## ${\bf Package\ 'bayes Cure Rate Model'}$

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Maintainer Panagiotis Papastamoulis <papapast@yahoo.gr></papapast@yahoo.gr>
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Author Panagiotis Papastamoulis [aut, cre] ( <a href="https://orcid.org/0000-0001-9468-7613">https://orcid.org/0000-0001-9468-7613</a> ), Fotios Milienos [aut] ( <a href="https://orcid.org/0000-0003-1423-7132">https://orcid.org/0000-0003-1423-7132</a> )
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Bayesian Cure Rate Modeling for Time-to-Event Data

#### **Description**

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A fully Bayesian approach in order to estimate a general family of cure rate models under the presence of covariates, see Papastamoulis and Milienos (2023) <doi:10.48550/arXiv.2310.06926>. The promotion time can be modelled (a) parametrically using typical distributional assumptions for time to event data (including the Weibull, Exponential, Gompertz, log-Logistic distributions), or (b) semiparametrically using finite mixtures of Gamma distributions. Posterior inference is carried out by constructing a Metropolis-coupled Markov chain Monte Carlo (MCMC) sampler, which combines Gibbs sampling for the latent cure indicators and Metropolis-Hastings steps with Langevin diffusion dynamics for parameter updates. The main MCMC algorithm is embedded within a parallel tempering scheme by considering heated versions of the target posterior distribution.

The main function of the package is cure\_rate\_MC3. See details for a brief description of the model.

#### **Details**

Let  $y = (y_1, \dots, y_n)$  denote the observed data, which correspond to time-to-event data or censoring times. Let also  $x_i = (x_{i1}, \dots, x_{x_{ip}})'$  denote the covariates for subject  $i, i = 1, \dots, n$ .

Assuming that the n observations are independent, the observed likelihood is defined as

$$L = L(\boldsymbol{\theta}; \boldsymbol{y}, \boldsymbol{x}) = \prod_{i=1}^{n} f_{P}(y_{i}; \boldsymbol{\theta}, \boldsymbol{x}_{i})^{\delta_{i}} S_{P}(y_{i}; \boldsymbol{\theta}, \boldsymbol{x}_{i})^{1-\delta_{i}},$$

where  $\delta_i = 1$  if the *i*-th observation corresponds to time-to-event while  $\delta_i = 0$  indicates censoring time. The parameter vector  $\boldsymbol{\theta}$  is decomposed as

$$\theta = (\alpha', \beta', \gamma, \lambda)$$

where

- $\alpha = (\alpha_1, \dots, \alpha_d)' \in \mathcal{A}$  are the parameters of the promotion time distribution whose cumulative distribution and density functions are denoted as  $F(\cdot, \alpha)$  and  $f(\cdot, \alpha)$ , respectively.
- $\beta \in \mathbf{R}^k$  are the regression coefficients with k denoting the number of columns in the design matrix (it may include a constant term or not).
- $\gamma \in \mathbf{R}$
- $\lambda > 0$ .

The population survival and density functions are defined as

$$S_P(y; \boldsymbol{\theta}) = \left(1 + \gamma \exp\{\boldsymbol{x}_i \boldsymbol{\beta}'\} c^{\gamma \exp\{\boldsymbol{x}_i \boldsymbol{\beta}'\}} F(y; \boldsymbol{\alpha})^{\lambda}\right)^{-1/\gamma}$$

whereas.

$$f_P(y; \boldsymbol{\theta}) = -\frac{\partial S_P(y; \boldsymbol{\theta})}{\partial y}.$$

Finally, the cure rate is affected through covariates and parameters as follows

$$p_0(\boldsymbol{x}_i; \boldsymbol{\theta}) = \left(1 + \gamma \exp\{\boldsymbol{x}_i \boldsymbol{\beta}'\} c^{\gamma \exp\{\boldsymbol{x}_i \boldsymbol{\beta}'\}}\right)^{-1/\gamma}$$

where  $c = e^{e^{-1}}$ .

The promotion time distribution can be a member of standard families (currently available are the following: Exponential, Weibull, Gamma, Lomax, Gompertz, log-Logistic) and in this case  $\alpha = (\alpha_1, \alpha_2) \in (0, \infty)^2$ . Also considered is the Dagum distribution, which has three parameters  $(\alpha_1, \alpha_2, \alpha_3) \in (0, \infty)^3$ . In case that the previous parametric assumptions are not justified, the promotion time can belong to the more flexible family of finite mixtures of Gamma distributions. For example, assume a mixture of two Gamma distributions of the form

$$f(y; \boldsymbol{\alpha}) = \alpha_5 f_{\mathcal{G}}(y; \alpha_1, \alpha_3) + (1 - \alpha_5) f_{\mathcal{G}}(y; \alpha_2, \alpha_4),$$

where

$$f_{\mathcal{G}}(y; \alpha, \beta) = \frac{\beta^{\alpha}}{\Gamma(\alpha)} y^{\alpha - 1} \exp\{-\beta y\}, y > 0$$

denotes the density of the Gamma distribution with parameters  $\alpha>0$  (shape) and  $\beta>0$  (rate). For the previous model, the parameter vector is

$$\alpha = (\alpha_1, \alpha_2, \alpha_3, \alpha_4, \alpha_5)' \in \mathcal{A}$$

where 
$$\mathcal{A} = (0, \infty)^4 \times (0, 1)$$
.

More generally, one can fit a mixture of K>2 Gamma distributions. The appropriate model can be selected according to information criteria such as the BIC.

The binary vector  $I = (I_1, \ldots, I_n)$  contains the (latent) cure indicators, that is,  $I_i = 1$  if the *i*-th subject is susceptible and  $I_i = 0$  if the *i*-th subject is cured.  $\Delta_0$  denotes the subset of  $\{1, \ldots, n\}$  containing the censored subjects, whereas  $\Delta_1 = \Delta_0^c$  is the (complementary) subset of uncensored subjects. The complete likelihood of the model is

$$L_c(\boldsymbol{\theta};\boldsymbol{y},\boldsymbol{I}) = \prod_{i \in \Delta_1} (1 - p_0(\boldsymbol{x}_i,\boldsymbol{\theta})) f_U(y_i;\boldsymbol{\theta},\boldsymbol{x}_i) \prod_{i \in \Delta_0} p_0(\boldsymbol{x}_i,\boldsymbol{\theta})^{1 - I_i} \{ (1 - p_0(\boldsymbol{x}_i,\boldsymbol{\theta})) S_U(y_i;\boldsymbol{\theta},\boldsymbol{x}_i) \}^{I_i}.$$

 $f_U$  and  $S_U$  denote the probability density and survival function of the susceptibles, respectively, that is

$$S_U(y_i; \boldsymbol{\theta}, \boldsymbol{x}_i) = \frac{S_P(y_i; \boldsymbol{\theta}, \boldsymbol{x}_i) - p_0(\boldsymbol{x}_i; \boldsymbol{\theta})}{1 - p_0(\boldsymbol{x}_i; \boldsymbol{\theta})}, f_U(y_i; \boldsymbol{\theta}, \boldsymbol{x}_i) = \frac{f_P(y_i; \boldsymbol{\theta}, \boldsymbol{x}_i)}{1 - p_0(\boldsymbol{x}_i; \boldsymbol{\theta})}.$$

## Index of help topics:

Summary method.

## Author(s)

Panagiotis Papastamoulis and Fotios S. Milienos

summary.bayesCureModel

Maintainer: Panagiotis Papastamoulis <papapast@yahoo.gr>

## References

Papastamoulis and Milienos (2023). Bayesian inference and cure rate modeling for event history data. arXiv:2310.06926

#### See Also

```
cure_rate_MC3
```

```
# TOY EXAMPLE (very small numbers... only for CRAN check purposes)
# simulate toy data
set.seed(1)
n = 4
stat = rbinom(n, size = 1, prob = 0.5)
x <- cbind(1, matrix(rnorm(n), n, 1))</pre>
```

```
y \leftarrow rexp(n)
# run a weibull model with default prior setup
# considering 2 heated chains
fit1 <- cure_rate_MC3(y = y, X = x, Censoring_status = stat,
promotion_time = list(distribution = 'weibull'),
nChains = 2,
nCores = 1,
mcmc\_cycles = 3, sweep=2)
# print method
fit1
# summary method
summary1 <- summary(fit1)</pre>
# WARNING: the following parameters
  mcmc_cycles, nChains
         should take _larger_ values. E.g. a typical implementation consists of:
#
         mcmc_cycles = 15000, nChains = 12
\# run a Gamma mixture model with K = 2 components and default prior setup
fit2 <- cure_rate_MC3(y = y, X = x, Censoring_status = stat,</pre>
promotion_time = list(
distribution = 'gamma_mixture',
        K = 2),
nChains = 8, nCores = 2,
mcmc\_cycles = 10)
summary2 <- summary(fit2)</pre>
```

complete\_log\_likelihood\_general

Logarithm of the complete log-likelihood for the general cure rate model.

#### **Description**

Compute the logarithm of the complete likelihood, given a realization of the latent binary vector of cure indicators  $I\_sim$  and current values of the model parameters g, lambda, b and promotion time parameters  $(\alpha)$  which yield log-density values (one per observation) stored to the vector  $log\_f$  and log-cdf values stored to the vector  $log\_F$ .

## Usage

```
complete_log_likelihood_general(y, X, Censoring_status,
g, lambda, log_f, log_F, b, I_sim, alpha)
```

# **Arguments**y

observed data (time-to-event or censored time)

X design matrix. Should contain a column of 1's if the model has a constant term. Censoring\_status

binary variables corresponding to time-to-event and censoring.

g The  $\gamma$  parameter of the model (real). lambda The  $\lambda$  parameter of the model (positive).

A vector containing the natural logarithm of the density function of the promotion time distribution per observation, for the current set of parameters. Its

length should be equal to the sample size.

log\_F A vector containing the natural logarithm of the cumulative density function of

the promotion time distribution per observation, for the current set of parame-

ters. Its length should be equal to the sample size.

b Vector of regression coefficients. Its length should be equal to the number of

columns of the design matrix.

I\_sim Binary vector of the current value of the latent cured status per observation. Its

length should be equal to the sample size. A time-to-event cannot be cured.

alpha A parameter between 0 and 1, corresponding to the temperature of the complete

posterior distribution.

#### **Details**

The complete likelihood of the model is

$$L_c(\boldsymbol{\theta}; \boldsymbol{y}, \boldsymbol{I}) = \prod_{i \in \Delta_1} (1 - p_0(\boldsymbol{x}_i, \boldsymbol{\theta})) f_U(y_i; \boldsymbol{\theta}, \boldsymbol{x}_i) \prod_{i \in \Delta_0} p_0(\boldsymbol{x}_i, \boldsymbol{\theta})^{1 - I_i} \{ (1 - p_0(\boldsymbol{x}_i, \boldsymbol{\theta})) S_U(y_i; \boldsymbol{\theta}, \boldsymbol{x}_i) \}^{I_i}.$$

 $f_U$  and  $S_U$  denote the probability density and survival function of the susceptibles, respectively, that is

$$S_U(y_i; \boldsymbol{\theta}, \boldsymbol{x}_i) = \frac{S_P(y_i; \boldsymbol{\theta}, \boldsymbol{x}_i) - p_0(\boldsymbol{x}_i; \boldsymbol{\theta})}{1 - p_0(\boldsymbol{x}_i; \boldsymbol{\theta})}, f_U(y_i; \boldsymbol{\theta}, \boldsymbol{x}_i) = \frac{f_P(y_i; \boldsymbol{\theta}, \boldsymbol{x}_i)}{1 - p_0(\boldsymbol{x}_i; \boldsymbol{\theta})}.$$

#### Value

A list with the following entries

cll the complete log-likelihood for the current parameter values.

logS Vector of logS values (one for each observation).

Vector of logP0 values (one for each observation).

## Author(s)

Panagiotis Papastamoulis

#### References

Papastamoulis and Milienos (2023). Bayesian inference and cure rate modeling for event history data. arXiv:2310.06926.

cure\_rate\_MC3 7

#### **Examples**

```
# simulate toy data
set.seed(1)
n = 4
stat = rbinom(n, size = 1, prob = 0.5)
x <- cbind(1, matrix(rnorm(n), n, 1))
y <- rexp(n)
lw <- log_weibull(y, a1 = 1, a2 = 1, c_under = 1e-9)
# compute complete log-likelihood
complete_log_likelihood_general(y = y, X = x,
Censoring_status = stat,
g = 1, lambda = 1,
log_f = lw$log_f, log_F = lw$log_F,
b = c(-0.5,0.5),
I_sim = stat, alpha = 1)</pre>
```

cure rate MC3

Main function of the package

#### **Description**

Runs a Metropolis Coupled MCMC (MC<sup>3</sup>) sampler in order to estimate the joint posterior distribution of the model.

#### **Usage**

```
cure_rate_MC3(y, X, Censoring_status, nChains = 12, mcmc_cycles = 15000,
alpha = NULL,nCores = 8, sweep = 5, mu_g = 1, s2_g = 1,
a_l = 2.1, b_l = 1.1, mu_b = rep(0, dim(X)[2]),
Sigma = 100 * diag(dim(X)[2]), g_prop_sd = 0.045,
lambda_prop_scale = 0.03, b_prop_sd = rep(0.022, dim(X)[2]),
initialValues = NULL, plot = TRUE, adjust_scales = FALSE,
verbose = FALSE, tau_mala = 1.5e-05, mala = 0.15,
promotion_time = list(distribution = "weibull",
prior_parameters = matrix(rep(c(2.1, 1.1), 2), byrow = TRUE, 2, 2),
prop_scale = c(0.1, 0.2)), single_MH_in_f = 0.2)
```

#### Arguments

y Observed data, that is, a vector of length n with positive entries.

X Design matrix with k > 1 columns. Should contain a column of 1's if the model has a constant term. Should be a matrix with dimension  $n \times k$ .

Censoring\_status

A vector  $\boldsymbol{\delta}=(\delta_1,\ldots,\delta_n)$  of binary variables corresponding to censoring indicators. The *i*-th observation is treated as a time-to-event if  $\delta_i=1$  or as a censoring time otherwise  $(\delta_i=0)$ .

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nChains	Positive integer corresponding to the number of heated chains in the $\mathrm{MC}^3$ scheme.
mcmc_cycles	Length of the generated MCMC sample. Default value: 15000. Note that each MCMC cycle consists of sweep (see below) usual MCMC iterations.
alpha	A decreasing sequence in $[1,0)$ of nChains temperatures (or heat values). The first value should always be equal to 1, which corresponds to the target posterior distribution (that is, the first chain).
nCores	The number of cores used for parallel processing.
sweep	The number of usual MCMC iterations per MCMC cycle. Default value: 10.
mu_g	Parameter $a_{\gamma}$ of the prior distribution of $\gamma$ .
s2_g	Parameter $b_{\gamma}$ of the prior distribution of $\gamma$ .
a_1	Shape parameter $a_{\lambda}$ of the Inverse Gamma prior distribution of $\lambda$ .
b_1	Scale parameter $b_{\lambda}$ of the Inverse Gamma prior distribution of $\lambda$ .
mu_b	Mean $(\mu)$ of the multivariate normal prior distribution of regression coefficients. Should be a vector whose length is equal to $k$ , i.e. the number of columns of the design matrix X. Default value: rep $(0, k)$ .
Sigma	Covariance matrix of the multivariate normal prior distribution of regression coefficients.
g_prop_sd	The scale of the proposal distribution for single-site updates of the $\gamma$ parameter.
lambda_prop_	scale
	The scale of the proposal distribution for single-site updates of the $\lambda$ parameter.
b_prop_sd	The scale of the proposal distribution for the update of the $\beta$ parameter (regression coefficients).
initialValue	
	A list of initial values for each parameter (optional).
plot	Plot MCMC sample on the run. Default: TRUE.
adjust_scale	
	Boolean. If TRUE the MCMC sampler runs an initial phase of a small number of iterations in order to tune the scale of the proposal distributions in the Metropolis-Hastings steps.
verbose	Print progress on the terminal if TRUE.
tau_mala	Scale of the Metropolis adjusted Langevin diffussion proposal distribution.
mala	A number between $[0,1]$ indicating the proportion of times the sampler attempts a MALA proposal. Thus, the probability of attempting a typical Metropolis-Hastings move is equal to 1 - mala.
promotion_ti	
	A list with details indicating the parametric family of distribution describing the promotion time and corresponding prior distributions. See 'details'.
single_MH_in	_f

The probability for attempting a series of single site updates in the typical Metropolis-Hastings move. Otherwise, with probability 1 - single\_MH\_in\_f a Metropolis-Hastings move will attempt to update all continuous parameters simultaneously.

It only makes sense when mala < 1.

cure\_rate\_MC3

#### **Details**

It is advised to scale all continuous explanatory variables in the design matrix, so their sample mean and standard deviations are equal to 0 and 1, respectively. The promotion\_time argument should be a list containing the following entries

distribution Character string specifying the family of distributions  $\{F(\cdot; \alpha); \alpha \in A\}$  describing the promotion time.

prior\_parameters Values of hyper-parameters in the prior distribution of the parameters  $\alpha$ . prop\_scale The scale of the proposal distributions for each parameter in  $\alpha$ .

dirichlet\_concentration\_parameter Relevant only in the case of the 'gamma\_mixture'. Positive scalar (typically, set to 1) determining the (common) concentration parameter of the Dirichlet prior distribution of mixing proportions.

The distribution entry should be one of the following: 'exponential', 'weibull', 'gamma', 'logLogistic', 'gompertz', 'lomax', 'dagum', 'gamma\_mixture'.

The joint prior distribution of  $\alpha=(\alpha_1,\ldots,\alpha_d)$  factorizes into products of inverse Gamma distributions for all (positive) parameters of F. Moreover, in the case of 'gamma\_mixture', the joint prior also consists of another term to the Dirichlet prior distribution on the mixing proportions.

The prop\_scale argument should be a vector with length equal to the length of vector d (number of elements in  $\alpha$ ), containing (positive) numbers which correspond to the scale of the proposal distribution. Note that these scale parameters are used only as initial values in case where adjust\_scales = TRUE.

#### Value

An object of class bayesCureModel, i.e. a list with the following entries

mcmc\_sample Object of class mcmc (see the **coda** package), containing the generated MCMC sample for the target chain. The column names correspond to

g\_mcmc Sampled values for parameter  $\gamma$ 

lambda\_mcmc Sampled values for parameter  $\lambda$ 

alphal\_mcmc... Sampled values for parameter  $\alpha_1$ ... of the promotion time distribution  $F(\cdot; \alpha_1, \ldots, \alpha_d)$ . The subsequent d-1 columns contain the sampled values for all remaining parameters,  $\alpha_2, \ldots, \alpha_d$ , where d depens on the family used in promotion\_time.

b0\_mcmc Sampled values for the constant term of the regression (present only in the case where the design matrix X contains a column of 1s).

b1\_mcmc... Sampled values for the regression coefficient for the first explanatory variable, and similar for all subsequent columns.

complete\_log\_likelihood

The complete log-likelihood for the target chain.

all\_cll\_values

The complete log-likelihood for all chains

latent\_status\_censored

A data frame with the simulated binary latent status for each censored item.

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#### Note

The core function is cure\_rate\_mcmc.

#### Author(s)

Panagiotis Papastamoulis

#### References

Papastamoulis and Milienos (2023). Bayesian inference and cure rate modeling for event history data. arXiv:2310.06926

#### See Also

```
cure_rate_mcmc
```

#### **Examples**

cure\_rate\_mcmc

The basic MCMC scheme.

## Description

This is core MCMC function. The continuous parameters of the model are updated using (a) single-site Metropolis-Hastings steps and (b) a Metropolis adjusted Langevin diffusion step. The binary latent variables of the model (cured status per censored observation) are updated according to a Gibbs step. This function is embedded to the main function of the package <a href="mailto:cure\_rate\_MC3">cure\_rate\_MC3</a> which runs parallel tempered MCMC chains.

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#### Usage

```
cure_rate_mcmc(y, X, Censoring_status, m, alpha = 1, mu_g = 1, s2_g = 1,
a_l = 2.1, b_l = 1.1, promotion_time = list(distribution = "weibull",
prior_parameters = matrix(rep(c(2.1, 1.1), 2), byrow = TRUE, 2, 2),
prop_scale = c(0.2, 0.03)), mu_b = NULL, Sigma = NULL, g_prop_sd = 0.045,
lambda prop scale = 0.03, b prop sd = NULL, initialValues = NULL,
plot = FALSE, verbose = FALSE, tau mala = 1.5e-05, mala = 0.15,
single MH in f = 0.5)
```

#### **Arguments**

observed data (time-to-event or censored time) У design matrix. Should contain a column of 1's if the model has a constant term. Χ Censoring\_status binary variables corresponding to time-to-event and censoring. number of MCMC iterations. m alpha A value between 0 and 1, corresponding to the temperature of the complete posterior distribution. The target posterior distribution corresponds to alpha = Parameter  $a_{\gamma}$  of the prior distribution of  $\gamma$ . mu\_g s2\_g Parameter  $b_{\gamma}$  of the prior distribution of  $\gamma$ . a\_l Shape parameter  $a_{\lambda}$  of the Inverse Gamma prior distribution of  $\lambda$ . b 1 Scale parameter  $b_{\lambda}$  of the Inverse Gamma prior distribution of  $\lambda$ . promotion\_time A list containing the specification of the promotion time distribution. See 'de-Mean  $\mu$  of the multivariate normal prior distribution of regression coefficients. mu\_b Should be a vector whose length is equal to the number of columns of the design matrix X. Covariance matrix of the multivariate normal prior distribution of regression Sigma coefficients. The scale of the proposal distribution for single-site updates of the  $\gamma$  parameter. q\_prop\_sd lambda\_prop\_scale The scale of the proposal distribution for single-site updates of the  $\lambda$  parameter. The scale of the proposal distribution for the update of the  $\beta$  parameter (regresb\_prop\_sd sion coefficients). initialValues A list of initial values for each parameter (optional). plot Boolean for plotting on the run.

Boolean for printing progress on the run. verbose

tau mala scale of the MALA proposal.

mala Propability of attempting a MALA step. Otherwise, a simple MH move is at-

tempted.

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```
single_MH_in_f
```

Probability of attempting a single-site MH move in the basic Metropolis-Hastings step. Otherwise, a joint update is attempted.

#### Value

A list containing the following entries

```
mcmc_sample The sampled MCMC values per parameter. See 'note'. complete log likelihood
```

Logarithm of the complete likelihood per MCMC iteration.

acceptance\_rates

The acceptance rate per move.

latent\_status\_censored

The MCMC sample of the latent status per censored observation.

log\_prior\_density

Logarithm of the prior density per MCMC iteration.

#### Note

In the case where the promotion time distribution is a Gamma mixture model, the mixing proportions  $w_1, \ldots, w_K$  are reparameterized according to the following transformation

$$w_j = \frac{\rho_j}{\sum_{i=1}^K \rho_i}, j = 1, \dots, K$$

where  $\rho_i > 0$  for  $i = 1, \dots, K-1$  and  $\rho_K = 1$ . The sampler returns the parameters  $\rho_1, \dots, \rho_{K-1}$ , not the mixing proportions.

#### Author(s)

Panagiotis Papastamoulis

#### References

Papastamoulis and Milienos (2023). Bayesian inference and cure rate modeling for event history data. arXiv:2310.06926

#### See Also

```
cure_rate_MC3
```

```
# simulate toy data just for cran-check purposes
    set.seed(1)
    n = 10
    stat = rbinom(n, size = 1, prob = 0.5)
    x <- cbind(1, matrix(rnorm(2*n), n, 2))
    y <- rexp(n)
# run a weibull model (including const. term)</pre>
```

log\_dagum 13

log\_dagum

PDF and CDF of the Dagum distribution

#### **Description**

The Dagum distribution as evaluated at the VGAM package.

#### Usage

```
log_dagum(y, a1, a2, a3, c_under = 1e-09)
```

#### **Arguments**

У	observed data
a1	scale parameter
a2	shape1.a parameter
a3	shape2.p parameter
c_under	A small positive value corresponding to the underflow threshold, e.g. c_under = 1e-9.

#### **Details**

The Dagum distribution is a special case of the 4-parameter generalized beta II distribution.

#### Value

A list containing the following entries

```
log_f natural logarithm of the pdf, evaluated at each datapoint.log_F natural logarithm of the CDF, evaluated at each datapoint.
```

#### Author(s)

Panagiotis Papastamoulis

log\_gamma

#### References

Thomas W. Yee (2015). Vector Generalized Linear and Additive Models: With an Implementation in R. New York, USA: Springer.

#### See Also

ddagum

#### **Examples**

```
log_dagum(y = 1:10, a1 = 1, a2 = 1, a3 = 1, c_under = 1e-9)
```

log\_gamma

PDF and CDF of the Gamma distribution

## Description

Computes the pdf and cdf of the Gamma distribution.

## Usage

```
log_gamma(y, a1, a2, c_under = 1e-09)
```

## **Arguments**

У	observed data
a1	shape parameter
a2	rate parameter
c_under	A small positive value corresponding to the underflow threshold, e.g. c_under = 1e-9.

#### Value

A list containing the following entries

```
log_fnatural logarithm of the pdf, evaluated at each datapoint.log_Fnatural logarithm of the CDF, evaluated at each datapoint.
```

## Author(s)

Panagiotis Papastamoulis

#### See Also

dgamma

```
log_gamma(y = 1:10, a1 = 1, a2 = 1, c_under = 1e-9)
```

log\_gamma\_mixture 15

```
log_gamma_mixture PDF and CDF of a Gamma mixture distribution
```

## Description

Computes the logarithm of the probability density function and cumulative density function per observation for each observation under a Gamma mixture model.

#### Usage

```
log_gamma_mixture(y, a1, a2, p, c_under = 1e-09)
```

## **Arguments**

У	observed data
a1	vector containing the shape parameters of each Gamma mixture component
a2	vector containing the rate parameters of each Gamma mixture component
р	vector of mixing proportions
c_under	threshold for underflows.

## Value

A list containing the following entries

```
log_f natural logarithm of the pdf, evaluated at each datapoint.log_F natural logarithm of the CDF, evaluated at each datapoint.
```

## Author(s)

Panagiotis Papastamoulis

```
y <- runif(10)
a1 <- c(1,2)
a2 <- c(1,1)
p <- c(0.9,0.1)
log_gamma_mixture(y, a1, a2, p)</pre>
```

log\_gompertz

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PDF and CDF of the Gompertz distribution

## Description

The Gompertz distribution as evaluated at the **flexsurv** package.

## Usage

```
log_gompertz(y, a1, a2, c_under = 1e-09)
```

## Arguments

У	observed data
a1	shape parameter
a2	rate parameter
c_under	A small positive value corresponding to the underflow threshold, e.g. c_under = 1e-9.

#### Value

A list containing the following entries

log\_f natural logarithm of the pdf, evaluated at each datapoint.log\_F natural logarithm of the CDF, evaluated at each datapoint.

## Author(s)

Panagiotis Papastamoulis

## References

Christopher Jackson (2016). flexsurv: A Platform for Parametric Survival Modeling in R. Journal of Statistical Software, 70(8), 1-33. doi:10.18637/jss.v070.i08

## See Also

```
dgompertz
```

```
log_gompertz(y = 1:10, a1 = 1, a2 = 1, c_under = 1e-9)
```

log\_logLogistic 17

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PDF and CDF of the log-Logistic distribution.

## Description

The log-Logistic distribution as evaluated at the **flexsurv** package.

## Usage

```
log_logLogistic(y, a1, a2, c_under = 1e-09)
```

## **Arguments**

У	observed data
a1	shape parameter
a2	scale parameter
c_under	A small positive value corresponding to the underflow threshold, e.g. c_under = 1e-9.

#### **Details**

The log-logistic distribution is the probability distribution of a random variable whose logarithm has a logistic distribution.

#### Value

A list containing the following entries

```
log_f natural logarithm of the pdf, evaluated at each datapoint.log_F natural logarithm of the CDF, evaluated at each datapoint.
```

#### Author(s)

Panagiotis Papastamoulis

#### References

Christopher Jackson (2016). flexsurv: A Platform for Parametric Survival Modeling in R. Journal of Statistical Software, 70(8), 1-33. doi:10.18637/jss.v070.i08

#### See Also

```
dllogis
```

```
log_log_log_istic(y = 1:10, a1 = 1, a2 = 1, c_under = 1e-9)
```

log\_lomax

loa	lomax

PDF and CDF of the Lomax distribution

## Description

The Lomax distribution as evaluated at the VGAM package.

## Usage

```
log_lomax(y, a1, a2, c_under = 1e-09)
```

## **Arguments**

У	observed data
a1	scale parameter
a2	shape parameter
c_under	A small positive value corresponding to the underflow threshold, e.g. c_under = 1e-9.

## **Details**

The Lomax distribution is a special case of the 4-parameter generalized beta II distribution.

## Value

A list containing the following entries

```
log_fnatural logarithm of the pdf, evaluated at each datapoint.log_Fnatural logarithm of the CDF, evaluated at each datapoint.
```

## Author(s)

Panagiotis Papastamoulis

## References

Thomas W. Yee (2015). Vector Generalized Linear and Additive Models: With an Implementation in R. New York, USA: Springer.

## See Also

dlomax

```
log_lomax(y = 1:10, a1 = 1, a2 = 1, c_under = 1e-9)
```

log\_weibull 19

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PDF and CDF of the Weibull distribution

## Description

Computes the log pdf and cdf of the weibull distribution.

## Usage

```
log_weibull(y, a1, a2, c_under)
```

## Arguments

У	observed data
a1	shape parameter
a2	rate parameter
c_under	A small positive value corresponding to the underflow threshold, e.g. c_under = 1e-9.

## Value

A list containing the following entries

```
log_f natural logarithm of the pdf, evaluated at each datapoint.log_F natural logarithm of the CDF, evaluated at each datapoint.
```

## Author(s)

Panagiotis Papastamoulis

```
log_weibull(y = 1:10, a1 = 1, a2 = 1, c_under = 1e-9)
```

20 plot.bayesCureModel

```
marriage_dataset Marriage data
```

## **Description**

The variable of interest (time) corresponds to the duration (in years) of first marriage for 1500 individuals. The available covariates are:

```
age age of respondent (in years) at the time of fist marriage.

kids factor: whether there were kids during the first marriage (1) or not (0).

race race of respondent decoded as: black (1), hispanic (2) and non-black/non-hispanic (4).
```

Among the 1500 observations, there are 1018 censoring times (censoring = 0) and 482 divorces (censoring = 1). Source: National Longitudinal Survey of Youth 1997 (NLSY97).

## Usage

```
data(marriage_dataset)
```

#### **Format**

Time-to-event data.

#### References

Bureau of Labor Statistics, U.S. Department of Labor. National Longitudinal Survey of Youth 1997 cohort, 1997-2022 (rounds 1-20). Produced and distributed by the Center for Human Resource Research (CHRR), The Ohio State University. Columbus, OH: 2023.

```
plot.bayesCureModel
```

Plot method

## Description

Plots and computes HDIs.

## Usage

```
## S3 method for class 'bayesCureModel'
plot(x, burn = NULL, alpha = 0.05, gamma_mix = TRUE,
K_gamma = 5, plot_graphs = TRUE, bw = "nrd0", what = NULL, p_cured_output = NULL,
index_of_main_mode = NULL,...)
```

plot.bayesCureModel 21

An object of class bayesCureModel

## **Arguments** Х

burn Number of iterations to discard as burn-in period. alpha A value between 0 and 1 in order to compute the 1- $\alpha$  Highest Posterior Density regions. Boolean. If TRUE, the density of the marginal posterior distribution of the  $\gamma$  pagamma\_mix rameter is estimated from the sampled MCMC values by fitting a normal mixture K\_gamma Used only when gamma\_mix = TRUE and corresponds to the number of normal mixture components used to estimate the marginal posterior density of the  $\gamma$  parameter. Boolean, if FALSE only HDIs are computed. plot\_graphs bandwidth bw Integer indicating which parameter should be plotted. If set to NULL (default), what all parameters are plotted one by one. If set to 'cured\_prob' the estimated cured probability is plotted, conditional on a set of covariates defined in the p\_cured\_output argument.

p\_cured\_output

Optional argument (list) which is required only when what = 'cured\_prob'. It is returned by the summary.bayesCureRateModel function.

index\_of\_main\_mode

If NULL (default), all modes are plotted. Otherwise, it is a subset of the retained MCMC iterations in order to identify the main mode of the posterior distribution, as returned by the index\_of\_main\_mode value of the summary.bayesCureRateModel function.

arguments passed by other methods.

#### Value

The function plots graphic output on the plot device if plot\_graphs = TRUE. Furthermore, a list of  $100(1-\alpha)\%$  Highest Density Intervals per parameter is returned.

#### Author(s)

Panagiotis Papastamoulis

```
# simulate toy data just for cran-check purposes
       set.seed(1)
       n = 4
        stat = rbinom(n, size = 1, prob = 0.5)
        # simulate design matrix
        # first column consists of 1s (const)
        # and second and third column contain
        # the values of two covariates
```

print.bayesCureModel

```
x \leftarrow cbind(1, matrix(rnorm(2*n), n, 2))
        colnames(x) <- c('const', 'x1', 'x2')</pre>
        y \leftarrow rexp(n)
fit1 <- cure_rate_MC3(y = y, X = x, Censoring_status = stat,
promotion_time = list(distribution = 'exponential'),
nChains = 2, nCores = 1,
mcmc\_cycles = 3, sweep = 2)
# plot the marginal posterior distribution of the first parameter in returned mcmc output
plot(fit1, what = 1, burn = 0)
# using 'cured_prob'
#compute cured probability for two individuals with
\# x1 = 0.2 \text{ and } x2 = -1
# and
\# x1 = -1 \text{ and } x2 = 0
covariate_levels1 <- rbind(c(1,0.2,-1), c(1,-1,0))
summary1 <- summary(fit1, covariate_levels = covariate_levels1, burn = 0)</pre>
plot(fit1, what='cured_prob', p_cured_output = summary1$p_cured_output,
  ylim = c(0,1))
```

print.bayesCureModel

Print method

#### **Description**

This function prints a summary of objects returned by the  ${\tt cure\_rate\_MC3}$  function.

#### Usage

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

## Arguments

x An object of class bayesCureModel, which is returned by the cure\_rate\_MC3 function.

... ignored.

#### **Details**

The function prints some basic information for a cure\_rate\_MC3, such as the MAP estimate of model parameters and the value of Bayesian information criterion.

## Value

No return value, called for side effects.

#### Author(s)

Panagiotis Papastamoulis

```
summary.bayesCureModel
```

Summary method.

## **Description**

This function produces all summaries after fitting a cure rate model.

## Usage

```
## S3 method for class 'bayesCureModel'
summary(object, burn = NULL, gamma_mix = TRUE,
K_gamma = 3, K_max = 3, fdr = 0.1,
covariate_levels = NULL, yRange = NULL, alpha = 0.1, ...)
```

## **Arguments**

object	An object of class bayesCureModel	
burn	Positive integer corresponding to the number of mcmc iterations to discard as burn-in period	
gamma_mix	Boolean. If TRUE, the density of the marginal posterior distribution of the $\gamma$ parameter is estimated from the sampled MCMC values by fitting a normal mixture model.	
K_gamma	Used only when gamma_mix = TRUE and corresponds to the number of normal mixture components used to estimate the marginal posterior density of the $\gamma$ parameter.	
K_max	Maximum number of components in order to cluster the (univariate) values of the joint posterior distribution across the MCMC run. Used to identify the main mode of the posterior distribution. See details.	
fdr	The target value for controlling the False Discovery Rate when classifying subjects as cured or not.	
covariate_levels		
	Optional levels for the covariates. It is only required when the user wishes to obtain a vector with the estimated posterior cured probabilities for a given combination of covariates. Include the value "1" in the case where the model contains constant term.	
yRange	Optional range (a vector of two non-negative values) for computing the sequence	

of posterior probabilities for the given values in covariate\_levels.

alpha Scalar between 0 and 1 corresponding to 1 - confidencel level for computing

Highest Density Intervals. If set to NULL, the confidence intervals are not com-

puted.

... ignored.

#### **Details**

The values of the posterior draws are clustered according to a (univariate) normal mixture model, and the main mode corresponds to the cluster with the largest mean. The maximum number of mixture components corresponds to the K\_max argument. The **mclust** library is used for this purpose. The inference for the latent cure status of each (censored) observation is based on the MCMC draws corresponding to the main mode of the posterior distribution. The FDR is controlled according to the technique proposed in Papastamoulis and Rattray (2018).

In case where covariate\_levels is set to TRUE, the summary function also returns a list named p\_cured\_output with the following entries

mcmc It is returned only in the case where the argument covariate\_values is not NULL. A vector of posterior cured probabilities for the given values in covariate\_values, per retained MCMC draw.

map It is returned only in the case where the argument covariate\_values is not NULL. The cured probabilities computed at the MAP estimate of the parameters, for the given values covariate values.

tau\_values tau values

covariate levels covariate levels

index\_of\_main\_mode the subset of MCMC draws allocated to the main mode of the posterior
distribution.

#### Value

A list with the following entries

map\_estimate Maximum A Posteriori (MAP) estimate of the parameters of the model.

highest\_density\_intervals

Highest Density Interval per parameter

latent\_cured\_status

Estimated posterior probabilities of the latent cure status per censored subject.

cured at given FDR

Classification as cured or not, at given FDR level.

p\_cured\_output

It is returned only in the case where the argument covariate\_values is not NULL. See details.

main mode index

The retained MCMC iterations which correspond to the main mode of the posterior distribution.

## Author(s)

Panagiotis Papastamoulis

#### References

Papastamoulis and Milienos (2023). Bayesian inference and cure rate modeling for event history data. arXiv:2310.06926.

Papastamoulis and Rattray (2018). A Bayesian Model Selection Approach for Identifying Differentially Expressed Transcripts from RNA Sequencing Data, Journal of the Royal Statistical Society Series C: Applied Statistics, Volume 67, Issue 1.

Scrucca L, Fraley C, Murphy TB, Raftery AE (2023). Model-Based Clustering, Classification, and Density Estimation Using mclust in R. Chapman and Hall/CRC. ISBN 978-1032234953

#### See Also

```
cure_rate_MC3
```

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