

Package ‘BayesSurvive’

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Title Bayesian Survival Models for High-Dimensional Data

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Description An implementation of Bayesian survival models with graph-structured selection priors for sparse identification of omics features predictive of survival (Madjar et al., 2021 <[doi:10.1186/s12859-021-04483-z](https://doi.org/10.1186/s12859-021-04483-z)>) and its extension to use a fixed graph via a Markov Random Field (MRF) prior for capturing known structure of omics features, e.g. disease-specific pathways from the Kyoto Encyclopedia of Genes and Genomes database.

URL <https://github.com/ocbe-uio/BayesSurvive>

BugReports <https://github.com/ocbe-uio/BayesSurvive/issues>

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BayesSurvive	<i>Fit Bayesian Cox Models</i>
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Description

This is the main function to fit a Bayesian Cox model with graph-structured selection priors for sparse identification of high-dimensional covariates.

Usage

```
BayesSurvive(
  survObj,
  model.type = "Pooled",
  MRF2b = FALSE,
  MRF.G = TRUE,
  g.ini = 0,
  hyperpar = NULL,
  initial = NULL,
  nIter = 1,
  burnin = 0,
  thin = 1,
  output_graph_para = FALSE,
  verbose = TRUE
)
```

Arguments

survObj	a list containing observed data from n subjects with components t , d_i , X . For graphical learning of the Markov random field prior, <code>survObj</code> should be a list of the list with survival and covariates data. For subgroup models with or without graphical learning, <code>survObj</code> should be a list of multiple lists with each component list representing each subgroup's survival and covariates data
---------	--

<code>model.type</code>	a method option from <code>c("Pooled", "CoxBVSSL", "Sub-struct")</code> . To enable graphical learning for "Pooled" model, please specify <code>list(survObj)</code> where <code>survObj</code> is the list of <code>t</code> , <code>di</code> and <code>X</code>
<code>MRF2b</code>	logical value. <code>MRF2b = TRUE</code> means two different hyperparameters <code>b</code> in MRF prior (values <code>b01</code> and <code>b02</code>) and <code>MRF2b = FALSE</code> means one hyperparameter <code>b</code> in MRF prior
<code>MRF.G</code>	logical value. <code>MRF.G = TRUE</code> is to fix the MRF graph which is provided in the argument <code>hyperpar</code> , and <code>MRF.G = FALSE</code> is to use graphical model for learning the MRF graph
<code>g.ini</code>	initial values for latent edge inclusion indicators in graph, should be a value in <code>[0,1]</code> . 0 or 1: set all random edges to 0 or 1; value in <code>(0,1)</code> : rate of indicators randomly set to 1, the remaining indicators are 0
<code>hyperpar</code>	a list containing prior parameter values
<code>initial</code>	a list containing prior parameters' initial values
<code>nIter</code>	the number of iterations of the chain
<code>burnin</code>	number of iterations to discard at the start of the chain. Default is 0
<code>thin</code>	thinning MCMC intermediate results to be stored
<code>output_graph_para</code>	allow (TRUE) or suppress (FALSE) the output for parameters 'G', 'V', 'C' and 'Sig' in the graphical model if <code>MRF.G = FALSE</code>
<code>verbose</code>	logical value to display the progress of MCMC

Value

An object of class `BayesSurvive` is saved as `obj_BayesSurvive.rda` in the output file, including the following components:

- `input` - a list of all input parameters by the user
- `output` - a list of the all output estimates:
 - `"gamma.p"` - a matrix with MCMC intermediate estimates of the indicator variables of regression coefficients.
 - `"beta.p"` - a matrix with MCMC intermediate estimates of the regression coefficients.
 - `"h.p"` - a matrix with MCMC intermediate estimates of the increments in the cumulative baseline hazard in each interval.
- `call` - the matched call.

