The European System for Cardiac Operative Risk Evaluation (EuroSCORE) is not appropriate for withholding surgery in high-risk patients with aortic stenosis: a retrospective cohort study

Dimitri Kalavrouziotis*, Debbie Li, Karen J Buth and Jean-Francois Légaré

Address: Department of Surgery, Division of Cardiac Surgery, Queen Elizabeth II Health Sciences Centre, Dalhousie University, 1796 Summer Street, room 2269, Halifax, Nova Scotia, B3H 3A7, Canada

Email: Dimitri Kalavrouziotis* - dkalavro@dal.ca; Debbie Li - debbie.x.li@gmail.com; Karen J Buth - kbuth@dal.ca; Jean-Francois Légaré - Jean.Legare@cdha.nshealth.ca
* Corresponding author

Abstract

Background: The European System for Cardiac Operative Risk Evaluation (EuroSCORE) is a widely used risk assessment tool in patients with severe aortic stenosis to determine operability and to select patients for alternative therapies such as transcatheter aortic valve implantation. The objective of this study was to determine the accuracy of the EuroSCORE in predicting mortality following aortic valve replacement (AVR).

Methods: The logistic EuroSCORE was determined for all consecutive patients that underwent conventional AVR between 1995 and 2005 at our institution. Provincial Vital Statistics were used to determine all-cause mortality. The accuracy of the prognostic risk prediction provided by logistic EuroSCORE was assessed by comparing observed and expected operative mortality.

Results: During the study period, a total of 1,421 patients underwent AVR including 237 patients (16.7%) that had a logistic EuroSCORE > 20. Among these patients, the mean predicted operative mortality was 38.7% (SD = 18.1). The actual mortality of these patients was significantly lower than that predicted by EuroSCORE (11.4% vs. 38.7%, observed/expected ratio 0.29, 95% CI 0.15–0.52, P < 0.05). The EuroSCORE overestimated mortality within all strata of predicted risk. Although medium-term mortality is significantly higher among patients with EuroSCORE > 20 (log rank P = 0.0001), approximately 60% are alive at five years.

Conclusion: Actual operative mortality in patients undergoing AVR is significantly lower than that predicted by the logistic EuroSCORE. Additionally, medium-term survival following AVR is acceptable in high-risk patients with EuroSCORE > 20. More accurate risk prediction models are needed for risk-stratifying patients with severe aortic stenosis.

Introduction

Aortic valve replacement (AVR) is established therapy for the treatment of severe, symptomatic aortic stenosis (AS) and has been shown to relieve symptoms and significantly improve survival [1-4]. Conventional AVR involves median sternotomy and the institution of cardiopulmonary bypass, excision of the stenotic native valve, and replacement with a biologic or mechanical prosthesis.
Recently, catheter-based transfemoral and transapical approaches to aortic valve implantation have been developed for patients with severe AS that are thought to be inoperable because of advanced age or significant comorbidities, with acceptable early results [5-13].

In these studies, patient selection for transcatheter therapy was determined largely on the basis of the EuroSCORE [14,26]. The score thus obtained corresponds exactly to the operative mortality predicted for a given patient, defined as death occurring anytime during the same postoperative hospital stay of any duration or within 30 days of surgery for patients discharged to home or to a secondary institution. The distribution of demographic, clinical, and operative variables was compared among patients with EuroSCORE ≤ 20 and those with EuroSCORE > 20. A EuroSCORE cut-off of 20, representing a predicted operative mortality of 20%, was chosen a priori to define low- and high-risk patient subgroups on the basis of previously published reports evaluating the safety, feasibility, and efficacy of transcatheter aortic valve implantation in which patients with EuroSCORE > 20 were considered to have too high a risk to be candidates for conventional AVR and were subsequently referred for catheter-based aortic valve implantation [5-13,27].

**Clinical outcomes**

The primary outcome of interest in this study was operative mortality, defined as death within 30 days of operation or within the same hospital admission. Because our institutional database captures in-hospital mortality only, we linked survivors to discharge to government vital statistics data to determine 30-day, 1, 3, and 5-year survival. Survival times were analyzed using Kaplan-Meier methodology and log-rank statistics were used to compare survival curves among patients with logistic EuroSCORE > 20 and those with logistic EuroSCORE ≤ 20 [28].

