



Enfermedades Infecciosas y Microbiología Clínica

www.elsevier.es/eimc



Brief report

Prediction of surgical risk in patients with endocarditis: Comparison of logistic EuroSCORE, EuroSCORE II and APOREI score



Stefano Urso^{a,*}, María Ángeles Tena^a, Iballe Horcajada^b, Federico Paredes^a, Miguel González-Barbeito^a, Francisco Portela^a, on behalf of the EME study group

^a Cardiac Surgery Department, Hospital Universitario Dr. Negrín, Las Palmas de Gran Canaria, Spain

^b Microbiology Department, Hospital Universitario Dr. Negrín, Las Palmas de Gran Canaria, Spain

ARTICLE INFO

Article history:

Received 31 March 2020

Accepted 5 May 2020

Available online 15 June 2020

Keywords:

Infective endocarditis

Risk scores

Cardiac surgery

ABSTRACT

Objectives: APOREI score is a new risk prediction model for patients with infective endocarditis. It has been recently validated on a Spanish multicentric national cohort of patients. The aim of the present study is to compare APOREI performances with logistic EuroSCORE and EuroSCORE II by testing calibration and discrimination on a local sample population underwent cardiac surgery because of endocarditis.

Methods: We tested three prediction scores on 111 patients underwent surgery from 2014 to 2020 at our Institution because of infective endocarditis. Area under the curves and Hosmer–Lemeshow test were used to analyze discrimination and calibration respectively of logistic EuroSCORE, EuroSCORE II and APOREI score.

Results: The overall observed one-month mortality rate was 21.6%. The observed-to-expected ratio was 1.27 for logistic EuroSCORE, 3.27 for EuroSCORE II and 0.94 for APOREI. The area under the curve (AUC) value of APOREI (0.88 ± 0.05) was significantly higher than that one of logistic EuroSCORE ($AUC 0.77 \pm 0.05$; $p 0.0001$) and of EuroSCORE II ($AUC 0.74 \pm 0.05$; $p 0.0005$). Hosmer–Lemeshow test showed better calibration performance of the APOREI, (logistic EuroSCORE: $p 0.19$; EuroSCORE II: $p 0.11$; APOREI: $p 0.56$).

Conclusion: APOREI risk score shows significantly higher performances in term of discrimination and calibration compared with both logistic EuroSCORE and EuroSCORE II.

© 2020 Elsevier España, S.L.U. and Sociedad Española de Enfermedades Infecciosas y Microbiología Clínica. All rights reserved.

Predicción de riesgo de mortalidad quirúrgica en pacientes con endocarditis infecciosa: comparación entre EuroSCORE logístico, EuroSCORE II y APOREI score

RESUMEN

Palabras clave:

Sistema de predicción de riesgo

Endocarditis infecciosa

Cirugía cardíaca

Objetivos: El APOREI score es un nuevo sistema de predicción de riesgo para pacientes con endocarditis infecciosa. El mismo ha sido recientemente validado en una cohorte de pacientes procedentes de un estudio nacional multicéntrico español. El objetivo del presente estudio es comparar la discriminación y la calibración del APOREI score con las del EuroSCORE logístico y del EuroSCORE II analizando una población sometida a cirugía cardíaca por endocarditis infecciosa.

Métodos: Analizamos las propiedades de 3 sistemas de predicción de riesgo sobre una población de 111 pacientes sometidos a cirugía cardíaca desde 2014 hasta 2020 en un único hospital terciario por endocarditis infecciosa. El área bajo las curvas y la prueba de Hosmer–Lemeshow se usaron para analizar la discriminación y la calibración, respectivamente, del EuroSCORE logístico, del EuroSCORE II y del APOREI score.

* Corresponding author.

E-mail address: stefano.urso@inwind.it (S. Urso).

Resultados: La mortalidad global observada a un mes fue del 21,6%. La relación mortalidad observada/mortalidad esperada fue de 1,27 para el EuroSCORE logístico, 3,27 para el EuroSCORE II y 0,94 para el APOREI score. El valor del área bajo la curva (AUC) del APOREI score ($0,88 \pm 0,05$) fue significativamente mayor que los del EuroSCORE logístico ($AUC 0,77 \pm 0,05$; $p 0,0001$) y del EuroSCORE II ($AUC 0,74 \pm 0,05$; $p 0,0005$). La prueba de Hosmer-Lemeshow mostró un mejor rendimiento en cuanto a calibración del APOREI score, (EuroSCORE logístico: $p 0,19$; EuroSCORE II: $p 0,11$; APOREI score: $p 0,56$).

Conclusión: El sistema de predicción de riesgo APOREI score muestra un rendimiento significativamente mejor en cuanto a discriminación y calibración en comparación con el EuroSCORE logístico y con el EuroSCORE II.

