

1. Introduction

- The Piecewise Exponential Model (PEM) can be utilised to approximate many distributions found in a survival setting, allowing for variation of hazard rates between intervals, while keeping hazards constant within intervals.

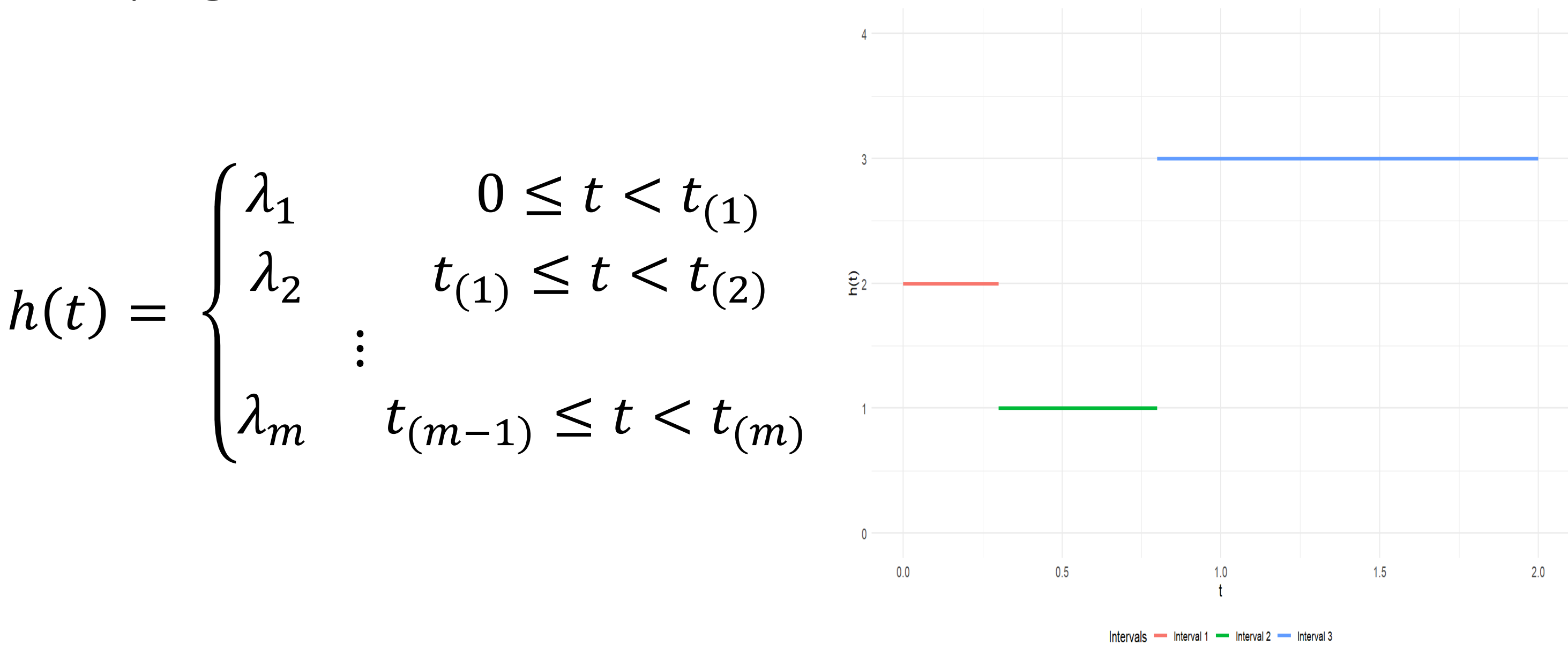


Figure 1. Piecewise exponential hazard, where Interval i , $I_i = [t_{(i-1)}, t_{(i)})$

2. Methods

- PEM can be extended to include covariates.
 $\lambda_m = e^{(\beta_{0m} + \beta_1 x_1 + \dots + \beta_p x_p)}$
- The above link reflects the variation between intervals via β_{0m} while ensuring covariate effects remain constant across intervals.
- The log-likelihood from this extension is as follows:

$$\log(L) = \sum_{m=1}^M \sum_{i \in I_m} \left\{ \delta_i(B_m X_i) - t_i e^{B_m X_i} - \sum_{j=1}^{m-1} t_{(j)} (e^{B_j X_i} - e^{B_{j+1} X_i}) \right\}$$

with $B_m = (\beta_{0m}, \beta_1, \dots, \beta_p)$

- Parameters were optimised using the Newton-Raphson method, implemented through nlm in the R package 'stats'.

