers, and patients, particularly in the preparation of patients for surgery.<sup>18-20</sup> In cardiovascular surgery, patient-centered preparation may reduce disparities in outcomes.<sup>21</sup> Our study reflects this important shift in priorities by including parameters of importance to patients in our definition of a good outcome post-TAVR (eg, survival, short LOS, no major complications, and no readmission).

This work is also notable because of the large dataset used for development and testing of the random forest model. In a recent review of ML prediction in cardiovascular diseases, many algorithms were developed based on sample sizes of less than 1,000.<sup>22</sup> We used a dataset of 8,332 cases after exclusions, which, to our knowledge, is the largest dataset used in machine-learning predictions of outcomes post-TAVR to date.<sup>5,23,24</sup> ML has previously been used to determine predictors of mortality,<sup>5,10,23</sup> risk of bleeding,<sup>24</sup> and need for pacemaker implantation<sup>25</sup> post-TAVR, but these identified only



nonmodifiable predictors such as patient comorbidities that are not amenable to care process improvement. Two previous studies identified potentially modifiable parameters as predictors of outcomes post-TAVR (fluid and electrolyte disturbance, procedures done before the weekend, valve type, and conscious sedation), but they were smaller in size than the current study and used traditional multivariate logistic regression.<sup>6,26</sup> ML has the potential to advance cardiovascular patient care by identifying novel patterns in large, complex datasets.<sup>27</sup> Our model identified several modifiable process characteristics discussed below and, similar to other studies,<sup>28-30</sup> had predictive power comparable to standard linear models. These results demonstrate that detecting patterns in the TAVR care process by applying ML algorithms to patient-level data can potentially improve patient



outcomes. The reporting of SHAP values may be particularly helpful in enhancing the interpretability of ML models for clinicians. Nevertheless, in the absence of significance testing for individual variables that traditional regression models provide, it is especially important to consider the potential clinical significance of individual predictors, both when choosing which predictors to include in an ML model and in interpreting the results.

The type of anesthesia used during the procedure was the most important predictor in our ML model, with general anesthesia predicting worse outcomes. This aligns with previous findings that showed use of moderate sedation was associated with lower health care resource utilization, reduced 30-day mortality, and increased odds of being discharged home while having the same procedural success as with general anesthesia. Findings from the 3M TAVR study also suggest that modification of certain aspects of the TAVR approach can increase the efficiency of the procedure as well as lead to cost savings.<sup>31</sup> There is a movement toward using a minimalist, risk-stratified clinical approach to TAVR that will maintain effectiveness and safety while reducing health care resource use and costs.<sup>32</sup> Such pathways may include use of local anesthesia rather than general anesthesia. These modifications have been associated with shorter procedure times, earlier discharge, and/or lower readmission rates among patients undergoing TAVR.<sup>32-36</sup>

Conscious sedation in patients receiving TAVR has increased over time, ranging from 33.4% in 2016 to

64.1% in 2019.<sup>37</sup> As the physical and hemodynamic stresses of the procedure have lessened, the intensity of anesthetic support required has declined as well. Transitioning from general anesthesia to conscious sedation may further enable TAVR cases to move from operating rooms to catheterization laboratories and improve postprocedural recovery times. Although the overall prevalence of moderate sedation increased in later years of the study period, its utilization still varies widely across hospitals, suggesting opportunities for improvement in many locations. Thus, findings from this study align with existing evidence emphasizing the value of conscious sedation during TAVR.

Our model also identified that patients who moved directly to a stepdown unit after their procedure rather than to the intensive care unit (ICU) were more likely to have excellent outcomes, controlling for potential risk of poor postprocedure clinical status. Interestingly, the correlation between STS risk score (measuring underlying risk) and direct-to-stepdown was quite small (-0.06). Other risk factors for ICU admission, including atrial fibrillation/flutter, presence of a conduction defect, and recent heart failure hospitalization, are also included in the model. Stepdown units provide an intermediate level of care for patients with needs falling between the general ward and the ICU.<sup>38,39</sup> Previous studies have shown benefits associated with availability of a stepdown unit, including improved efficiencies of patient flow,<sup>38</sup> reductions in ICU congestion,<sup>40</sup> improved



patient outcomes after ICU care,<sup>41</sup> and significantly reduced adjusted hospital mortality.<sup>42</sup> The results presented here provide new evidence supporting the benefits of a stepdown unit for patients undergoing TAVR.

