Package: BPrinStratTTE (via r-universe)

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Title Causal Effects in Principal Strata Defined by Antidrug Antibodies

Version 0.0.7

Description Bayesian models to estimate causal effects of biological treatments on time-to-event endpoints in clinical trials with principal strata defined by the occurrence of antidrug antibodies. The methodology is based on Frangakis and Rubin (2002) <doi:10.1111/j.0006-341x.2002.00021.x> and Imbens and Rubin (1997) <doi:10.1214/aos/1034276631>, and here adapted to a specific time-to-event setting.

License GPL (>= 3)

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BugReports https://github.com/Boehringer-Ingelheim/BPrinStratTTE/issues

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Contents

BPrinStratTTE-package 2
fit_mult_exp_covar 3
fit_mult_exp_nocovar 4
fit_single_exp_covar 6
fit_single_exp_nocovar 8
inv_logit 10
logit 11
ocs_exp_covar 12
ocs_exp_nocovar 14
run_sim_exp_covar 15
run_sim_exp_nocovar 17
sim_dat_mult_trials_exp_covar 18
sim_dat_mult_trials_exp_nocovar 19
sim_dat_one_trial_exp_covar 20
sim_dat_one_trial_exp_nocovar 22
true_vals_exp_covar 24
true_vals_exp_nocovar 25

Index **28**

`BPrinStratTTE-package` *The 'BPrinStratTTE' package.*

Description

Bayesian models to estimate causal effects of biological treatments on time-to-event endpoints in clinical trials with principal strata defined by the occurrence of antidrug antibodies. The methodology is based on Frangakis and Rubin (2002) doi:10.1111/j.0006-341x.2002.00021.x and Imbens and Rubin (1997) doi:10.1214/aos/1034276631, and intended to be applied to a specific time-to-event setting.#'

References

Stan Development Team (2022). RStan: the R interface to Stan. R package version 2.21.5. https://mc-stan.org

`fit_mult_exp_covar` *Fit multiple models to data from two-arm trials with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event*

Description

Fit multiple models to data from two-arm trials with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event

Usage

```
fit_mult_exp_covar(dat_mult_trials, params, seed = 23)
```

Arguments

`dat_mult_trials`	List generated by `sim_dat_mult_trials_exp_covar`.
`params`	List of model parameters as supplied to `fit_single_exp_covar`.
`seed`	Numeric value, seed for reproducibility.

Value

A list of objects generated by `fit_single_exp_covar`.

See Also

`sim_dat_mult_trials_exp_covar()`, `fit_single_exp_covar()`, `fit_mult_exp_nocovar()`

Examples

```
d_params_covar <- list(
  n = 1000,
  nt = 500,
  prob_X1 = 0.4,
  prob_ice_X1 = 0.5,
  prob_ice_X0 = 0.2,
  fu_max = 48*7,
  prop_cens = 0.15,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
 )
dat_mult_trials <- sim_dat_mult_trials_exp_covar(
  n_iter = 2,
  params = d_params_covar
)
m_params_covar <- list(
```

```
  tg = 48,
  p = 2,
  prior_delta = matrix(
    c(0, 5, 0, 5),
    nrow = 2, byrow = TRUE),
  prior_0N = c(1.5, 5),
  prior_1N = c(1.5, 5),
  prior_0T = c(1.5, 5),
  prior_1T = c(1.5, 5),
  t_grid =  seq(7, 7 * 48, 7) / 30,
  chains = 2,
  n_iter = 3000,
  warmup = 1500,
  cores = 2,
  open_progress = FALSE,
  show_messages = TRUE
)

fit_multiple <- fit_mult_exp_covar(
  dat_mult_trials = dat_mult_trials,
  params = m_params_covar,
  seed = 12
)
lapply(fit_multiple, dim)
head(fit_multiple[[1]])
```

`fit_mult_exp_nocovar` *Fit multiple models to data from two-arm trials with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event*

Description

Fit multiple models to data from two-arm trials with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event

Usage

```
fit_mult_exp_nocovar(dat_mult_trials, params, seed = 23)
```

Arguments

`dat_mult_trials`	List generated by `sim_dat_mult_trials_exp_nocovar`.
`params`	List of model parameters as supplied to `fit_single_exp_nocovar`.
`seed`	Numeric value, seed for reproducibility.

Value

A list of objects generated by `fit_single_exp_nocovar`.

