IMPROVING OUTCOME PREDICTIONS FOR PATIENTS RECEIVING MECHANICAL CIRCULATORY SUPPORT BY OPTIMIZING IMPUTATION OF MISSING VALUES

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ABSTRACT

Background Risk predictions play an important role in clinical decision making. When developing risk prediction models, practitioners often impute missing values to the mean. The purpose of this article is to evaluate the impact of applying different strategies to impute missing values on the prognostic accuracy of prediction models fitted to the imputed data. A secondary objective was to compare the accuracy of different imputation methods. To complete these objectives, we used data from the Interagency Registry for Mechanically Assisted Circulatory Support (INTERMACS).

Methods and Results We applied all pairwise combinations of different strategies to impute missing values and three different strategies to fit a risk prediction model for mortality and transplant after receiving mechanical circulatory support. Model performance was compared using Concordance (i.e. C-index), Brier Score, and Net reclassification index. Results indicated that multiple imputation consistently resulted in superior prognostic models compared to other missing data strategies, particularly imputation to the mean.

Conclusion Selecting an optimal strategy to handle missing values can have a substantial impact on improving model accuracy. In the current analysis, multiple imputation emerged as an optimal strategy to handle missing values in the INTERMACS data. The evaluation and selection of the optimal missing data strategy in this work has the potential to improve risk predictions for other longitudinal registries.

Keywords Missing Data, · INTERMACS, · imputation, · heart failure, · mortality, · risk prediction

1 Introduction

Heart disease is a leading cause of death in the United States. Heart failure, a primary component of heart disease, affects over 6 million Americans, and for ~10% of these patients medical management is no longer effective [1, 2]. Mechanical circulatory support (MCS) is a surgical intervention in which a mechanical device is implanted in parallel to the heart to improve circulation [3]. Typically, MCS is used while a patient waits for a heart transplant (bridge-to-transplant) or in some cases as an alternative to transplant (destination therapy) [4]. Over 250,000 patients could benefit from MCS [5]. However, less than 4,000 new patients receive a long-term MCS device each year, with widely heterogeneous outcomes [6]. The 2-year survival on MCS ranges from 61% for destination therapy to 78% for bridge-to-transplant [3]. Therefore, there is great need for reliable predictions of patient-specific probability to experience adverse events after receiving MCS. This information can be used to improve patient selection for MCS,inform the design of next generation pumps, and refine patient management strategies.

^{*}Source code available at https://github.com/bcjaeger/INTERMACS-missing-data

References

- [1] Emelia J Benjamin, Michael J Blaha, Stephanie E Chiuve, Mary Cushman, Sandeep R Das, Rajat Deo, J Floyd, M Fornage, C Gillespie, CR Isasi, et al. Heart disease and stroke statistics-2017 update: a report from the american heart association. *Circulation*, 135(10):e146–e603, 2017.
- [2] National Center for Health Statistics. *Health, United States, 2016, With chartbook on long-term trends in health.* Number 2017. Government Printing Office, 2017.
- [3] Chetan B Patel, Jennifer A Cowger, and Andreas Zuckermann. A contemporary review of mechanical circulatory support. *The Journal of Heart and Lung Transplantation*, 33(7):667–674, 2014.
- [4] Mark S Slaughter, Joseph G Rogers, Carmelo A Milano, Stuart D Russell, John V Conte, David Feldman, Benjamin Sun, Antone J Tatooles, Reynolds M Delgado III, James W Long, et al. Advanced heart failure treated with continuous-flow left ventricular assist device. *New England Journal of Medicine*, 361(23):2241–2251, 2009.
- [5] Leslie W Miller. Left ventricular assist devices are underutilized. Circulation, 123(14):1552–1558, 2011.
- [6] Garrick C Stewart and Lynne W Stevenson. Keeping left ventricular assist device acceleration on track. *Circulation*, 123(14):1559–1568, 2011.