

Chapter 31

A Semiparametric Bayesian Multivariate Model for Survival Probabilities After Acute Myocardial Infarction

Elena Prandoni, Alessandra Guglielmi, Francesca Ieva,
and Anna Maria Paganoni

Abstract In this work, a Bayesian semiparametric multivariate model is fitted to study data related to in-hospital and 60-day survival probabilities of patients admitted to a hospital with ST-elevation myocardial infarction diagnosis. We consider a hierarchical generalized linear model to predict survival probabilities and a process indicator (time of intervention). Poisson-Dirichlet process priors, generalizing the well-known Dirichlet process, are considered for modeling the random-effect distribution of the grouping factor which is the hospital of admission.

31.1 Introduction

The disease we are interested in is ST-elevation myocardial infarction (STEMI): it is caused by an occlusion of a coronary artery which causes an ischemia that, if untreated, can damage heart cells and make them die (infarction). It is very important that a reperfusion therapy could be done as quickly as possible, because its benefits decrease with delay in treatment; in our case, patients are treated with percutaneous transluminal coronary angioplasty. We consider data collected in the STEMI Archive [1], a multicenter observational prospective clinical study planned within the Strategic Program of Regione Lombardia. Data is recorded in a registry collecting clinical and process indicators, outcomes, and personal information on patients admitted to all hospitals of Regione Lombardia with STEMI diagnosis.

The regression model we introduce is multivariate, where the response has three components: door to balloon time (DB), i.e., the time between the admission to the hospital and angioplasty, on the logarithmic scale, in-hospital survival, and survival after 60 days from admission. The first term is an important indicator of

E. Prandoni (✉) • A. Guglielmi • F. Ieva • A.M. Paganoni
Politecnico di Milano, via Bonardi 9, 20133, Milano, Italy
e-mail: elenaprandoni@gmail.com

the efficiency of the health providers and plays a key role in the success of the therapy; the second one is the basic indicator of success or failure of the treatment, while the third one is a very important outcome, since doctors believe that it is in a 60-day period the effectiveness of the treatment in terms of survival and quality of life can be truly evaluated. We include the hospital random-effect parameters in the model and assume they are a sample from a Poisson-Dirichlet process a priori in order to eventually cluster the hospitals.

The main statistical aim of this work is prediction of both survival probabilities of new patients.

31.2 The Bayesian Model in a Nutshell

For each patient ($i = 1, \dots, 697$) let $\mathbf{Y}_i := (Y_{i1}, Y_{i2}, Y_{i3})$ be the response, where Y_1 is the logarithm of DB, Y_2 is the in-hospital survival, and Y_3 is the long-term survival. We assume that observations, given parameters and covariates, are independent and the law of the response can be factorized into three parts:

$$\mathcal{L}(\mathbf{Y}_i | par, cov) = \mathcal{L}(Y_{i1} | par_1, cov_1) \mathcal{L}(Y_{i2} | Y_{i1}, par_2, cov_2) \mathcal{L}(Y_{i3} | Y_{i2}, par_3, cov_3).$$

The likelihood can be expressed as

$$Y_{i1} | \mu_i, \sigma \stackrel{ind}{\sim} \mathcal{N}(\mu_i, \sigma^2), \quad \mu_i = \sum_{l=1}^4 \beta_l u_{il} + \beta_5 x_{i5} + \beta_6 x_{i6} \quad (1)$$

$$Y_{i2} | p_i, Y_{i1} \stackrel{ind}{\sim} Be(p_i), \quad \text{logit}(p_i) = \alpha_1 z_{i1} + \alpha_2 z_{i2} + \alpha_3 Y_{i1} + \sum_{l=4}^7 \alpha_l v_{il} + b_{\phi_{k[i]} k[i]} \quad (2)$$

$$Y_{i3} | r_i, Y_{i2} \stackrel{ind}{\sim} \begin{cases} Be(r_i) & \text{se } Y_{i2} = 1 \\ \delta_0 & \text{se } Y_{i2} = 0 \end{cases}, \quad \text{logit}(r_i) = \sum_{j=1}^4 \gamma_j s_{ij} + t_{\phi_{k[i]} k[i]}. \quad (3)$$

Here $k[i]$ denotes the hospital where the patient i is admitted to, while covariates include the type of rescue unit sent to the patient (u_{il} , $l = 1, \dots, 4$), the time of the first ECG (x_{i5}), the age (z_{i1}), the Killip class (v_{il} , $l = 1, \dots, 4$, which quantify in four categories the severity of infarction), other covariates measuring the health status of the patient, or if the treatment was successful or not. The indexes $\phi_{k[i]}$ of the random-effect parameters in (2) and (3) assume values 1 or 0, if the hospital is in Milano or not.

As far as the fixed-effects parameters are concerned, we considered a parametric prior; for the random-effect parameters we assume Poisson-Dirichlet process priors with parameters (f, g) [2]. This choice helps us to avoid dependency on parametric

assumptions and to increase flexibility in the prior and more robust inferences. We obtained posterior estimates of all the parameters through a Gibbs sampler algorithm implemented in JAGS [3] and with the support of R [4]; for this purpose we used a truncated stick-breaking representation of the nonparametric prior, which is a generalization of the well-known Dirichlet process [5].

Acknowledgements This work is within the Strategic Program “Exploitation, integration and study of current and future health databases in Lombardia for Acute Myocardial Infarction” supported by “Ministero del Lavoro, della Salute e delle Politiche Sociali” and by “Direzione Generale Sanità - Regione Lombardia.” The authors wish to thank the Working Group for Cardiac Emergency in Milano, the Cardiology Society, and the 118 Dispatch Center.

References

1. Ieva F (2013) Designing and mining a multicenter observational clinical registry concerning patients with Acute Coronary Syndromes. In: Grieco, N, Marzegalli M, Paganoni AM (eds) New diagnostic, therapeutic and organizational strategies for acute coronary syndromes patients. Springer, Heidelberg
2. Pitman J, Yor M (1997) The two-parameter Poisson Dirichlet distribution derived from a stable subordinator. *The Ann Probab* 2:855–900
3. Plummer M (2003) JAGS: A program for analysis of Bayesian graphical models using Gibbs sampling. In *Proceedings of the 3rd International Workshop on Distributed Statistical Computing*, pp 20–22
4. R Development Core Team (2009) *R: A Language and Environment for Statistical Computing*. R Foundation for Statistical Computing, Vienna.
5. Sethuraman J (1994) A constructive definition of Dirichlet process prior. *Statistica Sinica* 2: 639–650