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Psoas Muscle Size Predicts Risk-Adjusted Outcomes After Surgical Aortic Valve Replacement

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Abstract

Background—Frailty is an important predictor of outcomes after cardiac surgery, but utility is limited by difficult assessment and quantification. We hypothesize that sarcopenia defined as psoas muscle cross-sectional area is a useful predictor of surgical aortic valve replacement (SAVR) outcomes in moderate to high-risk patients.

Methods—Moderate to high-risk (predicted risk of mortality [PROM] >3%) patients who underwent SAVR with or without coronary bypass were extracted from an institutional database (2009–2016). Psoas index was calculated as the cross-sectional area of the psoas muscle at the L4 vertebral level normalized to body surface area. Patients were stratified by sarcopenia status, defined as <25th gender-specific percentile. Multivariable regression analysis identified risk-adjusted associations with psoas index using STS predicted risk scores.

Results—Of the 240 patients included, the median PROM was 6%, median age 80 years, and 40% were female. Patients with (33.3%) and without (66.7%) sarcopenia had equivalent baseline risk (median PROM 5.7% vs 6.0%, $p=0.29$). Patients with sarcopenia had higher 1-year mortality (31.9% vs 16.9% $p=0.03$). Psoas index significantly predicted risk-adjusted 1-year mortality (OR 0.84, $p=0.02$), long-term mortality (HR 0.92, $p=0.04$), as well as risk-adjusted major morbidity, prolonged ventilation, length of stay, discharge to a facility and hospital cost. Finally, psoas index measurements were highly reproducible (Pearson correlation coefficient 0.944).

Conclusions—Psoas index is an easily obtained and reproducible measure of frailty that predicts risk-adjusted resource utilization, morbidity, and long-term mortality. Psoas index may improve procedural selection and risk-adjustment in high-risk patients with aortic valve disease.

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Classifications

Aortic valve replacement; frailty; sarcopenia; resource utilization

The trend towards more complex and higher risk patients undergoing cardiac surgery has been underway for decades.[1] Patients undergoing surgical aortic valve replacement (SAVR) today have greater comorbid disease than in prior years.[2] Additionally, the population of the United States is aging, leading to an increase in the number of older patients undergoing cardiac surgery.[3] The inclusion of standard risk factors in current prediction models, such as the Society of Thoracic Surgeons and EUROScore, has resulted in highly accurate in risk prediction algorithms that are now the gold standard.[4] However, use of the so called “eyeball test” to determine patient appropriateness for surgery persists, with its importance highlighted by its inclusion as a selection criterion in the Placement of Aortic Transcatheter Valves (PARTNER) trials.[5]

Preoperative surgeon judgment of a patient attempts to assess frailty, the missing ingredient in current risk models. By definition, frailty is the diminished reserve across multiple organ systems resulting in a patient incapable of adapting to stressors.[5–7] Whether frailty is a phenotype or a series of accumulated deficits is a topic of debate, although the former tends to be more useful for cardiovascular purposes.[8] Comprehensive frailty assessments evaluate all aspects of this phenotype (weight loss, exhaustion, low energy, weakness, slowness) but are cumbersome and time consuming.[5, 9–11] While not the sole etiology of frailty, sarcopenia is a physical manifestation of frailty with significant overlap across the phenotypic model.[12] This can be observed, measured and objectively inserted into risk prediction models.

Psoas muscle size is a relatively new technique for quantifying sarcopenia and is recognized as a useful measure of frailty across multiple surgical specialties.[13–17] It is simple to obtain, highly reproducible, and validated. Sarcopenia has been shown to correlate with body fat percentage, lean muscle mass, grip strength, short physical performance battery scores and VO₂ max.[18–20] Psoas muscle size is predictive of mortality, major complications, and resource utilization in areas as diverse as emergency general surgery, abdominal aortic aneurysms, colorectal surgery and pancreatic surgery.[13–17]

The purpose of this study was to evaluate the utility of psoas muscle cross-sectional area as a quantitative measure of frailty in moderate to high-risk patients undergoing SAVR. Psoas muscle area is an easily quantified measure of sarcopenia that is typically available in higher risk SAVR patients due to procedural planning for potential transcatheter aortic valve replacement (TAVR). We hypothesized that patients with decreased psoas muscle cross-sectional area would have increased risk-adjusted morbidity, mortality and resource utilization following surgical aortic valve replacement.

