

# Supplementary material - code

Léa Orsini

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```
library(tidyverse)

## Warning: le package 'tidyverse' a été compilé avec la version R 4.2.3
## Warning: le package 'ggplot2' a été compilé avec la version R 4.2.3
## Warning: le package 'tibble' a été compilé avec la version R 4.2.3
## Warning: le package 'tidyr' a été compilé avec la version R 4.2.3
## Warning: le package 'readr' a été compilé avec la version R 4.2.3
## Warning: le package 'purrr' a été compilé avec la version R 4.2.3
## Warning: le package 'dplyr' a été compilé avec la version R 4.2.3
## Warning: le package 'stringr' a été compilé avec la version R 4.2.3
## Warning: le package 'forcats' a été compilé avec la version R 4.2.3
## Warning: le package 'lubridate' a été compilé avec la version R 4.2.3
library(pseudo)
library(geepack)
library(rstan)

## Warning: le package 'rstan' a été compilé avec la version R 4.2.3
## Warning: le package 'StanHeaders' a été compilé avec la version R 4.2.3
options(mc.cores = parallel::detectCores())

setwd("~/pseudoGMM_RMST")

source('MCMC_traceplot.R')

## Warning: le package 'plyr' a été compilé avec la version R 4.2.3
## Warning: le package 'coda' a été compilé avec la version R 4.2.2
## Warning: le package 'bayesplot' a été compilé avec la version R 4.2.3
## Warning: le package 'survminer' a été compilé avec la version R 4.2.3
## Warning: le package 'ggpubr' a été compilé avec la version R 4.2.3
## Warning: le package 'broom' a été compilé avec la version R 4.2.3
```

## Data generation

```
s = 1
n = 200
set.seed(s)
U <- runif(n)
Trt <- rbinom(n, 1, 0.5)
shape = 0.61*(Trt == 0) + 0.8*(Trt == 1)
scale = 0.28*(Trt == 0) + 0.18*(Trt == 1)

Z1 <- rnorm(n, 0, 1) # runif(n, 0, 2)
Z2 <- rbinom(n, 1, 0.5)
b1 <- log(2)
b2 <- log(1.5)

X1 <- rnorm(n)
X2 <- rbinom(n, 1, 0.5)

T_tilde = (-log(U))/((scale)^(1/shape)*exp(b1*Z1 + b2*Z2))^shape
C = runif(n, 0, 11)

time <- pmin(T_tilde, C, 8)
event <- as.numeric(time == T_tilde)

simu <- data.frame(time = time,
                   event = event,
                   patID = 1:n,
                   Trt = Trt,
                   Z1 = Z1,
                   Z2 = Z2,
                   X1 = X1,
                   X2 = X2)
```

## Data Analysis with the Bayesian GMM

Without covariable adjustment

```
tau = 5 # truncation time (5 years)

np = 2 #nb of parameters
n = length(simu$patID) #sample size
simu.tau = min(tau, min(max(simu$time[simu$Trt == 1]), max(simu$time[simu$Trt == 0])))
#compute the pseudo-observations
simu$rmst <- pseudomean(simu$time, simu$event, tmax = simu.tau)

X = matrix(c(rep(1, n), simu$Trt),
           nrow = n, ncol = np)

data <- list(X = X, Y = simu$rmst, n = n, N = n, np = np)

#fit a GMM with independence working matrix
GMM <- stan_model("GMM_ind_single_tau.stan")
fit <- sampling(GMM, data = data, chains = 3, iter = 2000, warmup = 1000, seed = 1, init_r = 1, cores
```

