

1. BayesSurv\_HReg : independent, univariate time-to-event data fit to a Cox PH model with Weibull baseline hazard
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12. BayesID\_HReg : cluster-correlated semi-competing risks data using an illness-death model with PEM baseline hazards

## Definition of the survival model

Let  $t_i$  denote the time-to-event of interest for individuals  $i = 1, \dots, n$ , subject to right censoring at time  $c_i$ . Let  $(y_i, \delta_i, x_i)$  denote independent observations, where  $y_i = \min(t_i, c_i)$ ,  $\delta_i = \mathbb{1}(t_i \leq c_i)$ , and  $x_i$  is a vector of covariates for individual  $i$ . The following Cox proportional hazards model is assumed

$$h(t_i|x_i) = h_0(t_i) \exp(x_i^\top \beta), \quad t_i > 0,$$

where the baseline hazard  $h_0$  is defined parametrically by a Weibull hazard,  $h_0(t) = \alpha \kappa t^{\alpha-1}$ .

In the Bayesian framework, priors must be specified for the regression parameter,  $\beta$ , and the shape and scale parameters of baseline hazard function,  $\alpha$  and  $\kappa$ , respectively. The following specifications are made

$$\begin{aligned} \pi(\beta) &\propto 1, \\ \pi(\alpha) &\sim \text{Gamma}(a, b), \\ \pi(\kappa) &\sim \text{Gamma}(c, d). \end{aligned}$$

## Hyperparameters

The hyperparameters  $a$  and  $b$  must be specified for the prior distribution of  $\alpha$  which is a Gamma distribution with mean  $ab$  and variance  $ab^2$ . Similarly, the hyperparameters  $c$  and  $d$  must be specified for the Gamma prior of  $\kappa$ .

## Arguments to specify

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### Model-related

**Formula** a **Formula** object that corresponds to the hazard  $h(t_i|x_i): y + \delta \sim x$ .  
**data** an  $(n \times q)$ -dimensional data.frame; the  $q$ -columns correspond to  $q$  covariate vectors named in the formula in **Formula**.

### Hyperparameters

**WB.ab** a 2-vector of positive hyperparameters  $a$  and  $b$  of the prior distribution for the shape parameter  $\alpha$  of the Weibull baseline hazard. Example: `WB.ab <- c(0.5, 0.01)`.  
**WB.cd** a 2-vector of positive hyperparameters  $c$  and  $d$  of the prior distribution for the scale parameter  $\kappa$  of the Weibull baseline hazard. Example: `WB.cd <- c(0.5, 0.05)`.

### MCMC Settings

**numReps** total number of scans  
**thin** extent of thinning, e.g. if `thin=10` retain every 10<sup>th</sup> sample.  
**burninPerc** the proportion of burn-in (samples to be discarded before analyzing the data).  
**mhProp\_alpha\_var** the shape parameter  $\alpha$  is updated using a Metropolis-Hastings random walk step generating proposals from a Gamma distribution with variance `mhProp_alpha_var`.

### Starting Values

**startValues** use `initiate.startValues_HReg(Formula, data, model, nChain, beta = NULL, WB.alpha = NULL, WB.kappa = NULL)` which initiates starting values for  $\beta$ ,  $\alpha$  and  $\kappa$  in the Metropolis-Hastings algorithm if left unspecified. Users may set non-null starting values for any of these parameters.

### Storage

**path** name of the directory where results are stored. Can leave unspecified.

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## Implementation

```
data(survData)
form <- Formula(time + event ~ cov1 + cov2)
##
WB.ab <- c(0.5, 0.01) # prior parameters for alpha
WB.cd <- c(0.5, 0.05) # prior parameters for kappa
hyperParams <- list(WB=list(WB.ab=WB.ab, WB.cd=WB.cd))
##
numReps <- 2000
burninPerc <- 0.5
thin <- 10
mhProp_alpha_var <- 0.01
mcmc <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
            tuning=list(mhProp_alpha_var=mhProp_alpha_var))
##
myModel <- "Weibull"
myPath <- "Output/01-Results-WB/"
startValues <- initiate.startValues_HReg(form, survData, model=myModel, nChain=2)
##
fit_WB <- BayesSurv_HReg(form, survData, id=NULL, model=myModel, hyperParams, startValues, mcmc, path=myPath)
summary(fit_WB)
pred_WB <- predict(fit_WB, tseq=seq(from=0, to=30, by=5))
plot(pred_WB, plot.est="Haz")
plot(pred_WB, plot.est="Surv")
```

## Definition of the survival model

Let  $t_i$  denote the time-to-event of interest for individuals  $i = 1, \dots, n$ , subject to right censoring at time  $c_i$ . Let  $(y_i, \delta_i, x_i)$  denote independent observations, where  $y_i = \min(t_i, c_i)$ ,  $\delta_i = \mathbb{1}(t_i \leq c_i)$ , and  $x_i$  is a vector of covariates for individual  $i$ . The following Cox proportional hazards model is assumed

$$h(t_i|x_i) = h_0(t_i) \exp(x_i^\top \beta), \quad t_i > 0.$$

The baseline hazard  $h_0$  is defined non-parametrically by a mixture of piecewise exponential functions as follows

$$\lambda_0(t) = \log h_0(t) = \sum_{k=1}^{K+1} \lambda_k \mathbb{1}\{t \in (s_{k-1}, s_k]\},$$

where  $\lambda_k$  is constant and the time interval between 0 and the largest observed failure time, denoted  $s_K$ , is partitioned into  $K + 1$  disjoint intervals:  $0 < s_1 < \dots < s_{K+1}$ .

In the Bayesian framework, priors must be specified for the regression parameter,  $\beta$ , the number of intervals,  $K$ , and the partition points  $(s_1, \dots, s_{K+1})$ , respectively. The following specifications are made

$$\begin{aligned} \pi(\beta) &\propto 1, \\ \lambda|K, \mu_\lambda, \sigma_\lambda^2 &\sim MVN_{K+1}(\mu_\lambda \mathbb{1}, \sigma_\lambda^2 \Sigma_\lambda) \\ K &\sim \text{Poisson}(\alpha), \\ \pi(s|K) &\propto \frac{(2K+1)! \prod_{k=1}^{K+1} (s_k - s_{k-1})}{(s_{K+1})^{(2K+1)}}, \\ \pi(\mu_\lambda) &\propto 1, \\ \sigma_\lambda^{-2} &\sim \text{Gamma}(a, b). \end{aligned}$$

The prior specification for  $\lambda$  follows a MVN-ICAR (see Supplemental Material to Lee, Haneuse, Schrag and Dominici, 2015). Note that  $K$  and  $s$  jointly form a time-homogeneous Poisson process prior for the partition.

## Hyperparameters

The hyperparameter  $\alpha$  must be specified for the prior distribution of  $K$ , as well as  $a$  and  $b$ , the rate and shape of the Gamma distributed hyperprior for  $\sigma_\lambda^{-2}$ .

## Arguments to specify

<b>Model-related</b>	
Formula	a <code>Formula</code> object that corresponds to the hazard $h(t_i x_i)$ : $y + \delta \sim x$ .
data	an $(n \times q)$ -dimensional <code>data.frame</code> ; the $q$ -columns correspond to $q$ covariate vectors named in the formula in <code>Formula</code> .
<b>Hyperparameters</b>	
PEM.ab	a 2-vector of positive hyperparameters $a$ and $b$ of the prior distribution for $\sigma_\lambda^{-2}$ . Example: <code>PEM.ab &lt;- c(0.7, 0.7)</code> .
PEM.alpha	hyperparameter $\alpha$ of the prior distribution for $K$ , which is one less than the number of partition points. Example: <code>PEM.alpha &lt;- 10</code> .
<b>MCMC Settings</b>	
numReps	total number of scans
thin	extent of thinning, e.g. if <code>thin=10</code> retain every $10^{th}$ sample.
burninPerc	the proportion of burn-in (samples to be discarded before analyzing the data).
C	a numeric value for the proportion that determines the sum of probabilities choosing the birth and death moves. <sup>1</sup>
delPert	the perturbation parameter in the birth updates; values must be between 0 and 0.5. <sup>1</sup>
rj.scheme	<code>rj.scheme=1</code> : the birth update will draw the proposal time split from $1 : s_{max}$ ; <code>rj.scheme=2</code> : the birth update will draw the proposal time split from uniquely ordered failure times in the data.
K_max	the number of splits allowed in each iteration of the Metropolis-Hastings-Green algorithm.
s_max	the largest observed failure time, given by <code>s_max &lt;- max(data\$time[data\$event==1])</code>
time_lambda	time points at which the $\lambda$ is monitored for convergence. Example: <code>time_lambda &lt;- seq(1, s_max, 1)</code> . The chains for these monitoring points can be found in <code>lambda.fin</code> in the chains of the <code>BayesSurv_HReg</code> object.
<b>Starting Values</b>	
startValues	use <code>initiate.startValues_HReg(Formula, data, model, nChain, beta = NULL)</code> which initiates all necessary starting values in the Metropolis-Hastings-Green algorithm. Users may set non-null starting values for <code>beta</code> .
<b>Storage</b>	
path	name of the directory where results are stored. Can leave unspecified.

<sup>1</sup>See Section A in Supplemental Material to Lee et al. (2015)

## Implementation

```
data(survData)
form <- Formula(time + event ~ cov1 + cov2)
##
PEM.ab <- c(0.7, 0.7) # prior parameters for 1/sigma^2
PEM.alpha <- 10 # prior parameters for K
hyperParams <- list(PEM=list(PEM.ab=PEM.ab, PEM.alpha=PEM.alpha))
##
numReps <- 2000
burninPerc <- 0.5
thin <- 10
C <- 0.2
delPert <- 0.5
rj.scheme <- 2
K_max <- 50
s_max <- max(survData$time[survData$event == 1])
time_lambda <- seq(1, s_max, 0.5)
mcmc <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
            tuning=list(C=C, delPert=delPert, rj.scheme=rj.scheme,
                       K_max=K_max, s_max=s_max, time_lambda=time_lambda) )
##
myModel <- "PEM"
myPath <- "Output/02-Results-PEM/"
startValues <- initiate.startValues_HReg(form, survData, model=myModel, nChain=2)
##
fit_PEM <- BayesSurv_HReg(form, survData, id=NULL, model=myModel,
                        hyperParams, startValues, mcmc, path=myPath)
summary(fit_PEM)
pred_PEM <- predict(fit_PEM)
plot(pred_PEM, plot.est="Haz")
plot(pred_PEM, plot.est="Surv")
```

### Definition of the survival model

Let  $t_i$  denote the time-to-event of interest for individuals  $i = 1, \dots, n$ . In the presence of interval censoring, the time-to-event for the  $i^{\text{th}}$  subject satisfies  $c_{ij} \leq t_i < c_{ij+1}$ . Let  $(c_{ij}, c_{ij+1}, L_i, x_i)$  denote independent observations, where  $L_i$  is the left-truncation time and  $x_i$  is a vector of covariates for individual  $i$ . The following AFT model is assumed

$$\log(t_i) = x_i^\top \beta + \epsilon_i, \quad t_i > 0.$$

We take  $\epsilon_i$  to follow the Normal( $\mu, \sigma^2$ ) distribution for  $\epsilon_i$  for the parametric AFT model. In the Bayesian framework, priors must be specified for  $\beta$ ,  $\mu$ , and  $\sigma^2$ . The following specifications are made

$$\begin{aligned} \pi(\beta, \mu) &\propto 1, \\ \sigma^2 &\sim \text{Inverse-Gamma}(a_\sigma, b_\sigma). \end{aligned}$$

### Hyperparameters

The hyperparameters,  $a_\sigma$  and  $b_\sigma$ , must be specified for the prior distribution of  $\sigma^2$ .

### Arguments to specify

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<b>Model-related</b>	
Formula	a Formula object that corresponds to $\log(t_i): L y_L + y_U \sim x$ .
data	an $(n \times q)$ -dimensional data.frame; the $q$ -columns correspond to $q$ covariate vectors named in the formula in Formula.
<b>Hyperparameters</b>	
LN.ab	a 2-vector of positive hyperparameters $a$ and $b$ of the prior distribution for $\sigma^2$ . Example: LN.ab <- c(0.7, 0.7).
<b>MCMC Settings</b>	
numReps	total number of scans
thin	extent of thinning, e.g. if thin=10 retain every 10 <sup>th</sup> sample.
burninPerc	the proportion of burn-in (samples to be discarded before analyzing the data).
beta.prop.var	the parameter $\beta$ is updated using a Metropolis-Hastings random walk step generating proposals from a Normal distribution with variance beta.prop.var.
mu.prop.var	the parameter $\mu$ is updated using a Metropolis-Hastings random walk step generating proposals from a Normal distribution with variance mu.prop.var.
zeta.prop.var	the parameter $\zeta = 1/\sigma^2$ is updated using a Metropolis-Hastings random walk step generating proposals from a log-Normal distribution with variance zeta.prop.var.
<b>Starting Values</b>	
startValues	use initiate.startValues_AFT(Formula, data, model, nChain, beta = NULL, y = NULL, LN.mu = NULL, LN.sigSq = NULL) which initiates all necessary starting values in the Metropolis-Hastings algorithm. Users may set non-null starting values for beta, y, LN.mu, LN.sigSq.
<b>Storage</b>	
path	name of the directory where results are stored. Can leave unspecified.