3. Simulation Study

- A simulation study was carried out to test the performance of the proportional hazard PEM and compare with the cox model.
- The data was simulated using a piecewise distribution with 3 true intervals (see figure 1), and 3 covariate effects.
- The PEM was fitted using 10 intervals, designed to keep the number of events consistent across intervals.

	True β_j	Mean $\hat{\beta}_j$	True SE($\hat{\beta}_j$)	Est SE($\hat{\beta}_j$)	Coverage
β_1	0.5	0.51	0.07	0.07	0.95
β_2	-0.5	-0.50	0.06	0.07	0.95
β_3	0.01	0.01	0.01	0.01	0.91

Table 1. Mean, Standard error and coverage of covariate effects calculated from 100 simulations

- The covariate estimates were also compared to the estimates obtained from the cox model.

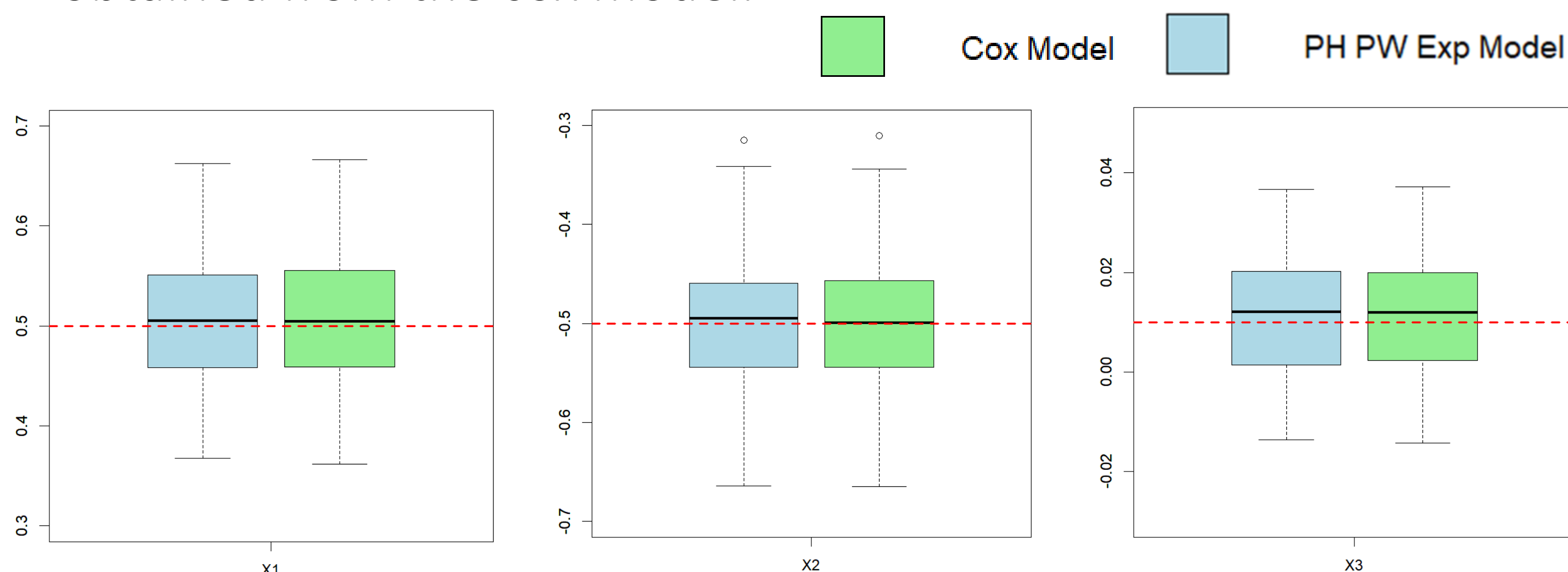


Figure 2. Boxplots of Covariate estimates, with true values shown by - - -

4. Case Study

- Next, we compare the performance of both models on a set of lung cancer data from the survival package in R.
- Significant covariates included both sex and ECOG score.