© 2020 Elsevier España, S.L.U. y Sociedad Española de Enfermedades Infecciosas y Microbiología Clínica. Todos los derechos reservados.

Introduction

Infective endocarditis (IE), especially in surgical setting, represents a complex condition associated with high early and long-term mortality and morbidity.¹ In fact, surgical mortality in patients with IE ranges from 15% to 45%.² To approach the IE complexity in terms of diagnosis and therapeutic strategies, building of multidisciplinary *ad-hoc* teams and use of statistical models for prognosis prediction have been extensively advocated³.

These are, in most cases, the product of sophisticated statistical models obtained from specific patients-based datasets such as logistic EuroSCORE⁴ and EuroSCORE II.⁵

Still, several concerns have been raised about the discrimination performances of risk models obtained from dataset where patients with IE are poorly represented.^{6,7}

Recently, a new prediction model tool, APOREI score, obtained from an extensive meta-analysis, has been reported to provide a precise and easy tool to stratify mortality risk in patients underwent surgery because of IE.^{8,9}

The aim of the present study is to analyze discrimination and calibration proprieties of the three risk scores, logistic EuroSCORE, EuroSCORE II and APOREI score, over a sample population undergoing cardiac surgery at our Institution because of IE.

Methods

The sample population was constituted by 111 patients underwent consecutively cardiac surgery because of infective endocarditis from January 2014 to March 2020 at our Institution, a tertiary-level hospital.

Data were collected prospectively through our local electronic database. Logistic EuroSCORE and EuroSCORE II were prospectively calculated using the specific online calculators (<http://www.euroscore.org/calculsp.html>). The APOREI score was retrospectively calculated for each patient using excel sheet published, as supplementary material, by the score developers⁹ (<https://academic.oup.com/ejcts/article/57/4/724/5647329#supplementary-data>). Continuous variables following a normal distribution were expressed as mean $\pm$ standard deviation (SD). Continuous variables not normally distributed were expressed as median (25th–75th percentiles). Dichotomous categorical variables were indicated as absolute frequency (percentage). Predicted surgical mortality was obtained for each score. Spearman rank correlation analyses were carried out between each score. Discrimination was analyzed by receive operating characteristic analyses from which area under the curve (AUC) $\pm$ standard error (SE) values were obtained. The method of DeLong¹⁰ was used for the calculation of the standard error (SE) of the AUC and of the difference between two AUCs. Univariable logistic regression analysis was built introducing “30-day mortality” as dependent variable and separately each of the three scores as independent variable. From this model the Hosmer–Lemeshow

Table 1

Pre and intra operative patients data of the whole sample population.

Patients characteristics	All sample population (N = 111)
Female	30 (27.0%)
Age	58.9 (± 13.7)
Body mass index	26.5 (± 4.9)
Diabetes mellitus	24 (21.6%)
Chronic lung disease	16 (14.4%)
Arterial hypertension	40 (36.0%)
Preoperative inotropic treatment	21 (18.9%)
Pulmonary hypertension (>60 mmHg)	15 (13.5%)
Congestive heart failure	54 (48.6%)
Invasive ventilation	25 (22.5%)
Left ventricle ejection fraction < 49%	12 (10.8%)
Cardiogenic or septic shock	29 (26.1%)
Emergency surgery	32 (28.8%)
Prosthetics endocarditis	30 (27.0%)
Peri annular abscess	27 (24.3%)
Nosocomial endocarditis	16 (14.4%)
Staphylococcus Aureus	21 (22.6%)
Logistic EuroSCORE	17.1 (7.8–41.7)
EuroSCORE II	6.8 (3.2–24.5)
APOREI score	46.4 (30.6–70.0)
APOREI predicted mortality	23.0 (13–39.8)
Location of endocarditis	
Aortic valve	48 (43.2%)
Mitral valve	32 (28.8%)
Aortic and mitral valves	23 (20.7%)
Mitral and tricuspid valves	2 (1.8%)
Pulmonary valve	2 (1.8%)
Interventricular septum	1 (0.9%)
Tricuspid valve	1 (0.9%)
Aortic, mitral and pulmonary valves	1 (0.9%)
Aortic, mitral and tricuspid valves	1 (0.9%)

test was obtained. The Hosmer–Lemeshow method, as test for goodness of fit for the three logistic regression models, was used to analyze the calibration of the scores. Statistical analyses were performed using MedCalc Statistical Software version 19.0.7 (MedCalc Software bvba, Ostend, Belgium; <https://www.medcalc.org>; 2019).

Results

Patients characteristics are listed in Table 1. Mean age was 58.9 years. Most frequent causative microorganism were *Staphylococcus aureus* (25.2%, $n = 28$), *Staphylococcus epidermidis* (14.4%, $n = 16$), and *Streptococcus viridans* group (11.7%, $n = 13$). The aortic valve was the most frequent localization (43.2%, $n = 48$), (Table 1).