See Also

`sim_dat_mult_trials_exp_nocovar()`, `fit_single_exp_nocovar()`, `fit_mult_exp_covar()`

Examples

```
d_params_nocovar <- list(
  n = 500L,
  nt = 250L,
  prob_ice = 0.5,
  fu_max = 336L,
  prop_cens = 0.15,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
)
dat_mult_trials <- sim_dat_mult_trials_exp_nocovar(
  n_iter = 2,
  params = d_params_nocovar
)
m_params_nocovar <- list(
  tg = 48L,
  prior_piT = c(0.5, 0.5),
  prior_0N = c(1.5, 5),
  prior_1N = c(1.5, 5),
  prior_0T = c(1.5, 5),
  prior_1T = c(1.5, 5),
  t_grid =  seq(7, 7 * 48, 7) / 30,
  chains = 2L,
  n_iter = 3000L,
  warmup = 1500L,
  cores = 2L,
  open_progress = FALSE,
  show_messages = TRUE
)

fit_multiple <- fit_mult_exp_nocovar(
  dat_mult_trials = dat_mult_trials,
  params = m_params_nocovar,
  seed = 12
)
lapply(fit_multiple, dim)
head(fit_multiple[[1]])
```

`fit_single_exp_covar` *Fit single model to data from a two-arm trial with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event*

Description

Fit single model to data from a two-arm trial with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event

Usage

```
fit_single_exp_covar(data, params, summarize_fit = TRUE)
```

Arguments

`data`	Data frame of a structure as generated by `sim_dat_one_trial_exp_covar()`.
`params`	List, containing model parameters:

- `tg` Positive integer value, number of intervals to calculate restricted mean survival time using the trapezoidal rule.
- `p` Positive integer value, number of predictors of the intercurrent event of interest (i.e. the event that determines. the principal stratum membership).
- `prior_delta` px2 matrix of positive numerical values, containing normal priors (mean and standard deviation) of the model parameter delta.
- `prior_0N` Numeric vector of length 2, containing parameters (alpha, beta) of the gamma prior on lambda_0N.
- `prior_1N` Numeric vector of length 2, containing parameters (alpha, beta) of the gamma prior on lambda_1N.
- `prior_0T` Numeric vector of length 2, containing parameters (alpha, beta) of the gamma prior on lambda_0T.
- `prior_1T` Numeric vector of length 2, containing parameters (alpha, beta) of the gamma prior on lambda_1T.
- `t_grid` Numeric vector of length `tg`, containing time points defining the time grid (in months) to calculate restricted mean survival time using the trapezoidal rule.
- `chains` Positive integer value, specifying the number of Markov chains.
- `n_iter` Positive integer value, specifying the number of iterations for each chain (including warmup).
- `warmup` Positive integer value, specifying the number of warmup (aka burnin) iterations per chain.
- `cores` Positive integer value, specifying the number of cores to use when executing the chains in parallel.
- `open_progress` Logical value, indicating whether the progress of the chains will be redirected to a file that is automatically opened for inspection.

- show_messages Logical value, indicating whether to print the summary of informational messages.

summarize_fit Logical, if TRUE (default), the output is restricted to a summary of results on key parameters over all chains, if FALSE, the complete stanfit object is returned.

Details

The data supplied as params are used either as priors (prior_delta, prior_0N, prior_1N, prior_1T), to inform the model setup (tg, p, t_grid), or as parameters to rstan::sampling() which is invoked internally (chains, n_iter, warmup, cores, open_progress, show_messages).

Value

tibble() containing a summary of results on key parameters, or a stanfit object (S4 class), depending on summarize_fit.

See Also

fit_single_exp_nocovar() and rstan::sampling()

Examples

```
d_params_covar <- list(
  n = 1000,
  nt = 500,
  prob_X1 = 0.4,
  prob_ice_X1 = 0.5,
  prob_ice_X0 = 0.2,
  fu_max = 48*7,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
 )
dat_single_trial <- sim_dat_one_trial_exp_covar(
  n = d_params_covar[["n"]],
  nt = d_params_covar[["nt"]],
  prob_X1 = d_params_covar[["prob_X1"]],
  prob_ice_X1 = d_params_covar[["prob_ice_X1"]],
  prob_ice_X0 = d_params_covar[["prob_ice_X0"]],
  fu_max = d_params_covar[["fu_max"]],
  T0T_rate = d_params_covar[["T0T_rate"]],
  T0N_rate = d_params_covar[["T0N_rate"]],
  T1T_rate = d_params_covar[["T1T_rate"]],
  T1N_rate = d_params_covar[["T1N_rate"]]
)
m_params_covar <- list(
  tg = 48,
  p = 2,
  prior_delta = matrix(
    c(0, 5, 0, 5),
    nrow = 2, byrow = TRUE),
```

```
  prior_0N = c(1.5, 5),
  prior_1N = c(1.5, 5),
  prior_0T = c(1.5, 5),
  prior_1T = c(1.5, 5),
  t_grid =  seq(7, 7 * 48, 7) / 30,
  chains = 2,
  n_iter = 3000,
  warmup = 1500,
  cores = 2,
  open_progress = FALSE,
  show_messages = FALSE
)

fit_single <- fit_single_exp_covar(
  data = dat_single_trial,
  params = m_params_covar,
  summarize_fit = FALSE
)
print(fit_single)
```

`fit_single_exp_nocovar` *Fit single model to data from a two-arm trial with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event*

Description

Fit single model to data from a two-arm trial with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event

Usage

```
fit_single_exp_nocovar(data, params, summarize_fit = TRUE)
```

Arguments

`data`	Data frame of a structure as generated by `sim_dat_one_trial_exp_nocovar()`.
`params`	List, containing model parameters: • `tg` Positive integer value, number of intervals to calculate restricted mean survival time using the trapezoidal rule. • `prior_piT` Numeric vector of length 2, containing parameters (alpha, beta) of the beta prior on pi, indicating the probability of belonging to the stratum of subjects developing the intercurrent event if given treatment. • `prior_0N` Numeric vector of length 2, containing parameters (alpha, beta) of the gamma prior on lambda_0N.