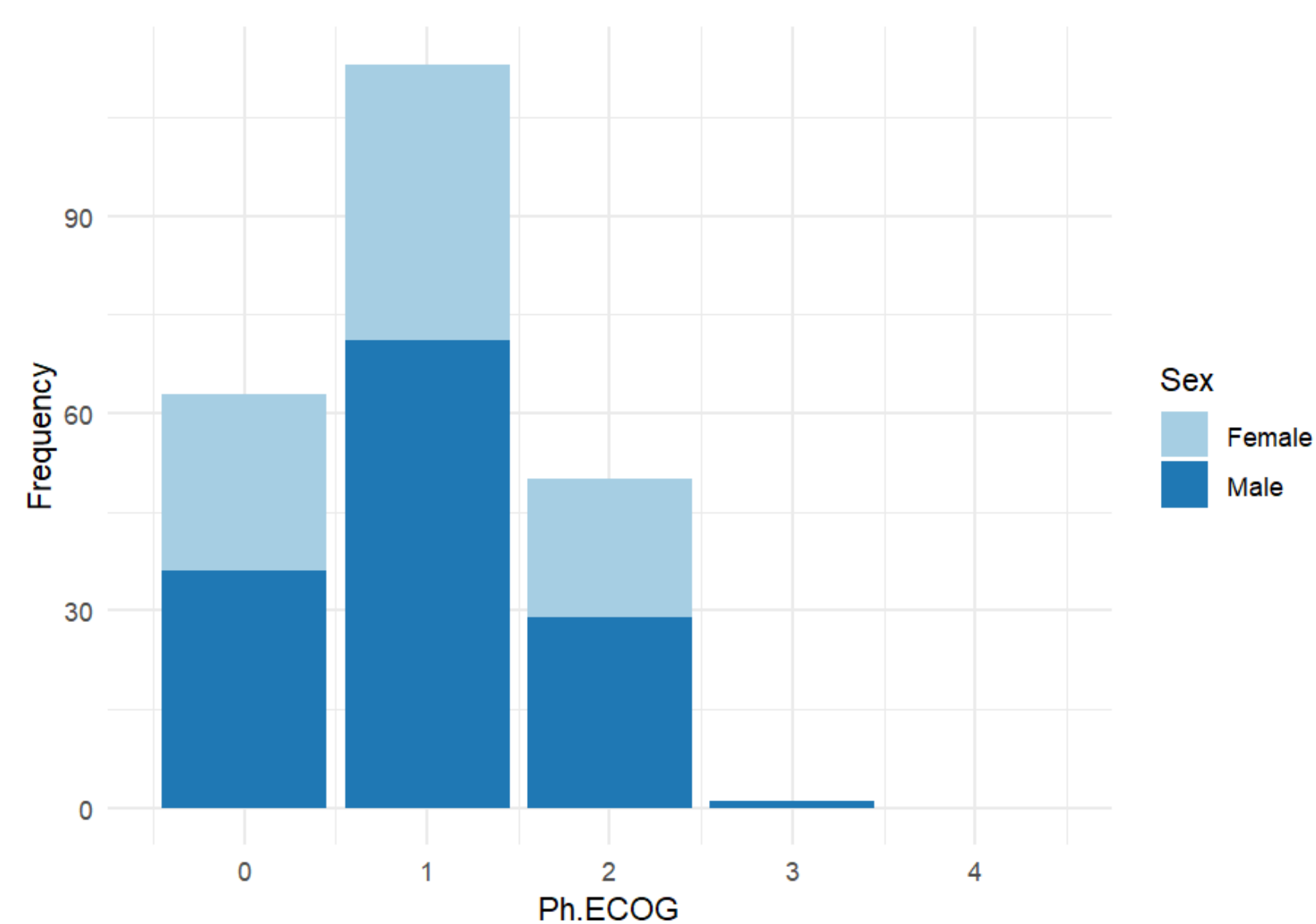


Figure 3. Cancer Ph.ECOG and Sex Frequency

- The resulting Cox Model and PEM were compared to the Kaplan-Meier curve for various subsets of the data.