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### Implementation

```
data(survData)
survData$yL <- survData$yU <- survData[,1]
survData$yU[which(survData[,2] == 0)] <- Inf
survData$LT <- rep(0, dim(survData)[1])
form <- Formula(LT | yL + yU ~ cov1 + cov2)
##
LN.ab <- c(0.3, 0.3)
hyperParams <- list(LN=list(LN.ab=LN.ab))
##
numReps <- 1000
thin <- 10
burninPerc <- 0.5
beta.prop.var <- 0.01
mu.prop.var <- 0.1
zeta.prop.var <- 0.1
mcmcParams <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
tuning=list(beta.prop.var=beta.prop.var, mu.prop.var=mu.prop.var,
zeta.prop.var=zeta.prop.var))
##
myModel <- "LN"
myPath <- "Output/01-Results-LN/"
startValues <- initiate.startValues_AFT(form, survData, model=myModel, nChain=2)
##
fit_LN <- BayesSurv_AFT(form, survData, model=myModel, hyperParams,
startValues, mcmcParams, path=myPath)

summary(fit_LN)
pred_LN <- predict(fit_LN, time = seq(0, 35, 1), tseq=seq(from=0, to=30, by=5))
plot(pred_LN, plot.est="Haz")
plot(pred_LN, plot.est="Surv")
```

**Definition of the survival model**

Let  $t_i$  denote the time-to-event of interest for individuals  $i = 1, \dots, n$ . Considering interval censoring, the time-to-event for the  $i^{\text{th}}$  subject satisfies  $c_{ij} \leq t_i < c_{i,j+1}$ . Let  $(c_{ij}, c_{i,j+1}, L_i, x_i)$  denote independent observations, where  $L_i$  is the left-truncation time and  $x_i$  is a vector of covariates for individual  $i$ . The following AFT model is assumed

$$\log(t_i) = x_i^\top \beta + \epsilon_i, \quad t_i > 0,$$

where  $\epsilon_i$  is assumed to be taken as draws from the DPM of normal distributions:

$$\begin{aligned} \epsilon_i | r_i &\sim \text{Normal}(\mu_{r_i}, \sigma_{r_i}^2), \\ (\mu_r, \sigma_r^2) &\sim G_0, \quad \text{for } r = 1, \dots, M, \\ r_i | p &\sim \text{Discrete}(r_i | p_1, \dots, p_M), \\ p &\sim \text{Dirichlet}(\tau/M, \dots, \tau/M). \end{aligned}$$

In the Bayesian framework, priors must be specified for the unknown parameters. We take the  $G_0$  as a normal distribution centered at  $\mu_0$  with a variance  $\sigma_0^2$  for  $\mu_r$  and an inverse-Gamma( $a_\sigma, b_\sigma$ ) for  $\sigma_r^2$ . For  $\beta$ , we adopt non-informative flat priors on the real line. Finally, we specify a Gamma( $a_\tau, b_\tau$ ) hyperprior for the precision parameter  $\tau$ .

**Hyperparameters**

The hyperparameter  $(\mu_0, \sigma_0^2, a_\sigma, b_\sigma)$  must be specified for the centering distribution  $G_0$ , as well as  $a_\tau$  and  $b_\tau$ , the rate and shape of the Gamma distributed hyperprior for  $\tau$ .

**Arguments to specify**


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<b>Model-related</b>	
Formula	a <code>Formula</code> object that corresponds to $\log(t_i)$ : $L y_L + y_U \sim x$ .
data	an $(n \times q)$ -dimensional data.frame; the $q$ -columns correspond to $q$ covariate vectors named in the formula in <code>Formula</code> .
<b>Hyperparameters</b>	
DPM.mu	a hyperparameter $\mu_0$ of the centering distribution $G_0$ .
DPM.sigSq	a positive-valued hyperparameter $\sigma_0^2$ of the centering distribution $G_0$ .
DPM.ab	a 2-vector of positive hyperparameters $a_\sigma$ and $b_\sigma$ of the centering distribution $G_0$ .
Tau.ab	a 2-vector of positive hyperparameters $a_\tau$ and $b_\tau$ of the hyperprior distribution for $\tau$ . Example: <code>Tau.ab &lt;- c(1.5, 0.0125)</code> .
<b>MCMC Settings</b>	
numReps	total number of scans
thin	extent of thinning, e.g. if <code>thin=10</code> retain every 10 <sup>th</sup> sample.
burninPerc	the proportion of burn-in (samples to be discarded before analyzing the data).
beta.prop.var	the parameter $\beta$ is updated using a Metropolis-Hastings random walk step generating proposals from a Normal distribution with variance <code>beta.prop.var</code> .
mu.prop.var	the parameter $\mu_r$ is updated using a Metropolis-Hastings random walk step generating proposals from a Normal distribution with variance <code>mu.prop.var</code> .
zeta.prop.var	the parameter $\zeta_r = 1/\sigma_r^2$ is updated using a Metropolis-Hastings random walk step generating proposals from a log-Normal distribution with variance <code>zeta.prop.var</code> .
<b>Starting Values</b>	
startValues	use <code>initiate.startValues_AFT(Formula, data, model, nChain, beta = NULL, y = NULL, DPM.class = NULL, DPM.mu = NULL, DPM.zeta=NULL, DPM.tau=NULL)</code> which initiates all necessary starting values in the Metropolis-Hastings algorithm. Users may set non-null starting values for <code>beta</code> , <code>y</code> , <code>DPM.class</code> , <code>DPM.mu</code> , <code>DPM.zeta</code> , <code>DPM.tau</code> .
<b>Storage</b>	
path	name of the directory where results are stored. Can leave unspecified.

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## Implementation

```
data(survData)
survData$yL <- survData$yU <- survData[,1]
survData$yU[which(survData[,2] == 0)] <- Inf
survData$LT <- rep(0, dim(survData)[1])
form <- Formula(LT | yL + yU ~ cov1 + cov2)
##
DPM.mu <- log(12)
DPM.sigSq <- 100
DPM.ab <- c(2, 1)
Tau.ab <- c(1.5, 0.0125)
hyperParams <- list(DPM=list(DPM.mu=DPM.mu, DPM.sigSq=DPM.sigSq, DPM.ab=DPM.ab, Tau.ab=Tau.ab))
##
numReps <- 1000
thin <- 10
burninPerc <- 0.5
beta.prop.var <- 0.01
mu.prop.var <- 0.1
zeta.prop.var <- 0.1
mcmcParams <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
tuning=list(beta.prop.var=beta.prop.var, mu.prop.var=mu.prop.var,
zeta.prop.var=zeta.prop.var))
##
myModel <- "DPM"
myPath <- "Output/02-Results-DPM/"
startValues <- initiate.startValues_AFT(form, survData, model=myModel, nChain=2)
##
fit_DPM <- BayesSurv_AFT(form, survData, model=myModel, hyperParams,
startValues, mcmcParams, path=myPath)

summary(fit_DPM)
pred_DPM <- predict(fit_DPM, time = seq(0, 35, 1), tseq=seq(from=0, to=30, by=5))
plot(pred_DPM, plot.est="Haz")
plot(pred_DPM, plot.est="Surv")
```

**Definition of the survival model**

Let  $t_{ji}$  denote the time-to-event of interest for individuals  $i = 1, \dots, n_j$  in cluster  $j = 1, \dots, J$ , subject to right censoring at time  $c_{ji}$ . Let  $(y_{ji}, \delta_{ji}, x_{ji})$  denote independent observations, where  $y_{ji} = \min(t_{ji}, c_{ji})$ ,  $\delta_{ji} = \mathbb{1}(t_{ji} \leq c_{ji})$ , and  $x_{ji}$  is a vector of covariates for individual  $i$ . The following Cox proportional hazards model is assumed

$$h(t_{ji}|x_{ji}) = h_0(t_{ji}) \exp(x_{ji}^T \beta + V_j), \quad t_{ji} > 0,$$

where the  $V_j$ 's are cluster-specific random effects and the baseline hazard  $h_0$  is defined parametrically by a Weibull hazard,  $h_0(t) = \alpha \kappa t^{\alpha-1}$ .

In the Bayesian framework, priors must be specified for the regression parameter,  $\beta$ , the cluster-specific random effects,  $V_j$ , and the shape and scale parameters of baseline hazard function,  $\alpha$  and  $\kappa$ , respectively. The prior distributions for  $\beta$ ,  $\alpha$  and  $\kappa$  are given below.

$$\begin{aligned} \pi(\beta) &\propto 1, \\ \pi(\alpha) &\sim \text{Gamma}(a, b), \\ \pi(\kappa) &\sim \text{Gamma}(c, d). \end{aligned}$$

We provide two possible prior specifications for the cluster-specific random effects below.

$$\begin{aligned} V_j &\sim \text{Normal}(0, \sigma^2), & V_j|m_j &\sim \text{Normal}(\mu_m, \sigma_m^2), \\ \zeta = \frac{1}{\sigma^2} &\sim \text{Gamma}(a_N, b_N), & (\mu_m, \sigma_m^2) &\sim G_0, \text{ for } m = 1, \dots, M, \\ & & m_j|p &\sim \text{Discrete}(m_j|p_1, \dots, p_M), \\ & & p &\sim \text{Dirichlet}(\tau/M, \dots, \tau/M), \\ & & \tau &\sim \text{Gamma}(a_\tau, b_\tau). \end{aligned}$$

In the first column, the individual specific-random effects are assumed to be  $iid N(0, \sigma^2)$ . In the second column, the cluster-specific random effects are drawn from a mixture of  $M$  normal distributions each with mean and variance  $(\mu_m, \sigma_m^2)$  which are distributed as a multivariate Normal/Inverse-Gamma (NIG), denoted by  $G_0$ ; we refer to this as the Dirichlet process mixture (DPM) prior. The probability density of  $G_0$  is defined by the product

$$f_{\text{NIG}}(\mu, \sigma^2 | \mu_0, \zeta_0, a_0, b_0) = f_{\text{Normal}}(\mu | \mu_0, 1/\zeta_0^2) \times f_{\text{Gamma}}(\zeta = 1/\sigma^2 | a_0, b_0).$$

We assume  $\mu_0 = 0$  and  $\zeta_0 = 1$ .

**Hyperparameters**

- $a, b$  : shape and rate of Gamma prior for  $\alpha$
- $c, d$  : shape and rate of Gamma prior for  $\kappa$
- $a_N, b_N$  : mean and variance of normal prior for  $V_j$
- $a_0, b_0$  : shape and rate of Gamma component of the prior distribution,  $G_0$ , of  $(\mu_m, \sigma_m^2)$  (DPM prior)
- $a_\tau, b_\tau$  : shape and rate of Gamma hyperprior for  $\tau$  (DPM prior)

**Arguments to specify****Model-related**

- Formula** a **Formula** object that corresponds to the hazard  $h(t_i|x_i): y + \delta \sim x$ .
- data** an  $(n \times q)$ -dimensional data.frame; the  $q$ -columns correspond to  $q$  covariate vectors named in the formula in **Formula**.
- model** a character vector that specifies the type of components in the model. Use **model** <- c("Weibull", "Normal") for Normal prior for  $V_j$  and use **model** <- c("Weibull", "DPM") for DPM prior.
- id** an  $n$ -vector of cluster information where cluster membership corresponds to one of the positive integers  $1, \dots, J$ .

**Hyperparameters**

- WB.ab** a 2-vector of positive hyperparameters  $a$  and  $b$  of the prior distribution for the shape parameter  $\alpha$  of the Weibull baseline hazard. Example: **WB.ab** <- c(0.5, 0.01).
- WB.cd** a 2-vector of positive hyperparameters  $c$  and  $d$  of the prior distribution for the scale parameter  $\kappa$  of the Weibull baseline hazard. Example: **WB.cd** <- c(0.5, 0.05).

**Normal prior for  $V_j$** 

- Normal.ab** a 2-vector of positive hyperparameters  $a_N$  and  $b_N$  of the prior for  $1/\sigma^2$ , the precision of the normally distributed cluster-specific random effects. Example: **Normal.ab** <- c(0.5, 0.01).

**DPM prior for  $V_j$** 

- DPM.ab** a 2-vector of positive hyperparameters  $a_0$  and  $b_0$  of the prior for  $(\mu_m, \sigma_m^2)$ , the parameters of the normally distributed cluster-specific random effects. Example: **DPM.ab** <- c(0.5, 0.01).
- aTau** a positive-valued hyperparameter corresponding to the shape parameter,  $a_\tau$ , of the Gamma prior of  $\tau$ .
- bTau** a positive-valued hyperparameter corresponding to the rate parameter,  $b_\tau$ , of the Gamma prior of  $\tau$ .

**MCMC Settings**

- numReps** total number of scans
- thin** extent of thinning, e.g. if **thin**=10 retain every 10<sup>th</sup> sample.
- burninPerc** the proportion of burn-in (samples to be discarded before analyzing the data).
- mhProp\_alpha\_var** the shape parameter  $\alpha$  is updated using a Metropolis-Hastings random walk algorithm which generates proposals from a Gamma distribution with variance **mhProp\_alpha\_var**.
- mhProp\_V\_var** the cluster-specific random effects,  $V_{ji}$ , are updated using a Metropolis-Hastings random walk algorithm which generates proposals from a Normal distribution with variance **mhProp\_V\_var**

**Starting Values**

- startValues** use **initiate.startValues\_HReg(Formula, data, model, id, nChain, beta = NULL, WB.alpha = NULL, WB.kappa = NULL, V.j = NULL, Normal.zeta = NULL, DPM.class = NULL, DPM.tau = NULL)** which initiates starting values for  $\beta$ ,  $\alpha$ ,  $\kappa$ ,  $V_j$ ,  $\zeta$  (in the DPM model for  $V_j$ ) and  $\tau$  in the Metropolis-Hastings-Green algorithm if left unspecified; **DPM.class** sets the starting value for class membership in the DPM model. Users may set non-null starting values for any of these parameters.



## Storage

path name of the directory where results are stored. Can leave unspecified.  
storeV a TRUE/FALSE logical constant indicating storage of  $V_j$  values.