Lower preprocedural LOS was also associated with a higher likelihood of excellent outcomes. Although the average preprocedure LOS observed in this study was a relatively short 0.49 days (**Table 1**), the standard deviation was 7.22 days, indicating that a substantial number of patients had very long stays before their TAVR procedure. These patients are the suggested targets of performance improvement efforts. Elective cardiac patients are at risk of physical deterioration in the preprocedural period.<sup>43,44</sup> In one study, 45% of patients reported that their health suffered in the wait period before cardiac surgery,<sup>44</sup> suggesting that a "time to therapy" concept might apply to TAVR. Further, patients with AS presenting with acute heart failure benefit from the prompt decrease in left

ventricular afterload known to occur post-TAVR,45 supporting the hypothesis that efficacy of TAVR may be time-dependent, especially in certain subgroups of patients. Various groups have shown that strategic use of the preprocedural period is associated with improved quality of life,<sup>46</sup> shorter postoperative LOS,43 lower likelihood of admission to rehabilitation,<sup>47</sup> and lower overall resource utilization.<sup>48</sup> The effect of preprocedural intervention has not been studied in TAVR patients, but a clinical trial is ongoing to investigate the safety and efficacy of prehabilitation in high-risk patients undergoing TAVR.<sup>49</sup> It is possible that the patients in this study benefited from being discharged to home prior to TAVR, as well as from avoiding a long period of hospitalization before their procedure. These results emphasize the need for health care providers to optimize the use of the preprocedural period. Decreasing the preprocedural LOS by discharging patients to home prior to surgery is a potential avenue for future research.

Existing evidence has shown that longer time from catheterization to TAVR (wait times) impacts post-TAVR prognosis, and the current study underscores this. A recent study found that increased wait times were associated with an increase in 1-year mortality by 2% per week after referral for TAVR.<sup>50</sup> Such evidence highlights the need for strategies to minimize delays in access to TAVR and prompt identification of high-risk patients who require faster processing. It has been suggested that treatment delay beyond 1 month should be avoided and that patients and physicians should proceed with aortic valve replacement on a semi-urgent rather than an elective basis.<sup>51</sup>

To help understand when during the care pathway the heart team can intervene, we identified process factors that can be potentially modified before the procedure, namely preprocedural LOS and time from catheterization to TAVR. Practice guidelines could be targeted to implement strategies to reduce the time from catheterization to TAVR, whereas hospitalspecific interventions are needed to target preprocedural LOS. Variables that can be modified during or after the procedure are anesthesia type and direct transfer to stepdown.

**STUDY LIMITATIONS.** Our study has several limitations, including the potential for unmeasured confounding. Specifically, patient outcomes post-TAVR are likely influenced by factors other than the 57 features included in this study, as we were limited to variables captured in the National Cardiovascular Data Registry and Transcatheter Valve Therapy Registry. We imputed missing values for two registry fields, which may reduce the potential predictive power of those variables in our models. We were unable to adjust for provider preference when studying predictors or account for variables that may have been influenced by hospital-specific protocols or provider preferences. Second, this was a retrospective study of all TAVRs performed at hospitals that shared data with Biome Analytics, so it does not represent all patients who underwent TAVR in the United States during the study period. The Biome dataset is derived from 21 hospitals, 13 of which are located on the West Coast, and thus the data are not necessarily geographically representative of the entire United States. Future external validation should be performed to assess the broader generalizability of these findings.

### CONCLUSIONS

Using ML and a large dataset of 8,332 TAVR cases, we identified 29 predictors of patient outcomes post-TAVR, including four highly impactful predictors that are potentially modifiable components of the care process. These findings suggest that ML algorithms can be leveraged in value-based care arrangements such as pay-for-performance and outcomes-based contracting, offering insights into risk stratification, quality of care, and process optimization. Future work is warranted to evaluate these findings in other datasets and to evaluate the clinical potential of ML models for the TAVR care process.

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#### PERSPECTIVES

**COMPETENCY IN MEDICAL KNOWLEDGE:** We used a large dataset and a ML approach to identify features of the care process that may be modified to optimize patient outcomes post-TAVR. This model was comparable to a traditional logistic regression model and showed high sensitivity and specificity, indicating that it could assist clinical decision-making to achieve patient-centered outcomes. **TRANSLATIONAL OUTLOOK:** Future studies are warranted to test these results on additional datasets, individually score each of the variables included in the composite outcome, or explore the potential for a percase risk score to aid in preprocedural planning.

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**KEY WORDS** artificial intelligence, cardiology, care process, patient-centered, random forest, TAVR

**APPENDIX** For supplemental tables and a figure, please see the online version of this paper.