Other postoperative adverse events examined were: dependence on a mechanical ventilator for a period exceeding 24 hours, stroke, the transfusion of banked red blood cells, new-onset atrial fibrillation, the implantation of a permanent pacemaker secondary to new, postoperative complete heart block, and length of hospitalization from the time of operation to discharge. Stroke was defined as a permanent neurologic deficit occurring after surgery with an acute change on intracranial computed tomography or magnetic resonance imaging.

Postoperative delirium and acute, reversible deficits were not included in the stroke end-point. Individual outcomes were univariately compared among patients with logistic EuroSCORE ≤ 20 and those with logistic EuroSCORE > 20. The accuracy of the prognostic risk prediction provided by EuroSCORE was assessed by comparing observed and expected operative mortality. Ninety-five percent confidence intervals (CI) for the observed-to-expected ratios (O/E) were generated using Byar’s approximation [29].
Statistical analysis
Statistical analysis for the study was performed using Statistical Analysis System (SAS) software version 8.2 (SAS Institute, Cary, North Carolina). Continuous variables are expressed as mean ± 1 standard deviation (SD). Continuous variables were compared using unpaired t-tests or non-parametric equivalents where appropriate. Proportions were compared using chi-square tests (or Fisher’s exact test where appropriate) and Mantel-Haenszel chi-square statistics with one degree of freedom were used to test for trends across ordered categories. Comparisons were considered statistically significant if the P value was less than 0.05. All P values were two-sided.

The study protocol was in accordance with the institutional clinical research ethics board and received full ethical approval.

Results
Baseline characteristics
Between 1995 and 2005, 1,421 patients underwent AVR at our institution. Among these patients, 237 (16.7%) had a logistic EuroSCORE > 20. Patients with logistic EuroSCORE > 20 were more likely to have a number of high-risk features such as advanced age, depressed systolic function of the left ventricle, and chronic renal insufficiency, although were not different from patients with logistic EuroSCORE ≤ 20 with respect to gender, or the prevalence of diabetes mellitus, hypertension, hyperlipidemia, and active or former cigarette smoking (Table 1). In addition, the acuity of operation, the type of prosthesis implanted, the need for concomitant revascularization or thoracic aortic repair, and operative durations also differed among the two groups (Table 2).

Early outcomes
The overall operative mortality, defined as death occurring anytime during the same postoperative hospital stay or within 30 days of surgery for patients discharged to home or to a secondary institution, for the entire cohort of patients was 4.6% (n = 66). The operative mortality was significantly higher among patients with logistic EuroSCORE > 20 compared to those with EuroSCORE ≤ 20 (11.4% vs. 3.2%, respectively, P < 0.0001). Of the pre-specified in-hospital adverse events following surgery examined, only the incidence of new-onset postoperative atrial fibrillation was similar between the two groups (Table 3).

Table 1: Baseline characteristics of patients with EuroSCORE > 20 compared to those with EuroSCORE ≤ 20

| Variable, % unless otherwise indicated | EuroSCORE > 20, n = 237 | EuroSCORE ≤ 20, n = 1,184 |
|----------------------------------------|--------------------------|---------------------------|
| Demographic data                       |                          |                           |
| Age                                    |                          |                           |
| <60 yrs                                | 11.4                     | 24.7                      |
| 60–69 yrs                              | 13.9                     | 25.4                      |
| 70–79 yrs                              | 42.6                     | 37.3                      |
| ≥ 80 yrs                               | 32                       | 12.6                      |
| Age, mean yrs ± SD                     | 73.2 ± 12.4              | 66.7 ± 12.8               |
| Female                                 | 35.0                     | 33.8                      |
| Cardiovascular risk factors/diseases   |                          |                           |
| Current or former cigarette smoking    | 57.0                     | 61.5                      |
| Body mass index, mean kg/m² ± SD       | 26.6 ± 4.6               | 28.2 ± 5.3                |
| Active endocarditis                    | 8.0                      | 1.1                       |
| Diabetes mellitus                      | 22.8                     | 21.5                      |
| Hypertension                           | 59.9                     | 53.3                      |
| Extracardiac arteriopathy              | 56.5                     | 18.6                      |
| Pulmonary hypertension                 | 31.2                     | 5.2                       |
| Previous cardiac surgery               | 30.4                     | 7.9                       |
| Left ventricular ejection fraction     |                          |                           |
| >0.50                                  | 37.6                     | 80.4                      |
| 0.30–0.50                              | 35.4                     | 16.4                      |
| <0.30                                  | 27.0                     | 3.2                       |
| Unstable angina                        | 14.4                     | 0.8                       |
| MI within 21 days of surgery           | 13.1                     | 2.6                       |
| Hyperlipidemia                         | 48.5                     | 49.1                      |
| Medical comorbidities                  |                          |                           |
| Hemoglobin, mean g/dL ± SD             | 12.2 ± 1.9               | 13.5 ± 1.7                |
| Serum creatinine > 200 μmol/L          | 15.2                     | 1.9                       |
| Chronic lung disease                   | 33.3                     | 15.0                      |