---

## Implementation

```
data(survData)
id=survData$cluster
form <- Formula(time + event ~ cov1 + cov2)
##
WB.ab <- c(0.5, 0.01) # prior parameters for alpha
WB.cd <- c(0.5, 0.05) # prior parameters for kappa
Normal.ab <- c(0.5, 0.01) # for Normal random effects
DPM.ab <- c(0.5, 0.01) # For DPM
  aTau <- 1.5
  bTau <- 0.0125
hyperParams.WB.Normal <- list(WB=list(WB.ab=WB.ab, WB.cd=WB.cd),
                             Normal=list(Normal.ab=Normal.ab))
hyperParams.WB.DPM <- list(WB=list(WB.ab=WB.ab, WB.cd=WB.cd),
                           DPM=list(DPM.ab=DPM.ab, aTau=aTau, bTau=bTau))
##
numReps <- 2000
burninPerc <- 0.5
thin <- 10
mhProp_alpha_var <- 0.01
mhProp_V_var <- 0.05
storeV <- TRUE
mcmc.WB <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
               storage=list(storeV=storeV),
               tuning=list(mhProp_alpha_var=mhProp_alpha_var, mhProp_V_var=mhProp_V_var))
##
myModel.WB.Normal <- c("Weibull","Normal")
myPath.WB.Normal <- "Output/03-Results-WB_Normal/"
startValues.WB.Normal <- initiate.startValues_HReg(form, survData, id, model=myModel.WB.Normal, nChain=2)
##
fit_WB_N <- BayesSurv_HReg(form, survData, id, model=myModel.WB.Normal, hyperParams.WB.Normal,
                          startValues.WB.Normal, mcmc.WB, path=myPath.WB.Normal)
summary(fit_WB_N)
pred_WB_N <- predict(fit_WB_N, tseq=seq(from=0, to=30, by=5))
plot(pred_WB_N, plot.est="Haz")
plot(pred_WB_N, plot.est="Surv")
##
myModel.WB.DPM <- c("Weibull","DPM")
myPath.WB.DPM <- "Output/04-Results-WB_DPM/"
startValues.WB.DPM <- initiate.startValues_HReg(form, survData, id, model=myModel.WB.DPM, nChain=2)
##
fit_WB_DPM <- BayesSurv_HReg(form, survData, id, model=myModel.WB.DPM, hyperParams.WB.DPM,
                          startValues.WB.DPM, mcmc.WB, path=myPath.WB.DPM)
summary(fit_WB_DPM)
pred_WB_DPM <- predict(fit_WB_DPM, tseq=seq(from=0, to=30, by=5))
plot(pred_WB_DPM, plot.est="Haz")
plot(pred_WB_DPM, plot.est="Surv")
```

**Definition of the survival model**

Let  $t_{ji}$  denote the time-to-event of interest for individuals  $i = 1, \dots, n_j$  in cluster  $j = 1, \dots, J$ , subject to right censoring at time  $c_{ji}$ . Let  $(y_{ji}, \delta_{ji}, x_{ji})$  denote independent observations, where  $y_{ji} = \min(t_{ji}, c_{ji})$ ,  $\delta_{ji} = \mathbb{1}(t_{ji} \leq c_{ji})$ , and  $x_{ji}$  is a vector of covariates for individual  $i$ . The following Cox proportional hazards model is assumed

$$h(t_{ji}|x_{ji}) = h_0(t_{ji}) \exp(x_{ji}^\top \beta + V_j), \quad t_{ji} > 0,$$

The baseline hazard  $h_0$  is defined non-parametrically by a mixture of piecewise exponential functions as follows

$$\lambda_0(t) = \log h_0(t) = \sum_{k=1}^{K+1} \lambda_k \mathbb{1}\{t \in (s_{k-1}, s_k]\},$$

where  $\lambda_k$  is constant and the time interval between 0 and the largest observed failure time, denoted  $s_k$ , is partitioned into  $K + 1$  disjoint intervals:  $0 < s_1 < \dots < s_{K+1}$ .

In the Bayesian framework, priors must be specified for the regression parameter,  $\beta$ , the number of intervals,  $K$ , and the partition points  $(s_1, \dots, s_{K+1})$ , respectively. The following specifications are made

$$\begin{aligned} \pi(\beta) &\propto 1, \\ \lambda|K, \mu_\lambda, \sigma_\lambda^2 &\sim MVN_{K+1}(\mu_\lambda \mathbb{1}, \sigma_\lambda^2 \Sigma_\lambda) \\ K &\sim \text{Poisson}(\alpha), \\ \pi(s|K) &\propto \frac{(2K+1)! \prod_{k=1}^{K+1} (s_k - s_{k-1})}{(s_{K+1})^{(2K+1)}}, \\ \pi(\mu_\lambda) &\propto 1, \\ \sigma_\lambda^{-2} &\sim \text{Gamma}(a, b). \end{aligned}$$

The prior specification for  $\lambda$  follows a MVN-ICAR (see Supplemental Material to Lee, Haneuse, Schrag and Dominici, 2015). Note that  $K$  and  $s$  jointly form a time-homogeneous Poisson process prior for the partition.

We provide two possible prior specifications for the cluster-specific random effects below.

$$\begin{aligned} V_j &\sim \text{Normal}(0, \sigma^2), & V_j|m_j &\sim \text{Normal}(\mu_m, \sigma_m^2), \\ \zeta = \frac{1}{\sigma^2} &\sim \text{Gamma}(a_N, b_N), & (\mu_m, \sigma_m^2) &\sim G_0, \text{ for } m = 1, \dots, M, \\ & & m_j|p &\sim \text{Discrete}(m_j|p_1, \dots, p_M), \\ & & p &\sim \text{Dirichlet}(\tau/M, \dots, \tau/M), \\ & & \tau &\sim \text{Gamma}(a_\tau, b_\tau). \end{aligned}$$

In the first column, the individual specific-random effects are assumed to be  $iid$   $N(0, \sigma^2)$ . In the second column, the cluster-specific random effects are drawn from a mixture of  $M$  normal distributions each with mean and variance  $(\mu_m, \sigma_m^2)$  which are distributed as a multivariate Normal/Inverse-Gamma (NIG), denoted by  $G_0$ ; we refer to this as the Dirichlet process mixture (DPM) prior. The probability density of  $G_0$  is defined by the product

$$f_{\text{NIG}}(\mu, \sigma^2|\mu_0, \zeta_0, a_0, b_0) = f_{\text{Normal}}(\mu|\mu_0, 1/\zeta_0^2) \times f_{\text{Gamma}}(\zeta = 1/\sigma^2|a_0, b_0).$$

We assume  $\mu_0 = 0$  and  $\zeta_0 = 1$ .

**Hyperparameters**

$\alpha$	: hyperparameter of $K$
$a, b$	: shape and rate of Gamma prior for $\sigma_\lambda^{-2}$
$a_N, b_N$	: mean and variance of normal prior for $V_j$
$a_0, b_0$	: shape and rate of Gamma component of the prior distribution, $G_0$ , of $(\mu_m, \sigma_m^2)$
$a_\tau, b_\tau$	: shape and rate of Gamma hyperprior for $\tau$

**Arguments to specify****Model-related**

Formula	a <code>Formula</code> object that corresponds to the hazard $h(t_i x_i)$ : $y + \delta \sim x$ .
data	an $(n \times q)$ -dimensional data.frame; the $q$ -columns correspond to $q$ covariate vectors named in the formula in <code>Formula</code> .
model	a character vector that specifies the type of components in the model. Use <code>model &lt;- c("PEM", "DPM")</code> .
id	an $n$ -vector of cluster information where cluster membership corresponds to one of the positive integers $1, \dots, J$ .

**Hyperparameters**

PEM.ab	a 2-vector of positive hyperparameters $a$ and $b$ of the prior distribution for $\sigma_\lambda^{-2}$ . Example: <code>PEM.ab &lt;- c(0.7, 0.7)</code> .
PEM.alpha	hyperparameter $\alpha$ of the prior distribution for $K$ , which is one less than the number of partition points. Example: <code>PEM.alpha &lt;- 10</code> .

**Normal prior for  $V_j$** 

Normal.ab	a 2-vector of positive hyperparameters $a_N$ and $b_N$ of the prior for $1/\sigma^2$ , the precision of the normally distributed cluster-specific random effects. Example: <code>Normal.ab &lt;- c(0.5, 0.01)</code> .
-----------	--

**DPM prior for  $V_j$** 

DPM.ab	a 2-vector of positive hyperparameters $a_0$ and $b_0$ of the prior for $(\mu_m, \sigma_m^2)$ , the parameters of the normally distributed cluster-specific random effects. Example: <code>DPM.ab &lt;- c(0.5, 0.01)</code> .
aTau	a positive-valued hyperparameter corresponding to the shape parameter, $a_\tau$ , of the Gamma prior of $\tau$ .
bTau	a positive-valued hyperparameter corresponding to the rate parameter, $b_\tau$ , of the Gamma prior of $\tau$ .

**MCMC Settings**

numReps	total number of scans
thin	extent of thinning, e.g. if <code>thin=10</code> retain every $10^{th}$ sample.
burninPerc	the proportion of burn-in (samples to be discarded before analyzing the data).

**mhProp\_V\_var** the cluster-specific random effects,  $V_{ji}$ , are updated using a Metropolis-Hastings random walk algorithm which generates proposals from a Normal distribution with variance **mhProp\_V\_var**  
**C** a numeric value for the proportion that determines the sum of probabilities choosing the birth and death moves.<sup>2</sup>  
**delPert** the perturbation parameter in the birth updates; values must be between 0 and 0.5.<sup>2</sup>  
**rj.scheme** **rj.scheme=1**: the birth update will draw the proposal time split from 1 :  $s_{max}$ ; **rj.scheme=2**: the birth update will draw the proposal time split from uniquely ordered failure times in the data.  
**K\_max** the number of splits allowed in each iteration of the Metropolis-Hastings-Green algorithm.  
**s\_max** the largest observed failure time, given by `s_max <- max(data$time[data$event==1])`  
**time\_lambda** time points at which the  $\lambda$  is monitored for convergence. Example: `time_lambda <- seq(1, s_max, 1)`. The chains for these monitoring points can be found in `lambda.fin` in the chains of the `BayesSurv` object.

#### Starting Values

**startValues** use `initiate.startValues_HReg(Formula, data, model, nChain, beta = NULL, V.j=NULL, Normal.zeta=NULL, DPM.class=NULL, DPM.tau=NULL)` which initiates starting values for  $\beta$ ,  $V_j$ ,  $\zeta$  (in the DPM model for  $V_j$ ) and  $\tau$  in the Metropolis-Hastings-Green algorithm if left unspecified; `DPM.class` sets the starting value for class membership in the DPM model. Users may set non-null starting values for any of these parameters.

#### Storage

**path** name of the directory where results are stored. Can leave unspecified.  
**storeV** a TRUE/FALSE logical constant indicating storage of  $V_j$  values.

## Implementation

```

data(survData)
id=survData$cluster
form <- Formula(time + event ~ cov1 + cov2)
##
PEM.ab <- c(0.7, 0.7) # prior parameters for 1/sigma^2
PEM.alpha <- 10 # prior parameters for K
Normal.ab <- c(0.5, 0.01) # for Normal random effects
DPM.ab <- c(0.5, 0.01) # For DPM
  aTau <- 1.5
  bTau <- 0.0125
hyperParams.PEM.Normal <- list(PEM=list(PEM.ab=PEM.ab, PEM.alpha=PEM.alpha),
  Normal=list(Normal.ab=Normal.ab))
hyperParams.PEM.DPM <- list(PEM=list(PEM.ab=PEM.ab, PEM.alpha=PEM.alpha),
  DPM=list(DPM.ab=DPM.ab, aTau=aTau, bTau=bTau))

##
numReps <- 2000
burninPerc <- 0.5
thin <- 10
mhProp_V_var <- 0.05
storeV <- TRUE
C <- 0.2
delPert <- 0.5
rj.scheme <- 2
K_max <- 50
s_max <- max(survData$time[survData$event == 1])
time_lambda <- seq(1, s_max, 0.5)
mcmc.PEM <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
  storage=list(storeV=storeV),
  tuning=list(mhProp_V_var=mhProp_V_var, C=C, delPert=delPert, rj.scheme=rj.scheme,
    K_max=K_max, s_max=s_max, time_lambda=time_lambda) )

##
myModel.PEM.Normal <- c("PEM", "Normal")
myPath.PEM.Normal <- "Output/05-Results-PEM_Normal/"
startValues.PEM.Normal <- initiate.startValues_HReg(form, survData, id, model=myModel.PEM.Normal, nChain=2)
##
fit_PEM_N <- BayesSurv_HReg(form, survData, id, model=myModel.PEM.Normal, hyperParams.PEM.Normal,
  startValues.PEM.Normal, mcmc.PEM, path=myPath.PEM.Normal)
summary(fit_PEM_N)
pred_PEM_N <- predict(fit_PEM_N)
plot(pred_PEM_N, plot.est="Haz")
plot(pred_PEM_N, plot.est="Surv")
##
myModel.PEM.DPM <- c("PEM", "DPM")
myPath.PEM.DPM <- "Output/06-Results-PEM_DPM/"
startValues.PEM.DPM <- initiate.startValues_HReg(form, survData, id, model=myModel.PEM.DPM, nChain=2)
##
fit_PEM_DPM <- BayesSurv_HReg(form, survData, id, model=myModel.PEM.DPM, hyperParams.PEM.DPM,
  startValues.PEM.DPM, mcmc.PEM, path=myPath.PEM.DPM)
pred_PEM_DPM <- predict(fit_PEM_DPM)
plot(pred_PEM_DPM, plot.est="Haz")
plot(pred_PEM_DPM, plot.est="Surv")

```

<sup>2</sup>See Section A in Supplemental Material to Lee et al. (2015)

**Definition of the survival model**

Let  $t_{i1}$  and  $t_{i2}$  denote the time to nonterminal event and terminal event from subject  $i = 1, \dots, n$ , subject to right censoring at time  $c_i$ . Let  $(y_{i1}, y_{i2}, \delta_{i1}, \delta_{i2}, x_i)$  denote independent observations, where  $y_{i1} = \min(t_{i1}, t_{i2}, c_i)$ ,  $\delta_{i1} = \mathbb{1}\{t_{i1} \leq \min(t_{i2}, c_i)\}$ ,  $y_{i2} = \min(t_{i2}, c_i)$ ,  $\delta_{i2} = \mathbb{1}\{t_{i2} \leq c_i\}$ , and  $x_i$  is a vector of covariates for individual  $i$ . The independent semi-competing risks data are assumed to arise from an illness-death model system with transitions that are modeled through the following three hazard functions:

$$h_1(t_{i1} | \gamma_{ji}, x_{i1}) = \gamma_{ji} h_{01}(t_{i1}) \exp(x_{i1}^\top \beta_1), \quad t_{i1} > 0, \quad (1)$$

$$h_2(t_{i2} | \gamma_{ji}, x_{i2}) = \gamma_{ji} h_{02}(t_{i2}) \exp(x_{i2}^\top \beta_2), \quad t_{i2} > 0, \quad (2)$$

$$h_3(t_{i2} | t_{i1}, \gamma_{ji}, x_{i3}) = \gamma_{ji} h_{03}(t_{i2}) \exp(x_{i3}^\top \beta_3), \quad t_{i2} > 0, \quad (3)$$

where  $\gamma_{ji}$  is a subject-specific frailty with vectors of covariates  $x_{i1}$ ,  $x_{i2}$  and  $x_{i3}$  which are subsets of  $x_i$ . The baseline hazard functions are defined parametrically by Weibull hazards of the form  $h_{0g}(t) = \alpha_g \kappa_g t^{\alpha_g - 1}$ , for  $g \in \{1, 2, 3\}$ . The baseline hazard function  $h_{03}$  is assumed to be Markov with respect to  $t_{i1}$ ; we will refer to the set of conditional hazard functions in (1)-(3) as the Markov model. Alternatively, we consider modeling  $h_3$  as follows:

$$h_3(t_{i2} | t_{i1}, \gamma_{ji}, x_{i3}) = \gamma_{ji} h_{03}(t_{i2} - t_{i1}) \exp(x_{i3}^\top \beta_3), \quad 0 < t_{i1} < t_{i2}. \quad (4)$$

We will refer to the set of conditional hazard functions in (1), (2) and (4) as the semi-Markov model.