Patients and Methods

Patient Data

This study was approved by the University of Virginia Institutional Review Board, #19762. All patients who underwent aortic valve replacement between January of 2009 and December of 2016 were extracted from an institutional Society of Thoracic Surgeons (STS) database. Inclusion criteria included first time aortic valve replacement for severe aortic stenosis, a STS predicted risk of mortality (PROM) >3% and a preoperative abdominal computed tomography (CT) scan available for review. Patients with endocarditis were excluded. Medical records were reviewed for 1,384 patients and of these, 240 met criteria for inclusion as demonstrated in the consort diagram (Supplemental Figure 1). The STS database contained clinical information that was paired with cost and long-term mortality information. The long-term mortality information was obtained from three separate sources including clinical records, the Virginia Department of Health and the Social Security Death Master File.

The cost data was abstracted from the Clinical Data Repository and is derived from finance department records. Each patient has charges identified by Current Procedural Terminology code and converted to costs based on monthly updates that include direct and indirect component costs. The sum cost was then adjusted for inflation to 2016 dollars using the market basket for the inpatient prospective payment system at the Centers for Medicare and Medicaid Services.

Preoperative CT scans were used to calculate the cross-sectional area of the psoas muscle at the level of the L4 vertebra using multiplanar reconstruction to account for rotation and kyphosis (Supplemental Figure 2). Three measurements were taken of the left and right psoas with the average used for analysis. This methodology has been described previously, and validated as a measure to estimate total body sarcopenia and cardiorespiratory fitness. [19–21] The mean psoas cross-sectional area was divided by the body surface area to obtain the psoas index. Sarcopenia was defined as a psoas index below the 25th gender specific percentile, based on definitions used in the original description of the frailty phenotype and prior analyses.[8, 14] Outcome measures evaluated included operative, one-year, and long-term mortality, in hospital complications, intensive care unit (ICU) or postoperative length of stay, and hospital cost.

Statistical Analyses

Categorical variables are presented as count (percent) while continuous variables are presented as mean \pm standard deviation (SD) or if skewed then median [interquartile range, Q1–Q3]. Normality was evaluated by Shapiro-Wilk Statistics. For univariate analysis, comparisons were made by Chi Square test, Independent T-test or Mann Whitney U test as appropriate. Multivariable logistic regression was used to evaluate psoas index as a predictive measure for categorical outcomes. Generalized linear models were used to evaluate hospital cost using a gamma distribution, length of stay using a negative binomial distribution, and albumin using a normal distribution.[22] Models were fit in linear form except for hospital cost which performed best as a logarithmic link making interpretation

more difficult. Risk-adjustment was performed using STS risk scores relevant to the outcome of interest, and if no specific risk model was available for a given outcome adjustment was performed using predicted risk of morbidity or mortality. Long-term mortality was compared by Kaplan-Meier survival analysis for comparison of sarcopenia status and by Cox proportional hazard analysis for prediction of risk adjusted psoas index on long-term survival. Inter-observer agreement for psoas size measurements read by two independent physician readers was assessed by Pearson correlation. All statistical analyses were performed using SAS Version 9.4 (SAS Institute, Cary, NC) with $p < 0.05$ defining statistical significance.