MI = myocardial infarction; SD = standard deviation.
Among the subgroup of patients with logistic EuroSCORE > 20, the mean predicted operative mortality by EuroSCORE was 38.7% ± 18.1%. The actual mortality of these patients was significantly lower than that predicted by EuroSCORE (11.4% vs. 38.7%, O/E 0.29, 95% CI 0.15 – 0.52, P < 0.05) (Figure 1). In the lower-risk subgroup of patients with logistic EuroSCORE ≤ 20, mean predicted operative mortality was 7.1% ± 4.8%. The observed mortality in these patients was 3.2%, although this difference was not statistically significant (O/E 0.47, 95% CI 0.10 – 1.30, P > 0.05). In addition, the EuroSCORE overestimated operative mortality within all strata of predicted risk, the magnitude of the discrepancy getting larger with increasing risk (Figure 2).

A simple linear regression model of actual mortality was fit as a function of EuroSCORE as the independent variable and compared to the predicted mortality given by EuroSCORE (Figure 3). Although there is a statistically significant association between EuroSCORE and operative mortality ($r^2 = 0.95$) such that increasing EuroSCORE is associated with increasing mortality, the two curves are widely divergent and the difference between them increases with increasing predicted risk.

### Late outcomes
Patients were linked to vital statistics databases to determine mortality from any cause. The Kaplan-Meier freedom from all-cause mortality was significantly lower for patients with logistic EuroSCORE > 20 compared to those with logistic EuroSCORE ≤ 20 at 1, 3, and 5 years: 80.9% vs. 93.5%, 68.5% vs. 87.9%, and 56.2% vs. 82.1%, respectively (log-rank P = 0.0001) (Figure 4).

### Discussion
The purpose of this study was to determine the early and medium-term outcomes of a contemporary cohort of patients with severe symptomatic AS undergoing AVR and to evaluate the accuracy of the EuroSCORE in predicting operative mortality. We have shown that in a tertiary care surgical practice in Canada a significant proportion of patients requiring AVR for severe symptomatic AS

### Table 2: Operative characteristics of patients with EuroSCORE > 20 compared to those with EuroSCORE ≤ 20

| Variable, % unless otherwise indicated | EuroSCORE > 20, n = 237 | EuroSCORE ≤ 20, n = 1,184 |
|----------------------------------------|-------------------------|---------------------------|
| Critical preoperative state*           | 24.5                    | 1.0                       |
| Emergency surgery                      | 10.1                    | 0.9                       |
| Concomitant thoracic aortic surgery    | 16.5                    | 5.7                       |
| Concomitant CABG                       | 58.7                    | 42.2                      |
| Type of prosthesis                    |                         |                           |
| Bioprosthesis (tissue valve)          | 84.0                    | 81.7                      |
| Mechanical valve                      | 13.5                    | 17.6                      |
| Homograft                              | 2.5                     | 0.8                       |
| Cardiopulmonary bypass time, mean min ± SD | 178.5 ± 64.4        | 148.8 ± 51.8              |
| Aortic clamp time, mean min ± SD      | 122.8 ± 47.0            | 106.7 ± 37.8              |
| Prosthetic valve indexed EOA§<0.75 cm²/m² | 4.7                      | 10.4                      |

*Includes: resuscitated cardiac arrest, dependence on mechanical ventilator, and cardiogenic shock requiring inotropic support and/or intra-aortic balloon counterpulsation.

§Effective orifice area was computed for each implanted valve type and size according to the measurements published by valve manufacturers and/or independent adjudicators and was indexed on patient body surface area.

CABG = coronary artery bypass graft surgery; EOA = effective orifice area; SD = standard deviation.

### Table 3: Early postoperative outcomes of patients with EuroSCORE > 20 compared to those with EuroSCORE ≤ 20

| Outcome, %                          | EuroSCORE > 20, n = 237 | EuroSCORE ≤ 20, n = 1,184 | P     |
|-------------------------------------|-------------------------|---------------------------|-------|
| Mortality (in-hospital or within 30 days of surgery) | 11.4                    | 3.2                       | <0.0001|
| In-hospital events                  |                         |                           |       |
| Ventilation > 24 hours              | 27.9                    | 9.3                       | <0.0001|
| Length of stay > 9 days             | 59.1                    | 28.1                      | <0.0001|
| Stroke                              | 5.1                     | 2.3                       | 0.02  |
| Atrial fibrillation (new onset)     | 31.7                    | 28.4                      | 0.31  |
| Erythrocyte transfusion             | 73.0                    | 34.5                      | <0.0001|
| Permanent pacemaker implantation    | 10.1                    | 5.4                       | 0.01  |
(16.7%) were found to have a high predicted operative mortality based on logistic EuroSCORE. We have arbitrarily chosen a logistic EuroSCORE > 20, which corresponds to a predicted in-hospital or 30-day mortality > 20%, as our cutoff of high risk based on published reports that have suggested that such patients may be considered inoperable and potentially eligible for alternative catheter-based therapy. Among the patients in our cohort with logistic EuroSCORE > 20, the mean operative mortality predicted by EuroSCORE was 38.7%. However, the actual mortality observed was 11.4%, significantly lower than that predicted by EuroSCORE (P < 0.05). In addition, the EuroSCORE appeared to overestimate operative mortality within all strata of predicted risk. The discrepancy logistic EuroSCORE > 20, the mean operative mortality predicted by EuroSCORE was 38.7%. However, the actual mortality observed was 11.4%, significantly lower than that predicted by EuroSCORE (P < 0.05). In addition, the EuroSCORE appeared to overestimate operative mortality within all strata of predicted risk. The discrepancy

Figure 1
Predicted and observed mean operative mortality among patients with EuroSCORE > 20 compared to those with EuroSCORE ≤ 20. Among high-risk patients with logistic EuroSCORE > 20, the actual operative mortality following AVR was significantly lower than that predicted by EuroSCORE (11.4% vs. 38.7%, respectively, O/E 0.29, 95% CI 0.15–0.52, P < 0.05).

Figure 2
Predicted and observed mean operative mortality within all subgroups of EuroSCORE. The EuroSCORE overestimated operative mortality in all categories of risk.

Figure 3
Simple linear regression model of observed mortality as a function of logistic EuroSCORE compared with predicted mortality. There is a statistically significant association between logistic EuroSCORE and actual operative mortality (r² = 0.95), although the empirical model is different from the theoretical model provided by EuroSCORE.

Figure 4
Five-year freedom from all-cause mortality. The Kaplan-Meier survival among patients with logistic EuroSCORE > 20 is significantly worse compared to those with logistic EuroSCORE ≤ 20.
between observed and expected mortality widened as pre-operative risk determined by EuroSCORE increased. Although long-term mortality from any cause is significantly higher among patients with EuroSCORE > 20 (log rank P = 0.0001), approximately 60% are alive at five years after operation. Similar findings were recently reported by Brown et al. However, unlike our study, no data are given on longitudinal outcomes.

The EuroSCORE predictive model for cardiac surgery has been extensively studied and found to accurately predict outcomes in a variety of jurisdictions and clinical practices. The initial additive EuroSCORE model was simple and reproducible and it could readily be used at the patient's bedside for rapid assessment of operative risk. However, what it gained in parsimony, interpretability, and ease of use, it lost in accuracy as it was found to have a tendency to over-predict mortality in low-risk patients and under-predict mortality in high-risk patients. The later-appearing logistic EuroSCORE, utilizing the complete regression equation of the multivariate model to predict mortality risk, was more stable at the extremes of risk and had greater predictive accuracy.

However, we have shown that the prognostic performance of the logistic EuroSCORE is poor in patients undergoing AVR, and progressively worsens as risk increases. Similar findings were reported by others in octogenarians undergoing valve operations and in patients undergoing coronary artery bypass graft surgery. It has been shown that subtle inter-institutional differences in the definitions of variables, in the sensitivity of risk factor screening tools, and in the potential presence of various degrees of interdependence and collinearity among covariates assumed to be independent may profoundly affect the performance of a risk prediction model such as the EuroSCORE. However, these influences are likely to be minor in our study given the availability of a nuanced dataset at our institution in which patient information on over 200 variables was prospectively collected at the time of surgery with minimal retrospective chart review, allowing us to adhere to the EuroSCORE definitions of variables as faithfully as possible and to score patients accurately. In no instance was risk factor status imputed secondary to incomplete or missing data fields.

Recent reports suggest that 30–40% of patients in whom AVR is indicated are denied surgical treatment, mostly secondary to advanced age and left ventricular dysfunction. Less invasive, catheter-based approaches to relieving the obstruction to left ventricular outflow in severe AS were developed in an attempt to target the subset of patients who were not felt to be candidates for conventional open AVR in order to potentially favorably alter their dismal prognosis. Initially, transcatheter AVR was offered to patients on compassionate grounds and was largely reserved for moribund patients with end-stage AS in whom risk factors were felt to be prohibitive for traditional surgery. As experience with transcatheter aortic valve implantation increased with attendant improvements in feasibility, safety, and efficacy profiles, patient selection criteria were expanded and investigators are increasingly relying on EuroSCORE to determine eligibility for catheter-based intervention on the aortic valve. Recent reports evaluating retrograde transfemoral and transapical aortic valve implantation have included patients with mean logistic EuroSCORE 11–35. In these studies, 30-day mortality rates range from 7 to 22% and the incidence of stroke ranges from 0 to 10%. However, these studies are limited by small numbers of patients and lack long-term data. In our study, the incidence of early stroke was 5.1% in the subgroup of patients with logistic EuroSCORE > 20; 30-day mortality in this high-risk group of patients was 11.4%. Furthermore, five-year survival was acceptable at nearly 60% despite a predicted operative mortality of approximately 40% in this cohort of patients. Taken further, this very same cohort of patients could have been potentially denied life-prolonging operation under current EuroSCORE criteria. However, this study is not a comparison of the effectiveness of conventional AVR and transcatheter aortic valve implantation, nor is it an analysis of the relative merits of one approach over the other. Importantly, our study does not attempt to question the value of important novel approaches to valvular heart disease but questions the validity of using EuroSCORE as a predictive model for high-risk patients with severe AS and suggests that caution should be exercised when using the EuroSCORE risk model to make important management decisions in this complex patient population. This is particularly important when one attempts to make inter-institutional comparisons, compare individual surgeons with different case mixes, predict risk for any single individual patient, or determine patient eligibility for surgery.

Study limitations
These data reflect the experience of a single center which may limit the study’s generalizability. However, our surgical approach to AVR is not likely to be substantially different from the practice at other centers. Also, our overall 30-day mortality of 4.6% following AVR and 11.4% in high-risk patients is within the lower range of post- AVR mortality data reported by the Society of Thoracic Surgeons who have examined more than 40,000 patients.

Another important limitation relates to the lack of data on the larger population of patients with severe aortic stenosis who are never presented to our multidisciplinary peer review conference, either because of an a priori judgment made by the health care provider as to a patient’s lack of
“operability,” or a patient’s own refusal to be considered for surgery. Therefore, we cannot comment on this wider denominator of untreated patients in the community. However, previous work from our group showed that 88.9% of patients presented to conference eventually undergo operation [23], which is higher than most published reports and is consistent with the existence of a more “aggressive” approach at our institution in accepting patients for surgery. Therefore, it is unlikely that a meaningful selection bias is present in this study such that truly high-risk patients are systematically turned away from surgery, and this limitation does not weaken our data or the study’s validity in reporting the outcome of 237 consecutive high-risk patients with severe aortic stenosis and a EuroSCORE of greater than 20. Moreover, in a recent study of patients refused for AVR, mostly by cardiac surgeons, no differences in EuroSCORE were identified between those patients who underwent operation compared to those who did not [37].

Conclusion

In our consecutive series of patients with severe AS undergoing AVR, we found that the logistic EuroSCORE was not an accurate risk assessment tool in all categories of risk but especially in high-risk patients. Therefore, this predictive model should not be used to determine procedural risk in patients with severe AS. Furthermore, the utilization of the logistic EuroSCORE in the assessment of operability in patients with severe AS may not be appropriate. More accurate risk prediction models are needed for risk-stratifying patients with severe AS.

Competing interests

The authors declare that they have no competing interests.

Authors’ contributions

DK and JFL conceived the research question, study design and coordination, and drafted the manuscript. DK, DL, and KJB participated in the design of the study and performed the data acquisition and statistical analysis. All authors read and approved the final manuscript.

Acknowledgements

This work was supported by a Canadian Institutes of Health Research (CIHR) Team Grant in Cardiovascular Outcomes Research (DK).

References

1. Schwarz F, Baumann P, Manthey J, Hoffmann M, Schuler G, Mehmel HC, Schmitz W, Kühbler W: The effects of aortic valve replacement on survival. Circulation 1982, 66:1105-1110.
2. Sharma UC, Barenbrug P, Pokharel S, Dassen WR, Pinto YM, Maessen JG: Systematic review of the outcome of aortic valve replacement in patients with aortic stenosis. Ann Thorac Surg 2004, 78:90-95.
3. Ross J, Braunwald E: Aortic stenosis. Circulation 1968, 38:61-67.
4. Bonow RO, Carabello BA, Kanu C, de Leon AC Jr, Faxon DP, Freed MD, Gaasch WH, Lytle BW, Nishimura RA, Oliva PT, O’Rourke RA, Otto CM, Shah PM, Shawcross JS, Smith SC Jr, Jacobs AK, Adams CD, Anderson JL, Antman EM, Faxon DP, Fuster V, Halperin JL, Hiratzka LF, Hunt SA, Lytle BW, Nishimura R, Page RL, Riegel B: ACC/AHA 2006 guidelines for the management of patients with valvular heart disease: a report of the American College of Cardiology/American Heart Association Task Force on Practice Guidelines (writing committee to revise the 1998 Guidelines for the Management of Patients With Valvular Heart Disease): developed in collaboration with the Society of Cardiovascular Anesthesiologists: endorsed by the Society for Cardiovascular Angiography and Interventions and the Society of Thoracic Surgeons. Circulation 2006, 114:e221.
5. Cribier A, Etchaninoff H, Bash A, Borenstein N, Tron C, Bauer F, Del Vallemaux G, Anseline F, Laborde F, Leon MS: Percutaneous transcatheater implantation of an aortic valve prosthesis for calcific aortic stenosis: first human case description. Circulation 2002, 106:3006-3008.
6. Grube E, Laborde JC, Gerckens U, Felderhoff T, Sauren B, Buellesfeld L, Mueller R, Menichelli M, Schmidt T, Zickmann B, Iversen S, Stone GW: Percutaneous implantation of the CoreValve self-expanding valve prosthesis in high-risk patients with aortic valve disease: the Siegburg first-in-man study. Circulation 2006, 114:1616-1624.
7. Wolner T, Falk V, Borger MA, Dewey T, Wimmer-Greinecker G, Schuler G, Mack M, Mohr FW: Minimally invasive transapical beating heart aortic valve implantation – proof of concept. Eur J Cardiothorac Surg 2007, 31:9-15.
8. Webb JG, Pasupati S, Humphries K, Thompson C, Altwegg L, Moss R, Sinhal A, Carere RG, Munt B, Ricci D, Ye J, Cheung A, Lichtenstein SV: Percutaneous transarterial aortic valve replacement in selected high-risk patients with aortic stenosis. Circulation 2007, 116:755-763.
9. Webb JG, Chandavimol M, Thompson CR, Ricci DR, Carere RG, Munt BI, Boller CE, Pasupati S, Lichtenstein S: Percutaneous aortic valve implantation retrograde from the femoral artery. Circulation 2006, 113:842-850.
10. Cribier A, Etchaninoff H, Tron C, Bauer F, Agatiello C, Nercolini D, Tapiero S, Litzler PY, Besouw JP, Babalarios V: Treatment of calcific aortic stenosis with the percutaneous heart valve: mid-term follow-up from the initial feasibility studies: the French experience. J Am Coll Cardiol 2006, 47:1214-1223.
11. Lichtenstein SV, Cheung A, Ye J, Thompson CR, Carere RG, Pasupati S, Webb JG: Transapical transcatheater aortic valve implantation in humans: initial clinical experience. Circulation 2006, 114:591-596.
12. Grube E, Schuler G, Buellesfeld L, Gerckens U, Linke A, Wenaweser P, Sauren B, Mohr FW, Wolter T, Zickmann B, Iversen S, Felderhoff T, Cartier R, Bonan R: Percutaneous aortic valve replacement for severe aortic stenosis in high-risk patients using the second- and current-generation self-expanding CoreValve prostheses: device success and 30-day clinical outcome. J Am Coll Cardiol 2007, 50:96-101.
13. Wolter T, Simon P, Dewey T, Wimmer-Greinecker G, Falk V, Kamirat MT, Doss M, Borger MA, Schuler G, Glogar D, Fehske W, Wolner E, Mohr FW, Mack M: Transapical minimally invasive aortic valve implantation: multicenter experience. Circulation 2007, 116(Suppl):I240-I245.
14. Nashraf SA, Roques F, Michel P, Gauducheau E, Lemoshow S, Salamon R: European System for Cardiac Operative Risk Evaluation (EuroSCORE). Eur J Cardiothorac Surg 1999, 16:9-13.
15. Roques F, Nashraf SAM, Michel P, Pinna Pintor P, David M, Baudet E, The EuroSCORE Study Group: Does EuroSCORE work in individual European countries? Eur J Cardiothorac Surg 2000, 18:27-30.
16. Nashraf SAM, Roques F, Hamill BG, Peterson ED, Michel P, Grover FL, Wyse RK, Ferguson TB, EuroSCORE Project Group: Validation of European System for Cardiac Operative Risk Evaluation (EuroSCORE) in North American cardiac surgery. Eur J Cardiothorac Surg 2002, 22:101-105.
17. Kurki TS, Jarvinen O, Kataja MJ, Laurikka J, Tarkka M: Performance of three preoperative risk indices: CABDEAL, EuroSCORE, and Cleveland models in a prospective coronary bypass database. Eur J Cardiothorac Surg 2002, 21:406-410.
18. Berman M, Samler A, Sahar G, Georgiou GP, Shariati E, Brauner R, Medalion B, Vidine BA, Kogan A: Validation of the 2000 Bernstein-Parsonnet Score versus the EuroSCORE as a prognostic tool in cardiac surgery. Ann Thorac Surg 2006, 81:537-540.
19. Florath I, Rosendahl UP, Mortasawi A, Bauer SF, Dalladaku F, Ennker IC, Ennker JC: Current determinants of operative mortality in 1400 patients requiring aortic valve replacement. J Thorac Cardiovasc Surg 2003, 126:75-83.

20. Ursu S, Sadaba R, Greco E, Pulitani I, Alvarez L, Juaristi A, Goiti JJ: One-hundred aortic valve replacements in octogenarians: outcomes and risk factors for early mortality. J Heart Valve Dis. 2007, 16(2):139-144.

21. Collart F, Feher H, Kerbaul F, Moully-Bandini A, Riberi A, Mesana TG, Metras D: Valvular surgery in octogenarians: operative risk factors, evaluation of EuroSCORE and long term results. Eur J Cardiothorac Surg 2005, 27:276-280.

22. Brown ML, Schaff HV, Sarano ME, Li Z, Sundt TM, Dearani JA, Mullany CJ, Orszulak TA: Is the European System for Cardiac Operative Risk Evaluation model valid for estimating the operative risk of patients considered for percutaneous aortic valve replacement? J Thorac Cardiovasc Surg 2008, 136:566-571.

23. Ray AA, Buth KJ, Sullivan JA, Johnstone DE, Hirsch GM, Ross DB, Sullivan JA: Waiting for cardiac surgery: results of a risk-stratified queuing process. Circulation. 2001, 104(12 Suppl 1):92-98.

24. Baskett RJ, Buth KJ, Legare JF, Hassan A, Hancock Friesen C, Hirsch GM, Ross DB, Sullivan JA: Is it safe to train residents to perform cardiac surgery? Ann Thorac Surg 2002, 74:1043-1048.

25. Shroyer AL, Plomondon ME, Grover FL, Edwards FH: The 1996 coronary artery bypass risk model: the Society of Thoracic Surgeons Adult Cardiac National Database. Ann Thorac Surg 1999, 67:1203-1208.

26. Roques F, Michel P, Goldstone AR, Nashef SA: The logistic EuroSCORE. Eur Heart J 2003, 24:881-882.

27. Kodali SK, O’Neill W, Moses JW, Kapadia S, Williams M, Hanzel G, Stewart A, Tuzcu ME, Collins M, Leon MB: Preliminary results from the Percutaneous Endovascular Implantation of Valves Trial in high risk patients with critical aortic stenosis [abstract]. Am J Cardiol 2006, 98(Suppl 1):48M.

28. Salzman K, Hillier SC, Todd K, Wainer W, Bhatnagar A, Bosmans J, Stewart R, Tsang S, Lundquist K, Lack M: The prospective use of the Society of Thoracic Surgeons Adult Cardiac National Database. J Thorac Cardiovasc Surg 2000, 119:1939-1945.

29. Urso S, Sadaba R, Greco E, Pulitani I, Alvarez L, Juaristi A, Goiti JJ: Current determinants of operative mortality in 1400 patients requiring aortic valve replacement. J Thorac Cardiovasc Surg 2003, 126:75-83.

30. Ursu S, Sadaba R, Greco E, Pulitani I, Alvarez L, Juaristi A, Goiti JJ: One-hundred aortic valve replacements in octogenarians: outcomes and risk factors for early mortality. J Heart Valve Dis. 2007, 16(2):139-144.

31. Collart F, Feher H, Kerbaul F, Moully-Bandini A, Riberi A, Mesana TG, Metras D: Valvular surgery in octogenarians: operative risk factors, evaluation of EuroSCORE and long term results. Eur J Cardiothorac Surg 2005, 27:276-280.

32. Brown ML, Schaff HV, Sarano ME, Li Z, Sundt TM, Dearani JA, Mullany CJ, Orszulak TA: Is the European System for Cardiac Operative Risk Evaluation model valid for estimating the operative risk of patients considered for percutaneous aortic valve replacement? J Thorac Cardiovasc Surg 2008, 136:566-571.

33. Ray AA, Buth KJ, Sullivan JA, Johnstone DE, Hirsch GM, Ross DB, Sullivan JA: Waiting for cardiac surgery: results of a risk-stratified queuing process. Circulation. 2001, 104(12 Suppl 1):92-98.

34. Baskett RJ, Buth KJ, Legare JF, Hassan A, Hancock Friesen C, Hirsch GM, Ross DB, Sullivan JA: Is it safe to train residents to perform cardiac surgery? Ann Thorac Surg 2002, 74:1043-1048.

35. Shroyer AL, Plomondon ME, Grover FL, Edwards FH: The 1996 coronary artery bypass risk model: the Society of Thoracic Surgeons Adult Cardiac National Database. Ann Thorac Surg 1999, 67:1203-1208.

36. Roques F, Michel P, Goldstone AR, Nashef SA: The logistic EuroSCORE. Eur Heart J 2003, 24:881-882.

37. Kodali SK, O’Neill W, Moses JW, Kapadia S, Williams M, Hanzel G, Stewart A, Tuzcu ME, Collins M, Leon MB: Preliminary results from the Percutaneous Endovascular Implantation of Valves Trial in high risk patients with critical aortic stenosis [abstract]. Am J Cardiol 2006, 98(Suppl 1):48M.

38. Salzman K, Hillier SC, Todd K, Wainer W, Bhatnagar A, Bosmans J, Stewart R, Tsang S, Lundquist K, Lack M: The prospective use of the Society of Thoracic Surgeons Adult Cardiac National Database. J Thorac Cardiovasc Surg 2000, 119:1939-1945.