In the Bayesian framework, priors must be specified for the regression parameter,  $\beta_g$ , the shape and scale parameters of baseline hazard function,  $\alpha_g$  and  $\kappa_g$ , and the frailty parameter,  $\gamma_{ji}$ , respectively, for  $g \in \{1, 2, 3\}$ . The following specifications are made

$$\begin{aligned} \pi(\beta_g) &\propto 1, \\ \alpha_g &\sim \text{Gamma}(a_g, b_g), \\ \kappa_g &\sim \text{Gamma}(c_g, d_g), \\ \gamma_{ji} | \theta &\sim \text{Gamma}(\theta^{-1}, \theta^{-1}), \\ \theta^{-1} &\sim \text{Gamma}(\psi, \omega). \end{aligned}$$

**Hyperparameters**

$a_g, b_g$  : shape and rate of Gamma prior for  $\alpha_g$  for  $g \in \{1, 2, 3\}$   
 $c_g, d_g$  : shape and rate of Gamma prior for  $\kappa_g$  for  $g \in \{1, 2, 3\}$   
 $\psi$  : the shape of Gamma prior for  $\theta^{-1}$   
 $\omega$  : the rate of Gamma prior for  $\theta^{-1}$

**Arguments to specify****Model-related**

**Formula** a `Formula` object that corresponds to  $h_g$ , for  $g \in \{1, 2, 3\}$ :  $y_1 + \delta_1 | y_2 + \delta_2 \sim x_1 | x_2 | x_3$ .  
**data** an  $(n \times q)$ -dimensional data.frame; the  $q$ -columns correspond to  $q$  covariate vectors named in the formula in **Formula** below.  
**model** c("Markov", "Weibull") for Markov definition of  $h_3$  in (3); c("semi-Markov", "Weibull") for semi-Markov definition of  $h_3$  in (4).

**Hyperparameters**

**WB.ab1** a 2-vector of positive hyperparameters  $a_1$  and  $b_1$  of the prior distribution for the shape parameter  $\alpha_1$  of the Weibull baseline hazard. Example: `WB.ab1 <- c(0.5, 0.01)`.  
**WB.ab2** a 2-vector of positive hyperparameters  $a_2$  and  $b_2$  of the prior for  $\alpha_2$ .  
**WB.ab3** a 2-vector of positive hyperparameters  $a_3$  and  $b_3$  of the prior for  $\alpha_3$ .  
**WB.cd1** a 2-vector of positive hyperparameters  $c_1$  and  $d_1$  of the prior distribution for the scale parameter  $\kappa_1$  of the Weibull baseline hazard. Example: `WB.cd1 <- c(0.5, 0.05)`.  
**WB.cd2** a 2-vector of positive hyperparameters  $c_2$  and  $d_2$  of the prior for  $\kappa_2$ .  
**WB.cd3** a 2-vector of positive hyperparameters  $c_3$  and  $d_3$  of the prior for  $\kappa_3$ .  
**theta** a 2-vector of positive hyperparameters  $\psi$  and  $\omega$  for the hyperprior  $\theta$ .

**MCMC Settings**

**numReps** total number of scans  
**thin** extent of thinning, e.g. if `thin=10` retain every  $10^{th}$  sample.  
**burninPerc** the proportion of burn-in (samples to be discarded before analyzing the data).  
**mhProp\_theta\_var** the parameter  $\theta$  is updated using a Metropolis-Hastings random walk step generating proposals from a Gamma distribution with variance `mhProp_theta_var`.  
**mhProp\_alphag\_var** a 3-vector which specifies the variances of the three random walk Metropolis-Hastings Gamma proposal distributions corresponding to  $\alpha_1$ ,  $\alpha_2$ ,  $\alpha_3$ .

**Starting Values**

**startValues** use `initiate.startValues_HReg(Formula, data, model, nChain, beta1 = NULL, beta2 = NULL, beta3 = NULL, gamma.ji=NULL, theta = NULL, WB.alpha = NULL, WB.kappa = NULL)` which initiates starting values for  $\beta_g$ ,  $\gamma_{ji}$ ,  $\theta$ ,  $\alpha_g$ , and  $\kappa_g$  in the Metropolis-Hastings algorithm if left unspecified. Users may set non-null starting values for any of these parameters.

**Storage**

**path** name of the directory where results are stored. Can leave unspecified.  
**nGam\_save** the number of  $\gamma$  to be stored.

## Implementation

```
data(scrData)
form <- Formula(time1 + event1 | time2 + event2 ~ x1 + x2 + x3 | x1 + x2 | x1 + x2)
##
WB.ab1 <- c(0.5, 0.01)
WB.ab2 <- c(0.5, 0.01)
WB.ab3 <- c(0.5, 0.01)
WB.cd1 <- c(0.5, 0.05)
WB.cd2 <- c(0.5, 0.05)
WB.cd3 <- c(0.5, 0.05)
theta <- c(0.7, 0.7) # prior params for 1/theta
hyperParams <- list(theta=theta,
                    WB=list(WB.ab1=WB.ab1, WB.ab2=WB.ab2, WB.ab3=WB.ab3,
                            WB.cd1=WB.cd1, WB.cd2=WB.cd2, WB.cd3=WB.cd3))
##
numReps <- 2000
thin <- 10
burninPerc <- 0.25
mhProp_theta_var <- 0.05
mhProp_alphag_var <- c(0.01, 0.01, 0.01)
nGam_save <- 0
mcmc.WB <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
               storage=list(nGam_save=nGam_save),
               tuning=list(mhProp_theta_var=mhProp_theta_var, mhProp_alphag_var=mhProp_alphag_var))
##
myModel <- c("semi-Markov", "Weibull")
myPath <- "Output/01-Results-WB/"
startValues <- initiate.startValues_HReg(form, scrData, model=myModel, nChain=2)
##
fit_WB <- BayesID_HReg(form, scrData, id=NULL, model=myModel,
                      hyperParams, startValues, mcmc.WB, path=myPath)
fit_WB
summary(fit_WB)
pred_WB <- predict(fit_WB, tseq=seq(from=0, to=30, by=5))
plot(pred_WB, plot.est="Haz")
plot(pred_WB, plot.est="Surv")
```

## Definition of the survival model

Let  $t_{i1}$  and  $t_{i2}$  denote the time to nonterminal event and terminal event from subject  $i = 1, \dots, n$ , subject to right censoring at time  $c_i$ . Let  $(y_{i1}, y_{i2}, \delta_{i1}, \delta_{i2}, x_i)$  denote independent observations, where  $y_{i1} = \min(t_{i1}, t_{i2}, c_i)$ ,  $\delta_{i1} = \mathbb{1}\{t_{i1} \leq \min(t_{i2}, c_i)\}$ ,  $y_{i2} = \min(t_{i2}, c_i)$ ,  $\delta_{i2} = \mathbb{1}\{t_{i2} \leq c_i\}$ , and  $x_i$  is a vector of covariates for individual  $i$ . The independent semi-competing risks data are assumed to arise from an illness-death model system with transitions that are modeled through the following three hazard functions:

$$h_1(t_{i1} | \gamma_{ji}, x_{i1}) = \gamma_{ji} h_{01}(t_{i1}) \exp(x_{i1}^\top \beta_1), \quad t_{i1} > 0, \quad (5)$$

$$h_2(t_{i2} | \gamma_{ji}, x_{i2}) = \gamma_{ji} h_{02}(t_{i2}) \exp(x_{i2}^\top \beta_2), \quad t_{i2} > 0, \quad (6)$$

$$h_3(t_{i2} | t_{i1}, \gamma_{ji}, x_{i3}) = \gamma_{ji} h_{03}(t_{i2}) \exp(x_{i3}^\top \beta_3), \quad t_{i2} > 0, \quad (7)$$

where  $\gamma_{ji}$  is a subject-specific frailty with vectors of covariates  $x_{i1}$ ,  $x_{i2}$  and  $x_{i3}$  which are subsets of  $x_i$ . The baseline hazard  $h_0$  is defined non-parametrically by a mixture of piecewise exponential functions as follows

$$\lambda_0(t) = \log h_0(t) = \sum_{k=1}^{K+1} \lambda_k \mathbb{1}\{t \in (s_{k-1}, s_k]\},$$

where  $\lambda_k$  is constant and the time interval between 0 and the largest observed failure time, denoted  $s_K$ , is partitioned into  $K + 1$  disjoint intervals:  $0 < s_1 < \dots < s_{K+1}$ . The baseline hazard function  $h_{03}$  is assumed to be Markov with respect to  $t_{i1}$ ; we will refer to the set of conditional hazard functions in (5)-(7) as the Markov model. Alternatively, we consider modeling  $h_3$  as follows:

$$h_3(t_{i2} | t_{i1}, \gamma_{ji}, x_{i3}) = \gamma_{ji} h_{03}(t_{i2} - t_{i1}) \exp(x_{i3}^\top \beta_3), \quad 0 < t_{i1} < t_{i2}. \quad (8)$$

We will refer to the set of conditional hazard functions in (5), (6) and (8) as the semi-Markov model.

In the Bayesian framework, priors must be specified for the regression parameter,  $\beta$ , the number of intervals,  $K$ , the partition points  $(s_1, \dots, s_{K+1})$ , and the frailty,  $\gamma_{ji}$ , respectively. The following specifications are made

$$\begin{aligned} \pi(\beta) &\propto 1, \\ \lambda | K, \mu_\lambda, \sigma_\lambda^2 &\sim MVN_{K+1}(\mu_\lambda \mathbb{1}, \sigma_\lambda^2 \Sigma_\lambda) \\ K &\sim \text{Poisson}(\alpha), \\ \pi(s | K) &\propto \frac{(2K+1)! \prod_{k=1}^{K+1} (s_k - s_{k-1})}{(s_{K+1})^{(2K+1)}}, \\ \pi(\mu_\lambda) &\propto 1, \\ \sigma_\lambda^{-2} &\sim \text{Gamma}(a, b), \\ \gamma_{ji} | \theta &\sim \text{Gamma}(\theta^{-1}, \theta^{-1}), \\ \theta^{-1} &\sim \text{Gamma}(\psi, \omega). \end{aligned}$$

The prior specification for  $\lambda$  follows a MVN-ICAR (see Supplemental Material to Lee, Haneuse, Schrag and Dominici, 2015). Note that  $K$  and  $s$  jointly form a time-homogeneous Poisson process prior for the partition.

## Hyperparameters

- $\alpha$  : parameter corresponding to the Poisson prior of  $K$
- $a, b$  : shape and rate of Gamma prior for  $\sigma_\lambda^{-2}$
- $\psi$  : the shape of Gamma prior for  $\theta^{-1}$
- $\omega$  : the rate of Gamma prior for  $\theta^{-1}$

## Arguments to specify

### Model-related

- Formula** : a `Formula` object that corresponds to  $h_g$ , for  $g \in \{1, 2, 3\}$ :  $y_1 + \delta_1 | y_2 + \delta_2 \sim x_1 | x_2 | x_3$ .
- data** : an  $(n \times q)$ -dimensional `data.frame`; the  $q$ -columns correspond to  $q$  covariate vectors named in the formula in **Formula** below.
- model** : `c("Markov", "PEM")` for Markov definition of  $h_3$  in (7); `c("semi-Markov", "PEM")` for semi-Markov definition of  $h_3$  in (8).

### Hyperparameters

- PEM.ab1** : a 2-vector of positive hyperparameters  $a_1$  and  $b_1$  which represent the shape and rate of the Gamma prior for  $\sigma_{\lambda,1}^{-2}$ . Example: `PEM.ab1 <- c(0.7, 0.7)`.
- PEM.ab2** : a 2-vector of positive hyperparameters  $a$  and  $b$  of the prior distribution for  $\sigma_{\lambda,2}^{-2}$ .
- PEM.ab3** : a 2-vector of positive hyperparameters  $a$  and  $b$  of the prior distribution for  $\sigma_{\lambda,3}^{-2}$ .
- PEM.alpha1** : hyperparameter  $\alpha$  of the prior distribution for  $K_1$ , which is one less than the number of partition points.
- PEM.alpha2** : hyperparameter  $\alpha$  of the prior distribution for  $K_2$ , which is one less than the number of partition points.
- PEM.alpha3** : hyperparameter  $\alpha$  of the prior distribution for  $K_3$ , which is one less than the number of partition points.
- theta** : a 2-vector of positive hyperparameters  $\psi$  and  $\omega$  for the hyperprior  $\theta$ .