```
save_warmup = T, refresh = 1000)
```

```
summary(fit)
```

```
## $summary
##               mean      se_mean      sd      2.5%      25%
## beta[1]      2.761974373 4.630210e-03 1.571415e-01 2.455701764 2.654427528
## beta[2]     -0.021878856 7.800103e-03 2.535319e-01 -0.527940051 -0.185899936
## loglik      -1.014736919 2.825121e-02 9.954402e-01 -3.744435603 -1.417752907
## Sigma_n[1,1] 0.014573341 3.082905e-06 1.108702e-04 0.014495986 0.014503876
## Sigma_n[1,2] 0.006177005 1.779891e-06 7.756490e-05 0.006111835 0.006131203
## Sigma_n[2,1] 0.006177005 1.779891e-06 7.756490e-05 0.006111835 0.006131203
## Sigma_n[2,2] 0.006175847 1.562189e-06 6.553504e-05 0.006130519 0.006134577
## C_n[1,1]     0.014645614 4.187748e-06 1.487087e-04 0.014499755 0.014540655
## C_n[1,2]     0.006206740 2.625527e-06 1.101429e-04 0.006130559 0.006137379
## C_n[2,1]     0.006206740 2.625527e-06 1.101429e-04 0.006130559 0.006137379
## C_n[2,2]     0.006206740 2.625527e-06 1.101429e-04 0.006130559 0.006137379
## lp__        -1.400633085 2.865194e-02 1.003363e+00 -4.148991904 -1.811698008
##               50%       75%       97.5%    n_eff    Rhat
## beta[1]      2.759159841 2.866637760 3.077192549 1151.808 1.001589
## beta[2]     -0.015956814 0.155459876 0.460018587 1056.488 1.001327
## loglik      -0.708035200 -0.303536264 -0.026374716 1241.527 1.000992
## Sigma_n[1,1] 0.014531151 0.014598050 0.014884848 1293.329 1.000972
## Sigma_n[1,2] 0.006148574 0.006193491 0.006397007 1899.082 1.000399
## Sigma_n[2,1] 0.006148574 0.006193491 0.006397007 1899.082 1.000399
## Sigma_n[2,2] 0.006150759 0.006189956 0.006371489 1759.868 1.000206
## C_n[1,1]     0.014599504 0.014705542 0.015051165 1260.989 1.000890
## C_n[1,2]     0.006164576 0.006230454 0.006535552 1759.868 1.000206
## C_n[2,1]     0.006164576 0.006230454 0.006535552 1759.868 1.000206
## C_n[2,2]     0.006164576 0.006230454 0.006535552 1759.868 1.000206
## lp__        -1.088415706 -0.683995852 -0.405034454 1226.332 1.000991
##
## $c_summary
## , , chains = chain:1
##
##               stats
## parameter      mean      sd      2.5%      25%      50%
## beta[1]      2.753127254 1.603578e-01 2.437558860 2.640370916 2.756190269
## beta[2]     -0.015288594 2.522506e-01 -0.519914464 -0.183104499 -0.012104625
## loglik      -1.026905159 1.029405e+00 -3.940753435 -1.463717873 -0.690024485
## Sigma_n[1,1] 0.014572490 1.076045e-04 0.014495984 0.014504577 0.014531087
## Sigma_n[1,2] 0.006175436 7.242916e-05 0.006113341 0.006131758 0.006149891
## Sigma_n[2,1] 0.006175436 7.242916e-05 0.006113341 0.006131758 0.006149891
## Sigma_n[2,2] 0.006174870 6.263442e-05 0.006130519 0.006134657 0.006151824
## C_n[1,1]     0.014647206 1.528224e-04 0.014500178 0.014540458 0.014596892
## C_n[1,2]     0.006205099 1.052679e-04 0.006130559 0.006137514 0.006166367
## C_n[2,1]     0.006205099 1.052679e-04 0.006130559 0.006137514 0.006166367
## C_n[2,2]     0.006205099 1.052679e-04 0.006130559 0.006137514 0.006166367
## lp__        -1.410365113 1.035466e+00 -4.362269841 -1.824250220 -1.068662873
##
##               stats
## parameter      75%      97.5%
## beta[1]      2.855696317 3.074991342
## beta[2]      0.158425113 0.460082842
## loglik      -0.302578844 -0.028769901
```

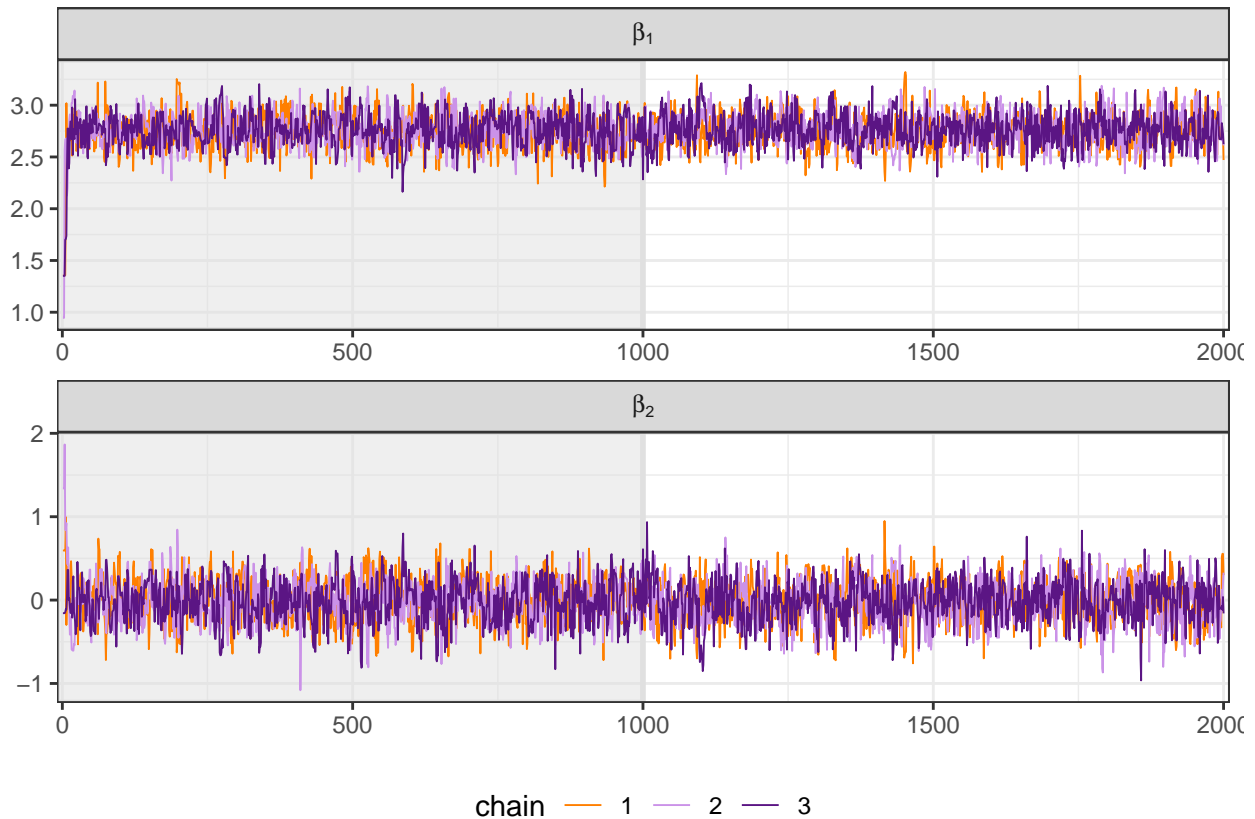
```