Results

Sarcopenia and Frailty

Patients included in the study were largely similar compared to those excluded for missing CT scans including rates of comorbidities and STS predicted risk of mortality, although there are some minor logical differences including being statistically older, more frequently elective, and having a higher rate of prior cardiac surgery (Supplemental Table 1). Of patients included in the study, the median psoas cross-sectional area was 18.9 cm^2 (14.3–23.3) and was significantly lower in females (15.0 vs 20.5 cm^2 , $p < 0.0001$). After adjusting for body surface area, the median psoas index was $9.6 \text{ cm}^2/\text{m}^2$ with both distributions by gender shown in Figure 1. The 25th percentile cutoff for sarcopenia was a psoas index of $6.96 \text{ cm}^2/\text{m}^2$ for women and $9.09 \text{ cm}^2/\text{m}^2$ for men. A subset of 145 (60%) patients had psoas measurements by both reviewers. This demonstrated high reproducibility with a Pearson correlation coefficient of 0.944 and by linear regression an R^2 of 0.892 (Figure 2).

There was no statistically significant correlation between psoas index and preoperative albumin level (parameter estimate = 0.012 [−0.006 – 0.031], $p = 0.195$), nor with prolonged 5 meter walk time (OR 1.22 (0.94–1.59), $p = 0.138$).

Baseline and Operative Characteristics

Baseline and operative characteristics for sarcopenic (33.3% [60]) and non-sarcopenic (66.7% [180]) patients are shown in Table 1. The only statistically significant baseline difference between patients with and without sarcopenia was the rate of smoking (6.7% vs 26.7%, $p = 0.001$). There were no differences in rates of comorbid disease, valve disease, or prior cardiac surgery. There was no significant difference between groups in STS PROM (5.7% vs 6.0%, $p = 0.29$). Similarly, there were no differences in rates of CABG, cross-clamp times or cardiopulmonary bypass times (Table 1).

Unadjusted Outcomes

While there was no difference in operative mortality rate between groups, sarcopenic patients had a higher one-year mortality rate (31.9% vs 16.9%, $p = 0.03$). There was a trend towards improved long-term survival in non-sarcopenic patients by Kaplan-Meier survival analysis (Figure 3). This trend becomes significant when utilizing the continuous psoas index instead of sarcopenia status (HR=0.916 [0.843–0.996], $p = 0.04$). The median survival for sarcopenic patients was 5.41 years while for nonsarcopenic patients was 5.49 years

($p=0.099$). There were no significant differences in rates of major or minor complications (Table 2). Finally, there were no significant differences in measures of resource utilization by sarcopenia status including readmission and ICU length of stay. There were trends towards sarcopenic patients having a higher rate of discharge to a facility (63.2% vs 49.1%, $p=0.07$) and longer median postoperative length of stay (7.5 vs 7 days, $p=0.06$).

Risk Adjusted Outcomes

Risk-adjusted outcomes for logistic regression models using psoas index can be found in Table 3. As a continuous variable, psoas index is predictive of risk-adjusted one-year mortality (OR 0.84, $p=0.02$). Similarly, psoas index remains a significant predictor of long-term survival even after risk-adjustment (HR=0.917 (0.845–0.996), $p=0.041$). Psoas index is a significant independent predictor of STS major morbidity, as well as prolonged ventilation.

Psoas index was also independently predictive of most measures of resource utilization including discharge to a facility (OR 0.88, $p=0.005$), postoperative length of stay (-0.46 , $p<0.0001$) and hospital cost ($-.03$, $p=0.001$). There was also a trend towards psoas index significantly predicting risk-adjusted ICU length of stay, although it was not associated with rates of readmission.

Comment

Using data already obtained in most moderate to high-risk aortic valve cases, we demonstrate that psoas muscle size is a useful measure of sarcopenia and predicts risk-adjusted morbidity, mortality and resource utilization. Sarcopenic patients had higher mortality at 1-year with a trend towards higher long-term mortality. When utilized as a continuous variable, psoas index is predictive of risk-adjusted major morbidity, 1-year mortality and long-term mortality. In addition, psoas index is independently associated with the resource utilization metrics of discharge to a facility, postoperative length of stay and hospital cost.