Thirty-day mortality was 21.6% ($n = 24$). Predicted surgical mortality by logistic EuroSCORE, EuroSCORE II and APOREI score were 17.1%, 6.6% and 23.0%, respectively.

Therefore, the observed-to-expected ratio was 1.27 for logistic EuroSCORE, 3.27 for EuroSCORE II and 0.94 for APOREI score.

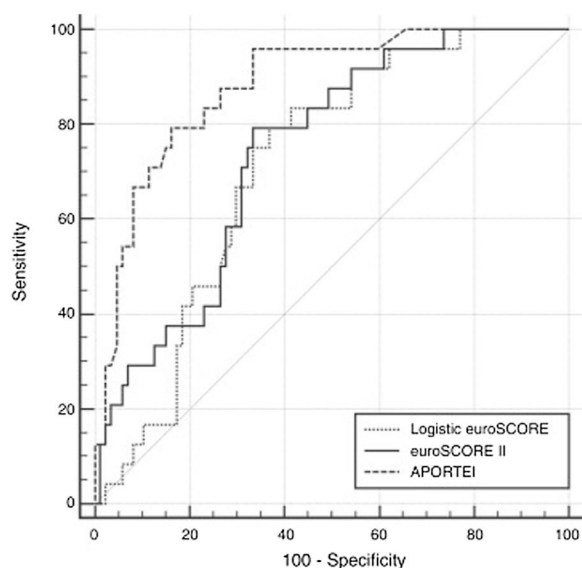


Fig. 1. ROC curves of Logistic EuroSCORE, EuroSCORE II and APOREI score.

According to Spearman's rank correlation analysis the highest coefficient of correlation was observed between logistic EuroSCORE and EuroSCORE II (ρ 0.88, $p < 0.0001$). Still, significant correlation was documented between logistic EuroSCORE and APOREI score (ρ 0.58, $p < 0.0001$) and between EuroSCORE II and APOREI score (ρ 0.63, $p < 0.0001$). According to the receiver operating characteristic analyses, AUC value of APOREI score was 0.88 ± 0.05 which was significantly higher than those of logistic EuroSCORE (AUC 0.77 ± 0.05 ; difference between areas 0.17; p 0.0001) and EuroSCORE II (AUC 0.74 ± 0.05 ; difference between areas 0.11; p 0.0005) (Fig. 1).

Hosmer–Lemeshow test of the regression logistic analysis showed better calibration performance of the APOREI score, (logistic EuroSCORE: chi-squared 12.30, p value: 0.19; EuroSCORE II: chi-squared 12.98, p value: 0.11; APOREI: chi-squared 6.77; p value: 0.56).

Discussion

The present study shows higher discrimination and calibration properties of APOREI score compared with logistic EuroSCORE and EuroSCORE II. The three scores were tested on a sample population of 111 patients underwent surgery because of IE.

In the last decade, several risk scores have been built to predict patients' outcomes after cardiac surgery.^{8,11} Risk models have been used also to identify specific risk factors, to allow comparison of postoperative results and to carry out risk stratification analysis.¹²

Patients undergoing cardiac surgery because of infective endocarditis represents a complex subgroup of patients, with higher surgical mortality than that one of the general populations undergoing cardiac surgery and specific risk factors.^{13–15} For this reason, several concerns have been raised about the properties of logistic EuroSCORE and EuroSCORE II when applied to IE populations.^{6,7} In fact, EuroSCORE I prediction models⁴ have been built from a database containing 19,030 patients underwent cardiac surgery, but, of these, only 1.1% (173) suffered from endocarditis. EuroSCORE II⁵ was built on a database of 22,381 patients undergoing major cardiac surgery. Of these only 2.2% (497) presented with endocarditis. In both cases prediction scores were obtained by logistic modellings.

APOREI score building process lied on a different methodology as it was not computed from a specific database of patients. Instead,

it was obtained by meta-analysis data consisted on 16 studies and 7484 patients with IE.⁸ Through overall pooled odds ratio (OR) calculation, 11 risk factors were identified and posteriorly tested on a national Spanish multicenter database containing 1338 patients with IE.⁹

Our results show clearly than logistic EuroSCORE and especially EuroSCORE II underestimates surgical mortality risk in patients with endocarditis. Also, receiver operating characteristic analyses demonstrated that APOREI score has the largest AUC value and, hence, the best performance in term of discrimination.

Even if the present study was carried out on a small sample population, its results confirm those of a previous study⁹ in terms of better performance of the APOREI score compared with logistic EuroSCORE. Also, our analysis shows that the APOREI score has higher discrimination and calibration properties even when compared with EuroSCORE II as prognostic and risk stratification tool for patients undergoing surgery because of IE.