### MCMC Settings

- numReps** : total number of scans
- thin** : extent of thinning, e.g. if `thin=10` retain every  $10^{th}$  sample.
- burninPerc** : the proportion of burn-in (samples to be discarded before analyzing the data).
- mhProp\_theta\_var** : the parameter  $\theta$  is updated using a Metropolis-Hastings random walk step generating proposals from a Gamma distribution with variance `mhProp_theta_var`.

<code>Cg</code>	a 3-vector for the proportion that determines the sum of probabilities choosing the birth and death moves for each of the baseline hazards, $h_{0g}$ , for $g \in \{1, 2, 3\}$ . <sup>3</sup>
<code>delPertg</code>	a 3-vector for the perturbation parameter in the birth updates for all three baseline hazard functions; values must be between 0 and 0.5. <sup>3</sup>
<code>rj.scheme</code>	<code>rj.scheme=1</code> : the birth update will draw the proposal time split from $1 : s_{max}$ ; <code>rj.scheme=2</code> : the birth update will draw the proposal time split from uniquely ordered failure times in the data.
<code>Kg_max</code>	a 3-vector for the number of splits allowed in each iteration of the Metropolis-Hastings-Green algorithm for the three baseline hazard functions.
<code>sg_max</code>	the largest observed failure time, given by <code>sg_max &lt;- c( max(data\$time1[data\$event1==1]), max(data\$time2[data\$event1==0 &amp; data\$event2==1]), max(data\$time2[data\$event1==1 &amp; data\$event2==1]))</code>
<code>time_lambda1</code>	time points at which the $\lambda_1$ is monitored for convergence. Example: <code>time_lambda1 &lt;- seq(1, sg_max[1], 1)</code> . The chains for these monitoring points can be found in <code>lambda.fin</code> in the chains of the <code>BayesID</code> object.
<code>time_lambda2</code>	time points at which the $\lambda_2$ is monitored for convergence. Example: <code>time_lambda2 &lt;- seq(1, sg_max[2], 1)</code> .
<code>time_lambda3</code>	time points at which the $\lambda_3$ is monitored for convergence. Example: <code>time_lambda3 &lt;- seq(1, sg_max[3], 1)</code> .
<b>Starting Values</b>	
<code>startValues</code>	use <code>initiate.startValues_HReg(form, data, model, nChain)</code> which initiates all necessary starting values. Users may set non-null starting values for any of the following: <code>beta1</code> , <code>beta2</code> , <code>beta3</code> , <code>gamma.ji</code> , <code>theta</code> .
<b>Storage</b>	
<code>path</code>	name of the directory where results are stored. Can leave unspecified.
<code>nGam_save</code>	the number of $\gamma$ to be stored.

---

## Implementation

```

data(scrData)
form <- Formula(time1 + event1 | time2 + event2 ~ x1 + x2 + x3 | x1 + x2 | x1 + x2)
##
theta <- c(0.7, 0.7)
PEM.ab1 <- c(0.7, 0.7) # prior parameters for 1/sigma_1^2
PEM.ab2 <- c(0.7, 0.7) # prior parameters for 1/sigma_2^2
PEM.ab3 <- c(0.7, 0.7) # prior parameters for 1/sigma_3^2
PEM.alpha1 <- 10 # prior parameters for K1
PEM.alpha2 <- 10 # prior parameters for K2
PEM.alpha3 <- 10 # prior parameters for K3
hyperParams <- list(theta=theta,
                    PEM=list(PEM.ab1=PEM.ab1, PEM.ab2=PEM.ab2, PEM.ab3=PEM.ab3,
                             PEM.alpha1=PEM.alpha1, PEM.alpha2=PEM.alpha2, PEM.alpha3=PEM.alpha3))
##
numReps <- 2000
thin <- 10
burninPerc <- 0.25
mhProp_theta_var <- 0.05
Cg <- c(0.2, 0.2, 0.2)
delPertg <- c(0.5, 0.5, 0.5)
rj.scheme <- 1
Kg_max <- c(50, 50, 50)
sg_max <- c(max(scrData$time1[scrData$event1 == 1]),
            max(scrData$time2[scrData$event1 == 0 & scrData$event2 == 1]),
            max(scrData$time2[scrData$event1 == 1 & scrData$event2 == 1]))
time_lambda1 <- seq(1, sg_max[1], 1)
time_lambda2 <- seq(1, sg_max[2], 1)
time_lambda3 <- seq(1, sg_max[3], 1)
nGam_save <- 0
mcmc.PEM <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
                storage=list(nGam_save=nGam_save),
                tuning=list(mhProp_theta_var=mhProp_theta_var,
                           Cg=Cg, delPertg=delPertg,
                           rj.scheme=rj.scheme, Kg_max=Kg_max, sg_max=sg_max,
                           time_lambda1=time_lambda1, time_lambda2=time_lambda2,
                           time_lambda3=time_lambda3))
##
myModel <- c("semi-Markov", "PEM")
myPath <- "Output/02-Results-PEM/"
startValues <- initiate.startValues_HReg(form, scrData, model=myModel, nChain=2)
##
fit_PEM <- BayesID_HReg(form, scrData, id=NULL, model=myModel,
                       hyperParams, startValues, mcmc.PEM, path=myPath)
fit_PEM
summ.fit_PEM <- summary(fit_PEM); names(summ.fit_PEM)
summ.fit_PEM
pred_PEM <- predict(fit_PEM)
plot(pred_PEM, plot.est="Haz")
plot(pred_PEM, plot.est="Surv")

```

<sup>3</sup>See Section A in Supplemental Material to Lee et al. (2015)

**Definition of the survival model**

Let  $t_{i1}$ ,  $t_{i2}$  denote time to non-terminal and terminal event from subject  $i = 1, \dots, n$ . The independent semi-competing risks data are assumed to arise from an illness-death model system with transitions that are modeled through the following three hazard functions:

$$\begin{aligned}\log(t_{i1}) &= \mathbf{x}_{i1}^\top \beta_1 + \gamma_i + \epsilon_{i1}, & t_{i1} > 0, \\ \log(t_{i2}) &= \mathbf{x}_{i2}^\top \beta_2 + \gamma_i + \epsilon_{i2}, & t_{i2} > 0, \\ \log(t_{i2} - t_{i1}) &= \mathbf{x}_{i3}^\top \beta_3 + \gamma_i + \epsilon_{i3}, & t_{i2} > t_{i1},\end{aligned}$$

where  $\gamma_i$  is a study participant-specific random effect,  $\mathbf{x}_{ig}$  is a vector of transition-specific covariates,  $\beta_g$  is a corresponding vector of transition-specific regression parameters, and  $\epsilon_{ig}$  is a transition-specific random variable whose distribution determines that of the corresponding transition time,  $g \in \{1, 2, 3\}$ . In the presence of interval censoring, the times-to-event for the  $i^{\text{th}}$  subject satisfy  $c_{ij} \leq t_{i1} < c_{ij+1}$  for some  $j$  and  $c_{ik} \leq t_{i2} < c_{ik+1}$  for some  $k$ . Let  $\{c_{ij}, c_{ij+1}, c_{ik}, c_{ik+1}, L_i, x_{i1}, x_{i2}, x_{i3}\}$  denote independent observations, where  $L_i$  is the left-truncation time.

For the parametric AFT illness-death model, we build on the log-Normal formulation and take the  $\epsilon_{ig}$  to follow independent  $\text{Normal}(\mu_g, \sigma_g^2)$  distributions,  $g=1,2,3$ . In the Bayesian framework, priors must be specified for the unknown parameters. The following specifications are made

$$\begin{aligned}\pi(\beta_g, \mu_g) &\propto 1, \\ \sigma_g^2 &\sim \text{inverse-Gamma}(a_{\sigma_g}, b_{\sigma_g}), \\ \gamma_i | \theta &\sim \text{Normal}(0, \theta), \\ \theta^{-1} &\sim \text{inverse-Gamma}(a_\theta, b_\theta).\end{aligned}$$

**Hyperparameters**

The hyperparameters  $a_{\sigma_g}$  and  $b_{\sigma_g}$  must be specified for the prior of  $\sigma_g^2$ , as well as  $a_\theta$  and  $b_\theta$ , the rate and shape of the inverse-Gamma distributed hyperprior for  $\theta$ .

**Arguments to specify****Model-related**

**Formula** a `Formula` object that corresponds to  $h_g$ , for  $g \in \{1, 2, 3\}$ :  $L|y_{1L} + y_{1U}|y_{2L} + y_{2U} \sim x_1|x_2|x_3$ .  
**data** an  $(n \times q)$ -dimensional data.frame; the  $q$ -columns correspond to  $q$  covariate vectors named in the formula in `Formula`.

**Hyperparameters**

**theta** a 2-vector of positive hyperparameters  $a_\theta$  and  $b_\theta$  for the hyperprior  $\theta$ .  
**LN.ab1** a 2-vector of positive hyperparameters  $a_{\sigma_1}$  and  $b_{\sigma_1}$  which represent the shape and rate of the inverse-Gamma prior for  $\sigma_1^2$ . Example: `LN.ab1 <- c(0.3, 0.3)`.  
**LN.ab2** a 2-vector of positive hyperparameters  $a_{\sigma_2}$  and  $b_{\sigma_2}$  which represent the shape and rate of the inverse-Gamma prior for  $\sigma_2^2$ . Example: `LN.ab2 <- c(0.3, 0.3)`.  
**LN.ab3** a 2-vector of positive hyperparameters  $a_{\sigma_3}$  and  $b_{\sigma_3}$  which represent the shape and rate of the inverse-Gamma prior for  $\sigma_3^2$ . Example: `LN.ab3 <- c(0.3, 0.3)`.

**MCMC Settings**

**numReps** total number of scans  
**thin** extent of thinning, e.g. if `thin=10` retain every 10<sup>th</sup> sample.  
**burninPerc** the proportion of burn-in (samples to be discarded before analyzing the data).  
**betag.prop.var** the parameter  $\beta_g$  is updated using a Metropolis-Hastings random walk step generating proposals from a Normal distribution with variance `betag.prop.var`.  
**gamma.prop.var** the parameter  $\gamma$  is updated using a Metropolis-Hastings random walk step generating proposals from a Normal distribution with variance `gamma.prop.var`.  
**mug.prop.var** the parameter  $\mu_g$  is updated using a Metropolis-Hastings random walk step generating proposals from a Normal distribution with variance `mug.prop.var`.  
**zetag.prop.var** the parameter  $\zeta_g = 1/\sigma_g^2$  is updated using a Metropolis-Hastings random walk step generating proposals from a log-Normal distribution with variance `zetag.prop.var`.

**Starting Values**

**startValues** use `initiate.startValues_AFT(Formula, data, model, nChain)` which initiates all necessary starting values. Users may set non-null starting values for any of the following: `beta1`, `beta2`, `beta3`, `gamma`, `theta`, `y1`, `y2`, `LN.mu`, `LN.sigSq`.

**Storage**

**nGam\_save** the number of  $\gamma$  to be stored  
**nY1\_save** the number of  $\log(t_1)$  to be stored  
**nY2\_save** the number of  $\log(t_2)$  to be stored  
**nY1.NA\_save** the number of  $\mathbb{1}\{t_1 > t_2\}$  to be stored  
**path** name of the directory where results are stored. Can leave unspecified.



## Implementation

```
data(scrData)
scrData$y1L <- scrData$y1U <- scrData[,1]
scrData$y1U[which(scrData[,2] == 0)] <- Inf
scrData$y2L <- scrData$y2U <- scrData[,3]
scrData$y2U[which(scrData[,4] == 0)] <- Inf
scrData$LT <- rep(0, dim(scrData)[1])
form <- Formula(LT | y1L + y1U | y2L + y2U ~ x1 + x2 + x3 | x1 + x2 | x1 + x2)
##
theta.ab <- c(0.5, 0.05)
LN.ab1 <- c(0.3, 0.3)
LN.ab2 <- c(0.3, 0.3)
LN.ab3 <- c(0.3, 0.3)
hyperParams <- list(theta=theta.ab,
LN=list(LN.ab1=LN.ab1, LN.ab2=LN.ab2, LN.ab3=LN.ab3))
##
numReps <- 300
thin <- 3
burninPerc <- 0.5
nGam_save <- 10
nY1_save <- 10
nY2_save <- 10
nY1.NA_save <- 10
betag.prop.var <- c(0.01,0.01,0.01)
mug.prop.var <- c(0.1,0.1,0.1)
zetag.prop.var <- c(0.1,0.1,0.1)
gamma.prop.var <- 0.01
mcmcParams <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
storage=list(nGam_save=nGam_save, nY1_save=nY1_save, nY2_save=nY2_save, nY1.NA_save=nY1.NA_save),
tuning=list(betag.prop.var=betag.prop.var, mug.prop.var=mug.prop.var, zetag.prop.var=zetag.prop.var,
gamma.prop.var=gamma.prop.var))
##
myModel <- "LN"
myPath <- "Output/01-Results-LN/"
startValues <- initiate.startValues_AFT(form, scrData, model=myModel, nChain=2)
##
fit_LN <- BayesID_AFT(form, scrData, model=myModel, hyperParams,
startValues, mcmcParams, path=myPath)
summary(fit_LN)
pred_LN <- predict(fit_LN, time = seq(0, 35, 1), tseq=seq(from=0, to=30, by=5))
plot(pred_LN, plot.est="Haz")
plot(pred_LN, plot.est="Surv")
```

**Definition of the survival model**

Let  $t_{i1}$ ,  $t_{i2}$  denote time to non-terminal and terminal event from subject  $i = 1, \dots, n$ . The independent semi-competing risks data are assumed to arise from an illness-death model system with transitions that are modeled through the following three hazard functions:

$$\begin{aligned}\log(t_{i1}) &= \mathbf{x}_{i1}^\top \beta_1 + \gamma_i + \epsilon_{i1}, & t_{i1} > 0, \\ \log(t_{i2}) &= \mathbf{x}_{i2}^\top \beta_2 + \gamma_i + \epsilon_{i2}, & t_{i2} > 0, \\ \log(t_{i2} - t_{i1}) &= \mathbf{x}_{i3}^\top \beta_3 + \gamma_i + \epsilon_{i3}, & t_{i2} > t_{i1},\end{aligned}$$

where  $\gamma_i$  is a study participant-specific random effect,  $\mathbf{x}_{ig}$  is a vector of transition-specific covariates,  $\beta_g$  is a corresponding vector of transition-specific regression parameters, and  $\epsilon_{ig}$  is a transition-specific random variable whose distribution determines that of the corresponding transition time,  $g \in \{1, 2, 3\}$ . In the presence of interval censoring, the times-to-event for the  $i^{\text{th}}$  subject satisfy  $c_{ij} \leq t_{i1} < c_{ij+1}$  for some  $j$  and  $c_{ik} \leq t_{i2} < c_{ik+1}$  for some  $k$ . Let  $\{c_{ij}, c_{ij+1}, c_{ik}, c_{ik+1}, L_i, x_{i1}, x_{i2}, x_{i3}\}$  denote independent observations, where  $L_i$  is the left-truncation time.