It has been well established that frailty increases the risk of mortality after cardiac surgery. Using a variety of frailty measures, it has been found to increase the risk of short and mid-term mortality.[23–25] This finding holds true for sarcopenia assessed by psoas muscle size in the TAVR population at 6 months, and the TAVR or SAVR population at 2 years.[21, 26] This analysis did not demonstrate an association between psoas index and operative mortality, but smaller psoas index was associated with increased odds of mortality at 1 year even after risk-adjustment. Moreover, this study is unique in finding that the mortality association persists long-term with psoas index predicting risk-adjusted long-term mortality. As quality metrics move past the 30-day threshold, psoas index could be a useful tool for accurate long-term risk prediction.

Frailty also appears to be an important driver of complications after cardiac surgery, although only a few studies have evaluated this question. One analysis demonstrated that slow gait speed in elderly cardiac surgery patients was an independent predictor of STS major morbidity or mortality.[27] The same group found that the addition of frailty and disability score improved the predictive abilities of multiple cardiac risk models including

the STS risk model for morbidity and mortality.[28] Finally, a recent systematic review found a strong relationship between frailty and major adverse cardiac and cerebrovascular events following cardiac surgery.[29] In this analysis we also find that frailty as measured by psoas index is independently associated with increased risk of major morbidity as well as its component complication of prolonged ventilation. There was also a trend towards a risk-adjusted association with atrial fibrillation.

Another important aspect of complications after cardiac surgery is that they are an important driver of increased resource utilization.[30, 31] The most consistent finding is that frailty is associated with an increased length of stay.[23] This includes prior analyses using psoas muscle size to define sarcopenia after both cardiac surgery and TAVR.[18, 32] In this analysis, a 1 point increase in psoas index was independently associated with almost a half day less in the hospital. Complications and length of stay can increase hospital costs, and frailty has previously been associated with an additional \$20,000.[33] This was corroborated in our analysis with a strong association between psoas index and hospital cost in a risk-adjusted model. Finally, with global billing efforts increasing discharges to a facility are going to be increasingly under scrutiny.[34] We found that increasing psoas index was strongly associated with a decreased risk of discharge to a facility (OR 0.88).

There are several limitations to this study including its single center, retrospective nature that inherently introduces some element of selection bias. The generalizability of the results is also limited by the prevalence of frailty in certain cardiac surgery populations and will be more useful in more frail cohorts. Psoas index, while a validated measure of sarcopenia that correlates with many different measures of frailty, is not a comprehensive frailty measure. This limited frailty assessment is going to be an inherent limitation of any simple, fast measurement. This is simply a tradeoff we believe is worth the downside if it enables inclusion in future risk models. Finally, risk adjustment was performed using the STS risk models. STS PROM was not designed to predict long-term mortality, although it has previously been validated to do so.[35]

Frailty and sarcopenia are known to be important predictors of morbidity, mortality and resource utilization in cardiac surgery. Here we demonstrate psoas index is a simple, reproducible measure of sarcopenia that is associated with short and long-term outcomes after SAVR. This makes it an ideal measure to incorporate into future risk models. As psoas size has demonstrated similar promise in the TAVR population, it would be useful in risk assessment across the spectrum of approaches currently available. In an era of emphasis on patient centered outcomes, autonomy and choice, the inclusion of frailty measures will be important to help patients and providers select the appropriate procedure. Additionally, with public reporting and value based payments the inclusion of a simple frailty metric such as psoas index could help improve longer-term risk prediction models.

Supplementary Material

Refer to Web version on PubMed Central for supplementary material.

