## ORIGINAL ARTICLE

# Probabilistic Model for Prediction of Prognostics in Myocardial Revascularization: Complications in Coronary Surgery

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#### **Abstract**

**Introduction:** Risk scores evaluate pre-operatory risk and present support for clinical decisions, however the performance of these tools in samples different from the original ones remains unclear.

**Objectives:** Investigate the external validity of risk scores (STS and Euroscore) in cardiac surgery and the predictive performance of clinical features derived from the sample.

**Methods:** Retrospective Cohort study conducted between October, 2010, and April, 2015. We used logistic regression to identify risk factors for hospital morbidity. The sample was divided for cross-validation, with 2/3 of the patients selected for model fitting and 1/3 for prediction testing. The performance of risk scores and clinical features was evaluated through AUROC and calibraton the Hosmer-Lemeshow test (H-L).

**Results:** Data was retrieved from 472 patients who underwent coronary cardiac surgery in Hospital Santa Izabel da Santa Casa, BA. Mean age was 62.8 years old and 32.5% of the sample were women. Traditional surgical risk scores did not present significant discriminative performance for this sample. Factors associated with the outcome after adjusting for covariates were: age, previous myocardial revascularization and pre-surgical creatinine levels. The adjusted model presented similar discrimination and calibration values during training (AUROC = 0,72; IC 95% 0,59-0,84; H-L valor p: 0,41) and validation (AUROC = 0,70; IC 95% 0,55 - 0,84; H-L valor p: 0,197).

**Conclusion:** Traditional scores may be inaccurate when applied to different environments. New risk scores with good predictive power can be developed using local clinical variables. (Int J Cardiovasc Sci. 2017;30(4):307-312)

**Keywords:** Thoracic Surgery / complications; Myocardial Infarction; Myocardial Revascularization; Risk Factors; Forecasting.

#### Introduction

Multivariate probabilistic models have been used in cardiac surgery to estimate the risk of fatal and nonfatal complications.<sup>1</sup> The goal is to evaluate the balance between risks and benefits of procedures for patients, with a better allocation of resources.<sup>2</sup> There are some risk scores of death and occurrence of complications in patients undergoing myocardial revascularization surgery such as EuroSCORE<sup>3</sup> and STS Score.<sup>4</sup> Assessing the prognosis related to the natural history of a clinical condition, predictive variables are reproducible

in different settings.<sup>5,6</sup> On the other hand, when predicting the success or complications of medical procedures, it is possible that predictor variables have a variable value depending on the environment in which the procedure is implemented. This is because differences in the way a treatment is applied can make patients more or less vulnerable to risk determinants. Using a retrospective cohort performed at a tertiary hospital in Salvador, the present study aimed to test the external validity of traditional risk scores for surgical myocardial revascularization, to identify risk markers for

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myocardial revascularization surgery, and to construct a regional prediction model for complications related to the procedure.

#### **Methods**

#### Study design

Observational study of a retrospective cohort, using a database retrieved from the institution's records, fed with variables recorded between preoperative and patient discharge, used for research, after approval by the hospital's ethics council, under the registry number 24304713.9.3001.5544.

#### Sample selection criteria

All patients submitted to myocardial revascularization surgery at our service at Santa Izabel Hospital between October 2010 and April 2015 were included in the study sample. Patients with associated surgeries or those performed at other institutions were excluded.

#### Variables studied

The variables included in the analysis were: Gender, age, weight, height, body mass index, chronic obstructive pulmonary disease (COPD) - use of bronchodilator or corticoid, peripheral arteriopathy intermittent claudication, carotid artery obstruction greater than 50%; left ventricular dysfunction moderate 30-50% and significant less than 30%; previous neurological dysfunction – motor dysfunction affecting ambulation or daily function; previous cardiac surgery – previous opening of the pericardium; pre-and postoperative serum creatinine; endocarditis – antibiotic therapy for endocarditis at the time of surgery; unstable angina - use of venous nitrate; recent infarction - less than 90 days; pulmonary hypertension – pulmonary artery systolic pressure greater than 60 mmHg; previous myocardial revascularization; post-infarction ventricular septal defect; diabetes - use of oral hypoglycemic or insulin; smoking; hypertension – antihypertensive use; dyslipidemia – total cholesterol greater than 200 mg / dl, hypertriglyceridemia greater than 150 mg / dl, HDL cholesterol less than 40 mg / dl women and less than 50 mg / dl men; number of coronary lesions greater than 75%; left coronary trunk lesion greater than 50%; preoperative hypoxemia - artery oxygen pressure lower than 60 mmHg, emergency / urgency surgery need for intervention within 48 hours due to imminent

risk of death or unstable clinical-hemodynamic status; hemodynamic instability – ventricular tachycardia, ventricular fibrillation, cardiac arrest, mechanical ventilation, intra-aortic balloon use.