For our semi-parametric AFT illness-death model,  $\epsilon_{ig}$  is assumed to be taken as draws from the independent DPM of normal distributions:

$$\begin{aligned}\epsilon_{ig} | r_i &\sim \text{Normal}(\mu_{r_i}, \sigma_{r_i}^2), \\ (\mu_{gr}, \sigma_{gr}^2) &\sim G_{g0}, \text{ for } r = 1, \dots, M_g, \\ r_i | p_g &\sim \text{Discrete}(r_i | p_{g1}, \dots, p_{gM_g}), \\ p_g &\sim \text{Dirichlet}(\tau_g / M_g, \dots, \tau_g / M_g).\end{aligned}$$

In the Bayesian framework, priors must be specified for the unknown parameters. We take the  $G_{g0}$  as a normal distribution centered at  $\mu_{g0}$  with a variance  $\sigma_{g0}^2$  for  $\mu_{gr}$  and an  $\text{IG}(a_{\sigma_g}, b_{\sigma_g})$  for  $\sigma_{gr}^2$ . For regression parameters  $\{\beta_1, \beta_2, \beta_3\}$ , we adopt non-informative flat priors on the real line. For  $\gamma$ , we assume that each  $\gamma_i$  is an independent random draw from a  $\text{Normal}(0, \theta)$  distribution. In the absence of prior knowledge on the variance component  $\theta$ , we adopt a conjugate inverse-Gamma hyperprior,  $\text{IG}(a_\theta, b_\theta)$ . Finally, we specify a  $\text{Gamma}(a_{\tau_g}, b_{\tau_g})$  hyperprior for the precision parameter  $\tau_g$ .

**Hyperparameters**

$a_\theta, b_\theta$  : the shape and rate of inverse-Gamma prior for  $\theta$   
 $\mu_{g0}, \sigma_{g0}^2, a_{\sigma_g}, b_{\sigma_g}$  : hyperparameters for  $G_{g0}$   
 $a_{\tau_g}, b_{\tau_g}$  : shape and rate of Gamma hyperprior for  $\tau_g$

**Arguments to specify****Model-related**

**Formula** : a `Formula` object that corresponds to  $h_g$ , for  $g \in \{1, 2, 3\}$ :  $L|y_{1L} + y_{1U}|y_{2L} + y_{2U} \sim x_1|x_2|x_3$ .  
**data** : an  $(n \times q)$ -dimensional `data.frame`; the  $q$ -columns correspond to  $q$  covariate vectors named in the formula in **Formula**.

**Hyperparameters**

**theta** : a 2-vector of positive hyperparameters  $a_\theta$  and  $b_\theta$  for the hyperprior  $\theta$ .  
**DPM.mu1** : a hyperparameter  $\mu_{10}$   
**DPM.mu2** : a hyperparameter  $\mu_{20}$   
**DPM.mu3** : a hyperparameter  $\mu_{30}$   
**DPM.sigSq1** : a hyperparameter  $\sigma_{10}^2$   
**DPM.sigSq2** : a hyperparameter  $\sigma_{20}^2$   
**DPM.sigSq3** : a hyperparameter  $\sigma_{30}^2$   
**DPM.ab1** : a 2-vector of positive hyperparameters  $a_{\sigma_1}, b_{\sigma_1}$   
**DPM.ab2** : a 2-vector of positive hyperparameters  $a_{\sigma_2}, b_{\sigma_2}$   
**DPM.ab3** : a 2-vector of positive hyperparameters  $a_{\sigma_3}, b_{\sigma_3}$   
**Tau.ab1** : a 2-vector of positive hyperparameters  $a_{\tau_1}, b_{\tau_1}$   
**Tau.ab2** : a 2-vector of positive hyperparameters  $a_{\tau_2}, b_{\tau_2}$   
**Tau.ab3** : a 2-vector of positive hyperparameters  $a_{\tau_3}, b_{\tau_3}$

**MCMC Settings**

**numReps** : total number of scans  
**thin** : extent of thinning, e.g. if **thin**=10 retain every 10<sup>th</sup> sample.  
**burninPerc** : the proportion of burn-in (samples to be discarded before analyzing the data).  
**betag.prop.var** : the parameter  $\beta_g$  is updated using a Metropolis-Hastings random walk step generating proposals from a Normal distribution with variance **betag.prop.var**.  
**gamma.prop.var** : the parameter  $\gamma$  is updated using a Metropolis-Hastings random walk step generating proposals from a Normal distribution with variance **gamma.prop.var**.  
**mug.prop.var** : the parameter  $\mu_{gr}$  is updated using a Metropolis-Hastings random walk step generating proposals from a Normal distribution with variance **mug.prop.var**.  
**zetag.prop.var** : the parameter  $\zeta_{gr} = 1/\sigma_{gr}^2$  is updated using a Metropolis-Hastings random walk step generating proposals from a log-Normal distribution with variance **zetag.prop.var**.

**Starting Values**

**startValues** : use `initiate.startValues_AFT(Formula, data, model, nChain)` which initiates all necessary starting values. Users may set non-null starting values for any of the following: **beta1**, **beta2**, **beta3**, **gamma**, **theta**, **y1**, **y2**, **DPM.class1**, **DPM.class2**, **DPM.class3**, **DPM.mu1**, **DPM.mu2**, **DPM.mu3**, **DPM.zeta1**, **DPM.zeta2**, **DPM.zeta3**, **DPM.tau**.

**Storage**

**nGam\_save** : the number of  $\gamma$  to be stored  
**nY1\_save** : the number of  $\log(t_1)$  to be stored  
**nY2\_save** : the number of  $\log(t_2)$  to be stored  
**nY1.NA\_save** : the number of  $\mathbb{1}\{t_1 > t_2\}$  to be stored  
**path** : name of the directory where results are stored. Can leave unspecified.

## Implementation

```
data(scrData)
scrData$y1L <- scrData$y1U <- scrData[,1]
scrData$y1U[which(scrData[,2] == 0)] <- Inf
scrData$y2L <- scrData$y2U <- scrData[,3]
scrData$y2U[which(scrData[,4] == 0)] <- Inf
scrData$LT <- rep(0, dim(scrData)[1])
form <- Formula(LT | y1L + y1U | y2L + y2U ~ x1 + x2 + x3 | x1 + x2 | x1 + x2)
##
theta.ab <- c(0.5, 0.05)
##
DPM.mu1 <- log(12)
DPM.mu2 <- log(12)
DPM.mu3 <- log(12)
DPM.sigSq1 <- 100
DPM.sigSq2 <- 100
DPM.sigSq3 <- 100
DPM.ab1 <- c(2, 1)
DPM.ab2 <- c(2, 1)
DPM.ab3 <- c(2, 1)
Tau.ab1 <- c(1.5, 0.0125)
Tau.ab2 <- c(1.5, 0.0125)
Tau.ab3 <- c(1.5, 0.0125)
hyperParams <- list(theta=theta.ab,
DPM=list(DPM.mu1=DPM.mu1, DPM.mu2=DPM.mu2, DPM.mu3=DPM.mu3, DPM.sigSq1=DPM.sigSq1,
DPM.sigSq2=DPM.sigSq2, DPM.sigSq3=DPM.sigSq3, DPM.ab1=DPM.ab1, DPM.ab2=DPM.ab2,
DPM.ab3=DPM.ab3, Tau.ab1=Tau.ab1, Tau.ab2=Tau.ab2, Tau.ab3=Tau.ab3))
##
numReps <- 300
thin <- 3
burninPerc <- 0.5
nGam_save <- 10
nY1_save <- 10
nY2_save <- 10
nY1.NA_save <- 10
betag.prop.var <- c(0.01,0.01,0.01)
mug.prop.var <- c(0.1,0.1,0.1)
zetag.prop.var <- c(0.1,0.1,0.1)
gamma.prop.var <- 0.01
mcmcParams <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
storage=list(nGam_save=nGam_save, nY1_save=nY1_save, nY2_save=nY2_save, nY1.NA_save=nY1.NA_save),
tuning=list(betag.prop.var=betag.prop.var, mug.prop.var=mug.prop.var,
zetag.prop.var=zetag.prop.var, gamma.prop.var=gamma.prop.var))
##
myModel <- "DPM"
myPath <- "Output/02-Results-DPM/"
startValues <- initiate.startValues_AFT(form, scrData, model=myModel, nChain=2)
##
fit_DPM <- BayesID_AFT(form, scrData, model=myModel, hyperParams,
startValues, mcmcParams, path=myPath)
summary(fit_DPM);
pred_DPM <- predict(fit_DPM, time = seq(0, 35, 1), tseq=seq(from=0, to=30, by=5))
plot(pred_DPM, plot.est="Haz")
plot(pred_DPM, plot.est="Surv")
```

### Definition of the survival model

Let  $t_{ji1}$  and  $t_{ji2}$  denote the time to nonterminal event and terminal event from subject  $i = 1, \dots, n_j$  in cluster  $j = 1, \dots, J$ , subject to right censoring at time  $c_{ji}$ . Let  $(y_{ji1}, y_{ji2}, \delta_{ji1}, \delta_{ji2}, x_{ji})$  denote independent observations, where  $y_{ji1} = \min(t_{ji1}, t_{ji2}, c_{ji})$ ,  $\delta_{ji1} = \mathbb{1}\{t_{ji1} \leq \min(t_{ji2}, c_{ji})\}$ ,  $y_{ji2} = \min(t_{ji2}, c_{ji})$ ,  $\delta_{ji2} = \mathbb{1}\{t_{ji2} \leq c_{ji}\}$ , and  $x_{ji}$  is a vector of covariates for individual  $i$ . The independent semi-competing risks data are assumed to arise from an illness-death model system with transitions that are modeled through the following three hazard functions:

$$h_1(t_{ji1} | \gamma_{ji}, x_{ji1}, V_{j1}) = \gamma_{ji} h_{01}(t_{ji1}) \exp(x_{ji1}^\top \beta_1 + V_{j1}), \quad t_{ji1} > 0, \quad (9)$$

$$h_2(t_{ji2} | \gamma_{ji}, x_{ji2}, V_{j2}) = \gamma_{ji} h_{02}(t_{ji2}) \exp(x_{ji2}^\top \beta_2 + V_{j2}), \quad t_{ji2} > 0, \quad (10)$$

$$h_3(t_{ji2} | t_{ji1}, \gamma_{ji}, x_{ji3}, V_{j3}) = \gamma_{ji} h_{03}(t_{ji2}) \exp(x_{ji3}^\top \beta_3 + V_{j3}), \quad t_{ji2} > 0, \quad (11)$$

where  $\gamma_{ji}$  is a subject-specific frailty,  $V_j = (V_{j1}, V_{j2}, V_{j3})$  is a vector of cluster-specific random effects, and  $x_{ji1}$ ,  $x_{ji2}$  and  $x_{ji3}$  which are subsets of  $x_i$  are vectors of covariates. The baseline hazard functions are defined parametrically by Weibull hazards of the form  $h_{0g}(t) = \alpha_g \kappa_g t^{\alpha_g - 1}$ , for  $g \in \{1, 2, 3\}$ . The baseline hazard function  $h_{03}$  is assumed to be Markov with respect to  $t_{ji1}$ ; we will refer to the set of conditional hazard functions in (9)-(11) as the Markov model. Alternatively, we consider modeling  $h_3$  as follows:

$$h_3(t_{ji2} | t_{ji1}, \gamma_{ji}, x_{ji3}, V_{j3}) = \gamma_{ji} h_{03}(t_{ji2} - t_{ji1}) \exp(x_{ji3}^\top \beta_3 + V_{j3}), \quad 0 < t_{ji1} < t_{ji2}. \quad (12)$$

We will refer to the set of conditional hazard functions in (9), (10) and (12) as the semi-Markov model.