Examples

```
library("BayesSurvive")
set.seed(123)

# Load the example dataset
data("simData", package = "BayesSurvive")

dataset <- list(
  "X" = simData[[1]]$X,
```

```

    "t" = simData[[1]]$time,
    "di" = simData[[1]]$status
  )

# Initial value: null model without covariates
initial <- list("gamma.ini" = rep(0, ncol(dataset$X)))
# Hyperparameters
hyperparPooled <- list(
  "c0" = 2, # prior of baseline hazard
  "tau" = 0.0375, # sd (spike) for coefficient prior
  "cb" = 20, # sd (spike) for coefficient prior
  "pi.ga" = 0.02, # prior variable selection probability for standard Cox models
  "a" = -4, # hyperparameter in MRF prior
  "b" = 0.1, # hyperparameter in MRF prior
  "G" = simData$G # hyperparameter in MRF prior
)

# run Bayesian Cox with graph-structured priors
fit <- BayesSurvive(
  survObj = dataset, hyperpar = hyperparPooled,
  initial = initial, nIter = 100
)

# show posterior mean of coefficients and 95% credible intervals
library("GGally")
plot(fit) +
  coord_flip() +
  theme(axis.text.x = element_text(angle = 90, size = 7))

```

coef.BayesSurvive

Create a dataframe of estimated coefficients

Description

Estimate regression coefficients with posterior mean/median, credible intervals, standard deviation, or MPM estimates, posterior gammas

Usage

```

## S3 method for class 'BayesSurvive'
coef(
  object,
  MPM = FALSE,
  type = "mean",
  CI = 95,
  SD = FALSE,

```

```

    subgroup = 1,
    ...
  )

```

Arguments

object	an object of class BayesSurvive
MPM	logical value to obtain MPM coefficients. Default: FALSE
type	type of point estimates of regression coefficients. One of c("mean", "median"). Default is mean
CI	size (level, as a percentage) of the credible interval to report. Default: 95, i.e. a 95% credible interval
SD	logical value to show each coefficient's standard deviation over MCMC iterations
subgroup	index of the subgroup for visualizing posterior coefficients
...	other arguments

Value

dataframe object

Examples

```

library("BayesSurvive")
set.seed(123)

# Load the example dataset
data("simData", package = "BayesSurvive")

dataset <- list(
  "X" = simData[[1]]$X,
  "t" = simData[[1]]$time,
  "di" = simData[[1]]$status
)

# Initial value: null model without covariates
initial <- list("gamma.ini" = rep(0, ncol(dataset$X)))
# Hyperparameters
hyperparPooled <- list(
  "c0" = 2, # prior of baseline hazard
  "tau" = 0.0375, # sd for coefficient prior
  "cb" = 20, # sd for coefficient prior
  "pi.ga" = 0.02, # prior variable selection probability for standard Cox models
  "a" = -4, # hyperparameter in MRF prior
  "b" = 0.1, # hyperparameter in MRF prior
  "G" = simData$G # hyperparameter in MRF prior
)

# run Bayesian Cox with graph-structured priors

```

```

fit <- BayesSurvive(
  survObj = dataset, hyperpar = hyperparPooled,
  initial = initial, nIter = 100
)

# show posterior coefficients
betas <- coef(fit)
head(betas)

```

func_MCMC

Function to run MCMC sampling

Description

This an internal function for MCMC sampling

Usage

```

func_MCMC(
  survObj,
  hyperpar,
  initial,
  nIter,
  thin,
  burnin,
  S,
  method,
  MRF_2b,
  MRF_G,
  output_graph_para,
  verbose
)

```

Arguments

survObj	a list containing observed data from n subjects; t, di, X. See details for more information
hyperpar	a list containing prior parameter values
initial	a list containing prior parameters' initial values
nIter	the number of iterations of the chain
thin	thinning MCMC intermediate results to be stored
burnin	number of iterations to discard at the start of the chain. Default is 0
S	the number of subgroups
method	a method option from c("Pooled", "CoxBVSSL", "Sub-struct")