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Abbreviations

CT	computed tomography
ICU	intensive care unit
PARTNER	Placement of AoRTic TraNscathetER Valve Trial
PROM	predicted risk of mortality
SAVR	surgical aortic valve replacement
SD	standard deviation

STS Society of Thoracic Surgeons
TAVR transcatheter aortic valve replacement

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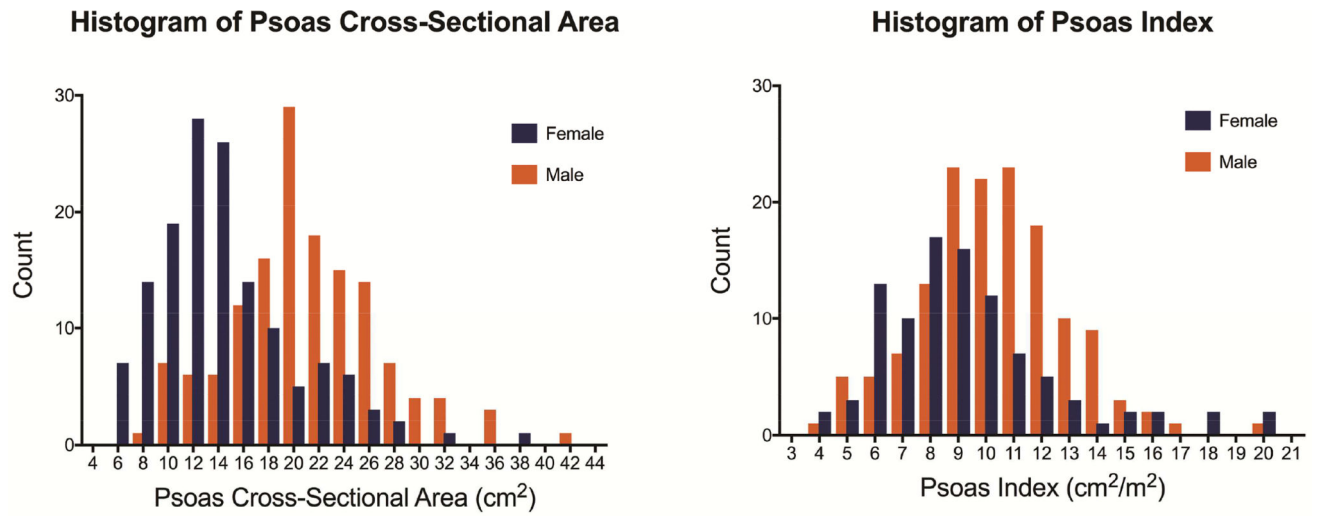


Figure 1. Histogram of (A) psoas cross sectional area and (B) psoas index by sex.