#### **Definition of outcome**

The main analysis was performed considering the composite outcome of major morbidity, including: stroke, stroke (central neurological deficit persisting for more than 72 hours); Prolonged intubation (more than 48 hours); Reoperation (tamponade or hemostasis); Mediastinitis (need for surgical reintervention, plus antibiotic therapy with or without positive culture), and death within 30 days after the surgical procedure. The events that made up the outcome were chosen based on models developed and validated from previous studies in cardiovascular surgery.<sup>7</sup>

#### Statistical analysis

Three logistic regression models were adjusted to test the predictive power of the scores in the sample: EuroSCORE, STS Mortality, and STS Morbidity. Each model was adjusted using the points of the respective score as the only independent variable. A proper model was adjusted following the algorithm proposed by Hosmer and Lemeshow<sup>8</sup> considering results of bivariate analysis and biological plausibility. The sample was divided into two parts: cohort derivation, intended for bivariate analysis and fit of the models (2/3 of the original sample, randomly selected); Cohort validation to test the obtained model (1/3 of the original sample, randomly selected). After obtaining the coefficients from the sample used for derivation, the model was tested using the validation sample. The Area under ROC Curve (AUROC) and model adequacy statistics are presented for comparison purposes. The analyzes were conducted using the programming language and R development environment.

## Results

Between October 2010 and April 2015, information from 472 consecutive patients submitted to myocardial revascularization was included in the database. No patient was excluded because of lack of information. The mean age was  $63 \pm 8.6$  years, 22.5% were women, 37% were diabetics, 32% were smokers, 18% had left ventricular dysfunction and 29% had left coronary artery

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disease. The incidence of the composite outcome was 37 cases (7.8%), resulting in death 12 (2.5%), stroke 15 (3.2%), cardiac tamponade 4 (0.8%), reoperation for revision of hemostasis 9 (1.9%), mediastinitis 1 (0.2%) and prolonged intubation 21 (4.4%).

### Prognostic value of traditional risk scores

The EuroSCORE did not show accuracy for prediction of surgical complications, with C-statistic of 0.507 (95% CI 0.415 - 0.599, p=0.310). The same was observed with the STS score, with STS Morbidity C-statistic of 0.568 (95% CI 0.473-0.665, p=0.160) and STS Mortality of 0.550 (95% CI 0.452-0.643, p=0.860). – Figure 1.

#### Derivation of the proper model

The variables predictor candidates were selected in 2/3 random of the total sample through univariate analysis, considering statistical significance (p < 0.20). Table 1. Multiple logistic regression analysis according to Hosmer-Lemeshow algorithm: cerebrovascular

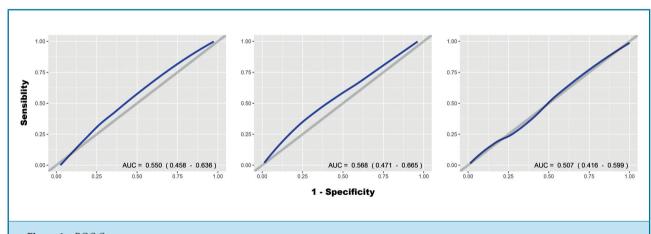
disease, previous myocardial revascularization, previous creatinine, age and ejection fraction.

The independent predictors, related to outcome, identified in the final model according to their statistical significance were: previous myocardial revascularization (OR 8.519 95% CI 1.026-59.381 p = 0.029), previous creatinine (OR 2,217 95% CI 0.815-6.018 p = 0.095) and age (OR 1.081 95% CI 1.025-1.145 p = 0.006). Table 2. In the shunt sample (2/3 of the total sample) the ROC curve was 0.72 (95% CI 0.60-0.84, p < 0.001). The resulting logistics model followed the formula below:

$$P(y) = \frac{1}{1 + e^{(8.422 + 0.07*Age + 2.14*MR_{prev} + 0.07*Cr_{prev})}}$$

## **AgePrevious**

Validation was performed in 168 patients (1/3 of the total sample) randomly selected. The score was discriminated by the area under the ROC curve of 0.70 (95% CI: 0.55-0.84); P = 0.008. Calibration by the Hosmer-Lemeshow test (p = 0.197).



**Figure 1 –** ROC Curves. A: STS Mortality; B: STS Morbidity; C: Euroscore.

## Discussion

The use of multivariate models in the form of scores represents the most accurate mean to predict risk, being superior to that predicted subjectively by clinical impression. And even showing good accuracy in different populations, especially in the clinical context, the results of the present study suggest greater caution regarding the external validity of these scores in the field of cardiac surgery. In addition, a

score developed in our local sample demonstrated good accuracy in an independent validation cohort, which may be the predictor in places with different characteristics of the traditional score validator centers. Models not readjusted to the local context may present bias in predicting risk in cardiac surgery and should be systematically compared with regional models. Among the various scores used to predict death and occurrence of complications in cardiac surgery, EuroSCORE<sup>3</sup> and STSscore<sup>4</sup> are the most widespread