MRF_2b	two different b in MRF prior for subgraphs G _{ss} and G _{rs}
MRF_G	logical value. MRF_G = TRUE is to fix the MRF graph which is provided in the argument hyperpar, and MRF_G = FALSE is to use graphical model for learning the MRF graph
output_graph_para	allow (TRUE) or suppress (FALSE) the output for parameters 'G', 'V', 'C' and 'Sig' in the graphical model if MRF_G = FALSE
verbose	logical value to display the progress of MCMC

Value

A list object saving the MCMC results with components including 'gamma.p', 'beta.p', 'h.p', 'gamma.margin', 'beta.margin', 's', 'eta0', 'kappa0', 'c0', 'pi.ga', 'tau', 'cb', 'accept.RW', 'log.jpost', 'log.like', 'post.gamma'

func_MCMC_graph	<i>Function to learn MRF graph</i>
-----------------	------------------------------------

Description

This an internal function for MCMC sampling

Usage

```
func_MCMC_graph(sobj, hyperpar, ini, S, method, MRF_2b)
```

Arguments

sobj	a list containing observed data from n subjects; t, di, X. See details for more information
hyperpar	a list containing prior parameter values
ini	a list containing prior parameters' ini values
S	the number of subgroups
method	a method option from c("Pooled", "CoxBVSSL", "Sub-struct")
MRF_2b	two different b in MRF prior for subgraphs G _{ss} and G _{rs}

Value

A list object with components "Sig" the updated covariance matrices, "G.ini" the updated graph, "V.ini" the updated variances for precision matrices in all subgroups, "C.ini" the updated precision matrices omega for each subgroup

plot.BayesSurvive *Create a plot of estimated coefficients*

Description

Plot point estimates of regression coefficients and 95% credible intervals

Usage

```
## S3 method for class 'BayesSurvive'
plot(x, type = "mean", interval = TRUE, subgroup = 1, ...)
```

Arguments

x	an object of class BayesSurvive or a matrix. If x is a matrix, use BayesSurvive:::plot.BayesSurvive
type	type of point estimates of regression coefficients. One of c("mean", "median"). Default is mean
interval	logical argument to show 95% credible intervals. Default is TRUE
subgroup	index of the subgroup for visualizing posterior coefficients
...	additional arguments sent to ggplot2::geom_point()

Value

ggplot object

Examples

```
library("BayesSurvive")
set.seed(123)

# Load the example dataset
data("simData", package = "BayesSurvive")

dataset <- list(
  "X" = simData[[1]]$X,
  "t" = simData[[1]]$time,
  "di" = simData[[1]]$status
)

# Initial value: null model without covariates
initial <- list("gamma.ini" = rep(0, ncol(dataset$X)))
# Hyperparameters
hyperparPooled <- list(
  "c0" = 2, # prior of baseline hazard
  "tau" = 0.0375, # sd for coefficient prior
  "cb" = 20, # sd for coefficient prior
  "pi.ga" = 0.02, # prior variable selection probability for standard Cox models
  "a" = -4, # hyperparameter in MRF prior
```



```

    "b"      = 0.1, # hyperparameter in MRF prior
    "G"      = simData$G # hyperparameter in MRF prior
  )

# run Bayesian Cox with graph-structured priors
fit <- BayesSurvive(
  survObj = dataset, hyperpar = hyperparPooled,
  initial = initial, nIter = 100
)

# show posterior mean of coefficients and 95% credible intervals
library("GGally")
plot(fit) +
  coord_flip() +
  theme(axis.text.x = element_text(angle = 90, size = 7))

```

plotBrier

Time-dependent Brier scores

Description

Predict time-dependent Brier scores based on Cox regression models

Usage

```

plotBrier(
  object,
  survObj.new = NULL,
  method = "mean",
  times = NULL,
  subgroup = 1
)

```

Arguments

object	fitted object obtained with BayesSurvive
survObj.new	a list containing observed data from new subjects with components t, di, X
method	option to use the posterior mean ("mean") of coefficients for prediction or Bayesian model averaging ("BMA") for prediction
times	maximum time point to evaluate the prediction
subgroup	index of the subgroup in survObj.new for prediction. Default value is 1

Value

A ggplot2::ggplot object. See ?ggplot2::ggplot for more details of the object.