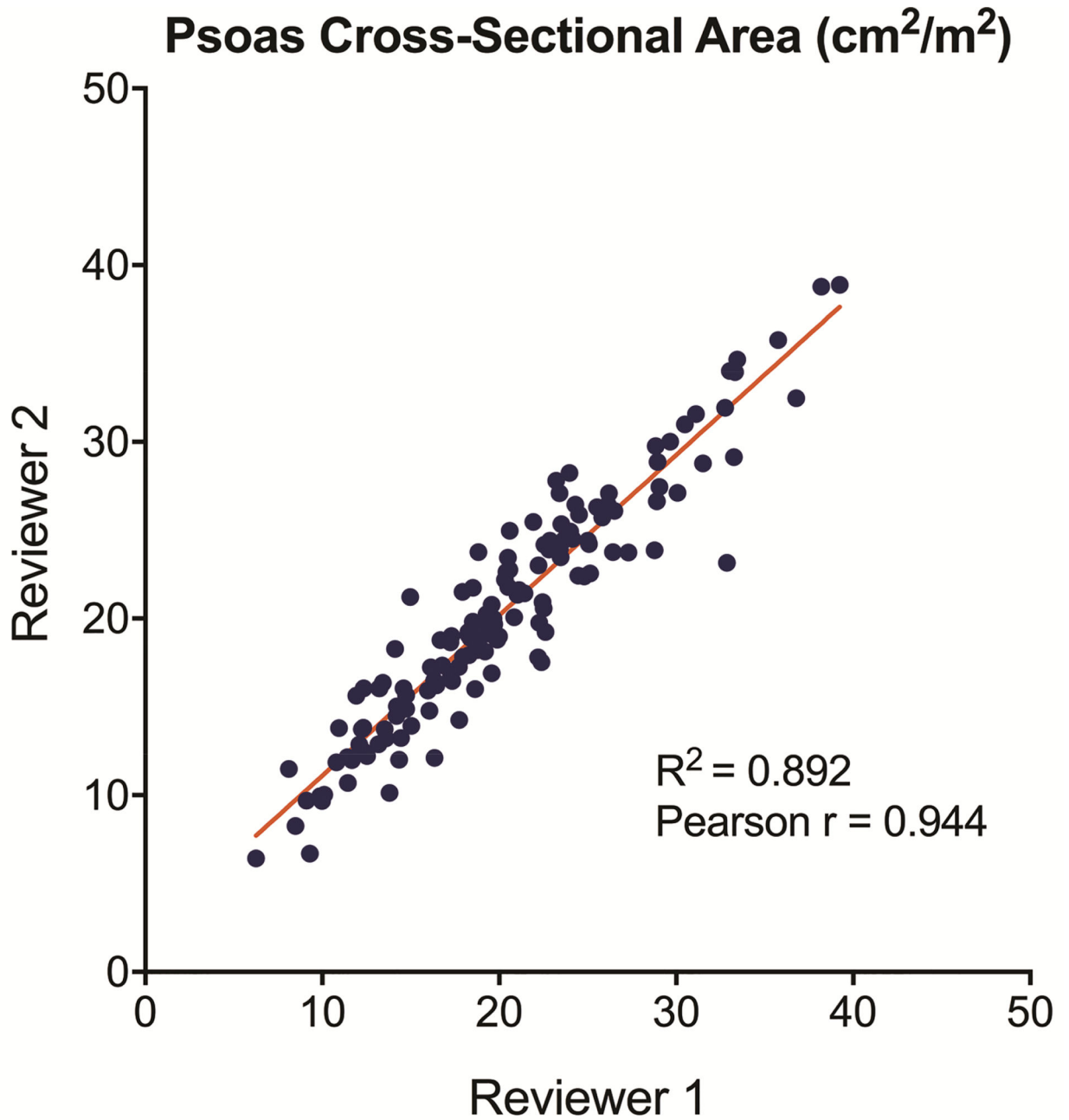


Figure 2.

Plot of psoas cross-sectional area measurements for both reviewers with Pearson correlation coefficient and linear regression results.