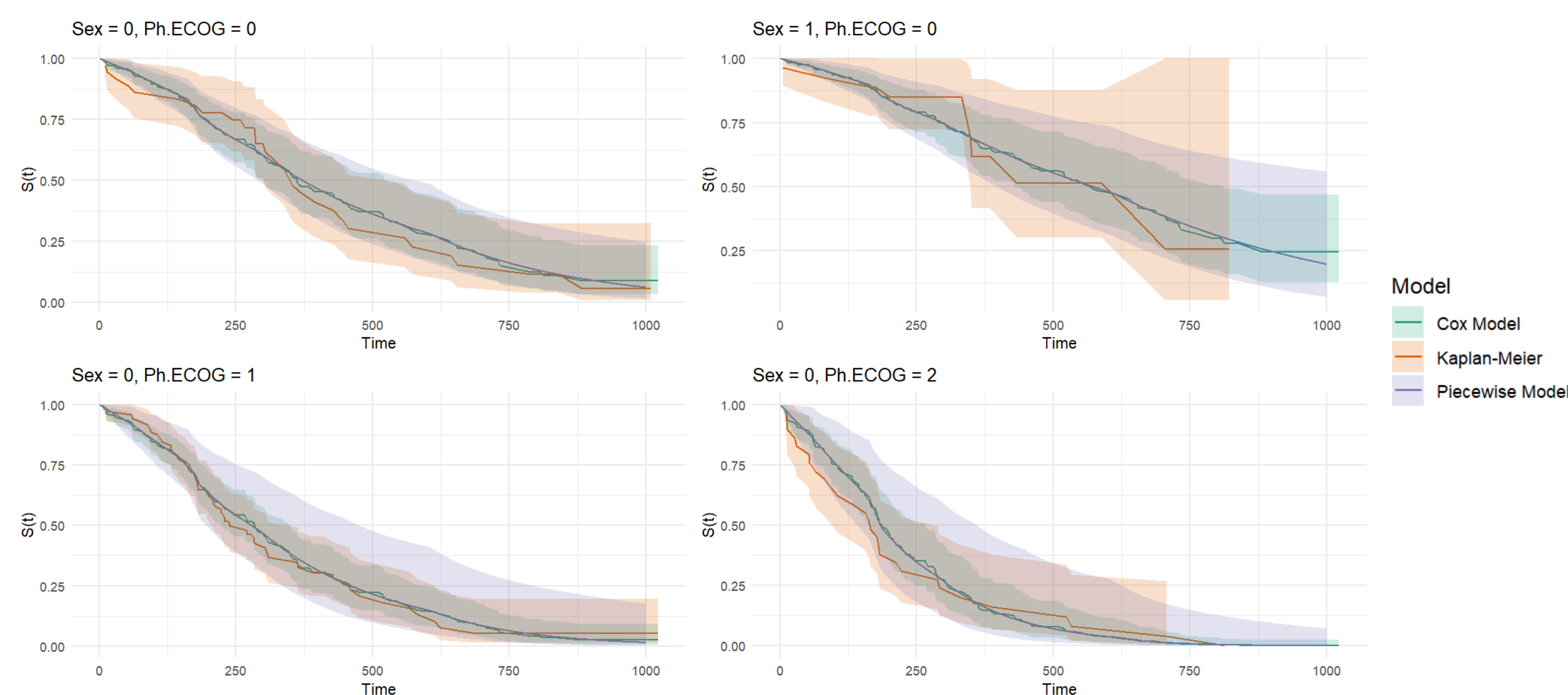


Figure 4. Comparison of Survival Curves

5. Future Work

- We can further extend PEM by allowing our covariate effects to change between intervals. This can be achieved by redefining $B_m = (\beta_{0m}, \beta_{1m}, \dots, \beta_{pm})$.
- Further exploration on how the number of intervals affect this model.

6. Acknowledgements

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7. References

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- Bender, Andreas & Scheipl, Fabian. (2018). pammttools: Piece-wise exponential Additive Mixed Modeling tools.

8. Contact Details

- Email:
moore.jack@ul.ie

- Poster Download:

