# Package 'survextrap'

July 23, 2025

**Title** Bayesian Flexible Parametric Survival Modelling and Extrapolation

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#### **Description**

Survival analysis using a flexible Bayesian model for individual-level right-censored data, optionally combined with aggregate data on counts of survivors in different periods of time. An M-spline is used to describe the hazard function, with a prior on the coefficients that controls overfitting. Proportional hazards or flexible non-proportional hazards models can be used to relate survival to predictors. Additive hazards (relative survival) models, waning treatment effects, and mixture cure models are also supported. Priors can be customised and calibrated to substantive beliefs. Posterior distributions are estimated using 'Stan', and outputs are arranged in a tidy format. See Jackson (2023) <doi:10.1186/s12874-023-02094-1>.

```
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URL https://github.com/chjackson/survextrap,
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BugReports https://github.com/chjackson/survextrap/issues
```

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# **Description**

For an introduction to and overview of the survextrap package, and full documentation, see https://chjackson.github.io/survextrap/.

For details of the methods, see the paper by Jackson (2023).

#### Author(s)

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- Michael Sweeting (ORCID) [contributor]

#### References

Jackson, C. (2023) survextrap: a package for flexible and transparent survival extrapolation. BMC Medical Research Methodology 23:282. doi:10.1186/s12874023020941

Timmins I, Torabi F, Jackson C, Lambert P, Sweeting M J. (2025) Simulation-based assessment of a Bayesian survival model with flexible baseline hazard and time-dependent effects. doi:10.48550/arXiv.2503.21388.

#### See Also

Useful links:

- https://github.com/chjackson/survextrap
- https://chjackson.github.io/survextrap/
- Report bugs at https://github.com/chjackson/survextrap/issues

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cetux

Datasets for evaluation of cetuximab in head and neck cancer

#### **Description**

Datasets for evaluation of cetuximab in head and neck cancer, as previously analysed by Guyot et al. (2017) to demonstrate models for survival extrapolation with Bayesian evidence synthesis.

#### Usage

```
cetux
cetux_seer
cetux_bh
```

#### **Format**

cetux contains synthetic individual-level survival data, generated (using the method of Guyot et al 2012) to be consistent with the Kaplan-Meier estimates of survival published by Bonner et al. (2006). Columns months, years, give survival in months or years since the date of diagnosis of head and neck cancer. d is a numeric vector where 0 indicates censoring, and 1 indicates death at this time. treat is a factor indicating the treatment group (Cetuximab or Control); both groups also received radiotherapy.

cetux\_seer Estimates of conditional survival from registry data, matched to the Bonner trial population by age, gender, cancer site, and date of diagnosis. From the "Surveillance Epidemiology and End Results" (SEER) database. Each line gives counts of r survivors up to stop years, given n people alive at start. haz is the corresponding constant hazard estimate over this period, computed as  $-\log(r/n)$ . There are also 95% interval estimates for the hazard based on the data on one period at a time, derived from Bayesian principles as  $-\log(qbeta(c(0.975, 0.025), r, n-r))$ .

cetux\_bh Mortality rates for the population of the USA, matched by age and sex to the patients from the Bonner trial. 80% are male, and the median age is 57 (range 34 to 83). Hence the ith row is a weighted average of the male and and female mortality rates for age 57 + i - 1.

See Guyot et al. (2017) for more details of each of these.

An object of class data. frame with 424 rows and 4 columns.

An object of class data. frame with 21 rows and 8 columns.

An object of class data. frame with 54 rows and 2 columns.

# References

Guyot, P., Ades, A.E., Beasley, M., Lueza, B., Pignon, J.P. and Welton, N.J., (2017). Extrapolation of survival curves from cancer trials using external information. Medical Decision Making, 37(4), pp.353-366.

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Bonner, J.A., Harari, P.M., Giralt, J., Azarnia, N., Shin, D.M., Cohen, R.B., Jones, C.U., Sur, R., Raben, D., Jassem, J. and Ove, R., (2006) Radiotherapy plus cetuximab for squamous-cell carcinoma of the head and neck. New England Journal of Medicine, 354(6), pp.567-578.

Guyot, P., Ades, A.E., Ouwens, M.J. and Welton, N.J., (2012) Enhanced secondary analysis of survival data: reconstructing the data from published Kaplan-Meier survival curves. BMC Medical Research Methodology, 12, pp.1-13.

colons

Colon cancer survival data

## **Description**

Survival times of 191 patients from a trial of chemotherapy for colon cancer. This is the data provided in the survival package as colon, but this is restricted to outcome of recurrence-free survival, artificially censored at 3 years, and taking a 20% random sample.

#### Usage

colons

#### **Format**

Documented in colon.

#### Source

See colon.

#### References

See colon.

cumhaz

Estimates of cumulative hazard from a survextrap model

#### **Description**

Estimates of the cumulative hazard at particular times, from a survextrap model

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#### Usage

```
cumhaz(
  newdata = NULL,
  t = NULL
  tmax = NULL.
  niter = NULL,
  summ_fns = NULL,
  sample = FALSE,
  newdata0 = NULL,
  wane_period = NULL,
 wane_nt = 10
)
```

#### **Arguments**

х

A fitted model object as returned by survextrap

newdata

Data frame of covariate values to compute the output for. If there are covariates in the model and this is not supplied, the following default is used:

- (a) if the only covariate is one factor variable, then the output is computed for each level of this factor.
- (b) if there are multiple covariates, or any numeric covariates, then the output is computed at the mean of each numeric covariate in the original data, and at the baseline level of each factor covariate.

Note: caution is required about how treatment groups (for example) are stored in your data. If these are coded as numeric (0/1), then if newdata is not specified, only one output will be shown. This relates to the average value of this numeric variable over the data, which doesn't correspond to either of the treatment groups. To avoid this, a treatment group should be stored as a factor.

t

Vector of times at which to compute the estimates.

tmax

Maximum time at which to compute the estimates. If t is supplied, then this is ignored. If t is not supplied, then t is set to a set of 100 equally spaced time points from 0 to tmax. If both tmax and t are not supplied, then tmax is set to the maximum follow up time in the data.

niter

Number of MCMC iterations to use to compute credible intervals. Set to a low value to make this function quicker, at the cost of some approximation error (which may not be important for plotting or model development).

summ\_fns

A list of functions to use to summarise the posterior sample. This is passed to posterior::summarise\_draws. By default this is list(median=median,  $\neg$ quantile(.x, probs=c(0.025, 0.975))). If the list is named, then the names will be used for the columns of the output.

sample

If TRUE then the MCMC samples are returned instead of being summarised as a median and 95% credible intervals.

newdata0

Data frame of covariate values defining the "untreated" group for use in treatment waning models. See Survmspline\_wane.

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wane\_period Vector of two numbers, defining the time period over which the hazard is inter-

polated between the hazard of the "treated" group (taken from newdata) and the hazard of the "untreated" group (taken from newdata0). Optional - if this is not

supplied, then no waning is assumed.

wane\_nt Number of intervals defining the piecewise constant approximation to the hazard

during the waning period.

#### Value

A data frame (tibble) giving posterior summary statistics, or (if sample=TRUE) an array giving samples from the posterior.

curedata

Simulated data for testing mixture cure models

# **Description**

Survival times of 200 fake people. The cure probability is 0.5 for x=0, and the log odds ratio for cure is 0.5, so that the cure probability is about 0.62 for x=1. The uncured population have survival times distributed as a Weibull with shape 1.5 and scale 1.2.

# Usage

curedata

### **Format**

t Survival times, right-censored at 10 years.

status. 1 for an observed death and 0 for censoring.

x. A numeric variable with value of 0 for 100 individuals, and 1 for the other 100.

#### Source

Simulated.

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get\_draws

Posterior draws from a survextrap model

# Description

Return the matrix of draws from the posterior distribution of parameters in a survextrap model, with all chains collapsed.

# Usage

```
get_draws(x)
```

#### **Arguments**

Х

A fitted model object as returned by survextrap

# Value

A matrix of draws in the draws\_matrix format of the **posterior** package.

hazard

Estimates of hazard from a survextrap model

# **Description**

Estimates of the hazard function at particular times, from a survextrap model

# Usage

```
hazard(
    x,
    newdata = NULL,
    t = NULL,
    tmax = NULL,
    niter = NULL,
    summ_fns = NULL,
    sample = FALSE,
    newdata0 = NULL,
    wane_period = NULL,
    wane_nt = 10
)
```

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#### **Arguments**

A fitted model object as returned by survextrap Х

newdata Data frame of covariate values to compute the output for. If there are covariates in the model and this is not supplied, the following default is used:

> (a) if the only covariate is one factor variable, then the output is computed for each level of this factor.

> (b) if there are multiple covariates, or any numeric covariates, then the output is computed at the mean of each numeric covariate in the original data, and at the baseline level of each factor covariate.

> Note: caution is required about how treatment groups (for example) are stored in your data. If these are coded as numeric (0/1), then if newdata is not specified, only one output will be shown. This relates to the average value of this numeric variable over the data, which doesn't correspond to either of the treatment groups. To avoid this, a treatment group should be stored as a factor.

Vector of times at which to compute the estimates.

Maximum time at which to compute the estimates. If t is supplied, then this is ignored. If t is not supplied, then t is set to a set of 100 equally spaced time points from 0 to tmax. If both tmax and t are not supplied, then tmax is set to

the maximum follow up time in the data.

Number of MCMC iterations to use to compute credible intervals. Set to a low value to make this function quicker, at the cost of some approximation error

(which may not be important for plotting or model development).

summ\_fns A list of functions to use to summarise the posterior sample. This is passed

> to posterior::summarise\_draws. By default this is list(median=median,  $\neg$ quantile(.x, probs=c(0.025, 0.975))). If the list is named, then the names

will be used for the columns of the output.

If TRUE then the MCMC samples are returned instead of being summarised as a

median and 95% credible intervals.

Data frame of covariate values defining the "untreated" group for use in treatnewdata0

ment waning models. See Survmspline\_wane.

wane\_period Vector of two numbers, defining the time period over which the hazard is inter-

> polated between the hazard of the "treated" group (taken from newdata) and the hazard of the "untreated" group (taken from newdata0). Optional - if this is not

supplied, then no waning is assumed.

wane\_nt Number of intervals defining the piecewise constant approximation to the hazard

during the waning period.

#### Value

A data frame (tibble) giving posterior summary statistics, or (if sample=TRUE) an array giving samples from the posterior.

t

tmax

niter

sample

10 hazard\_ratio

hazard\_ratio

Hazard ratio against time in a survextrap model

#### **Description**

Compute the hazard ratio at a series of time points, estimated from a survextrap model. Intended for use with non-proportional hazards models (survextrap(...,nonprop=TRUE)). In proportional hazards models (which survextrap fits by default) the hazard ratio is constant with time.

## Usage

```
hazard_ratio(
    x,
    newdata = NULL,
    t = NULL,
    tmax = NULL,
    niter = NULL,
    summ_fns = NULL,
    sample = FALSE
)
```

#### **Arguments**

x A fitted model object as returned by survextrap

newdata A data frame with two rows. The hazard ratio will be defined as hazard(second row) divided by hazard(first row). If the only covariate in the model is a factor with two levels, then newdata defaults to a data frame containing the levels of

this factor, so that the hazard ratio for the second level versus the first level is computed. For any other models, newdata must be supplied explicitly.

Standardisation (with standardise\_to) is not supported. This might be done by hand by using hazard(...,sample=TRUE) to obtain posterior samples for the two standardised hazards separately, then summarising by hand.

t Vector of times at which to compute the estimates.

Maximum time at which to compute the estimates. If t is supplied, then this is

ignored. If t is not supplied, then t is set to a set of 100 equally spaced time points from 0 to tmax. If both tmax and t are not supplied, then tmax is set to

the maximum follow up time in the data.

niter Number of MCMC iterations to use to compute credible intervals. Set to a low

value to make this function quicker, at the cost of some approximation error

(which may not be important for plotting or model development).

summ\_fns A list of functions to use to summarise the posterior sample. This is passed

to posterior::summarise\_draws. By default this is list(median=median,  $\neg$ quantile(.x, probs=c(0.025, 0.975))). If the list is named, then the names

will be used for the columns of the output.

sample If TRUE then the MCMC samples are returned instead of being summarised as a

median and 95% credible intervals.

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#### Value

A data frame (tibble) with each row containing posterior summary statistics for different times.

Or if sample=TRUE, an array with dimensions length(t), niter, and 1, giving the incremental RMST evaluated at different times and MCMC iterations respectively.

hrtime

Hazard ratio between high and low values of the hazard over time

#### **Description**

This is intended as an intuitive single-number measure of how much a hazard function changes over time. The hazard is computed on an equally-spaced fine grid between the boundary knots. The ratio between a "high" and "low" one of these hazard values is computed. For example, if the hazard is constant over time, then this hazard ratio will be 1.

# Usage

```
hrtime(
   x,
   newdata = NULL,
   niter = NULL,
   summ_fns = NULL,
   hq = c(0.1, 0.9),
   sample = FALSE
)
```

# Arguments

newdata

A fitted model object as returned by survextrap

Data frame of covariate values to compute the output for. If there are covariates in the model and this is not supplied, the following default is used:

- (a) if the only covariate is one factor variable, then the output is computed for each level of this factor.
- (b) if there are multiple covariates, or any numeric covariates, then the output is computed at the mean of each numeric covariate in the original data, and at the baseline level of each factor covariate.

Note: caution is required about how treatment groups (for example) are stored in your data. If these are coded as numeric (0/1), then if newdata is not specified, only one output will be shown. This relates to the average value of this numeric variable over the data, which doesn't correspond to either of the treatment groups. To avoid this, a treatment group should be stored as a factor.

niter

Number of MCMC iterations to use to compute credible intervals. Set to a low value to make this function quicker, at the cost of some approximation error (which may not be important for plotting or model development).

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summ\_fns A list of functions to use to summarise the posterior sample. This is passed to posterior::summarise\_draws. By default this is list(median=median, ~quantile(.x, probs=c(0.025, 0.975))). If the list is named, then the names

will be used for the columns of the output.

hq Quantiles which define the "low" and "high" values of a time-varying quan-

tity (hazard in prior\_haz\_sd and the hazard ratio in prior\_hr\_sd). The ratio between the high and low values will be summarised, as a measure of time-dependence. By default, this is c(0.1, 0.9), so that the 10% and 90% quantiles

are used respectively.

sample If TRUE then the MCMC samples are returned instead of being summarised as a

median and 95% credible intervals.

#### Value

A summary of the posterior distribution of this hazard ratio from the fitted model, as a data frame with one row per covariate value requested in newdata, and one column for each posterior summary statistic.

Or if sample=TRUE, an array with dimensions 1, niter, and nrow(newdata), giving the incremental RMST evaluated at different MCMC iterations and covariate values respectively.

irmst

Incremental restricted mean survival time

# **Description**

Compute the difference in the restricted mean survival times between two covariate values (e.g. treatment groups).

#### Usage

```
irmst(
    x,
    t,
    newdata = NULL,
    newdata0 = NULL,
    wane_period = NULL,
    wane_nt = 10,
    niter = NULL,
    summ_fns = NULL,
    sample = FALSE,
    disc_rate = 0,
    method = "gl",
    gl_nodes = 100
)
```

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# Arguments

x	A fitted model object as returned by survextrap
t	Vector of times. The restricted mean survival time up to each one of these times will be computed.
newdata	A data frame with two rows. The result will be the restricted mean for the covariates in the second row, minus the restricted mean for the covariates in the first row. If newdata is omitted for models where the only covariate is a factor with two levels, then this is taken from these levels. Otherwise newdata must be supplied explicitly.
newdata0	Data frame of covariate values defining the "untreated" group for use in treatment waning models. See Survmspline_wane.
wane_period	Vector of two numbers, defining the time period over which the hazard is interpolated between the hazard of the "treated" group (taken from newdata) and the hazard of the "untreated" group (taken from newdata0). Optional - if this is not supplied, then no waning is assumed.
wane_nt	Number of intervals defining the piecewise constant approximation to the hazard during the waning period.
niter	Number of MCMC iterations to use to compute credible intervals. Set to a low value to make this function quicker, at the cost of some approximation error (which may not be important for plotting or model development).
summ_fns	A list of functions to use to summarise the posterior sample. This is passed to posterior::summarise_draws. By default this is list(median=median, ~quantile(.x, probs=c(0.025, 0.975))). If the list is named, then the names will be used for the columns of the output.
sample	If TRUE then an MCMC sample is returned from the posterior of the output, rather than summary statistics.
disc_rate	Discounting rate used to calculate the discounted mean or restricted mean survival time, using an exponential discounting function.
method	Method of numerical integration to obtain the restricted mean survival time from the survival function.
	The default is method="gl", a Gauss-Legendre method with 100 nodes between zero and the maximum of t.
	method="adaptive" uses the base R integrate function, which is much slower, but potentially more robust for badly-behaved survival functions.
gl_nodes	Number of nodes for the Gauss-Legendre method.

# **Details**

The posterior distribution is obtained by calling rmst for each group, obtaining each posterior sample from the "sample" attribute, and taking the difference to get a posterior sample for the difference.

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#### Value

A data frame (tibble) with each row containing posterior summary statistics for a particular time and covariate value.

Or if sample=TRUE, an array with dimensions length(t), niter, nrow(newdata), giving the incremental RMST evaluated at different times, MCMC iterations and covariate values respectively.

mean.survextrap

Mean survival time

## **Description**

Compute the mean survival time from a model fitted with survextrap. Defined as the integral of the fitted survival curve from zero to infinity. This relies on numerical integration, which is done for every parameter in the MCMC sample, so it may be slow.

#### Usage

```
## $3 method for class 'survextrap'
mean(
    X,
    newdata = NULL,
    newdata0 = NULL,
    wane_period = NULL,
    wane_nt = 10,
    disc_rate = 0,
    niter = NULL,
    summ_fns = NULL,
    sample = FALSE,
    ...
)
```

# **Arguments**

X

A fitted model object as returned by survextrap

newdata

Data frame of covariate values to compute the output for. If there are covariates in the model and this is not supplied, the following default is used:

- (a) if the only covariate is one factor variable, then the output is computed for each level of this factor.
- (b) if there are multiple covariates, or any numeric covariates, then the output is computed at the mean of each numeric covariate in the original data, and at the baseline level of each factor covariate.

Note: caution is required about how treatment groups (for example) are stored in your data. If these are coded as numeric (0/1), then if newdata is not specified, only one output will be shown. This relates to the average value of this numeric variable over the data, which doesn't correspond to either of the treatment groups. To avoid this, a treatment group should be stored as a factor.

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newdata0	Data frame of covariate values defining the "untreated" group for use in treatment waning models. See Survmspline_wane.
wane_period	Vector of two numbers, defining the time period over which the hazard is interpolated between the hazard of the "treated" group (taken from newdata) and the hazard of the "untreated" group (taken from newdata0). Optional - if this is not supplied, then no waning is assumed.
wane_nt	Number of intervals defining the piecewise constant approximation to the hazard during the waning period.
disc_rate	Discounting rate used to calculate the discounted mean or restricted mean survival time, using an exponential discounting function.
niter	Number of MCMC iterations to use to compute credible intervals. Set to a low value to make this function quicker, at the cost of some approximation error (which may not be important for plotting or model development).
summ_fns	A list of functions to use to summarise the posterior sample. This is passed to posterior::summarise_draws. By default this is list(median=median, ~quantile(.x, probs=c(0.025, 0.975))). If the list is named, then the names will be used for the columns of the output.
sample	If TRUE then an MCMC sample is returned from the posterior of the output, rather than summary statistics.
	Other options (currently unused).

#### **Details**

Additionally for some models, the integration up to infinity may not converge, giving an error message. This typically occurs if there is a substantial probability of high survival times or zero hazards at later times. The restricted mean survival time can usually be computed in these situations with rmst, but the model should also be investigated to ensure the posterior distributions are realistic, and simplified or supplemented with external data or informative priors if appropriate.

#### Value

A data frame with each row containing posterior summary statistics for a particular covariate value. An attribute "sample" is also returned, containing a matrix of samples from the posterior distribution of the RMST.

msplinemodel_init	Create an M-spline survival model, both structure and parameters.

#### **Description**

mspline\_init is first used to create the M-spline model structure, including knot positions. Parameters including basis coefficients and scale are either supplied or set to a default that defines a constant hazard model.

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#### Usage

```
msplinemodel_init(
  df = 10,
  degree = 3,
  bsmooth = TRUE,
  knots = NULL,
  bknot = 10,
  obstimes = NULL,
  coefs = NULL,
  hscale = 1
)
```

#### **Arguments**

df Desired number of basis terms, or '	"degrees of freedom" in the spline. If knots
--	--

is not supplied, the number of knots is then chosen to satisfy this.

degree Spline polynomial degree. Can only be changed from the default of 3 if bsmooth

is FALSE.

bsmooth If TRUE then the function is constrained to also have zero derivative and second

derivative at the boundary.

knots Vector of knot locations. If not supplied, df has to be specified. One of two

rules is then used to choose the knot locations. If bknot is specified, a set of equally spaced knots between zero and bknot is used. Otherwise if obstimes is

supplied, the knots are chosen as equally spaced quantiles of obstimes.

The number of knots (excluding zero) is df - degree + 1 if bsmooth is TRUE, or

df - degree - 1 otherwise.

bknot Location of the final spline knot.

obstimes Vector of observation times whose quantiles will be used to choose knot loca-

tions

coefs Basis coefficients

hscale Hazard scale parameter

#### **Details**

This function is not for fitting models to data, but for setting up a theoretical M-spline model for illustration.

#### Value

A list defining the M-spline, with any omitted list components set to defaults. See mspline\_init for details. The parameters are included as the coefs and hscale components.

mspline\_basis 17

mspline_basis	Evaluate an M-spline basis matrix at the specified times.	

# Description

Evaluate an M-spline basis matrix at the specified times. Extrapolation beyond the boundary knots is done by assuming that each basis term is constant beyond the boundary.

#### Usage

```
mspline_basis(times, knots, degree = 3, integrate = FALSE, bsmooth = TRUE)
```

#### **Arguments**

times A numeric vector of times at which to evaluate the basis.

knots Spline knots degree Spline degree

integrate If TRUE, then the integrated M-spline (I-spline) basis is returned.

bsmooth If TRUE then the function is constrained to also have zero derivative and second

derivative at the boundary, which improves smoothness at the boundary.

#### **Details**

The lower boundary is fixed to zero, and each basis term is assumed to be zero at times less than zero, since these models are used for hazard functions in survival data.

#### Value

A two-dimensional array. Rows are the times, and columns are the basis terms.

#### References

The splines2 package is used.

```
mspline_constant_coefs
```

Determine M-spline basis coefficients which give a constant function.

## **Description**

This works by obtaining the coefficients of the corresponding B-spline basis, which are equal if the B-spline is a constant function.

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#### Usage

```
mspline_constant_coefs(mspline, logit = FALSE)
```

# Arguments

mspline A list with components knots (vector of knots), degree (polynomial degree)

and bsmooth (logical for smoothness constraint at boundary), defining an M-

spline configuration.

logit If TRUE then the multinomial logit transform of the coefficients is returned. This

is a vector of length one less than the number of coefficients, with the rth element

defined by log(coefs[r+1]/coefs[1]).

#### Value

A numeric vector of the coefficients.

#### References

Ramsay, J. O. (1988). Monotone regression splines in action. Statistical Science, 3(4): 425-441.

mspline\_init

Create a default M-spline model structure

#### **Description**

Create a default M-spline model structure

# Usage

```
mspline_init(
  df = 10,
  degree = 3,
  bsmooth = TRUE,
  knots = NULL,
  bknot = 10,
  obstimes = NULL)
```

#### **Arguments**

df Desired number of basis terms, or "degrees of freedom" in the spline. If knots

is not supplied, the number of knots is then chosen to satisfy this.

degree Spline polynomial degree. Can only be changed from the default of 3 if bsmooth

is FALSE.

bsmooth If TRUE then the function is constrained to also have zero derivative and second

derivative at the boundary.

mspline\_list\_init 19

knots Vector of knot locations. If not supplied, df has to be specified. One of two

rules is then used to choose the knot locations. If bknot is specified, a set of equally spaced knots between zero and bknot is used. Otherwise if obstimes is supplied, the knots are chosen as equally spaced quantiles of obstimes.

The number of knots (excluding zero) is df - degree + 1 if bsmooth is TRUE, or

df - degree - 1 otherwise.

bknot Location of the final spline knot.

obstimes Vector of observation times whose quantiles will be used to choose knot loca-

tions

#### Value

A list with fundamental components knots, degree, and bsmooth, as documented above.

The component df is also included, and derived as a consequence of the fundamental components.

basis\_means gives the "mean" of each basis term (i.e. the mean of a random variable whose probability density function is given by the basis function)

basis\_spans and sqrt\_wt are quantities used in the construction of random walk prior distributions for the basis coefficients (following https://arxiv.org/abs/2401.12640 and https://arxiv.org/abs/2201.06808).

mspline\_list\_init Validate an M-spline object supplied as a list, choosing defaults if needed.

#### **Description**

Validate an M-spline object supplied as a list, choosing defaults if needed.

# Usage

```
mspline_list_init(mspline, obstimes = NULL)
```

#### **Arguments**

mspline A list with any or none of the following components: df, degree, bsmooth,

knots, bknot, as documented in mspline\_init.

obstimes Vector of observation times whose quantiles will be used to choose knot loca-

tions

#### Value

A list defining the M-spline, with any omitted list components set to defaults. See mspline\_init for details.

If mspline\$knots is not supplied, giving knot locations, then either mspline\$bknot or obstimes must be specified, so that default locations can be obtained.

20 mspline\_plotdata

mspline_plotdata	Data for plotting an M-spline function, showing how it is built up from its basis
------------------	---

# Description

Data for plotting an M-spline function, showing how it is built up from its basis

# Usage

```
mspline_plotdata(
  knots = NULL,
  bknot = 10,
  df = 10,
  degree = 3,
  bsmooth = TRUE,
  coefs = NULL,
  scale = 1,
  tmin = 0,
  tmax = 10
)
```

# Arguments

knots	Vector of knot locations. If not supplied, df has to be specified. One of two rules is then used to choose the knot locations. If bknot is specified, a set of equally spaced knots between zero and bknot is used. Otherwise if obstimes is supplied, the knots are chosen as equally spaced quantiles of obstimes.  The number of knots (excluding zero) is df - degree + 1 if bsmooth is TRUE, or df - degree - 1 otherwise.
bknot	Location of the final spline knot.
df	Desired number of basis terms, or "degrees of freedom" in the spline. If knots is not supplied, the number of knots is then chosen to satisfy this.
degree	Spline polynomial degree. Can only be changed from the default of 3 if b smooth is FALSE.
bsmooth	If TRUE then the function is constrained to also have zero derivative and second derivative at the boundary.
coefs	Coefficients of the spline basis terms. These are normalised internally to sum to 1, if they do not already sum to 1.
scale	Scale parameter. After computing the standard M-spline function as a weighted sum of the basis terms, the function is multiplied by scale. The log of the scale is the parameter called alpha in the results of a survextrap model, the intercept of the linear model on the log hazard.
tmin	Minimum plotting time. Defaults to zero.
tmax	Maximum plotting time. Defaults to the highest knot.

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# Value

A data frame with values of the hazard at different times for different basis functions.

mspline\_plotsetup Get basis for an illustration of an M-spline with given knots.

# Description

Get basis for an illustration of an M-spline with given knots.

# Usage

```
mspline_plotsetup(
  knots,
  bknot = 10,
  tmin = NULL,
  tmax = NULL,
  degree = 3,
  df = 10,
  bsmooth = TRUE
)
```

# Arguments

knots	Vector of knot locations. If not supplied, df has to be specified. One of two rules is then used to choose the knot locations. If bknot is specified, a set of equally spaced knots between zero and bknot is used. Otherwise if obstimes is supplied, the knots are chosen as equally spaced quantiles of obstimes.  The number of knots (excluding zero) is df - degree + 1 if bsmooth is TRUE, or df - degree - 1 otherwise.
bknot	Location of the final spline knot.
tmin	Minimum plotting time. Defaults to zero.
tmax	Maximum plotting time. Defaults to the highest knot.
degree	Spline polynomial degree. Can only be changed from the default of 3 if bsmooth is FALSE.
df	Desired number of basis terms, or "degrees of freedom" in the spline. If knots is not supplied, the number of knots is then chosen to satisfy this.
bsmooth	If TRUE then the function is constrained to also have zero derivative and second derivative at the boundary.

# Value

Data frame containing the basis, as returned by mspline\_basis.

22 mspline\_spec

mspline\_spec

Make default M-spline knot specification given a survival dataset.

# **Description**

Choose default M-spline knot locations given a dataset and desired number of spline parameters. Assumes a cubic spline, and knots based on quantiles of event times observed in the individual data.

#### Usage

```
mspline_spec(
  formula,
  data,
  cure = FALSE,
  nonprop = NULL,
  backhaz = NULL,
  backhaz_strata = NULL,
  external = NULL,
  df = 10,
  add_knots = NULL,
  degree = 3,
 bsmooth = TRUE
)
```

#### **Arguments**

formula

A survival formula in standard R formula syntax, with a call to Surv() on the left hand side.

Covariates included on the right hand side of the formula with be modelled with proportional hazards, or if nonprop is TRUE then a non-proportional hazards is used.

If data is omitted, so that the model is being fitted to external aggregate data alone, without individual data, then the formula should not include a Surv() call. The left-hand side of the formula will then be empty, and the right hand side specifies the covariates as usual. For example, formula = ~1 if there are no covariates.

data

Data frame containing variables in formula. Variables should be in a data frame, and not in the working environment.

This may be omitted, in which case external must be supplied. This allows a model to be fitted to external aggregate data alone, without any individual-level data.

cure

If TRUE, a mixture cure model is used, where the "uncured" survival is defined by the M-spline model, and the cure probability is estimated.

nonprop

Non-proportional hazards model specification. This is achieved by modelling the spline basis coefficients in terms of the covariates. See the methods vignette for more details.

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> If TRUE, then all covariates are modelled with non-proportional hazards, using the same model formula as formula.

> If this is a formula, then this is assumed to define a model for the dependence of the basis coefficients on the covariates.

> IF this is NULL or FALSE (the default) then any covariates are modelled with proportional hazards.

backhaz

Background hazard, that is, for causes of death other than the cause of interest. This defines a "relative survival" or "additive hazards" model. The overall hazard that describes the all-cause survival data (given in the data and/or external argument) is then modelled as the sum of a cause-specific hazard and a background hazard.

The background hazard is assumed to be known, and the cause-specific hazard is modelled with the flexible parametric model.

The background hazard can be supplied in two forms. The meaning of predictions from the model depends on which of these is used.

- (a) A data frame with columns "hazard" and "time", specifying the background hazard at all times as a piecewise-constant (step) function. Each row gives the background hazard between the specified time and the next time. The first element of "time" should be 0, and the final row specifies the hazard at all times greater than the last element of "time". Predictions from the model fitted by survextrap will then include this background hazard, because it is known at all times.
- (b) The (quoted) name of a variable in the data giving the background hazard. For censored cases, the exact value does not matter. The predictions from survextrap will then describe the excess hazard or survival on top of this background. The overall hazard cannot be predicted in general, because the background hazard is only specified over a limited range of time.

If there is external data, and backhaz is supplied in form (b), then the user should also supply the background survival at the start and stop points in columns of the external data named "backsurv\_start" and "backsurv\_stop". That is, the probability of survival up to each of these times for someone alive at time 0. This should describe the same reference population as backhaz, though the package does not check for consistency between these.

If there are stratifying variables specified in backhaz\_strata, then there should be multiple rows giving the background hazard value for each time period and stratifying variable.

If backhaz is NULL (the default) then no background hazard component is included in the model.

backhaz\_strata A character vector of names of variables that appear in backhaz that indicate strata, e.g. backhaz\_strata = c("agegroup", "sex"). This allows different background hazard values to be used for different subgroups. These variables must also appear in the datasets being modelled, that is, in data, external or both. Each row of those datasets should then have a corresponding row in backhaz which has the same values of the stratifying variables.

> This is NULL by default, indicating no stratification of the background hazard. If stratification is done, then backhaz must be supplied in form (a), as a data frame rather than a variable in the data.

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external External data as a data frame of aggregate survival counts with columns named:

start: Start time stop: Follow-up time

n: Number of people alive at start

r: Number of those people who are still alive at stop

If there are covariates in formula, then the values they take in the external data must be supplied as additional columns in external. Therefore if there are external data, the covariates in formula and data should not be named

start, stop, n or r.

df Desired number of basis terms, or "degrees of freedom" in the spline. If knots

is not supplied, the number of knots is then chosen to satisfy this.

add\_knots Additional knots, other than those determined from the quantiles of the individ-

ual data. Typically used to add a maximum knot at the time that we want to

extrapolate to.

degree Spline polynomial degree. Can only be changed from the default of 3 if bsmooth

is FALSE.

bsmooth If TRUE then the function is constrained to also have zero derivative and second

derivative at the boundary.

#### **Details**

If there are also external data, then these are based on quantiles of a vector defined by concatenating the event times in the individual data with the unique start and stop times in the external data.

This is designed to have the same arguments as survextrap. It is intended for use when we want to fit a set of survextrap models with the same spline specification.

See also mspline\_list\_init and mspline\_init, which have lower-level interfaces, and are designed for use without data, e.g. when illustrating a theoretical M-spline model.

#### Value

A list with components

knots Knot locations. The number of knots will be equal to df + degree + 2. degree Spline polynomial degree (i.e. 3) nvars Number of basis variables (an alias for df)

plot.survextrap Plot method for survextrap model objects

#### **Description**

Intended as a quick check of the results of a model fit. See plot\_survival and plot\_hazard for the functions behind each panel, and survival and hazard to extract the data being plotted here to enable custom plots like these.

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#### Usage

```
## S3 method for class 'survextrap'
plot(x, type = "hazsurv", newdata = NULL, ...)
```

#### **Arguments**

Х

A fitted model object as returned by survextrap

type

"survival" for a plot of the survival function, "hazard" for the hazard function, against time.

newdata

Data frame of covariate values to compute the output for. If there are covariates in the model and this is not supplied, the following default is used:

- (a) if the only covariate is one factor variable, then the output is computed for each level of this factor.
- (b) if there are multiple covariates, or any numeric covariates, then the output is computed at the mean of each numeric covariate in the original data, and at the baseline level of each factor covariate.

Note: caution is required about how treatment groups (for example) are stored in your data. If these are coded as numeric (0/1), then if newdata is not specified, only one output will be shown. This relates to the average value of this numeric variable over the data, which doesn't correspond to either of the treatment groups. To avoid this, a treatment group should be stored as a factor.

groups: 10 avera and, a a can

Additional arguments, passed on to plot\_hazard and plot\_survival.

#### Value

A ggplot2 plot object.

plot\_hazard

Plot hazard curves from a survextrap model

#### **Description**

Plot hazard curves from a survextrap model. This function is intended as a quick check of a fitted model, so it deliberately has limited options for customisation. The data behind these plots can be extracted with hazard into a tidy data frame to enable custom plots to be constructed with **ggplot2**. See the case study vignette for some examples.

#### Usage

```
plot_hazard(
    x,
    newdata = NULL,
    t = NULL,
    tmax = NULL,
    niter = NULL,
```

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```
newdata0 = NULL,
wane_period = NULL,
wane_nt = 10,
ci = NULL,
xlab = "Time",
ylab = "Hazard",
line_size = 1.5,
ci_alpha = 0.2,
show_knots = FALSE
)
```

#### **Arguments**

Χ

A fitted model object as returned by survextrap

newdata

Data frame of covariate values to compute the output for. If there are covariates in the model and this is not supplied, the following default is used:

(a) if the only covariate is one factor variable, then the output is computed for each level of this factor.

(b) if there are multiple covariates, or any numeric covariates, then the output is computed at the mean of each numeric covariate in the original data, and at the baseline level of each factor covariate.

Note: caution is required about how treatment groups (for example) are stored in your data. If these are coded as numeric (0/1), then if newdata is not specified, only one output will be shown. This relates to the average value of this numeric variable over the data, which doesn't correspond to either of the treatment groups. To avoid this, a treatment group should be stored as a factor.

t

Vector of times at which to compute the estimates.

tmax

Maximum time at which to compute the estimates. If t is supplied, then this is ignored. If t is not supplied, then t is set to a set of 100 equally spaced time points from 0 to tmax. If both tmax and t are not supplied, then tmax is set to the maximum follow up time in the data.

niter

Number of MCMC iterations to use to compute credible intervals. Set to a low value to make this function quicker, at the cost of some approximation error (which may not be important for plotting or model development).

newdata0

Data frame of covariate values defining the "untreated" group for use in treatment waning models. See Survmspline\_wane.

wane\_period

Vector of two numbers, defining the time period over which the hazard is interpolated between the hazard of the "treated" group (taken from newdata) and the hazard of the "untreated" group (taken from newdata0). Optional - if this is not supplied, then no waning is assumed.

wane\_nt

Number of intervals defining the piecewise constant approximation to the hazard during the waning period.

ci

If TRUE then credible intervals are drawn. Defaults to drawing the intervals if the plot shows the curve for only one covariate value.

xlab

X-axis label

plot\_hazard\_ratio 27

ylab Y-axis label

line\_size Passed to geom\_line

ci\_alpha Transparency for the credible interval ribbons

show\_knots Show the locations of the spline knots as vertical lines

#### Details

If the model has a single factor covariate (excluding background hazard strata), then curves are produced for each level of this factor if newdata requests this (or is left to its default). Otherwise, only a single curve is produced, illustrating the corresponding output from hazard.

#### Value

A ggplot2 plot object.

plot\_hazard\_ratio

Plot hazard ratio against time from a survextrap model

#### **Description**

For use with non-proportional hazards models (survextrap(...,nonprop=TRUE)). Intended as a quick check of a model fit, so there are limited customisation options. The underlying data can be extracted with hazard\_ratio.

#### Usage

```
plot_hazard_ratio(
    x,
    newdata = NULL,
    t = NULL,
    tmax = NULL,
    niter = NULL,
    ci = TRUE,
    xlab = "Time",
    ylab = "Hazard ratio",
    line_size = 1.5,
    ci_alpha = 0.2
)
```

# Arguments

Χ

A fitted model object as returned by survextrap

newdata

A data frame with two rows. The hazard ratio will be defined as hazard(second row) divided by hazard(first row). If the only covariate in the model is a factor with two levels, then newdata defaults to a data frame containing the levels of this factor, so that the hazard ratio for the second level versus the first level is computed. For any other models, newdata must be supplied explicitly.

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Standardisation (with standardise\_to) is not supported. This might be done by hand by using hazard(...,sample=TRUE) to obtain posterior samples for the two standardised hazards separately, then summarising by hand.

t Vector of times at which to compute the estimates.

tmax Maximum time at which to compute the estimates. If t is supplied, then this is

ignored. If t is not supplied, then t is set to a set of 100 equally spaced time points from 0 to tmax. If both tmax and t are not supplied, then tmax is set to

the maximum follow up time in the data.

niter Number of MCMC iterations to use to compute credible intervals. Set to a low

value to make this function quicker, at the cost of some approximation error

(which may not be important for plotting or model development).

ci If TRUE then credible intervals are drawn. Defaults to drawing the intervals if the

plot shows the curve for only one covariate value.

xlab X-axis label ylab Y-axis label

line\_size Passed to geom\_line

ci\_alpha Transparency for the credible interval ribbons

#### Value

A ggplot2 plot object.

plot\_mspline

Plot a M-spline function, showing how it is built up from its basis

# Description

See mspline\_plotdata for the data behind the plot

#### Usage

```
plot_mspline(
  knots = NULL,
  bknot = 10,
  df = 10,
  degree = 3,
  bsmooth = TRUE,
  coefs = NULL,
  scale = 1,
  tmin = 0,
  tmax = 10,
  show_knots = FALSE,
  show_means = FALSE
```

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# Arguments

knots	Vector of knot locations. If not supplied, df has to be specified. One of two rules is then used to choose the knot locations. If bknot is specified, a set of equally spaced knots between zero and bknot is used. Otherwise if obstimes is supplied, the knots are chosen as equally spaced quantiles of obstimes.  The number of knots (excluding zero) is df - degree + 1 if bsmooth is TRUE, or df - degree - 1 otherwise.
bknot	Location of the final spline knot.
df	Desired number of basis terms, or "degrees of freedom" in the spline. If knots is not supplied, the number of knots is then chosen to satisfy this.
degree	Spline polynomial degree. Can only be changed from the default of 3 if bsmooth is FALSE.
bsmooth	If TRUE then the function is constrained to also have zero derivative and second derivative at the boundary.
coefs	Coefficients of the spline basis terms. These are normalised internally to sum to 1, if they do not already sum to 1.
scale	Scale parameter. After computing the standard M-spline function as a weighted sum of the basis terms, the function is multiplied by scale. The log of the scale is the parameter called alpha in the results of a survextrap model, the intercept of the linear model on the log hazard.
tmin	Minimum plotting time. Defaults to zero.
tmax	Maximum plotting time. Defaults to the highest knot.
show_knots	Show the positions of the knots, including the upper boundary
show_means	Show the "mean" around which each basis term is centred (defined as the mean of a random variable whose PDF is defined by the basis term).

# Value

A ggplot2 plot object.

plot_survival	Plot survival curves from a survextrap model
· <del>-</del>	, i

# Description

Plot survival curves from a survextrap model. This function is intended as a quick check of a fitted model, so it deliberately has limited options for customisation. The data behind these plots can be extracted with survival into a tidy data frame to enable custom plots to be constructed with **ggplot2**. See the case study vignette for some examples.

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#### Usage

```
plot_survival(
  Х,
  newdata = NULL,
  t = NULL
  tmax = NULL,
  km = NULL.
  niter = NULL,
  newdata0 = NULL,
 wane_period = NULL,
 wane_nt = 10,
  ci = NULL,
  xlab = "Time",
  ylab = "Survival",
  line\_size = 1.5,
  ci_alpha = 0.2,
  show_knots = FALSE
)
```

#### **Arguments**

x A fitted model object as returned by survextrap

newdata

Data frame of covariate values to compute the output for. If there are covariates in the model and this is not supplied, the following default is used:

- (a) if the only covariate is one factor variable, then the output is computed for each level of this factor.
- (b) if there are multiple covariates, or any numeric covariates, then the output is computed at the mean of each numeric covariate in the original data, and at the baseline level of each factor covariate.

Note: caution is required about how treatment groups (for example) are stored in your data. If these are coded as numeric (0/1), then if newdata is not specified, only one output will be shown. This relates to the average value of this numeric variable over the data, which doesn't correspond to either of the treatment groups. To avoid this, a treatment group should be stored as a factor.

t

Vector of times at which to compute the estimates.

tmax

Maximum time at which to compute the estimates. If t is supplied, then this is ignored. If t is not supplied, then t is set to a set of 100 equally spaced time points from 0 to tmax. If both tmax and t are not supplied, then tmax is set to the maximum follow up time in the data.

km

If TRUE then a Kaplan-Meier curve of the observed data is plotted, using the results of survival::survfit() on the formula originally used for the survextrap fit. By default, this is only done when there are no covariates or one factor covariate.

The Kaplan-Meier estimates are returned in the km component of the fitted model object returned by survextrap, for use in hand-crafted plots like these.

print.survextrap 31

Number of MCMC iterations to use to compute credible intervals. Set to a low niter value to make this function quicker, at the cost of some approximation error (which may not be important for plotting or model development). Data frame of covariate values defining the "untreated" group for use in treatnewdata0 ment waning models. See Survmspline\_wane. wane\_period Vector of two numbers, defining the time period over which the hazard is interpolated between the hazard of the "treated" group (taken from newdata) and the hazard of the "untreated" group (taken from newdata0). Optional - if this is not supplied, then no waning is assumed. wane\_nt Number of intervals defining the piecewise constant approximation to the hazard during the waning period. If TRUE then credible intervals are drawn. Defaults to drawing the intervals if the ci plot shows the curve for only one covariate value. xlab X-axis label Y-axis label ylab Passed to geom\_line line\_size ci\_alpha Transparency for the credible interval ribbons

#### Details

show\_knots

If the model has a single factor covariate (excluding background hazard strata), then curves are produced for each level of this factor if newdata requests this (or is left to its default). Otherwise, only a single curve is produced, illustrating the corresponding output from hazard.

Show the locations of the spline knots as vertical lines

# Value

A ggplot2 plot object.

print.survextrap Print a fitted survextrap model

#### **Description**

Print a fitted survextrap model

#### Usage

```
## S3 method for class 'survextrap' print(x, ...)
```

# Arguments

x A fitted model object as returned by survextrap

... Other arguments (currently unused).

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# Value

No value is returned.

This prints a summary of the data, a statement of the fitted model, the prior distributions, and a table of summary statistics of the posterior distributions of the model parameters. For descriptions of the parameters in this summary table, see <a href="mailto:summary.survextrap">survextrap</a>.

print\_priors

Print the priors used in a fitted survextrap model

# **Description**

Print the priors used in a fitted survextrap model

## Usage

```
print_priors(x)
```

# **Arguments**

Х

A fitted model object as returned by survextrap

# Value

No explicit value is returned.

priors

Prior distributions and options

# **Description**

The functions described on this page are used to specify the prior distributions for the parameters in a survextrap model.

#### Usage

```
p_normal(location = 0, scale = 2.5)

p_t(location = 0, scale = 2.5, df = 1)

p_beta(shape1 = 1, shape2 = 1)

p_gamma(shape = 2, rate = 1)
```

# **Arguments**

location	Prior location. For the normal distribution, this is the mean. Defaults to 0
scale	Prior scale. For the normal distribution, this is the standard deviation. Defaults to 2.5.
df	Prior degrees of freedom (only for Student t distribution).
shape1	First shape parameter (for Beta distribution, defaults to 1).
shape2	Second shape parameter (for Beta distribution, defaults to 1).
shape	Shape parameter (for Gamma distribution, defaults to 2).
rate	Rate parameter (for Gamma distribution, defaults to 1).

#### Value

A named list with the prior distribution name and parameters, to be used internally by the survextrap model fitting functions.

#### See Also

survextrap.

prior\_haz Determine priors for time-varying hazards and hazard ratios

### **Description**

Computes consequences of priors chosen for the parameters hsd and hrsd in a flexible hazard model survextrap on an interpretable scale. This can be used to calibrate Gamma priors for these parameters to match interpretable beliefs.

# Usage

```
prior_haz_sd(
   mspline,
   coefs_mean = NULL,
   prior_hsd = p_gamma(2, 1),
   prior_hscale = p_normal(0, 20),
   smooth_model = "exchangeable",
   prior_loghr = NULL,
   formula = NULL,
   cure = NULL,
   nonprop = NULL,
   newdata = NULL,
   prior_hrsd = NULL,
   tmin = 0,
   tmax = NULL,
   nsim = 1000,
```

```
hq = c(0.1, 0.9),
 quantiles = c(0.025, 0.5, 0.975)
)
prior_hr_sd(
  mspline,
  coefs_mean = NULL,
  prior_hsd = p_gamma(2, 1),
  prior_hscale = p_normal(0, 20),
  smooth_model = "exchangeable",
  prior_loghr = NULL,
  formula = NULL,
  cure = NULL,
  nonprop = NULL,
  newdata = NULL,
  newdata0 = NULL,
  prior_hrsd = NULL,
  tmin = 0,
  tmax = 10,
  nsim = 100,
 hq = c(0.1, 0.9),
  quantiles = c(0.025, 0.5, 0.975)
)
```

### Arguments

mspline

A list of control parameters defining the spline model.

knots: Spline knots. If this is not supplied, then the number of knots is taken from df, and their location is taken from equally-spaced quantiles of the observed event times in the individual-level data.

add\_knots: This is intended to be used when there are external data included in the model. External data are typically outside the time period covered by the individual data. add\_knots would then be chosen to span the time period covered by the external data, so that the hazard trajectory can vary over that time.

If there are external data, and both knots and add\_knots are omitted, then a default set of knots is chosen to span both the individual and external data, by taking the quantiles of a vector defined by concatenating the individual-level event times with the start and stop times in the external data.

df: Degrees of freedom, i.e. the number of parameters (or basis terms) intended to result from choosing knots based on quantiles of the data. The total number of parameters will then be df plus the number of additional knots specified in add\_knots. df defaults to 10. This does not necessarily overfit, because the function is smoothed through the prior.

degree: Polynomial degree used for the basis function. The default is 3, giving a cubic. This can only be changed from 3 if bemooth is FALSE.

bsmooth: If TRUE (on by default) the spline is smoother at the highest knot, by defining the derivative and second derivative at this point to be zero.

coefs\_mean

Spline basis coefficients that define the prior mean for the hazard function. By default, these are set to values that define a constant hazard function (see mspline\_constant\_coefs). They are normalised to sum to 1 internally (if they do not already).

prior\_hsd

Gamma prior for the standard deviation that controls the variability over time (or smoothness) of the hazard function. This should be a call to p\_gamma(). The default is p\_gamma(2,1). See prior\_haz\_sd for a way to calibrate this to represent a meaningful belief.

prior\_hscale

Prior for the baseline log hazard scale parameter (alpha or log(eta)). This should be a call to a prior constructor function, such as  $p_normal(0,1)$  or  $p_t(0,2,2)$ . Supported prior distribution families are normal (parameters mean and SD) and t distributions (parameters location, scale and degrees of freedom). The default is a normal distribution with mean 0 and standard deviation 20.

Note that eta is not in itself a hazard, but it is proportional to the hazard (see the vignette for the full model specification).

"Baseline" is defined by the continuous covariates taking a value of zero and factor covariates taking their reference level. To use a different baseline, the data should be transformed appropriately beforehand, so that a value of zero has a different meaning. For continuous covariates, it helps for both computation and interpretation to define the value of zero to denote a typical value in the data, e.g. the mean.

smooth\_model

The default "random\_walk", specifies a random walk prior for the multinomial-logit spline coefficients, based on logistic distributions. See the methods vignette for full details.

The alternative "exchangeable" uses independent logistic priors on the multinomial-logit spline coefficients, conditionally on a common smoothing variance parameter. Note this is the method explained in the original survextrap paper (Jackson, BMC Med Res 2023). The random walk model is shown to perform better in Timmins et al (2025).

In non-proportional hazards models, setting smooth\_model also determines whether an exchangeable or random walk model is used for the non-proportionality parameters ( $\delta$ ).

prior\_loghr

Priors for log hazard ratios. This should be a call to  $p_normal()$  or  $p_t()$ . A list of calls can also be provided, to give different priors to different coefficients, where the name of each list component matches the name of the coefficient, e.g.  $list("age45-59" = p_normal(0,1), "age60+" = p_t(0,2,3))$ 

The default is  $p_normal(0, 2.5)$  for all coefficients.

formula

A survival formula in standard R formula syntax, with a call to Surv() on the left hand side.

Covariates included on the right hand side of the formula with be modelled with proportional hazards, or if nonprop is TRUE then a non-proportional hazards is used.

If data is omitted, so that the model is being fitted to external aggregate data alone, without individual data, then the formula should not include a Surv() call. The left-hand side of the formula will then be empty, and the right hand side specifies the covariates as usual. For example, formula = ~1 if there are no covariates.

cure If TRUE, a mixture cure model is used, where the "uncured" survival is defined

by the M-spline model, and the cure probability is estimated.

nonprop Non-proportional hazards model specification. This is achieved by modelling

the spline basis coefficients in terms of the covariates. See the methods vignette

for more details.

If TRUE, then all covariates are modelled with non-proportional hazards, using

the same model formula as formula.

If this is a formula, then this is assumed to define a model for the dependence of

the basis coefficients on the covariates.

IF this is NULL or FALSE (the default) then any covariates are modelled with

proportional hazards.

newdata A data frame with one row, containing variables in the model formulae. Samples

will then be drawn, for any covariate-dependent parameters, with covariates set

to the values given here.

prior\_hrsd Prior for the standard deviation parameters that smooth the non-proportionality

effects over time in non-proportional hazards models. This should be a call to p\_gamma() or a list of calls to p\_gamma() with one component per covariate, as in prior\_loghr. See prior\_hr\_sd for a way to calibrate this to represent a

meaningful belief.

tmin Minimum plotting time. Defaults to zero.

tmax Maximum plotting time. Defaults to the highest knot.

nsim Number of simulations to draw

hq Quantiles which define the "low" and "high" values of a time-varying quan-

tity (hazard in prior\_haz\_sd and the hazard ratio in prior\_hr\_sd). The ratio between the high and low values will be summarised, as a measure of time-dependence. By default, this is c(0.1, 0.9), so that the 10% and 90% quantiles

are used respectively.

quantiles Quantiles used to summarise the implied prior distributions of the simulated

quantities.

newdata0 A data frame with one row, containing "reference" values of variables in the

model formulae. The hazard ratio between the hazards at newdata and newdata0

will be returned.

#### **Details**

The spline model in survextrap allows the hazard to change over time in an arbitrarily flexible manner. The prior distributions on the parameters of this model have implications for how much we expect the hazard to plausibly vary over time. These priors are hard to interpret directly, but this function can be used to compute their implications on a more easily-understandable scale.

This is done by:

- (1) simulating a set of parameters from their prior distributions
- (2) computing the hazard at a fine grid of equally-spaced points spanning the boundary knots
- (3) calculating the empirical standard deviation of the set of hazards at these points

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(4) repeatedly performing steps 1-3, and summarising the distribution of the resulting standard deviations. This is the implied prior for the hazard variability.

prior\_haz\_sd computes the SD of the hazard, and the SD of the inverse hazard is also computed. The inverse hazard at time t is the expected time to the event given survival to t. The hazard ratio between a high and low value (defined by quantiles of values at different times) is also computed.

prior\_hr\_sd computes the SD of the hazard ratio between two covariate values supplied by the

All of these SDs refer to the variability over time, e.g. a SD of 0 indicates that the hazard (or inverse hazard, or hazard ratio) is constant with time.

#### Value

A data frame with columns sd\_haz (SD of the hazard), sd\_mean (SD of the inverse hazard) and hr (ratio between high/low hazards) (for prior\_haz\_sd), and rows giving prior quantiles of these.

In prior\_hr\_sd, sd\_hr is the SD of hazard ratios over time, and hrr is the ratio between high/low hazard ratios.

prior\_haz\_const

Summarises the prior for the constant hazard implied by a particular prior on the hazard scale parameter and spline specification.

## **Description**

Summarises the prior for the constant hazard implied by a particular prior on the hazard scale parameter and M-spline specification, when the spline coefficients are fixed to define a constant hazard using mspline\_constant\_coefs.

## Usage

```
prior_haz_const(
  mspline,
  prior_hscale = p_normal(0, 20),
  quantiles = c(0.025, 0.5, 0.975)
)
```

## Arguments

mspline

A list of control parameters defining the spline model.

knots: Spline knots. If this is not supplied, then the number of knots is taken from df, and their location is taken from equally-spaced quantiles of the observed event times in the individual-level data.

add\_knots: This is intended to be used when there are external data included in the model. External data are typically outside the time period covered by the individual data. add\_knots would then be chosen to span the time period covered by the external data, so that the hazard trajectory can vary over that time.

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If there are external data, and both knots and add\_knots are omitted, then a default set of knots is chosen to span both the individual and external data, by taking the quantiles of a vector defined by concatenating the individual-level event times with the start and stop times in the external data.

df: Degrees of freedom, i.e. the number of parameters (or basis terms) intended to result from choosing knots based on quantiles of the data. The total number of parameters will then be df plus the number of additional knots specified in add\_knots. df defaults to 10. This does not necessarily overfit, because the function is smoothed through the prior.

degree: Polynomial degree used for the basis function. The default is 3, giving a cubic. This can only be changed from 3 if bsmooth is FALSE.

bsmooth: If TRUE (on by default) the spline is smoother at the highest knot, by defining the derivative and second derivative at this point to be zero.

prior\_hscale

Prior for the baseline log hazard scale parameter (alpha or log(eta)). This should be a call to a prior constructor function, such as  $p\_normal(0,1)$  or  $p\_t(0,2,2)$ . Supported prior distribution families are normal (parameters mean and SD) and t distributions (parameters location, scale and degrees of freedom). The default is a normal distribution with mean 0 and standard deviation 20.

Note that eta is not in itself a hazard, but it is proportional to the hazard (see the vignette for the full model specification).

"Baseline" is defined by the continuous covariates taking a value of zero and factor covariates taking their reference level. To use a different baseline, the data should be transformed appropriately beforehand, so that a value of zero has a different meaning. For continuous covariates, it helps for both computation and interpretation to define the value of zero to denote a typical value in the data, e.g. the mean.

quantiles

Quantiles used to summarise the implied prior distributions of the simulated quantities.

#### Value

A data frame with columns haz for the hazard and mean for the inverse hazard (on the mean survival scale) and rows for three summary quantiles.

## See Also

p\_meansurv, mspline\_constant\_coefs

prior\_hr

Summarises the prior for the hazard ratio implied by a particular prior on the log hazard ratio

#### **Description**

Summarises the prior for the hazard ratio implied by a particular prior on the log hazard ratio. Simply applies an exponential transform to quantiles of the given prior.

39 prior\_pred

## Usage

```
prior_hr(prior_loghr = p_normal(0, 2.5), quantiles = c(0.025, 0.5, 0.975))
```

## **Arguments**

prior\_loghr

Priors for log hazard ratios. This should be a call to p\_normal() or p\_t(). A list of calls can also be provided, to give different priors to different coefficients, where the name of each list component matches the name of the coefficient, e.g.

list("age45-59" =  $p_normal(0,1)$ , "age60+" =  $p_t(0,2,3)$ )

The default is  $p_normal(0, 2.5)$  for all coefficients.

quantiles Quantiles used to summarise the implied prior distributions of the simulated

quantities.

#### Value

A vector of quantiles.

prior\_pred

Simulate a dataset from the prior predictive distribution of survival times in an M-spline survival model.

## Description

Simulate a dataset from the prior predictive distribution of survival times in an M-spline survival model. Additive hazards models not currently supported.

## Usage

```
prior_pred(
 n,
  fix_prior = FALSE,
 mspline,
  censtime = Inf,
  coefs_mean = NULL,
  prior_hscale = p_normal(0, 20),
 prior_hsd = p_gamma(2, 1),
  newdata = NULL,
  formula = NULL,
  prior_loghr = NULL,
 prior_hrsd = NULL,
  prior_cure = NULL
)
```

40 prior\_pred

#### **Arguments**

n

Sample size of the simulated dataset. Each observation in the dataset is generated from a model with the same parameters. These parameters are generated from a single simulation from the prior distribution.

fix\_prior

If TRUE, then one value of the parameter vector is drawn from the prior, followed by n individual-level times given this common prior value. If FALSE, then to produce each sampled individual time, a different sample from the prior is used.

mspline

A list of control parameters defining the spline model.

knots: Spline knots. If this is not supplied, then the number of knots is taken from df, and their location is taken from equally-spaced quantiles of the observed event times in the individual-level data.

add\_knots: This is intended to be used when there are external data included in the model. External data are typically outside the time period covered by the individual data. add\_knots would then be chosen to span the time period covered by the external data, so that the hazard trajectory can vary over that time.

If there are external data, and both knots and add\_knots are omitted, then a default set of knots is chosen to span both the individual and external data, by taking the quantiles of a vector defined by concatenating the individual-level event times with the start and stop times in the external data.

df: Degrees of freedom, i.e. the number of parameters (or basis terms) intended to result from choosing knots based on quantiles of the data. The total number of parameters will then be df plus the number of additional knots specified in add\_knots. df defaults to 10. This does not necessarily overfit, because the function is smoothed through the prior.

degree: Polynomial degree used for the basis function. The default is 3, giving a cubic. This can only be changed from 3 if bsmooth is FALSE.

bsmooth: If TRUE (on by default) the spline is smoother at the highest knot, by defining the derivative and second derivative at this point to be zero.

censtime

Right-censoring time to impose on the simulated event times.

coefs\_mean

Spline basis coefficients that define the prior mean for the hazard function. By default, these are set to values that define a constant hazard function (see mspline\_constant\_coefs). They are normalised to sum to 1 internally (if they do not already).

prior\_hscale

Prior for the baseline log hazard scale parameter (alpha or log(eta)). This should be a call to a prior constructor function, such as  $p_normal(0,1)$  or  $p_t(0,2,2)$ . Supported prior distribution families are normal (parameters mean and SD) and t distributions (parameters location, scale and degrees of freedom). The default is a normal distribution with mean 0 and standard deviation 20.

Note that eta is not in itself a hazard, but it is proportional to the hazard (see the vignette for the full model specification).

"Baseline" is defined by the continuous covariates taking a value of zero and factor covariates taking their reference level. To use a different baseline, the data should be transformed appropriately beforehand, so that a value of zero has a different meaning. For continuous covariates, it helps for both computation

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	and interpretation to define the value of zero to denote a typical value in the data, e.g. the mean.
prior_hsd	Gamma prior for the standard deviation that controls the variability over time (or smoothness) of the hazard function. This should be a call to p_gamma(). The default is p_gamma(2,1). See prior_haz_sd for a way to calibrate this to represent a meaningful belief.
newdata	A data frame with one row, containing variables in the model formulae. Samples will then be drawn, for any covariate-dependent parameters, with covariates set to the values given here.
formula	A model formula with no response, defining the covariates on the hazard scale.
prior_loghr	Priors for log hazard ratios. This should be a call to p_normal() or p_t(). A list of calls can also be provided, to give different priors to different coefficients, where the name of each list component matches the name of the coefficient, e.g. list("age45-59" = p_normal(0,1), "age60+" = p_t(0,2,3))  The default is p_normal(0,2.5) for all coefficients.
prior_hrsd	Prior for the standard deviation parameters that smooth the non-proportionality effects over time in non-proportional hazards models. This should be a call to p_gamma() or a list of calls to p_gamma() with one component per covariate, as in prior_loghr. See prior_hr_sd for a way to calibrate this to represent a meaningful belief.
prior_cure	Prior for the baseline cure probability. This should be a call to p_beta(). The default is a uniform prior, p_beta(1,1). Baseline is defined by the mean of continuous covariates and the reference level of factor covariates.

# Value

A data frame with columns time (simulated time) and event (indicator for whether the time is an event time, as opposed to a right-censoring time). The prior parameters are returned in the prior attribute as a list with components alpha (baseline log hazard) and coefs (spline coefficients).

# See Also

prior\_sample

# Description

Draws a sample from the joint prior distribution of the parameters governing a survextrap model for given covariates. This is used, for example, in prior\_sample\_hazard, to visualise the prior distribution around hazard curves implied by a particular M-spline model and parameter priors.

42 prior\_sample

## Usage

```
prior_sample(
 mspline,
  coefs_mean = NULL,
  prior_hsd = p_gamma(2, 1),
  prior_hscale,
  smooth_model = "exchangeable",
  prior_loghr = NULL,
  formula = NULL,
  cure = NULL,
  nonprop = NULL,
  newdata = NULL,
  newdata0 = NULL,
  prior_hrsd = NULL,
  prior_cure = NULL,
  prior_logor_cure = NULL,
  nsim = 100
)
```

#### **Arguments**

mspline

A list of control parameters defining the spline model.

knots: Spline knots. If this is not supplied, then the number of knots is taken from df, and their location is taken from equally-spaced quantiles of the observed event times in the individual-level data.

add\_knots: This is intended to be used when there are external data included in the model. External data are typically outside the time period covered by the individual data. add\_knots would then be chosen to span the time period covered by the external data, so that the hazard trajectory can vary over that time.

If there are external data, and both knots and add\_knots are omitted, then a default set of knots is chosen to span both the individual and external data, by taking the quantiles of a vector defined by concatenating the individual-level event times with the start and stop times in the external data.

df: Degrees of freedom, i.e. the number of parameters (or basis terms) intended to result from choosing knots based on quantiles of the data. The total number of parameters will then be df plus the number of additional knots specified in add\_knots. df defaults to 10. This does not necessarily overfit, because the function is smoothed through the prior.

degree: Polynomial degree used for the basis function. The default is 3, giving a cubic. This can only be changed from 3 if bsmooth is FALSE.

bsmooth: If TRUE (on by default) the spline is smoother at the highest knot, by defining the derivative and second derivative at this point to be zero.

coefs\_mean

Spline basis coefficients that define the prior mean for the hazard function. By default, these are set to values that define a constant hazard function (see mspline\_constant\_coefs). They are normalised to sum to 1 internally (if they do not already).

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prior\_hsd

Gamma prior for the standard deviation that controls the variability over time (or smoothness) of the hazard function. This should be a call to p\_gamma(). The default is p\_gamma(2,1). See prior\_haz\_sd for a way to calibrate this to represent a meaningful belief.

prior\_hscale

Prior for the baseline log hazard scale parameter (alpha or log(eta)). This should be a call to a prior constructor function, such as p\_normal(0,1) or  $p_t(0, 2, 2)$ . Supported prior distribution families are normal (parameters mean and SD) and t distributions (parameters location, scale and degrees of freedom). The default is a normal distribution with mean 0 and standard deviation 20.

Note that eta is not in itself a hazard, but it is proportional to the hazard (see the vignette for the full model specification).

"Baseline" is defined by the continuous covariates taking a value of zero and factor covariates taking their reference level. To use a different baseline, the data should be transformed appropriately beforehand, so that a value of zero has a different meaning. For continuous covariates, it helps for both computation and interpretation to define the value of zero to denote a typical value in the data, e.g. the mean.

smooth\_model

The default "random\_walk", specifies a random walk prior for the multinomiallogit spline coefficients, based on logistic distributions. See the methods vignette for full details.

The alternative "exchangeable" uses independent logistic priors on the multinomiallogit spline coefficients, conditionally on a common smoothing variance parameter. Note this is the method explained in the original survextrap paper (Jackson, BMC Med Res 2023). The random walk model is shown to perform better in Timmins et al (2025).

In non-proportional hazards models, setting smooth\_model also determines whether an exchangeable or random walk model is used for the non-proportionality parameters ( $\delta$ ).

prior\_loghr

Priors for log hazard ratios. This should be a call to p\_normal() or p\_t(). A list of calls can also be provided, to give different priors to different coefficients, where the name of each list component matches the name of the coefficient, e.g. list("age45-59" =  $p_normal(0,1)$ , "age60+" =  $p_t(0,2,3)$ )

The default is  $p_normal(0, 2.5)$  for all coefficients.

formula A model formula with no response, defining the covariates on the hazard scale.

A model formula with no response, giving any covariates on the cure proportion.

A model formula with no response, defining any covariates affecting the spline basis coefficients, which gives a nonproportional hazards model.

A data frame with one row, containing variables in the model formulae. Samples will then be drawn, for any covariate-dependent parameters, with covariates set to the values given here.

A data frame with one row, containing "reference" values of variables in the

model formulae. The hazard ratio between the hazards at newdata and newdata0 will be returned.

Prior for the standard deviation parameters that smooth the non-proportionality effects over time in non-proportional hazards models. This should be a call to

cure nonprop

newdata

newdata0

prior\_hrsd

44 prior\_sample\_hazard

p\_gamma() or a list of calls to p\_gamma() with one component per covariate, as in prior\_loghr. See prior\_hr\_sd for a way to calibrate this to represent a meaningful belief.

prior\_cure Prior for the baseline cure probability. This should be a call to p\_beta(). The

default is a uniform prior, p\_beta(1,1). Baseline is defined by the mean of continuous covariates and the reference level of factor covariates.

prior\_logor\_cure

Priors for log odds ratios on cure probabilities. This should be a call to  $p_normal()$  or  $p_t()$ . The default is  $p_normal(\emptyset, 2.5)$ .

nsim Number of simulations to draw

#### Value

prior\_sample\_hazard returns a data frame of the samples, and plot\_prior\_hazard generates a plot. No customisation options are provided for the plot function, which is just intended as a quick check.

A list with components:

alpha: Baseline log hazard scale parameter (log(eta) in the notation of the manual). For models with covariates, this is at the covariate values supplied in X, or at zero if X is not supplied.

hscale: Baseline hazard scale parameter (eta).

coefs: Spline coefficients. For non-proportional hazards model with covariates, these are returned at the suppled value of X, or at values of zero if X is not supplied.

beta: Multinomial logit-transformed spline coefficients.

hsd: Smoothing standard deviation for spline coefficients.

If X0 is supplied, then alpha0, hscale0, beta0, coefs0 are also returned, representing reference covariate values.

pcure is returned in cure models (the cure probability).

prior\_sample\_hazard Generate and/or plot a sample from the prior distribution of M-spline hazard curves

## **Description**

Generates and/or plots the hazard curves (as functions of time) implied by a prior mean for the spline coefficients (a constant hazard by default) and particular priors for the baseline log hazard and smoothness standard deviation.

prior\_sample\_hazard

## Usage

```
prior_sample_hazard(
  knots = NULL,
  df = 10,
  degree = 3,
  bsmooth = TRUE,
  coefs_mean = NULL,
  prior_hsd = p_gamma(2, 1),
  prior_hscale = NULL,
  smooth_model = "exchangeable",
  prior_loghr = NULL,
  formula = NULL,
  cure = NULL,
  nonprop = NULL,
  newdata = NULL,
  newdata0 = NULL,
  prior_hrsd = NULL,
  tmin = 0,
  tmax = 10,
  nsim = 10
)
plot_prior_hazard(
  knots = NULL,
  df = 10,
  degree = 3,
  bsmooth = TRUE,
  coefs_mean = NULL,
  prior_hsd = p_gamma(2, 1),
  prior_hscale = p_normal(0, 20),
  smooth_model = "exchangeable",
  prior_loghr = NULL,
  formula = NULL,
  cure = NULL,
  nonprop = NULL,
  newdata = NULL,
  prior_hrsd = p_gamma(2, 1),
  tmin = 0,
  tmax = NULL,
  nsim = 10
)
```

# Arguments

knots

Vector of knot locations. If not supplied, df has to be specified. One of two rules is then used to choose the knot locations. If bknot is specified, a set of equally spaced knots between zero and bknot is used. Otherwise if obstimes is supplied, the knots are chosen as equally spaced quantiles of obstimes.

The number of knots (excluding zero) is df - degree + 1 if bsmooth is TRUE, or df - degree - 1 otherwise.

df

Desired number of basis terms, or "degrees of freedom" in the spline. If knots is not supplied, the number of knots is then chosen to satisfy this.

degree

Spline polynomial degree. Can only be changed from the default of 3 if bsmooth is FALSE.

bsmooth

If TRUE then the function is constrained to also have zero derivative and second derivative at the boundary.

coefs\_mean

Spline basis coefficients that define the prior mean for the hazard function. By default, these are set to values that define a constant hazard function (see mspline\_constant\_coefs). They are normalised to sum to 1 internally (if they do not already).

prior\_hsd

Gamma prior for the standard deviation that controls the variability over time (or smoothness) of the hazard function. This should be a call to p\_gamma(). The default is p\_gamma(2,1). See prior\_haz\_sd for a way to calibrate this to represent a meaningful belief.

prior\_hscale

Prior for the baseline log hazard scale parameter (alpha or log(eta)). This should be a call to a prior constructor function, such as  $p_normal(0,1)$  or  $p_t(0,2,2)$ . Supported prior distribution families are normal (parameters mean and SD) and t distributions (parameters location, scale and degrees of freedom). The default is a normal distribution with mean 0 and standard deviation 20.

Note that eta is not in itself a hazard, but it is proportional to the hazard (see the vignette for the full model specification).

"Baseline" is defined by the continuous covariates taking a value of zero and factor covariates taking their reference level. To use a different baseline, the data should be transformed appropriately beforehand, so that a value of zero has a different meaning. For continuous covariates, it helps for both computation and interpretation to define the value of zero to denote a typical value in the data, e.g. the mean.

smooth\_model

The default "random\_walk", specifies a random walk prior for the multinomial-logit spline coefficients, based on logistic distributions. See the methods vignette for full details.

The alternative "exchangeable" uses independent logistic priors on the multinomial-logit spline coefficients, conditionally on a common smoothing variance parameter. Note this is the method explained in the original survextrap paper (Jackson, BMC Med Res 2023). The random walk model is shown to perform better in Timmins et al (2025).

In non-proportional hazards models, setting smooth\_model also determines whether an exchangeable or random walk model is used for the non-proportionality parameters ( $\delta$ ).

prior\_loghr

Priors for log hazard ratios. This should be a call to  $p_normal()$  or  $p_t()$ . A list of calls can also be provided, to give different priors to different coefficients, where the name of each list component matches the name of the coefficient, e.g.  $list("age45-59" = p_normal(0,1), "age60+" = p_t(0,2,3))$ 

The default is  $p_normal(0, 2.5)$  for all coefficients.

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formula

A survival formula in standard R formula syntax, with a call to Surv() on the left hand side.

Covariates included on the right hand side of the formula with be modelled with proportional hazards, or if nonprop is TRUE then a non-proportional hazards is used.

If data is omitted, so that the model is being fitted to external aggregate data alone, without individual data, then the formula should not include a Surv() call. The left-hand side of the formula will then be empty, and the right hand side specifies the covariates as usual. For example, formula = ~1 if there are no covariates.

cure

If TRUE, a mixture cure model is used, where the "uncured" survival is defined by the M-spline model, and the cure probability is estimated.

nonprop

Non-proportional hazards model specification. This is achieved by modelling the spline basis coefficients in terms of the covariates. See the methods vignette for more details.

If TRUE, then all covariates are modelled with non-proportional hazards, using the same model formula as formula.

If this is a formula, then this is assumed to define a model for the dependence of the basis coefficients on the covariates.

IF this is NULL or FALSE (the default) then any covariates are modelled with proportional hazards.

newdata

A data frame with one row, containing variables in the model formulae. Samples will then be drawn, for any covariate-dependent parameters, with covariates set to the values given here.

newdata0

A data frame with one row, containing "reference" values of variables in the model formulae. The hazard ratio between the hazards at newdata and newdata0 will be returned.

prior\_hrsd

Prior for the standard deviation parameters that smooth the non-proportionality effects over time in non-proportional hazards models. This should be a call to p\_gamma() or a list of calls to p\_gamma() with one component per covariate, as in prior\_loghr. See prior\_hr\_sd for a way to calibrate this to represent a meaningful belief.

tmin Minimum plotting time. Defaults to zero.

tmax Maximum plotting time. Defaults to the highest knot.

nsim Number of simulations to draw

# Value

A tidy data frame containing values of the hazard at a sequence of times, for different draws from the prior distribution.

48 p\_meansurv

p_hr	Derive a normal prior for the log hazard ratio parameter based on a guess at the hazard ratio

## Description

Derive a normal prior for the log hazard ratio parameter based on a guess at the hazard ratio. This assumes that the log upper limit is 2 standard deviations away from the log median.

#### Usage

```
p_hr(median, upper)
```

## **Arguments**

median Best guess (prior median) for a typical hazard ratio upper Upper limit of 95% credible interval for hazard ratio

#### Value

A normal prior in the format returned by p\_normal, which can be passed directly to the prior\_loghr argument in survextrap.

p_meansurv	Derive a normal prior for the log hazard scale parameter based on a guess at survival times

# Description

Derive a normal prior for the log hazard scale parameter based on a guess at survival times. The scale parameter is hard to interpret, and depends on the spline knots. However for any scale parameter, we can determine the spline coefficients that give a constant hazard (mspline\_constant\_coefs). Therefore if we can guess a typical survival time, we can guess a typical hazard (as 1 divided by the survival time) and deduce the scale parameter. The prior is then constructed by assuming normality on the log scale, and assuming the log upper credible limit is two SDs away from the log median.

## Usage

```
p_meansurv(median, upper, mspline = NULL)
```

# Arguments

median	Best guess (prior median) for a typical survival time
upper	Upper limit of 95% credible interval for a survival time

mspline A list with components knots (vector of knots), degree (polynomial degree)

and bsmooth (logical for smoothness constraint at boundary), defining an M-

spline configuration.

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## Value

A normal prior in the format returned by p\_normal, which can be passed directly to the prior\_hscale argument in survextrap.

## See Also

```
prior_haz_const, mspline_constant_coefs
```

rmst

Restricted mean survival time

## **Description**

Compute the restricted mean survival time from a model fitted with survextrap. Defined as the integral of the fitted survival curve up to a specified time.

## Usage

```
rmst(
    x,
    t,
    newdata = NULL,
    newdata0 = NULL,
    wane_period = NULL,
    wane_nt = 10,
    disc_rate = 0,
    method = "gl",
    gl_nodes = 100,
    niter = NULL,
    summ_fns = NULL,
    sample = FALSE
)
```

# Arguments

x A fitted model object as returned by survextrap

t Vector of times. The restricted mean survival time up to each one of these times will be computed.

newdata

Data frame of covariate values to compute the output for. If there are covariates in the model and this is not supplied, the following default is used:

- (a) if the only covariate is one factor variable, then the output is computed for each level of this factor.
- (b) if there are multiple covariates, or any numeric covariates, then the output is computed at the mean of each numeric covariate in the original data, and at the baseline level of each factor covariate.

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Note: caution is required about how treatment groups (for example) are stored in your data. If these are coded as numeric (0/1), then if newdata is not specified, only one output will be shown. This relates to the average value of this numeric variable over the data, which doesn't correspond to either of the treatment groups. To avoid this, a treatment group should be stored as a factor.

Data frame of covariate values defining the "untreated" group for use in treatment waning models. See Survmspline\_wane.

wane\_period

newdata0

Vector of two numbers, defining the time period over which the hazard is interpolated between the hazard of the "treated" group (taken from newdata) and the hazard of the "untreated" group (taken from newdata0). Optional - if this is not supplied, then no waning is assumed.

wane\_nt

Number of intervals defining the piecewise constant approximation to the hazard during the waning period.

disc\_rate

Discounting rate used to calculate the discounted mean or restricted mean survival time, using an exponential discounting function.

method

Method of numerical integration to obtain the restricted mean survival time from

the survival function.

The default is method="gl", a Gauss-Legendre method with 100 nodes between

zero and the maximum of t.

method="adaptive" uses the base R integrate function, which is much slower, but potentially more robust for badly-behaved survival functions.

gl\_nodes

Number of nodes for the Gauss-Legendre method.

niter

Number of MCMC iterations to use to compute credible intervals. Set to a low value to make this function quicker, at the cost of some approximation error (which may not be important for plotting or model development).

summ\_fns

A list of functions to use to summarise the posterior sample. This is passed to posterior::summarise\_draws. By default this is list(median=median, ~quantile(.x, probs=c(0.025, 0.975))). If the list is named, then the names will be used for the columns of the output.

sample

If TRUE then an MCMC sample is returned from the posterior of the output, rather than summary statistics.

## Value

A data frame (tibble) with each row containing posterior summary statistics for a particular time and covariate value.

Or if sample=TRUE, an array with dimensions length(t), niter, nrow(newdata), giving the RMST evaluated at different times, MCMC iterations and covariate values respectively.

## **Examples**

```
mod <- survextrap(Surv(years, status) ~ rx, data=colons, fit_method="opt")
rmst(mod, t=3, niter=100)
rmst(mod, t=3, summ_fns=list(mean=mean), niter=100)</pre>
```

51 standardise\_to

standardise_to	Constructor for a standardising population used for survextrap out-
	puts

#### **Description**

Standardised outputs are outputs from models with covariates, that are defined by marginalising (averaging) over covariate values in a given population, rather than being conditional on a given covariate value.

## **Usage**

```
standardise_to(newdata, nstd = 1, random = FALSE)
standardize_to(newdata, nstd = 1, random = FALSE)
```

# **Arguments**

Data frame describing a population. newdata

nstd Number of draws from the population distribution used per MCMC sample from

> the parameters when random=TRUE. With the default of 1, the value of the covariate vector X is essentially treated as if it were an additional parameter in the Bayesian model, drawn by Monte Carlo independently of the remaining param-

eters.

By default this is FALSE, indicating that standardised samples should be obtained

by concatenating the posterior samples for each covariate value in the standard population. The sample from the standardised posterior of parameters then has size niter times the number of rows in newdata, where niter is the number of MCMC iterations used in the original survextrap fit. Computing the resulting output function (e.g. RMST which uses numerical integration) can then be

computationally intensive if this sample size is large.

A quicker alternative is to sample a random row of the standard population for each MCMC iteration. The standardised sample from the posterior then has size niter. This is specified by using random=TRUE. If this is used, then the result depends on the random number seed, and it should be checked that the results are stable to within the required number of significant figures. If not, run survextrap with more MCMC iterations or increase nstd here.

#### **Details**

These are produced by generating a Monte Carlo sample from the joint distribution of parameters  $\theta$  and covariate values X,  $p(X,\theta) = p(\theta|X)p(X)$ , where p(X) is defined by the empirical distribution of covariates in the standard population.

Hence applying a vectorised output function g() (such as the RMST or survival probability) to this sample produces a sample from the posterior of  $\int g(\theta|X)dX$ : the average RMST (say) for a heterogeneous population.

See the Examples vignette for some examples and notes on computation.

random

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#### Value

A copy of newdata, but with attributes added to indicate that this should be used as a standard population. When this newdata is passed to survextrap's output functions, the outputs will then be presented as an average over the empirical distribution of covariate values described by newdata, rather than as one output per row of newdata (distinct covariate values).

## **Examples**

```
rxph_mod <- survextrap(Surv(years, status) ~ rx, data=colons, fit_method="opt")
ref_pop <- data.frame(rx = c("Obs","Lev+5FU"))

# covariate-specific outputs
survival(rxph_mod, t = c(5,10), newdata = ref_pop)

# standardised outputs
survival(rxph_mod, t = c(5,10), newdata = standardise_to(ref_pop))</pre>
```

summary.survextrap

Posterior summary statistics for parameters of survextrap models

## Description

Posterior summary statistics for parameters of survextrap models. The summary statistics presented by default include the posterior median and 95% credible intervals, alongside the Rhat convergence diagnostic and the bulk effective sample size (as computed by the posterior package). For models fitted by optimisation rather than MCMC, the posterior mode is always returned.

Any other posterior summary can be computed if the appropriate function to compute it is supplied in summ\_fns.

#### Usage