## Sigma_n[1,1] 0.014596834 0.014871045
## Sigma_n[1,2] 0.006192701 0.006362028
## Sigma_n[2,1] 0.006192701 0.006362028
## Sigma_n[2,2] 0.006186637 0.006365776
## C_n[1,1] 0.014710891 0.015071707
## C_n[1,2] 0.006224875 0.006525949
## C_n[2,1] 0.006224875 0.006525949
## C_n[2,2] 0.006224875 0.006525949
## lp__ -0.679691829 -0.406294921
##
## , , chains = chain:2
##
## stats
## parameter mean sd 2.5% 25% 50%
## beta[1] 2.767293608 1.535082e-01 2.456994438 2.661763489 2.760499368
## beta[2] -0.028189179 2.586651e-01 -0.517326638 -0.207381193 -0.027352536
## loglik -1.029983844 9.678091e-01 -3.646808824 -1.429370078 -0.742517298
## Sigma_n[1,1] 0.014576571 1.086648e-04 0.014496026 0.014503628 0.014533576
## Sigma_n[1,2] 0.006181088 7.906222e-05 0.006111784 0.006131630 0.006149451
## Sigma_n[2,1] 0.006181088 7.906222e-05 0.006111784 0.006131630 0.006149451
## Sigma_n[2,2] 0.006179436 6.895324e-05 0.006130507 0.006135564 0.006152588
## C_n[1,1] 0.014648319 1.454152e-04 0.014499151 0.014541980 0.014605518
## C_n[1,2] 0.006212773 1.158878e-04 0.006130539 0.006139039 0.006167650
## C_n[2,1] 0.006212773 1.158878e-04 0.006130539 0.006139039 0.006167650
## C_n[2,2] 0.006212773 1.158878e-04 0.006130539 0.006139039 0.006167650
## lp__ -1.417438367 9.756595e-01 -4.043385714 -1.827572608 -1.125481023
##
## stats
## parameter 75% 97.5%
## beta[1] 2.870645322 3.081782323
## beta[2] 0.152231322 0.471562579
## loglik -0.316388851 -0.022096603
## Sigma_n[1,1] 0.014606040 0.014870053
## Sigma_n[1,2] 0.006205470 0.006405016
## Sigma_n[2,1] 0.006205470 0.006405016
## Sigma_n[2,2] 0.006197261 0.006383871
## C_n[1,1] 0.014709776 0.015044652
## C_n[1,2] 0.006242730 0.006556361
## C_n[2,1] 0.006242730 0.006556361
## C_n[2,2] 0.006242730 0.006556361
## lp__ -0.690861861 -0.400099200
##
## , , chains = chain:3
##
## stats
## parameter mean sd 2.5% 25% 50%
## beta[1] 2.765502258 1.572616e-01 2.468785017 2.656711838 2.760207416
## beta[2] -0.022158796 2.496848e-01 -0.542037104 -0.179255658 -0.014451337
## loglik -0.987321755 9.885467e-01 -3.704690726 -1.367735376 -0.689311837
## Sigma_n[1,1] 0.014570962 1.161791e-04 0.014495974 0.014502998 0.014529103
## Sigma_n[1,2] 0.006174490 8.086211e-05 0.006112062 0.006130771 0.006145969
## Sigma_n[2,1] 0.006174490 8.086211e-05 0.006112062 0.006130771 0.006145969
## Sigma_n[2,2] 0.006173234 6.476740e-05 0.006130529 0.006133848 0.006148535
## C_n[1,1] 0.014641317 1.478461e-04 0.014499735 0.014539281 0.014596186
## C_n[1,2] 0.006202349 1.088528e-04 0.006130577 0.006136154 0.006160838

```

```
## C_n[2,1]      0.006202349 1.088528e-04 0.006130577 0.006136154 0.006160838
## C_n[2,2]      0.006202349 1.088528e-04 0.006130577 0.006136154 0.006160838
## lp__          -1.374095776 9.985205e-01 -3.998752409 -1.751546780 -1.066314184
##
## stats
## parameter      75%      97.5%
## beta[1]        2.871909900 3.077367699
## beta[2]        0.157225692 0.442571404
## loglik         -0.291204092 -0.026803616
## Sigma_n[1,1]   0.014590052 0.014891892
## Sigma_n[1,2]   0.006184275 0.006396439
## Sigma_n[2,1]   0.006184275 0.006396439
## Sigma_n[2,2]   0.006186363 0.006342046
## C_n[1,1]       0.014697506 0.015041670
## C_n[1,2]       0.006224415 0.006486067
## C_n[2,1]       0.006224415 0.006486067
## C_n[2,2]       0.006224415 0.006486067
## lp__          -0.675224422 -0.407062807
```

```
MCMC_traceplot(fit, c('beta[1]', 'beta[2]'))
```



With covariates adjustments

```
np = 4 #nb of parameters
n = length(simu$patID)
simu.tau = min(tau, min(max(simu$time[simu$Trt == 1]), max(simu$time[simu$Trt == 0])))
#compute the pseudo-observations
```

```

simu$rmst<-pseudomean(simu$time, simu$event, tmax = simu.tau)

X = matrix(c(rep(1,n), simu$Trt, simu$Z1, simu$Z2),
           nrow = n, ncol = np)

data <- list(X = X, Y = simu$rmst, n = n, N = n, np = np)

#fit a GMM with independence working matrix
GMM <- stan_model("GMM_ind_single_tau.stan")
fit <- sampling(GMM, data = data, chains = 3, iter = 2000, warmup = 1000, seed = 1, init_r = 1, cores = 
               save_warmup = T, refresh = 1000)

summary(fit, c('beta[1]', 'beta[2]', 'beta[3]', 'beta[4]'))

## $summary
##           mean      se_mean      sd      2.5%      25%      50%
## beta[1]  2.9540974  0.005018810  0.1861586  2.5955115  2.82719933  2.9535350
## beta[2]  0.1732505  0.005050838  0.2287608 -0.2774066  0.02054917  0.1765520
## beta[3] -0.7709282  0.002110713  0.1058734 -0.9780248 -0.84351363 -0.7696884
## beta[4] -0.5276528  0.005201831  0.2127319 -0.9491606 -0.67131191 -0.5261116
##           75%      97.5%    n_eff    Rhat
## beta[1]  3.0785239  3.3278200 1375.830 1.0007288
## beta[2]  0.3343891  0.6147117 2051.333 1.0004189
## beta[3] -0.7004957 -0.5701211 2516.026 0.9997914
## beta[4] -0.3820136 -0.1227919 1672.448 1.0015541
##
## $c_summary
## , , chains = chain:1
##
##           stats
## parameter      mean      sd      2.5%      25%      50%      75%
## beta[1]  2.9549072  0.1887723  2.5911437  2.82798828  2.9556018  3.0765087
## beta[2]  0.1714923  0.2276161 -0.2665657  0.01894498  0.1695766  0.3270774
## beta[3] -0.7681728  0.1067679 -0.9774929 -0.84473781 -0.7653101 -0.6939492
## beta[4] -0.5321434  0.2182482 -0.9627798 -0.68163811 -0.5263027 -0.3838414
##           stats
## parameter      97.5%
## beta[1]  3.3325290
## beta[2]  0.6195108
## beta[3] -0.5741417
## beta[4] -0.1101182
##
## , , chains = chain:2
##
##           stats
## parameter      mean      sd      2.5%      25%      50%      75%
## beta[1]  2.9526095  0.1870667  2.6023559  2.820663545  2.9482229  3.0806480
## beta[2]  0.1622171  0.2282766 -0.2787305  0.005698821  0.1627164  0.3299335
## beta[3] -0.7713113  0.1051982 -0.9777381 -0.844526925 -0.7719289 -0.7015365
## beta[4] -0.5263961  0.2114555 -0.9427642 -0.675811834 -0.5245449 -0.3817802
##           stats
## parameter      97.5%
## beta[1]  3.3123095

```

```
## beta[2] 0.5910822
## beta[3] -0.5648475
## beta[4] -0.1289649
##
## , , chains = chain:3
##
##      stats
## parameter      mean      sd      2.5%      25%      50%      75%
## beta[1]  2.9547756 0.1827627 2.5959493 2.83238963 2.9569740 3.0727130
## beta[2]  0.1860422 0.2299808 -0.2745304 0.03600071 0.1921720 0.3465467
## beta[3] -0.7733004 0.1056907 -0.9796632 -0.84050701 -0.7700801 -0.7038971
## beta[4] -0.5244188 0.2085120 -0.9333820 -0.65712731 -0.5273655 -0.3813274
##      stats
## parameter      97.5%
## beta[1]  3.3344582
## beta[2]  0.6170123
## beta[3] -0.5718877
## beta[4] -0.1328086
```

```
MCMC_traceplot(fit, c('beta[1]', 'beta[2]', 'beta[3]', 'beta[4]'))
```