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Table 1 – Bivariate analysis for combined outcome in derivation sub sample

	Combined outcome			
	Total (n = 304)	No (n = 281)	Yes (n = 23)	Value p
Age §	$62.4 \pm 8.9$	$61.9 \pm 8.9$	$67.6 \pm 8.2$	0.003
Male gender	228 (75.0)	209 (74.4)	19 (82.6)	0.531
$BMI(Kg/m^2) * \S$	$26.6 \pm 3.6$	$26.6 \pm 3.6$	$26.8 \pm 3.9$	0.788
Previous clearance //	83 (68 - 103)	84 (68 – 104)	81 (69 - 91)	0.454
Previous creatinine //	0,9 (0.8 - 1.1)	0.9 (0.8 - 1.1)	1.0 (0.9 - 1.3)	0.031
COPD †	6 (2.0)	6 (2.1)	0 (0.0)	1.000
Peripheral artery disease	13 (4.3)	11 (3.9)	2 (8.7)	0.619
Diabetes (In use of Hypoglycemic agent + Insulin)	109 (35.9)	100 (35.6)	9 (39.1)	0.909
Vascular Brain Disease	2 (0.7)	1 (0.4)	1 (4.3)	0.149
‡ Previous CABG	5 (1.6)	3 (1.1)	2 (8.7)	0.044
Emergency	19 (6.2)	18 (6.4)	1 (4.3)	1.000
Smoker	94 (30.9)	88 (31.3)	6 (26.1)	0.900
Antyhipertensive drug	263 (86.5)	242 (86.1)	21 (91.3)	0.728
Dyslipedemia	263 (86.5)	243 (86.5)	20 (87.0)	1.000
Coronary trunk lesion	87 (28.6)	82 (29.2)	5 (21.7)	0.597
Ejection Fraction				
> 50	238 (78.3)	221 (78.6)	17 (73.9)	
50-30	52 (17.1)	47 (16.7)	5 (21.7)	0.099
< 30	2 (0.7)	1 (0.4)	1 (4.3)	

<sup>\*</sup> BMI: body mass index; † COPD: chronic obstructive coronary disease; ‡ MR: myocardial revascularization.

All data presented with n (%), except stated otherwise. §Mean +- Standard deviation. // Average (percentile 25 – percentile 75).

and validated. Since the end of the nineties, several centers have used EuroSCORE to validate it. 11-13 In the United States, it was more accurate compared to other predictor models when validated in the database with more than 500.000 patients of the Society of Thorac Surgery. 14 However, a systematic review evaluating the performance of the EuroSCORE concluded that the model overestimates surgical risk based on five studies of different nationalities. 15 The present study corroborates the findings, finding unsatisfactory results for the predictive capacity of the evaluated scores, in contrast to a good performance of the locally adjusted model. The development of local risk scores presents growth, finding difficulties

in implementation, but presenting progressive improvements and contributing to the identification of risk factors. <sup>16-18</sup> A recent body of results shows a better performance of models fitted with local data in relation to EuroSCORE, Parsonet Score and Ontario Risk Score. <sup>19</sup> Other studies in cardiovascular surgery suggest that most information on prognosis is contained in a few clinical variables, showing that simple models are as effective as complex models. <sup>20</sup> Although it is better suited than traditional scores, the score derived from the sample in this study is not intended for use in other services. Performing in only one center limits the external validity, where characteristics of the patients and the care body of the institution may vary.

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Table 2 –	Coefficients for adjusted models
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Suitable model*	Odds Ratio (IC 95%)	Coefficient	Default error	p Value of the coefficient (Statistics Wald)
Previous myocardial revascularization	8.519 (1.026 - 59.381)	2.142	0.979	0.029
Previous creatinine	2.217 (0.815 - 6.018)	0.796	0.476	0.095
Age	1.081 (1.025 - 1.145)	0.077	0.028	0.006
FE ≤ 50	1.251 (0.360 – 3.617)	0.224	0.577	0.698
Vascular Brain Disease	1.398 (0.506 – 397.795)	2.637	1.489	0.077

<sup>\*</sup> Constant Value for Model: -8,422; Default error: 1,955; p Value: < 0,001

#### Conclusion

Risk scores in cardiovascular surgery should be revalidated locally and the development of simple local predictive models may present better results in a delimited environment.

#### **Author contributions**

Conception and design of the research: Viana VJC, Ribeiro NAM, Silva Junior AF. Acquisition of data: Viana VJC, Ribeiro NAM, Silva Junior AF. Analysis and interpretation of the data: Argolo FC, Correia LCL. Statistical analysis: Argolo FC. Writing of the manuscript: Viana VJC. Critical revision of the manuscript for intellectual content: Correia LCL.

# **Potential Conflict of Interest**

No potential conflict of interest relevant to this article was reported.

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## **Study Association**

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