Examples

```

library("BayesSurvive")
set.seed(123)

# Load the example dataset
data("simData", package = "BayesSurvive")

dataset <- list(
  "X" = simData[[1]]$X,
  "t" = simData[[1]]$time,
  "di" = simData[[1]]$status
)

# Initial value: null model without covariates
initial <- list("gamma.ini" = rep(0, ncol(dataset$X)))
# Hyperparameters
hyperparPooled <- list(
  "c0" = 2, # prior of baseline hazard
  "tau" = 0.0375, # sd for coefficient prior
  "cb" = 20, # sd for coefficient prior
  "pi.ga" = 0.02, # prior variable selection probability for standard Cox models
  "a" = -4, # hyperparameter in MRF prior
  "b" = 0.1, # hyperparameter in MRF prior
  "G" = simData$G # hyperparameter in MRF prior
)

# run Bayesian Cox with graph-structured priors
fit <- BayesSurvive(
  survObj = dataset, hyperpar = hyperparPooled,
  initial = initial, nIter = 100
)
# predict survival probabilities of the train data
plotBrier(fit, survObj.new = dataset)

```

predict.BayesSurvive *Predict survival risk*

Description

Predict survival probability, (cumulative) hazard or (integrated) Brier scores based on Cox regression models

Usage

```

## S3 method for class 'BayesSurvive'
predict(

```

```

    object,
    survObj.new,
    type = "brier",
    method = "mean",
    times = NULL,
    subgroup = 1,
    verbose = TRUE,
    ...
  )

```

Arguments

object	fitted object obtained with BayesSurvive
survObj.new	a list containing observed data from new subjects with components t, di, x. If type is among c("hazard", "cumhazard", "survival"), only survObj.new\$X is needed.
type	option to chose for predicting brier scores with type="brier" or one of type=c("brier", "hazard", "cumhazard", "survival")
method	option to use the posterior mean ("mean") of coefficients for prediction or Bayesian model averaging ("BMA") for prediction
times	time points at which to evaluate the risks. If NULL (default), the event/censoring times are used. If type="brier", the largest one of the times is used
subgroup	index of the subgroup in survObj.new for prediction. Default value is 1
verbose	logical value to print IBS of the NULL model and the Bayesian Cox model
...	not used

Value

A list object including seven components with the first component as the specified argument type. The other components of the list are "se", "band", "type", "diag", "baseline" and "times", see function `riskRegression::predictCox` for details

Examples

```

library("BayesSurvive")
set.seed(123)

# Load the example dataset
data("simData", package = "BayesSurvive")

dataset <- list(
  "X" = simData[[1]]$X,
  "t" = simData[[1]]$time,
  "di" = simData[[1]]$status
)

# Initial value: null model without covariates
initial <- list("gamma.ini" = rep(0, ncol(dataset$X)))

```

```

# Hyperparameters
hyperparPooled <- list(
  "c0" = 2, # prior of baseline hazard
  "tau" = 0.0375, # sd for coefficient prior
  "cb" = 20, # sd for coefficient prior
  "pi.ga" = 0.02, # prior variable selection probability for standard Cox models
  "a" = -4, # hyperparameter in MRF prior
  "b" = 0.1, # hyperparameter in MRF prior
  "G" = simData$G # hyperparameter in MRF prior
)

# run Bayesian Cox with graph-structured priors
fit <- BayesSurvive(
  survObj = dataset, hyperpar = hyperparPooled,
  initial = initial, nIter = 100
)
# predict survival probabilities of the train data
predict(fit, survObj.new = dataset)

```

simData	<i>Simulated survival data</i>
---------	--------------------------------

Description

Simulated data set for a quick test. The data set is a list with six components: covariates "X", survival times "time", event status "status". The R code for generating the simulated data is given in the Examples.