Funding

This research received no funding.

Conflict of interest

None.

Acknowledgments

The present authors sincerely thank all the remaining members of the Equipo Multidisciplinar de Endocarditis (EME) of the Hospital Universitario Dr. Negrín:

Eloy Gomez, Oscar Sanz, Rafael Granados, Xerach Bosh (Unidad de Enfermedades Infecciosas), Ana Bordes, Raúl Gilarranz (Microbiología), José Luis Romero, Adela Hernández, Sílvia Hipola, José María Gómez, (Medicina Intensiva), Verónica Quevedo, Marta Blanco, Irene Menduiña (Cardiología), Luis Ríos, Raquel Bellot (Cirugía Cardíaca).

References

- Varela Barca L, López-Menéndez J, Navas Elorza E, Moya Mur JL, Centella Hernández T, et al. Long-term prognosis after surgery for infective endocarditis: distinction between predictors of early and late survival. *Enferm Infecc Microbiol Clin*. 2019;37:435–40.
- Murdoch DR, Corey GR, Hoen B, Miró JM, Fowler VG, Bayer AS, et al. Clinical presentation, etiology, and outcome of infective endocarditis in the 21st century: the International Collaboration on Endocarditis-Prospective Cohort Study. *Arch Intern Med*. 2009;169:463–73.
- Habib G, Lancellotti P, Antunes MJ, Bongioni MG, Casalta JP, Del Zotti F, et al. ESC Scientific Document Group. 2015 ESC Guidelines for the management of infective endocarditis: The Task Force for the Management of Infective Endocarditis of the European Society of Cardiology (ESC). *Eur Heart J*. 2015;36:3075–128.
- Nashef SA, Roques F, Michel P, Gauducheau E, Lemeshow S, Salamon R. European system for cardiac operative risk evaluation (EuroSCORE). *Eur J Cardiothorac Surg*. 1999;16:9–13.
- Nashef SA, Roques F, Sharples LD, Nilsson J, Smith C, Goldstone AR, et al. EuroSCORE II. *Eur J Cardiothorac Surg*. 2012;41:734–44.
- Fernández-Hidalgo N, Ferrer-González I, Marsal JR, Ribera A, Aznar ML, de Alarcón A, et al. A pragmatic approach for mortality prediction after surgery in infective endocarditis: optimizing and refining EuroSCORE. *Clin Microbiol Infect*. 2018;24, 1102.e7–1102.e15.
- Varela L, López-Menéndez J, Redondo A, Fajardo ER, Miguelena J, Centella T, et al. Mortality risk prediction in infective endocarditis surgery: reliability analysis of specific scores. *Eur J Cardiothorac Surg*. 2018;53:1049–54.
- Varela Barca L, Navas Elorza E, Fernández-Hidalgo N, Moya Mur JL, Muriel García A, Fernández-Felix BM, et al. Prognostic factors of mortality after surgery in infective endocarditis: systematic review and meta-analysis. *Infection*. 2019;47:879–95.

9. Varela Barca L, Fernández-Felix BM, Navas Elorza E, Mestres CA, Muñoz P, Cuerpo-Caballero G, et al. Members of GAMES. Prognostic assessment of valvular surgery in active infective endocarditis: multicentric nationwide validation of a new score developed from a meta-analysis. *Eur J Cardiothorac Surg*. 2020;57:724–31.
10. DeLong ER, DeLong DM, Clarke-Pearson DL. Comparing the areas under two or more correlated receiver operating characteristic curves: a nonparametric approach. *Biometrics*. 1988;44:837–45.
11. Gaca JG, Sheng S, Daneshmand MA, O'Brien S, Rankin JS, Brennan JM, et al. Outcomes for endocarditis surgery in North America: a simplified risk scoring system. *J Thorac Cardiovasc Surg*. 2011;141:98–106.
12. Ad N, Holmes SD, Patel J, Pritchard G, Shuman DJ, Halpin L. Comparison of uroSCORE II, Original EuroSCORE, and the society of thoracic surgeons risk score in cardiac surgery patients. *Ann Thorac Surg*. 2016;102:573–9.
13. Cahill TJ, Prendergast BD. Infective endocarditis. *Lancet*. 2016;387:882–93.
14. Fernández-Hidalgo N, Tornos Mas P. Epidemiology of infective endocarditis in Spain in the last 20 years. *Rev Esp Cardiol (Engl Ed)*. 2013;66:728–33.
15. Mestres CA, Paré JC, Miró JM, Working Group on Infective Endocarditis of the ospital Clínic de Barcelona. Organization and Functioning of a Multidisciplinary Team for the Diagnosis and Treatment of Infective Endocarditis: A 30-year Perspective (1985–2014). *Rev Esp Cardiol (Engl Ed)*. 2015;68:363–8.