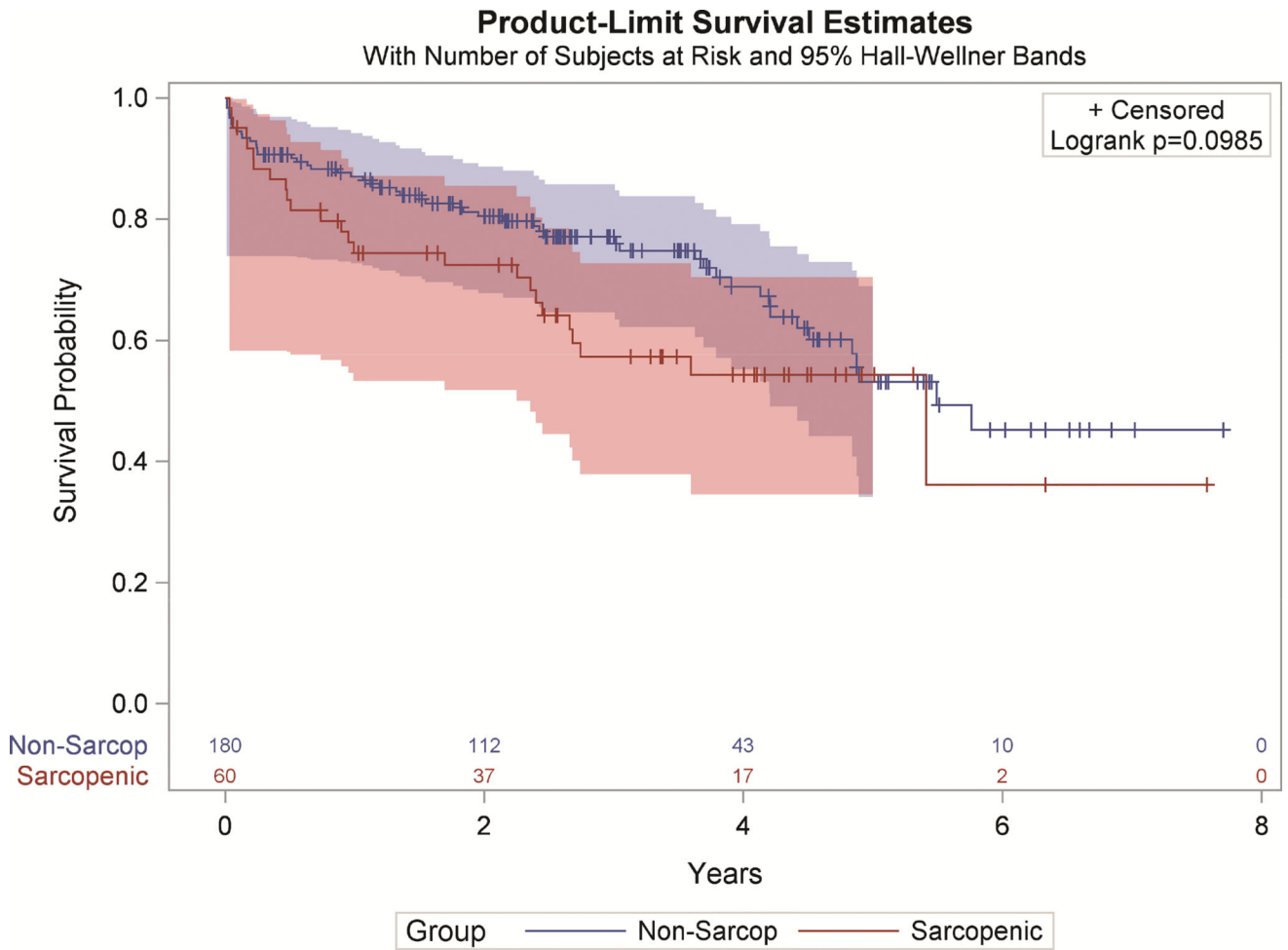


Figure 3. Kaplan Meier Survival analysis by Sarcopenia status. Number at risk is displayed below the figure and 95% confidence intervals are displayed by the thin lines in the corresponding color.