Usage

```
simData
```

Format

An object of class list of length 3.

UpdateGamma	<i>Subfunctions to update parameters</i>
-------------	--

Description

This contains subfunctions to update parameters gammas, betas, baseline hazard and graph learning parameters

Usage

```
UpdateGamma(sobj, hyperpar, ini, S, method, MRF_G, MRF_2b)
```

Arguments

sobj	a list containing observed data
hyperpar	a list containing prior parameter values
ini	a list containing prior parameters' initial values
S	the number of subgroups
method	a method option from c("Pooled", "CoxBVSSL", "Sub-struct", "Subgroup")
MRF_G	logical value. MRF_G = TRUE is to fix the MRF graph which is provided in the argument hyperpar, and MRF_G = FALSE is to use graphical model for learning the MRF graph
MRF_2b	two different b in MRF prior for subgraphs G _{ss} and G _{rs}

Value

A list object with two components for the latent variable selection indicators gamma with either independent Bernoulli prior

 UpdateRplee11

Update coefficients of Bayesian Cox Models

Description

This an internal function to update coefficients of the Bayesian Cox Lasso Model

Usage

```
UpdateRplee11(sobj, hyperpar, ini, S, method, MRF_G)
```

Arguments

sobj	a list containing observed data
hyperpar	a list containing prior parameter values
ini	a list containing prior parameters' initial values
S	the number of subgroups
method	a method option from c("Pooled", "CoxBVSSL", "Sub-struct")
MRF_G	logical value. MRF_G = TRUE is to fix the MRF graph which is provided in the argument hyperpar, and MRF_G = FALSE is to use graphical model for learning the MRF graph

Value

A list object with component 'beta.ini' for the updated coefficients and component 'acceptlee' for the MCMC acceptance rate

VS *Function to perform variable selection*

Description

Perform variable selection using the 95 neighborhood criterion (SNC), median probability model (MPM) or Bayesian false discovery rate (FDR). Note that the Bayesian FDR only applies for each subgroup if there are subgroups.

Usage

```
VS(x, method = "FDR", threshold = NA, subgroup = 1)
```

Arguments

x	fitted object obtained with BayesSurvive
method	variable selection method to choose from c("CI", "SNC", "MPM", "FDR"). Default is "FDR"
threshold	SNC threshold value (default 0.5) or the Bayesian expected false discovery rate threshold (default 0.05)
subgroup	index of the subgroup for visualizing posterior coefficients

Value

A boolean vector of selected (= TRUE) and rejected (= FALSE) variables

References

Lee KH, Chakraborty S, Sun J (2015). Survival prediction and variable selection with simultaneous shrinkage and grouping priors. *Statistical Analysis and Data Mining*, 8:114-127

Newton MA, Noueiry A, Sarkar D, Ahlquist P (2004). Detecting differential gene expression with a semiparametric hierarchical mixture method. *Biostatistics*, 5(2), 155-76

Examples

```
library("BayesSurvive")
set.seed(123)

# Load the example dataset
data("simData", package = "BayesSurvive")

dataset <- list(
  "X" = simData[[1]]$X,
  "t" = simData[[1]]$time,
  "di" = simData[[1]]$status
)
```

```
# Initial value: null model without covariates
initial <- list("gamma.ini" = rep(0, ncol(dataset$X)))
# Hyperparameters
hyperparPooled <- list(
  "c0" = 2, # prior of baseline hazard
  "tau" = 0.0375, # sd for coefficient prior
  "cb" = 20, # sd for coefficient prior
  "pi.ga" = 0.02, # prior variable selection probability for standard Cox models
  "a" = -4, # hyperparameter in MRF prior
  "b" = 0.1, # hyperparameter in MRF prior
  "G" = simData$G # hyperparameter in MRF prior
)

# run Bayesian Cox with graph-structured priors
fit <- BayesSurvive(
  survObj = dataset, hyperpar = hyperparPooled,
  initial = initial, nIter = 100
)
# show variable selection
VS(fit, method = "FDR")
```

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