In the Bayesian framework, priors must be specified for the regression parameter,  $\beta_g$ , the shape and scale parameters of baseline hazard function,  $\alpha_g$  and  $\kappa_g$ , and the frailty parameter,  $\gamma_{ji}$ , respectively, for  $g \in \{1, 2, 3\}$ . The following specifications are made

$$\begin{aligned} \pi(\beta_g) &\propto 1, \\ \alpha_g &\sim \text{Gamma}(a_g, b_g), \\ \kappa_g &\sim \text{Gamma}(c_g, d_g), \\ \gamma_{ji} | \theta &\sim \text{Gamma}(\theta^{-1}, \theta^{-1}), \\ \theta^{-1} &\sim \text{Gamma}(\psi, \omega). \end{aligned}$$

We provide two possible prior specifications for the cluster-specific random effects below.

$$\begin{array}{ll} V_j \sim \text{MVN}(0, \Sigma_V), & V_j | m_j \sim \text{MVN}(\mu_{m_j}, \Sigma_{m_j}), \\ \Sigma_V \sim \text{Inverse-Wishart}(\Psi_V, \rho_V), & (\mu_m, \Sigma_m) \sim G_0, \text{ for } m = 1, \dots, M, \\ & m_j | p \sim \text{Discrete}(m_j | p_1, \dots, p_M), \\ & p \sim \text{Dirichlet}(\tau/M, \dots, \tau/M), \\ & \tau \sim \text{Gamma}(a_\tau, b_\tau). \end{array}$$

In the first column, the individual specific-random effects are assumed to be *iid*  $\text{MVN}(0, \Sigma_V)$ . In the second column, the cluster-specific random effects are drawn from a mixture of  $M$  multivariate normal distributions each with mean vector and covariance matrix  $(\mu_m, \Sigma_m)$  which are distributed as a multivariate Normal/Inverse-Wishart (NIW), denoted by  $G_0$ ; we refer to this as the Dirichlet process mixture (DPM) prior. The probability density of  $G_0$  is defined by the product

$$f_{\text{NIW}}(\mu, \Sigma | \Psi_0, \rho_0) = f_{\text{MVN}}(\mu | 0, \Sigma) \times f_{\text{Inv-Wish}}(\Sigma | \Psi_0, \rho_0).$$

### Hyperparameters

- $a_g, b_g$  : shape and rate of Gamma prior for  $\alpha_g$  for  $g \in \{1, 2, 3\}$
- $c_g, d_g$  : shape and rate of Gamma prior for  $\kappa_g$  for  $g \in \{1, 2, 3\}$
- $\psi$  : the shape of Gamma prior for  $\theta^{-1}$
- $\omega$  : the rate of Gamma prior for  $\theta^{-1}$
- $\Psi_0, \rho_0$  : shape and scale of Inverse-Wishart component of the prior distribution,  $G_0$ , of  $(\mu_m, \Sigma_m)$  (DPM prior)
- $a_\tau, b_\tau$  : shape and rate of Gamma hyperprior for  $\tau$  (DPM prior)

### Arguments to specify

---

#### Model-related

- Formula** : a `Formula` object that corresponds to  $h_g$ , for  $g \in \{1, 2, 3\}$ :  $y_1 + \delta_1 | y_2 + \delta_2 \sim x_1 | x_2 | x_3$ .
- data** : an  $(n \times q)$ -dimensional data.frame; the  $q$ -columns correspond to  $q$  covariate vectors named in the formula in `Formula`.
- model** : `c("Markov", "Weibull")` for Markov definition of  $h_3$  in (11); `c("semi-Markov", "Weibull")` for semi-Markov definition of  $h_3$  in (12).
- id** : an  $n$ -vector of cluster information where cluster membership corresponds to one of the positive integers  $1, \dots, J$ .

#### Hyperparameters

- WB.ab1** : a 2-vector of positive hyperparameters  $a_1$  and  $b_1$  of the prior distribution for the shape parameter  $\alpha_1$  of the Weibull baseline hazard. Example: `WB.ab1 <- c(0.5, 0.01)`.
- WB.ab2** : a 2-vector of positive hyperparameters  $a_2$  and  $b_2$  of the prior for  $\alpha_2$ .
- WB.ab3** : a 2-vector of positive hyperparameters  $a_3$  and  $b_3$  of the prior for  $\alpha_3$ .
- WB.cd1** : a 2-vector of positive hyperparameters  $c_1$  and  $d_1$  of the prior distribution for the scale parameter  $\kappa_1$  of the Weibull baseline hazard. Example: `WB.cd1 <- c(0.5, 0.05)`.
- WB.cd2** : a 2-vector of positive hyperparameters  $c_2$  and  $d_2$  of the prior for  $\kappa_2$ .
- WB.cd3** : a 2-vector of positive hyperparameters  $c_3$  and  $d_3$  of the prior for  $\kappa_3$ .
- theta** : a 2-vector of positive hyperparameters  $\psi$  and  $\omega$  for the hyperprior  $\theta$ .

#### MVN prior for $V_{ji}$

- Psi\_v** : a positive-definite scale matrix of the Inverse-Wishart prior for the cluster random effects,  $V_{ji}$ . Example: `Psi_v <- diag(1,3)`.
- rho\_v** : the degrees of freedom of the Inverse-Wishart prior for  $V_{ji}$ . Example: `rho_v <- 100`.

<b>DPM prior for <math>V_{ji}</math></b>	
Psi0	a positive-definite scale matrix of the Inverse-Wishart component of $G_0$ . Example: <code>Psi0 &lt;- diag(1,3)</code> .
rho0	the degrees of freedom of the Inverse-Wishart component of $G_0$ . Example: <code>rho0 &lt;- 10</code> .
aTau	a positive-valued hyperparameter corresponding to the shape parameter, $a_\tau$ , of the Gamma prior of $\tau$ .
bTau	a positive-valued hyperparameter corresponding to the rate parameter, $b_\tau$ , of the Gamma prior of $\tau$ .
<b>MCMC Settings</b>	
numReps	total number of scans
thin	extent of thinning, e.g. if <code>thin=10</code> retain every 10 <sup>th</sup> sample.
burninPerc	the proportion of burn-in (samples to be discarded before analyzing the data).
mhProp_theta_var	the parameter $\theta$ is updated using a Metropolis-Hastings random walk step generating proposals from a Gamma distribution with variance <code>mhProp_theta_var</code> .
mhProp_alphag_var	a 3-vector which specifies the variances of the three random walk Metropolis-Hastings Gamma proposal distributions corresponding to $\alpha_1$ , $\alpha_2$ , $\alpha_3$ .
mhProp_Vg_var	a 3-vector which specifies the variances of the three random walk Metropolis-Hastings proposals from normal distributions with the same variance <code>mhProp_Vg_var</code> .
<b>Starting Values</b>	
startValues	use <code>initiate.startValues_HReg(Formula, data, model, id, nChain)</code> which initiates all necessary starting values. Users may set non-null starting values for: <code>beta1</code> , <code>beta2</code> , <code>beta3</code> , <code>theta</code> , <code>WB.alpha</code> , <code>WB.kappa</code> , <code>gamma.ji</code> , <code>V.j1</code> , <code>V.j2</code> , <code>V.j3</code> , <code>MVN.SigmaV</code> , <code>DPM.tau</code> , <code>DPM.class</code> .
<b>Storage</b>	
path	name of the directory where results are stored. Can leave unspecified.
nGam_save	the number of $\gamma$ to be stored.
storeV	a 3-vector of TRUE/FALSE logical constants indicating storage of $V_{ji}$ values for $g = 1, 2, 3$ . Example: <code>storeV &lt;- rep(TRUE, 3)</code> .

---

## Implementation

```

data(scrData)
id=scrData$cluster
form <- Formula(time1 + event1 | time2 + event2 ~ x1 + x2 + x3 | x1 + x2 | x1 + x2)
##
WB.ab1 <- c(0.5, 0.01)
WB.ab2 <- c(0.5, 0.01)
WB.ab3 <- c(0.5, 0.01)
WB.cd1 <- c(0.5, 0.05)
WB.cd2 <- c(0.5, 0.05)
WB.cd3 <- c(0.5, 0.05)
theta <- c(0.7, 0.7) # prior params for 1/theta
Psi_v <- diag(1, 3) # MVN cluster-specific random effects
rho_v <- 100
Psi0 <- diag(1, 3) # DPM cluster-specific random effects
rho0 <- 10
aTau <- 1.5
bTau <- 0.0125
hyperParams.WB.MVN <- list(theta=theta,
  WB=list(WB.ab1=WB.ab1, WB.ab2=WB.ab2, WB.ab3=WB.ab3,
    WB.cd1=WB.cd1, WB.cd2=WB.cd2, WB.cd3=WB.cd3),
  MVN=list(Psi_v=Psi_v, rho_v=rho_v))
hyperParams.WB.DPM <- list(theta=theta,
  WB=list(WB.ab1=WB.ab1, WB.ab2=WB.ab2, WB.ab3=WB.ab3,
    WB.cd1=WB.cd1, WB.cd2=WB.cd2, WB.cd3=WB.cd3),
  DPM=list(Psi0=Psi0, rho0=rho0, aTau=aTau, bTau=bTau))
##
numReps <- 2000
thin <- 10
burninPerc <- 0.25
mhProp_theta_var <- 0.05
mhProp_alphag_var <- c(0.01, 0.01, 0.01)
mhProp_Vg_var <- c(0.05, 0.05, 0.05)
nGam_save <- 0
storeV <- rep(TRUE, 3)
mcmc.WB <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
  storage=list(nGam_save=nGam_save, storeV=storeV),
  tuning=list(mhProp_theta_var=mhProp_theta_var, mhProp_alphag_var=mhProp_alphag_var,
    mhProp_Vg_var =mhProp_Vg_var))
##
Sigma_V <- diag(0.1, 3)
Sigma_V[1,2] <- Sigma_V[2,1] <- -0.05
Sigma_V[1,3] <- Sigma_V[3,1] <- -0.06
Sigma_V[2,3] <- Sigma_V[3,2] <- 0.07
##
myModel <- c("semi-Markov", "Weibull", "MVN")
myPath <- "Output/03-Results-WB_MVN/ "
startValues <- initiate.startValues_HReg(form, scrData, model=myModel, id, nChain=2)
##

```

```
fit_WB_MVN <- BayesID_HReg(form, scrData, id, model=myModel,
                          hyperParams.WB.MVN, startValues, mcmc.WB, path=myPath)

fit_WB_MVN
summ.fit_WB_MVN <- summary(fit_WB_MVN); names(summ.fit_WB_MVN)
summ.fit_WB_MVN
pred_WB_MVN <- predict(fit_WB_MVN, tseq=seq(from=0, to=30, by=5))
plot(pred_WB_MVN, plot.est="Haz")
plot(pred_WB_MVN, plot.est="Surv")
##
myModel <- c("semi-Markov", "Weibull", "DPM")
myPath <- "Output/04-Results-WB_DPM/"
startValues <- initiate.startValues_HReg(form, scrData, model=myModel, id, nChain=2)
fit_WB_DPM <- BayesID_HReg(form, scrData, id, model=myModel,
                          hyperParams.WB.DPM, startValues, mcmc.WB, path=myPath)

fit_WB_DPM
summ.fit_WB_DPM <- summary(fit_WB_DPM); names(summ.fit_WB_DPM)
summ.fit_WB_DPM
pred_WB_DPM <- predict(fit_WB_MVN, tseq=seq(from=0, to=30, by=5))
plot(pred_WB_DPM, plot.est="Haz")
plot(pred_WB_DPM, plot.est="Surv")
```

### Definition of the survival model

Let  $t_{ji1}$  and  $t_{ji2}$  denote the time to nonterminal event and terminal event from subject  $i = 1, \dots, n_j$  in cluster  $j = 1, \dots, J$ , subject to right censoring at time  $c_{ji}$ . Let  $(y_{ji1}, y_{ji2}, \delta_{ji1}, \delta_{ji2}, x_{ji})$  denote independent observations, where  $y_{ji1} = \min(t_{ji1}, t_{ji2}, c_{ji})$ ,  $\delta_{ji1} = \mathbb{1}\{t_{ji1} \leq \min(t_{ji2}, c_{ji})\}$ ,  $y_{ji2} = \min(t_{ji2}, c_{ji})$ ,  $\delta_{ji2} = \mathbb{1}\{t_{ji2} \leq c_{ji}\}$ , and  $x_{ji}$  is a vector of covariates for individual  $i$ . The independent semi-competing risks data are assumed to arise from an illness-death model system with transitions that are modeled through the following three hazard functions:

$$h_1(t_{ji1} | \gamma_{ji}, x_{ji1}, V_{j1}) = \gamma_{ji} h_{01}(t_{ji1}) \exp(x_{ji1}^\top \beta_1 + V_{j1}), \quad t_{ji1} > 0, \quad (13)$$

$$h_2(t_{ji2} | \gamma_{ji}, x_{ji2}, V_{j2}) = \gamma_{ji} h_{02}(t_{ji2}) \exp(x_{ji2}^\top \beta_2 + V_{j2}), \quad t_{ji2} > 0, \quad (14)$$

$$h_3(t_{ji2} | t_{ji1}, \gamma_{ji}, x_{ji3}, V_{j3}) = \gamma_{ji} h_{03}(t_{ji2}) \exp(x_{ji3}^\top \beta_3 + V_{j3}), \quad t_{ji2} > 0, \quad (15)$$

where  $\gamma_{ji}$  is a subject-specific frailty,  $V_j = (V_{j1}, V_{j2}, V_{j3})$  is a vector of cluster-specific random effects, and  $x_{ji1}$ ,  $x_{ji2}$  and  $x_{ji3}$  which are subsets of  $x_i$  are vectors of covariates. The baseline hazard  $h_0$  is defined non-parametrically by a mixture of piecewise exponential functions as follows

$$\lambda_0(t) = \log h_0(t) = \sum_{k=1}^{K+1} \lambda_k \mathbb{1}\{t \in (s_{k-1}, s_k]\},$$

where  $\lambda_k$  is constant and the time interval between 0 and the largest observed failure time, denoted  $s_k$ , is partitioned into  $K + 1$  disjoint intervals:  $0 < s_1 < \dots < s_{K+1}$ . The baseline hazard function  $h_{03}$  is assumed to be Markov with respect to  $t_{i1}$ ; we will refer to the set of conditional hazard functions in (9)-(11) as the Markov model. Alternatively, we consider modeling  $h_3$  as follows:

$$h_3(t_{ji2} | t_{ji1}, \gamma_{ji}, x_{ji3}, V_{j3}) = \gamma_{ji} h_{03}(t_{ji2} - t_{ji1}) \exp(x_{ji3}^\top \beta_3 + V_{j3}), \quad 0 < t_{ji1} < t_{ji2}. \quad (16)$$

We will refer to the set of conditional hazard functions in (9), (10) and (12) as the semi-Markov model.

In the Bayesian framework, priors must be specified for the regression parameter,  $\beta$ , the number of intervals,  $K$ , the partition points  $(s_1, \dots, s_{K+1})$ , and the frailty,  $\gamma_{ji}$ , respectively. The following specifications are made

$$\begin{aligned} \pi(\beta) &\propto 1, \\ \lambda | K, \mu_\lambda, \sigma_\lambda^2 &\sim MVN_{K+1}(\mu_\lambda \mathbb{1}, \sigma_\lambda^2 \Sigma_\lambda) \\ K &\sim \text{Poisson}(\alpha), \\ \pi(s | K) &\propto \frac{(2K+1)! \prod_{k=1}^{K+1} (s_k - s_{k-1})}{(s_{K+1})^{(2K+1)}}, \\ \pi(\mu_\lambda) &\propto 1, \\ \sigma_\lambda^{-2} &\sim \text{Gamma}(a, b), \\ \gamma_{ji} | \theta &\sim \text{Gamma}(\theta^{-1}, \theta^{-1}), \\ \theta^{-1} &\sim \text{Gamma}(\psi, \omega). \end{aligned}$$

The prior specification for  $\lambda$  follows a MVN-ICAR (see Supplemental Material to Lee, Haneuse, Schrag and Dominici, 2015). Note that  $K$  and  $s$  jointly form a time-homogeneous Poisson process prior for the partition.

We provide two possible prior specifications for the cluster-specific random effects below.

$$\begin{array}{ll} V_j \sim MVN(0, \Sigma_V), & V_j | m_j \sim MVN(\mu_{m_j}, \Sigma_{m_j}), \\ \Sigma_V \sim \text{Inverse-Wishart}(\Psi_V, \rho_V). & (\mu_m, \Sigma_m) \sim G_0, \text{ for } m = 1, \dots, M, \\ & m_j | p \sim \text{Discrete}(m_j | p_1, \dots, p_M), \\ & p \sim \text{Dirichlet}(\tau/M, \dots, \tau/M), \\ & \tau \sim \text{Gamma}(a_\tau, b_\tau). \end{array}$$

In the first column, the individual specific-random effects are assumed to be  $iid$   $MVN(0, \Sigma_V)$ . In the second column, the cluster-specific random effects are drawn from a mixture of  $M$  multivariate normal distributions each with mean vector and covariance matrix  $(\mu_m, \Sigma_m)$  which are distributed as a multivariate Normal/Inverse-Wishart (NIW), denoted by  $G_0$ ; we refer to this as the Dirichlet process mixture (DPM) prior. The probability density of  $G_0$  is defined by the product

$$f_{NIW}(\mu, \Sigma | \Psi_0, \rho_0) = f_{MVN}(\mu | 0, \Sigma) \times f_{Inv-Wish}(\Sigma | \Psi_0, \rho_0).$$

### Hyperparameters

- $\alpha$  : parameter corresponding to the Poisson prior of  $K$
- $a, b$  : shape and rate of Gamma prior for  $\sigma_\lambda^{-2}$
- $\psi$  : the shape of Gamma prior for  $\theta^{-1}$
- $\omega$  : the rate of Gamma prior for  $\theta^{-1}$
- $\Psi_0, \rho_0$  : shape and scale of Inverse-Wishart component of the prior distribution,  $G_0$ , of  $(\mu_m, \Sigma_m)$  (DPM prior)
- $a_\tau, b_\tau$  : shape and rate of Gamma hyperprior for  $\tau$  (DPM prior)

### Arguments to specify

#### Model-related

- Formula** : a **Formula** object that corresponds to  $h_g$ , for  $g \in \{1, 2, 3\}$ :  $y_1 + \delta_1 | y_2 + \delta_2 \sim x_1 | x_2 | x_3$ .
- data** : an  $(n \times q)$ -dimensional data.frame; the  $q$ -columns correspond to  $q$  covariate vectors named in the formula in **Formula**.
- model** : c("Markov", "PEM") for Markov definition of  $h_3$  in (11); c("semi-Markov", "PEM") for semi-Markov definition of  $h_3$  in (12).
- id** : an  $n$ -vector of cluster information where cluster membership corresponds to one of the positive integers  $1, \dots, J$ .

## Hyperparameters

PEM.ab1	a 2-vector of positive hyperparameters $a_1$ and $b_1$ which represent the shape and rate of the Gamma prior for $\sigma_{\lambda_1}^{-2}$ . Example: <code>PEM.ab1 &lt;- c(0.7, 0.7)</code> .
PEM.ab2	a 2-vector of positive hyperparameters $a$ and $b$ of the prior distribution for $\sigma_{\lambda_2}^{-2}$ .
PEM.ab3	a 2-vector of positive hyperparameters $a$ and $b$ of the prior distribution for $\sigma_{\lambda_3}^{-2}$ .
PEM.alpha1	hyperparameter $\alpha$ of the prior distribution for $K_1$ , which is one less than the number of partition points.
PEM.alpha2	hyperparameter $\alpha$ of the prior distribution for $K_2$ , which is one less than the number of partition points.
PEM.alpha3	hyperparameter $\alpha$ of the prior distribution for $K_3$ , which is one less than the number of partition points.
theta	a 2-vector of positive hyperparameters $\psi$ and $\omega$ for the hyperprior $\theta$ .
<b>MVN prior for <math>V_{ji}</math></b>	
Psi_v	a positive-definite scale matrix of the Inverse-Wishart prior for the cluster random effects, $V_{ji}$ . Example: <code>Psi_v &lt;- diag(1,3)</code> .
rho_v	the degrees of freedom of the Inverse-Wishart prior for $V_{ji}$ . Example: <code>rho_v &lt;- 100</code> .
<b>DPM prior for <math>V_{ji}</math></b>	
Psi0	a positive-definite scale matrix of the Inverse-Wishart component of $G_0$ . Example: <code>Psi0 &lt;- diag(1,3)</code> .
rho0	the degrees of freedom of the Inverse-Wishart component of $G_0$ . Example: <code>rho0 &lt;- 10</code> .
aTau	a positive-valued hyperparameter corresponding to the shape parameter, $a_\tau$ , of the Gamma prior of $\tau$ .
bTau	a positive-valued hyperparameter corresponding to the rate parameter, $b_\tau$ , of the Gamma prior of $\tau$ .
<b>MCMC Settings</b>	
numReps	total number of scans
thin	extent of thinning, e.g. if <code>thin=10</code> retain every $10^{th}$ sample.
burninPerc	the proportion of burn-in (samples to be discarded before analyzing the data).
mhProp_theta_var	the parameter $\theta$ is updated using a Metropolis-Hastings random walk step generating proposals from a Gamma distribution with variance <code>mhProp_theta_var</code> .
mhProp_Vg_var	3-vector which specifies the variances of the three random walk Metropolis-Hastings proposals from normal distributions with the same variance <code>mhProp_Vg_var</code> .
Cg	a 3-vector for the proportion that determines the sum of probabilities choosing the birth and death moves for each of the baseline hazards, $h_{0g}$ , for $g \in \{1, 2, 3\}$ . <sup>4</sup>
delPertg	a 3-vector for the perturbation parameter in the birth updates for all three baseline hazard functions; values must be between 0 and 0.5. <sup>4</sup>
rj.scheme	<code>rj.scheme=1</code> : the birth update will draw the proposal time split from $1 : s_{max}$ ; <code>rj.scheme=2</code> : the birth update will draw the proposal time split from uniquely ordered failure times in the data.
Kg_max	a 3-vector for the number of splits allowed in each iteration of the Metropolis-Hastings-Green algorithm for the three baseline hazard functions.
sg_max	the largest observed failure time, given by <code>sg_max &lt;- c( max(data\$time1[data\$event1==1]),               max(data\$time2[data\$event1==0 &amp; data\$event2==1]),               max(data\$time2[data\$event1==1 &amp; data\$event2==1]) )</code>
time_lambda1	time points at which the $\lambda_1$ is monitored for convergence. Example: <code>time_lambda1 &lt;- seq(1, sg_max[1], 1)</code> . The chains for these monitoring points can be found in <code>lambda.fin</code> in the chains of the <code>BayesID_HReg</code> object.
time_lambda2	time points at which the $\lambda_2$ is monitored for convergence. Example: <code>time_lambda2 &lt;- seq(1, sg_max[2], 1)</code> .
time_lambda3	time points at which the $\lambda_3$ is monitored for convergence. Example: <code>time_lambda3 &lt;- seq(1, sg_max[3], 1)</code> .
<b>Starting Values</b>	
startValues	use <code>initiate.startValues_HReg(Formula, data, model, id, nChain)</code> which initiates all necessary starting values. Users may set non-null starting values for any of the following: <code>beta1</code> , <code>beta2</code> , <code>beta3</code> , <code>gamma.ji</code> , <code>theta</code> , <code>V.j1</code> , <code>V.j2</code> , <code>V.j3</code> , <code>MVN.SigmaV</code> , <code>DPM.tau</code> , <code>DPM.class</code> .
<b>Storage</b>	
path	name of the directory where results are stored. Can leave unspecified.
nGam_save	the number of $\gamma$ to be stored.
storeV	a 3-vector of TRUE/FALSE logical constants indicating storage of $V_{ji}$ values for $g = 1, 2, 3$ . Example: <code>storeV &lt;- rep(TRUE, 3)</code> .

## Implementation

```
data(scrData)
id=scrData$cluster
form <- Formula(time1 + event1 | time2 + event2 ~ x1 + x2 + x3 | x1 + x2 | x1 + x2)
##
theta <- c(0.7, 0.7)
PEM.ab1 <- c(0.7, 0.7) # prior parameters for 1/sigma_1^2
PEM.ab2 <- c(0.7, 0.7) # prior parameters for 1/sigma_2^2
PEM.ab3 <- c(0.7, 0.7) # prior parameters for 1/sigma_3^2
PEM.alpha1 <- 10 # prior parameters for K1
PEM.alpha2 <- 10 # prior parameters for K2
PEM.alpha3 <- 10 # prior parameters for K3
Psi_v <- diag(1, 3) # MVN cluster-specific random effects
rho_v <- 100
Psi0 <- diag(1, 3) # DPM cluster-specific random effects
rho0 <- 10
aTau <- 1.5
bTau <- 0.0125
```

<sup>4</sup>See Section A in Supplemental Material to Lee et al. (2015)



```

hyperParams.PEM.MVN <- list(theta=theta,
  PEM=list(PEM.ab1=PEM.ab1, PEM.ab2=PEM.ab2, PEM.ab3=PEM.ab3,
    PEM.alpha1=PEM.alpha1, PEM.alpha2=PEM.alpha2, PEM.alpha3=PEM.alpha3),
  MVN=list(Psi_v=Psi_v, rho_v=rho_v))
hyperParams.PEM.DPM <- list(theta=theta,
  PEM=list(PEM.ab1=PEM.ab1, PEM.ab2=PEM.ab2, PEM.ab3=PEM.ab3,
    PEM.alpha1=PEM.alpha1, PEM.alpha2=PEM.alpha2, PEM.alpha3=PEM.alpha3),
  DPM=list(Psi0=Psi0, rho0=rho0, aTau=aTau, bTau=bTau))

##
numReps <- 2000
thin <- 10
burninPerc <- 0.25
mhProp_theta_var <- 0.05
mhProp_Vg_var <- c(0.05, 0.05, 0.05)
Cg <- c(0.2, 0.2, 0.2)
delPertg <- c(0.5, 0.5, 0.5)
rj.scheme <- 1
Kg_max <- c(50, 50, 50)
sg_max <- c(max(scrData$time1[scrData$event1 == 1]),
  max(scrData$time2[scrData$event1 == 0 & scrData$event2 == 1]),
  max(scrData$time2[scrData$event1 == 1 & scrData$event2 == 1]))
time_lambda1 <- seq(1, sg_max[1], 1)
time_lambda2 <- seq(1, sg_max[2], 1)
time_lambda3 <- seq(1, sg_max[3], 1)
nGam_save <- 0
storeV <- rep(TRUE, 3)
mcmc.PEM <- list(run=list(numReps=numReps, thin=thin, burninPerc=burninPerc),
  storage=list(nGam_save=nGam_save, storeV=storeV),
  tuning=list(mhProp_theta_var=mhProp_theta_var, mhProp_Vg_var=mhProp_Vg_var,
    Cg=Cg, delPertg=delPertg,
    rj.scheme=rj.scheme, Kg_max=Kg_max, sg_max=sg_max,
    time_lambda1=time_lambda1, time_lambda2=time_lambda2,
    time_lambda3=time_lambda3))

##
Sigma_V <- diag(0.1, 3)
Sigma_V[1,2] <- Sigma_V[2,1] <- -0.05
Sigma_V[1,3] <- Sigma_V[3,1] <- -0.06
Sigma_V[2,3] <- Sigma_V[3,2] <- 0.07
##
myModel <- c("semi-Markov", "PEM", "MVN")
myPath <- "Output/05-Results-PEM_MVN/"
startValues <- initiate.startValues_HReg(form, scrData, model=myModel, id, nChain=2)
##
fit_PEM_MVN <- BayesID_HReg(form, scrData, id, model=myModel,
  hyperParams.PEM.MVN, startValues, mcmc.PEM, path=myPath)

fit_PEM_MVN
summ.fit_PEM_MVN <- summary(fit_PEM_MVN); names(summ.fit_PEM_MVN)
summ.fit_PEM_MVN
pred_PEM_MVN <- predict(fit_PEM_MVN)
plot(pred_PEM_MVN, plot.est="Haz")
plot(pred_PEM_MVN, plot.est="Surv")
##
myModel <- c("semi-Markov", "PEM", "DPM")
myPath <- "Output/06-Results-PEM_DPM/"
startValues <- initiate.startValues_HReg(form, scrData, model=myModel, id, nChain=2)
##
fit_PEM_DPM <- BayesID_HReg(form, scrData, id, model=myModel,
  hyperParams.PEM.DPM, startValues, mcmc.PEM, path=myPath)

fit_PEM_DPM
summ.fit_PEM_DPM <- summary(fit_PEM_DPM); names(summ.fit_PEM_DPM)
summ.fit_PEM_DPM
pred_PEM_DPM <- predict(fit_PEM_DPM)
plot(pred_PEM_DPM, plot.est="Haz")
plot(pred_PEM_DPM, plot.est="Surv")

```