Table 1

Baseline and operative characteristics by sarcopenic status

<i>Baseline Characteristics</i>	Sarcopenic (n = 60)	Non-Sarcopenic (n = 180)	p value
Psoas Index (cm ² /m ²)	6.7 [5.7–7.9]	10.6 [9.4–12.2]	<0.0001
Age (years)	81 [77–85]	80 [75–85]	0.165
Body surface area (m ²)	1.96 ± 0.29	1.92 ± 0.27	0.350
Female	24 (40.0%)	73 (40.6%)	0.940
Smoker	4 (6.7%)	48 (26.7%)	0.001
Hypertension	51 (85.0%)	162 (90.0%)	0.289
Diabetes	26 (43.3%)	82 (45.6%)	0.764
Dialysis dependent renal failure	2 (3.3%)	1 (0.6%)	0.094
Prior stroke	9 (15.0%)	18 (10.0%)	0.289
Chronic lung disease, moderate/severe	10 (16.7%)	47 (26.3%)	0.131
Prior myocardial infarction	20 (33.3%)	65 (36.1%)	0.697
Heart failure within 2 weeks	45 (75.0%)	154 (85.6%)	0.060
Ejection fraction (%)	57 [50–63]	57 [43–63]	0.160
Aortic insufficiency (moderate/severe)	4 (6.7%)	19 (10.6%)	0.376
Mitral insufficiency (moderate/severe)	15 (26.3%)	40 (23.3%)	0.639
Predicted risk of mortality (%)	5.7% [3.9–8.4%]	6.0% [4.2–7.9%]	0.285
<i>Operative Characteristics</i>			
Prior cardiac surgery	15 (46.9%)	45 (45.9%)	0.925
Urgent or Emergent status	14 (23.3%)	48 (26.7%)	0.610
Coronary artery bypass grafting	17 (28.3%)	67 (37.2%)	0.211
Cross clamp time (min)	69 [61.5–85]	73 [60–90]	0.375
Cardiopulmonary bypass time (min)	105.5 [89.5–123]	103.5 [87–124]	0.363

Table 2

Outcomes by sarcopenic status

	Sarcopenic (n = 60)	Non-Sarcopenic (n = 180)	p value
STS operative mortality	3 (5.0%)	10 (5.6%)	0.869
One-year mortality (n=183)	15 (31.9%)	23 (16.9%)	0.029
STS major morbidity	14 (23.3%)	36 (20.0%)	0.582
Permanent stroke	3 (5.0%)	5 (2.8%)	0.406
Cardiac arrest	2 (3.3%)	3 (1.7%)	0.434
Prolonged ventilation	11 (18.3%)	19 (10.6%)	0.115
Renal failure requiring dialysis	4 (6.7%)	6 (3.3%)	0.263
Deep sternal wound infection	0 (0%)	1 (0.6%)	0.563
Atrial fibrillation	20 (33.3%)	42 (23.3%)	0.125
Pneumonia	5 (8.3%)	7 (3.9%)	0.171
Transfusion, packed red blood cells	34 (56.7%)	101 (56.1%)	0.940
Reoperation for any reason	7 (11.7%)	15 (8.3%)	0.438
Readmission	8 (13.6%)	28 (16.2%)	0.631
Discharge to facility	36 (63.2%)	85 (49.1%)	0.066
Hospital cost (median)	\$51905	\$51787	0.328
Length of stay (days; median, IQR)	7.5 (6–12)	7 (5–9)	0.056
ICU stay (hrs; median, IQR)	46.7 [30.0–139.5]	87 [41.4–125.7]	0.208

ICU = intensive care unit; STS = Society of Thoracic Surgeons

Table 3

Risk-adjusted outcomes for psoas index

	Odds Ratio	95% Confidence Limit	p-value	c-statistic	
STS operative mortality	0.94	0.77	1.15	0.568	0.690
One-year mortality (n=183)	0.84	0.73	0.97	0.017	0.647
STS major morbidity	0.86	0.76	0.98	0.020	0.705
Prolonged ventilation	0.81	0.69	0.95	0.012	0.730
Dialysis dependent renal failure	0.84	0.65	1.10	0.200	0.770
Permanent stroke	0.89	0.67	1.17	0.401	0.728
Atrial fibrillation	0.90	0.81	1.00	0.058	0.574
Readmission	0.95	0.84	1.08	0.444	0.610
Discharge to a facility	0.88	0.80	0.96	0.005	0.612

	Estimate	95% Confidence Limit	p-value	χ^2	
Hospital cost (\$; log form)	-0.03	-0.05	-0.01	0.001	69
Postoperative length of stay (d)	-0.46	-0.65	-0.27	<0.0001	407
ICU length of stay (hr)	-2.41	-5.40	0.57	0.113	477

ICU = intensive care unit; STS = Society of Thoracic Surgeons