```
## S3 method for class 'survextrap'
summary(object, summ_fns = NULL, ...)
```

## **Arguments**

object A fitted model object as returned by survextrap

summ\_fns A list of functions to calculate different posterior summaries from the MCMC

sample. This is passed to posterior::summarise\_draws. If the list is named,

then the names will be used for the columns of the output. See the examples below for different ways this can be used.

Defaults to list(median = median, ~quantile(.x, probs=c(0.025, 0.975)),

sd = sd, rhat = posterior::rhat, ess\_bulk = posterior::ess\_bulk)

Many useful such functions are provided with the posterior package.

Summary functions can also be supplied in separate arguments here. They will

then be added to those supplied in summ\_fns.

#### Value

A data frame (actually a tibble) of summary statistics for the model parameters.

The parameters, as indicated in the variable column, are:

alpha: Baseline log hazard scale. If there are covariates, this describes the log hazard scale with continuous covariates set to zero, and factor covariates set to their baseline levels. Note that this is not the log hazard, which also depends on the spline coefficients and basis. See hazard to extract the actual hazard.

coefs: Coefficients of the M-spline basis terms. If a non-proportional hazards model was fitted, these are with covariates set to zero or baseline levels.

loghr: Log hazard ratios for each covariate in the model. For cure models, this refers to covariates on survival for uncured people. For non-proportional hazards models, these are the multiplicative effects of covariates on the hazard scale parameter. See the methods vignette for a full description of this model.

hr: Hazard ratios (the exponentials of loghr).

pcure: Probability of cure (for cure models only). If there are covariates on cure, this parameter describes the probability of cure with continuous covariates set to zero, and factor covariates set to their baseline levels.

logor\_cure: Log odds ratio of cure for each covariate on cure.

or\_cure: Odds ratios of cure (the exponentials of logor\_cure).

nperr: Standardised departures from proportional hazards in the non-proportional hazards model, defined as  $b_{ks}^{(np)}/\sigma_s^{(np)}$  (see the methods vignette for definitions of these).

hrsd: Smoothness standard deviations  $\tau_s$  for the non-proportionality effects.

## **Examples**

```
mod <- survextrap(Surv(years, status) ~ rx, data=colons, fit_method="opt")
summary(mod)
summary(mod, mean=mean)
summary(mod, list(mean=mean))
summary(mod, list(mean=mean, ess_tail=posterior::ess_tail))
summary(mod, mean=mean, ess_tail=posterior::ess_tail)</pre>
```

survextrap

Flexible Bayesian parametric survival models

## **Description**

Flexible Bayesian parametric survival models. Individual data are represented using M-splines and a proportional hazards or flexible non-proportional hazards model. External aggregate data can be included, for example, to enable extrapolation outside the individual data. A fixed background hazard can also be included in an additive hazards (relative survival) model. Mixture cure versions of these models can also be used.

## Usage

```
survextrap(
  formula,
  data = NULL,
  external = NULL,
  cure = FALSE,
  nonprop = FALSE,
  prior_hscale = p_normal(0, 20),
  prior_loghr = NULL,
  prior_hsd = p_gamma(2, 1),
  prior_cure = p_beta(1, 1),
  prior_logor_cure = NULL,
  prior_hrsd = p_gamma(2, 1),
  backhaz = NULL,
  backhaz_strata = NULL,
  mspline = NULL,
  add_knots = NULL,
  smooth_model = "random_walk",
  hsd = "bayes",
  coefs_mean = NULL,
  fit_method = "mcmc",
  loo = (fit_method == "mcmc"),
)
```

# **Arguments**

formula

A survival formula in standard R formula syntax, with a call to Surv() on the left hand side.

Covariates included on the right hand side of the formula with be modelled with proportional hazards, or if nonprop is TRUE then a non-proportional hazards is used.

If data is omitted, so that the model is being fitted to external aggregate data alone, without individual data, then the formula should not include a Surv() call. The left-hand side of the formula will then be empty, and the right hand side specifies the covariates as usual. For example, formula = ~1 if there are no covariates.

data

Data frame containing variables in formula. Variables should be in a data frame, and not in the working environment.

This may be omitted, in which case external must be supplied. This allows a model to be fitted to external aggregate data alone, without any individual-level

external

External data as a data frame of aggregate survival counts with columns named:

start: Start time

stop: Follow-up time n: Number of people alive at start

r: Number of those people who are still alive at stop

If there are covariates in formula, then the values they take in the external data must be supplied as additional columns in external. Therefore if there are external data, the covariates in formula and data should not be named start, stop, n or r.

cure

If TRUE, a mixture cure model is used, where the "uncured" survival is defined by the M-spline model, and the cure probability is estimated.

nonprop

Non-proportional hazards model specification. This is achieved by modelling the spline basis coefficients in terms of the covariates. See the methods vignette for more details.

If TRUE, then all covariates are modelled with non-proportional hazards, using the same model formula as formula.

If this is a formula, then this is assumed to define a model for the dependence of the basis coefficients on the covariates.

IF this is NULL or FALSE (the default) then any covariates are modelled with proportional hazards.

prior\_hscale

Prior for the baseline log hazard scale parameter (alpha or log(eta)). This should be a call to a prior constructor function, such as  $p\_normal(0,1)$  or  $p\_t(0,2,2)$ . Supported prior distribution families are normal (parameters mean and SD) and t distributions (parameters location, scale and degrees of freedom). The default is a normal distribution with mean 0 and standard deviation 20.

Note that eta is not in itself a hazard, but it is proportional to the hazard (see the vignette for the full model specification).

"Baseline" is defined by the continuous covariates taking a value of zero and factor covariates taking their reference level. To use a different baseline, the data should be transformed appropriately beforehand, so that a value of zero has a different meaning. For continuous covariates, it helps for both computation and interpretation to define the value of zero to denote a typical value in the data, e.g. the mean.

prior\_loghr

Priors for log hazard ratios. This should be a call to p\_normal() or p\_t(). A list of calls can also be provided, to give different priors to different coefficients, where the name of each list component matches the name of the coefficient, e.g. list("age45-59" = p\_normal(0,1), "age60+" = p\_t(0,2,3))

The default is  $p_normal(0, 2.5)$  for all coefficients.

prior\_hsd

Gamma prior for the standard deviation that controls the variability over time (or smoothness) of the hazard function. This should be a call to p\_gamma(). The default is p\_gamma(2,1). See prior\_haz\_sd for a way to calibrate this to represent a meaningful belief.

prior\_cure

Prior for the baseline cure probability. This should be a call to p\_beta(). The default is a uniform prior, p\_beta(1,1). Baseline is defined by the mean of continuous covariates and the reference level of factor covariates.

prior\_logor\_cure

Priors for log odds ratios on cure probabilities. This should be a call to  $p_normal()$  or  $p_t()$ . The default is  $p_normal(0,2.5)$ .

prior\_hrsd

Prior for the standard deviation parameters that smooth the non-proportionality effects over time in non-proportional hazards models. This should be a call to

> p\_gamma() or a list of calls to p\_gamma() with one component per covariate, as in prior\_loghr. See prior\_hr\_sd for a way to calibrate this to represent a meaningful belief.

backhaz

Background hazard, that is, for causes of death other than the cause of interest. This defines a "relative survival" or "additive hazards" model. The overall hazard that describes the all-cause survival data (given in the data and/or external argument) is then modelled as the sum of a cause-specific hazard and a background hazard.

The background hazard is assumed to be known, and the cause-specific hazard is modelled with the flexible parametric model.

The background hazard can be supplied in two forms. The meaning of predictions from the model depends on which of these is used.

- (a) A data frame with columns "hazard" and "time", specifying the background hazard at all times as a piecewise-constant (step) function. Each row gives the background hazard between the specified time and the next time. The first element of "time" should be 0, and the final row specifies the hazard at all times greater than the last element of "time". Predictions from the model fitted by survextrap will then include this background hazard, because it is known at all times.
- (b) The (quoted) name of a variable in the data giving the background hazard. For censored cases, the exact value does not matter. The predictions from survextrap will then describe the excess hazard or survival on top of this background. The overall hazard cannot be predicted in general, because the background hazard is only specified over a limited range of time.

If there is external data, and backhaz is supplied in form (b), then the user should also supply the background survival at the start and stop points in columns of the external data named "backsurv\_start" and "backsurv\_stop". That is, the probability of survival up to each of these times for someone alive at time 0. This should describe the same reference population as backhaz, though the package does not check for consistency between these.

If there are stratifying variables specified in backhaz\_strata, then there should be multiple rows giving the background hazard value for each time period and stratifying variable.

If backhaz is NULL (the default) then no background hazard component is included in the model.

backhaz\_strata A character vector of names of variables that appear in backhaz that indicate strata, e.g. backhaz\_strata = c("agegroup", "sex"). This allows different background hazard values to be used for different subgroups. These variables must also appear in the datasets being modelled, that is, in data, external or both. Each row of those datasets should then have a corresponding row in backhaz which has the same values of the stratifying variables.

This is NULL by default, indicating no stratification of the background hazard.

If stratification is done, then backhaz must be supplied in form (a), as a data frame rather than a variable in the data.

mspline

A list of control parameters defining the spline model.

knots: Spline knots. If this is not supplied, then the number of knots is taken from df, and their location is taken from equally-spaced quantiles of the observed event times in the individual-level data.

add\_knots: This is intended to be used when there are external data included in the model. External data are typically outside the time period covered by the individual data. add\_knots would then be chosen to span the time period covered by the external data, so that the hazard trajectory can vary over that time.

If there are external data, and both knots and add\_knots are omitted, then a default set of knots is chosen to span both the individual and external data, by taking the quantiles of a vector defined by concatenating the individual-level event times with the start and stop times in the external data.

df: Degrees of freedom, i.e. the number of parameters (or basis terms) intended to result from choosing knots based on quantiles of the data. The total number of parameters will then be df plus the number of additional knots specified in add\_knots. df defaults to 10. This does not necessarily overfit, because the function is smoothed through the prior.

degree: Polynomial degree used for the basis function. The default is 3, giving a cubic. This can only be changed from 3 if bsmooth is FALSE.

bsmooth: If TRUE (on by default) the spline is smoother at the highest knot, by defining the derivative and second derivative at this point to be zero.

add\_knots

Any extra knots beyond those chosen from the individual-level data (or supplied in knots). All other spline specifications are set to their defaults, as described in mspline. For example, add\_knots = 10 is a shorthand for mspline = list(add\_knots = 10).

smooth\_model

The default "random\_walk", specifies a random walk prior for the multinomial-logit spline coefficients, based on logistic distributions. See the methods vignette for full details.

The alternative "exchangeable" uses independent logistic priors on the multinomial-logit spline coefficients, conditionally on a common smoothing variance parameter. Note this is the method explained in the original survextrap paper (Jackson, BMC Med Res 2023). The random walk model is shown to perform better in Timmins et al (2025).

In non-proportional hazards models, setting smooth\_model also determines whether an exchangeable or random walk model is used for the non-proportionality parameters ( $\delta$ ).

hsd

Smoothing variance parameter estimation.

"bayes": the smoothing parameter is estimated by full Bayes (the default).

"eb": empirical Bayes is used.

Alternatively, if a number is supplied here, then the smoothing parameter is fixed to this number.

coefs\_mean

Spline basis coefficients that define the prior mean for the hazard function. By default, these are set to values that define a constant hazard function (see mspline\_constant\_coefs). They are normalised to sum to 1 internally (if they do not already).

fit\_method

Method from rstan used to fit the model.

If fit\_method="mcmc" then a sample from the posterior is drawn using Markov Chain Monte Carlo sampling, via **rstan**'s rstan::sampling() function. This is the default. It is the most accurate but the slowest method.

If fit\_method="opt", then instead of an MCMC sample from the posterior, survextrap returns the posterior mode calculated using optimisation, via **rstan**'s rstan::optimizing() function. A sample from a normal approximation to the (real-line-transformed) posterior distribution is drawn in order to obtain credible intervals. This is useful for model development, while using MCMC for the "final answer".

If fit\_method="vb", then variational Bayes methods are used, via **rstan**'s **rstan**: vb() function. This is labelled as "experimental" by **rstan**. It might give a better approximation to the posterior than fit\_method="opt", and is faster than MCMC, in particular for large datasets, but has not been investigated in depth for these models.

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Compute leave-one-out cross-validation statistics. This is done by default. Set to FALSE to not compute them. If these statistics are computed, then they are returned in the loo and loo\_external components of the object returned by survextrap. loo describes the fit of the model to the individual-level data, and loo\_external describes fit to the external data.

See the "examples" vignette for more explanation of these.

. . .

Additional arguments to supply to control the Stan fit, passed to the appropriate **rstan** function, depending on which is chosen through the fit\_method argument.

## Value

A list of objects defining the fitted model. These are not intended to be extracted directly by users. Instead see summary.survextrap for summarising the parameter estimates, and the functions hazard, survival, rmst and others for computing interesting summaries of the fitted survival distribution.

The object returned by rstan containing samples from the fitted model is returned in the stanfit component. See the rstan documentation. The function get\_draws is provided to convert this to a simple matrix of posterior samples with all chains collapsed.

#### References

Jackson, C. (2023) survextrap: a package for flexible and transparent survival extrapolation. BMC Medical Research Methodology 23:282. doi:10.1186/s12874023020941

Timmins I, Torabi F, Jackson C, Lambert P, Sweeting M J. (2025) Simulation-based assessment of a Bayesian survival model with flexible baseline hazard and time-dependent effects. doi:10.48550/arXiv.2503.21388.

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survival

Estimates of survival from a survextrap model

## **Description**

Estimates of survival probabilities at particular times, from a survextrap model

# Usage

```
survival(
    x,
    newdata = NULL,
    t = NULL,
    tmax = NULL,
    niter = NULL,
    summ_fns = NULL,
    sample = FALSE,
    newdata0 = NULL,
    wane_period = NULL,
    wane_nt = 10
)
```

## **Arguments**

Х

A fitted model object as returned by survextrap

newdata

Data frame of covariate values to compute the output for. If there are covariates in the model and this is not supplied, the following default is used:

- (a) if the only covariate is one factor variable, then the output is computed for each level of this factor.
- (b) if there are multiple covariates, or any numeric covariates, then the output is computed at the mean of each numeric covariate in the original data, and at the baseline level of each factor covariate.

Note: caution is required about how treatment groups (for example) are stored in your data. If these are coded as numeric (0/1), then if newdata is not specified, only one output will be shown. This relates to the average value of this numeric variable over the data, which doesn't correspond to either of the treatment groups. To avoid this, a treatment group should be stored as a factor.

t

Vector of times at which to compute the estimates.

tmax

Maximum time at which to compute the estimates. If t is supplied, then this is ignored. If t is not supplied, then t is set to a set of 100 equally spaced time points from 0 to tmax. If both tmax and t are not supplied, then tmax is set to the maximum follow up time in the data.

niter

Number of MCMC iterations to use to compute credible intervals. Set to a low value to make this function quicker, at the cost of some approximation error (which may not be important for plotting or model development).

summ_fns	A list of functions to use to summarise the posterior sample. This is passed to posterior::summarise_draws. By default this is list(median=median, ~quantile(.x, probs=c(0.025, 0.975))). If the list is named, then the names will be used for the columns of the output.
sample	If TRUE then the MCMC samples are returned instead of being summarised as a median and 95% credible intervals.
newdata0	Data frame of covariate values defining the "untreated" group for use in treatment waning models. See Survmspline_wane.
wane_period	Vector of two numbers, defining the time period over which the hazard is interpolated between the hazard of the "treated" group (taken from newdata) and the hazard of the "untreated" group (taken from newdata0). Optional - if this is not supplied, then no waning is assumed.
wane_nt	Number of intervals defining the piecewise constant approximation to the hazard during the waning period.

# Value

A data frame (tibble) giving posterior summary statistics, or (if sample=TRUE) an array giving samples from the posterior.

Survmspline

M-spline survival distribution

# Description

Probability density, distribution, quantile, random generation, hazard, cumulative hazard, mean and restricted mean functions for the M-spline time-to-event model. This can optionally be combined with a cure probability, and / or with a known background hazard trajectory that is a piecewise-constant function of time.

# Usage

```
psurvmspline(
   q,
   alpha,
   coefs,
   knots,
   degree = 3,
   lower.tail = TRUE,
   log.p = FALSE,
   pcure = 0,
   offsetH = 0,
   backhaz = NULL,
   bsmooth = TRUE
)
```

```
Hsurvmspline(
  х,
  alpha,
  coefs,
  knots,
  degree = 3,
  log = FALSE,
  pcure = 0,
 offsetH = 0,
 backhaz = NULL,
 bsmooth = TRUE
)
hsurvmspline(
  Х,
  alpha,
  coefs,
  knots,
  degree = 3,
  log = FALSE,
  pcure = 0,
  offseth = 0,
 backhaz = NULL,
 bsmooth = TRUE
)
dsurvmspline(
  х,
  alpha,
  coefs,
  knots,
  degree = 3,
  log = FALSE,
  pcure = 0,
 offseth = 0,
 offsetH = 0,
 backhaz = NULL,
 bsmooth = TRUE
qsurvmspline(
  р,
  alpha,
  coefs,
  knots,
  degree = 3,
  lower.tail = TRUE,
  log.p = FALSE,
```

```
pcure = 0,
 offsetH = 0,
 backhaz = NULL,
 bsmooth = TRUE
)
rsurvmspline(
 n,
 alpha,
 coefs,
  knots,
 degree = 3,
 pcure = 0,
 offsetH = 0,
 backhaz = NULL,
 bsmooth = TRUE
)
rmst_survmspline(
  t,
 alpha,
 coefs,
 knots,
 degree = 3,
 pcure = 0,
 offsetH = 0,
 backhaz = NULL,
 bsmooth = TRUE,
 disc_rate = 0,
 method = "gl",
 gl_nodes = 100
)
mean_survmspline(
  alpha,
  coefs,
 knots,
 degree = 3,
 pcure = 0,
 offsetH = 0,
 backhaz = NULL,
 bsmooth = TRUE,
 disc_rate = 0
)
```

# **Arguments**

alpha Log hazard scale parameter.

coefs	Spline basis coefficients. These should sum to 1, otherwise they are normalised internally to sum to 1. Supplied either as a vector with one element per basis term, or a matrix with one column per basis term, and rows for alternative values of the coefficients (in vectorised usage of this function). If an array is supplied, it is collapsed into a matrix with number of columns equal to the final dimension of the array.
knots	Locations of knots on the axis of time, supplied in increasing order. These include the two boundary knots.
	In vectorised usage of these functions, the knots and degree must be the same for all alternative times and parameter values.
degree	Spline polynomial degree. Can only be changed from the default of 3 if bsmooth is FALSE.
lower.tail	logical; if TRUE (default), probabilities are $P(X \le x)$ , otherwise, $P(X > x)$ .
pcure	Probability of "cure", which defaults to zero. If this is non-zero, this defines a "mixture cure" version of the M-spline model.
offsetH	Constant to be added to the cumulative hazard.
backhaz	A data frame that defines the background hazard as a piecewise-constant function of time. This should have two columns, named "time" and "hazard". Each row gives the "background" hazard between the specified time and the next time. The first element of "time" should be 0, and the final row specifies the hazard at all times greater than the last element of "time".
bsmooth	If TRUE then the function is constrained to also have zero derivative and second derivative at the boundary.
x, q, t	Vector of times.
log, log.p	Return log density or probability.
offseth	Constant to be added to the hazard, e.g. representing a "background" risk of death from causes other than the cause of interest.
p	Vector of probabilities.
n	Number of random numbers to simulate.
disc_rate	Discounting rate used to calculate the discounted mean or restricted mean survival time, using an exponential discounting function.
method	Method of numerical integration to obtain the restricted mean survival time from the survival function.
	The default is method="gl", a Gauss-Legendre method with 100 nodes between zero and the maximum of t.
	method="adaptive" uses the base R integrate function, which is much slower, but potentially more robust for badly-behaved survival functions.
gl_nodes	Number of nodes for the Gauss-Legendre method.

# **Details**

These are the same as the M-splines used to model survival data in rstanarm, except that an additional assumption is made that the hazard is constant beyond the boundary knots at its value at the boundary. This gives a continuous but non-smooth function.

The "cure" model can be interpreted in two different ways. These result in identical probability distribution functions for the event time, hence they are indistinguishable from data:

(a) a model where everyone follows the same hazard trajectory that is decreasing through time, with a higher rate of decrease for higher poure.

(b) a model where a person either has a constant zero hazard at all times, or a hazard that follows a parametric model (M-spline in this case). The probability that a person has a zero hazard is pcure. This is the "mixture model" interpretation.

In the "background hazard" model, the overall hazard is defined as a sum of the background hazard and a cause-specific hazard. The cause-specific hazard is modelled with the M-spline model, and the background hazard is assumed to be a known piecewise-constant function defined by backhaz.

If both backhaz and poure are supplied, then the cure probability applies only to the cause-specific hazard. That is, the cause-specific hazard decreases to zero, and the overall hazard converges towards the background hazard, as time increases.

#### Value

dsurvmspline gives the density, psurvmspline gives the distribution function, hsurvmspline gives the hazard and Hsurvmspline gives the cumulative hazard.

qsurvmspline gives the quantile function, which is computed by numerical inversion.

rsurvmspline generates random survival times by using qsurvmspline on a sample of uniform random numbers.

#### Author(s)

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## References

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Survmspline\_wane

M-spline survival distribution under treatment effect waning

## **Description**

This defines the CDF, cumulative hazard and hazard of a survival distribution defined by combining the hazards of two different groups (e.g. "treated" and "untreated") each defined by a standard M-spline model. The log hazards of one group and the other are interpolated over a defined period of time. This may be used for models where the treatment effect wanes, over a period of time, between an estimated hazard ratio and zero.

# Usage

```
Hsurvmspline_wane(
  alpha1,
  alpha0,
  coefs1,
  coefs0,
  knots,
  degree = 3,
 wane_period,
 wane_nt = 10,
 pcure1 = 0,
  pcure0 = 0,
 offsetH = 0,
  backhaz = NULL,
 bsmooth = TRUE,
  log = FALSE
)
psurvmspline_wane(
  q,
  alpha1,
  alpha0,
  coefs1,
  coefs0,
  knots,
  degree = 3,
  bsmooth = TRUE,
 wane_period,
 wane_nt = 10,
  lower.tail = TRUE,
  pcure1 = 0,
  pcure0 = 0,
  offsetH = 0,
  backhaz = NULL,
  log.p = FALSE
)
hsurvmspline_wane(
  Х,
  alpha1,
  alpha0,
  coefs1,
  coefs0,
  knots,
  degree = 3,
  bsmooth = TRUE,
 wane_period,
```

```
wane_nt = 10,
 pcure1 = 0,
 pcure0 = 0,
 offseth = 0,
 backhaz = NULL,
 log = FALSE
)
dsurvmspline_wane(
  alpha1,
 alpha0,
  coefs1,
  coefs0,
  knots,
  degree = 3,
  bsmooth = TRUE,
 wane_period,
 wane_nt = 10,
 pcure1 = 0,
 pcure0 = 0,
 offseth = 0,
 offsetH = 0,
 backhaz = NULL,
 log = FALSE
)
qsurvmspline_wane(
 р,
  alpha1,
 alpha0,
  coefs1,
  coefs0,
  knots,
  degree = 3,
 bsmooth = TRUE,
  lower.tail = TRUE,
 log.p = FALSE,
  pcure1 = 0,
 pcure0 = 0,
 offsetH = 0,
 backhaz = NULL,
 wane_period,
 wane_nt = 10
)
rsurvmspline_wane(
 n,
```

```
alpha1,
  alpha0,
  coefs1,
  coefs0,
  knots,
  degree = 3,
  bsmooth = TRUE,
  pcure1 = 0,
  pcure0 = 0,
 offsetH = 0,
 backhaz = NULL,
 wane_period,
 wane_nt = 10
)
rmst_survmspline_wane(
  t,
  alpha1,
  alpha0,
  coefs1,
  coefs0,
  knots,
  degree = 3,
  pcure1 = 0,
  pcure0 = 0,
  offsetH = 0,
  backhaz = NULL,
  bsmooth = TRUE,
 wane_period,
 wane_nt = 10,
  disc_rate = 0,
 method = "gl",
  gl_nodes = 100
)
mean_survmspline_wane(
  alpha1,
  alpha0,
  coefs1,
  coefs0,
  knots,
  degree = 3,
  pcure1 = 0,
  pcure0 = 0,
  backhaz = NULL,
  bsmooth = TRUE,
  wane_period,
 wane_nt = 10,
```

```
disc_rate = 0
)
```

#### **Arguments**

x, q, t Vector of times.

alpha1, coefs1, pcure1

log hazard intercept, spline coefficients and cure probability before the start of the waning period ("treated")

alpha0, coefs0, pcure0

log hazard intercept, spline coefficients and cure probability after the end of the

waning period ("untreated")

knots Locations of knots on the axis of time, supplied in increasing order. These

include the two boundary knots.

In vectorised usage of these functions, the knots and degree must be the same

for all alternative times and parameter values.

degree Spline polynomial degree. Can only be changed from the default of 3 if bsmooth

is FALSE.

wane\_period vector of two components giving start and stop of waning period

wane\_nt time resolution for piecewise constant hazard approximation in the waning pe-

riod. If this is not specified, defaults to dividing the waning period into 10

pieces.

offsetH Constant to be added to the cumulative hazard.

backhaz A data frame that defines the background hazard as a piecewise-constant func-

tion of time. This should have two columns, named "time" and "hazard". Each row gives the "background" hazard between the specified time and the next time. The first element of "time" should be 0, and the final row specifies the hazard

at all times greater than the last element of "time".

bsmooth If TRUE then the function is constrained to also have zero derivative and second

derivative at the boundary.

log, log.p Return log density or probability.

lower.tail logical; if TRUE (default), probabilities are  $P(X \le x)$ , otherwise, P(X > x).

offseth Constant to be added to the hazard, e.g. representing a "background" risk of

death from causes other than the cause of interest.

p Vector of probabilities.

n Number of random numbers to simulate.

disc\_rate Discounting rate used to calculate the discounted mean or restricted mean sur-

vival time, using an exponential discounting function.

method Method of numerical integration to obtain the restricted mean survival time from

the survival function.

The default is method="g1", a Gauss-Legendre method with 100 nodes between

zero and the maximum of t.

method="adaptive" uses the base R integrate function, which is much slower,

but potentially more robust for badly-behaved survival functions.

#### **Details**

This distribution is defined as follows.

• Between time 0 and wane\_period[1], the "treated" hazard is used, as defined by an M-spline with intercept alpha1.

- Between wane\_period[1] and wane\_period[2], the log hazard is defined by linear interpolation. The waning period is divided into a number of discrete pieces in which the hazard is assumed to be constant, defined by the hazard at the start of the piece. These hazard values are obtained from the spline model, using an intercept parameter alpha (log scale parameter) defined by linearly interpolating between alpha1 and alpha0 over the waning period. The cumulative hazard at any time can then be deduced by adding up contributions on each piece.
- After wane\_period[2], the "untreated" hazard is used, as defined by an M-spline with intercept alpha0.

See the methods vignette for more details and examples.

This can be used to predict the hazard of a person treated with a treatment whose short-term effect is estimated from shorter-term data, but we wish to extrapolate this model over a longer period where the effect is assumed to diminish.

This may not work if the hazard is zero or infinite at any point in the waning period (thus the log hazard is indeterminate). This might typically happen at time 0.

#### Value

psurvmspline\_wane gives the CDF, Hsurvmspline\_wane gives the cumulative hazard, hsurvmspline\_wane gives the hazard, dsurvmspline\_wane gives the PDF, qsurvmspline\_wane gives the quantiles, and rsurvmspline\_wane generates random numbers from the distribution.

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