- prior_1N Numeric vector of length 2, containing parameters (alpha, beta) of the gamma prior on lambda_1N.
- prior_0T Numeric vector of length 2, containing parameters (alpha, beta) of the gamma prior on lambda_0T.
- prior_1T Numeric vector of length 2, containing parameters (alpha, beta) of the gamma prior on lambda_1T.
- t_grid Numeric vector of length tg, containing time points defining the time grid (in months) to calculate restricted mean survival time using the trapezoidal rule.
- chains Positive integer value, specifying the number of Markov chains.
- n_iter Positive integer value, specifying the number of iterations for each chain (including warmup).
- warmup Positive integer value, specifying the number of warmup (aka burnin) iterations per chain.
- cores Positive integer value, specifying the number of cores to use when executing the chains in parallel.
- open_progress Logical value, indicating whether the progress of the chains will be redirected to a file that is automatically opened for inspection.
- show_messages Logical value, indicating whether to print the summary of informational messages.

summarize_fit Logical, if TRUE (default), the output is restricted to a summary of results on key parameters over all chains, if FALSE, the complete stanfit object is returned.

Details

The data supplied as params are used either as priors (prior_delta, prior_0N, prior_1N, prior_1T), to inform the model setup (tg, p, t_grid), or as parameters to rstan::sampling() which is invoked internally (chains, n_iter, warmup, cores, open_progress, show_messages).

Value

tibble() containing a summary of results on key parameters, or a stanfit object, depending on summarize_fit.

See Also

fit_single_exp_covar() and rstan::sampling()

Examples

```
d_params_nocovar <- list(
  n = 500L,
  nt = 250L,
  prob_ice = 0.5,
  fu_max = 336L,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
```

```
  T1N_rate = 0.1
)
dat_single_trial <- sim_dat_one_trial_exp_nocovar(
  n = d_params_nocovar[["n"]],
  nt = d_params_nocovar[["nt"]],
  prob_ice = d_params_nocovar[["prob_ice"]],
  fu_max = d_params_nocovar[["fu_max"]],
  T0T_rate = d_params_nocovar[["T0T_rate"]],
  T0N_rate = d_params_nocovar[["T0N_rate"]],
  T1T_rate = d_params_nocovar[["T1T_rate"]],
  T1N_rate = d_params_nocovar[["T1N_rate"]]
)
m_params_nocovar <- list(
  tg = 48L,
  prior_piT = c(0.5, 0.5),
  prior_0N = c(1.5, 5),
  prior_1N = c(1.5, 5),
  prior_0T = c(1.5, 5),
  prior_1T = c(1.5, 5),
  t_grid =  seq(7, 7 * 48, 7) / 30,
  chains = 2L,
  n_iter = 3000L,
  warmup = 1500L,
  cores = 2L,
  open_progress = FALSE,
  show_messages = TRUE
)

fit_single <- fit_single_exp_nocovar(
  data = dat_single_trial,
  params = m_params_nocovar,
  summarize_fit = TRUE
)
print(fit_single)
```

`inv_logit` *Inverse logit function*

Description

Inverse logit function

Usage

```
inv_logit(x)
```

Arguments

`x`	Numeric value (usually a logarithm of odds).

Details

The inverse logit function is also known as logistic function.

Value

Numeric value on the interval $[0, 1]$, result of `log(pi/(1-pi))`.

Numeric value, result of `exp(x)/(1+exp(x))`.

See Also

`logit()`

Examples

```
# probabilities
prob_ICE_base <- 0.3
prob_ICE_risk <- 0.6
# model coefficients
(beta1 <- logit(prob_ICE_base))
(beta2 <- logit(prob_ICE_risk) - logit(prob_ICE_base))
# linear predictor
logit(prob_ICE_base); (lin_pred1 <- beta1 + beta2*0)
logit(prob_ICE_risk); (lin_pred2 <- beta1 + beta2*1)
# inverse logit of linear predictor
(inv_logit(lin_pred1)) # prob for X1 = 0
(inv_logit(lin_pred2)) # prob for X1 = 1
```

`logit` *Logit function*

Description

Logit function

Usage

```
logit(pi)
```

Arguments

`pi`	Numeric value on the interval $[0, 1]$ (usually a probability).

Value

Numeric value, result of `log(pi/(1-pi))`.

See Also

`inv_logit()`

Examples

```
# probabilities
prob_ICE_base <- 0.3
prob_ICE_risk <- 0.6
# model coefficients
(beta1 <- logit(prob_ICE_base))
(beta2 <- logit(prob_ICE_risk) - logit(prob_ICE_base))
# linear predictor
logit(prob_ICE_base); (lin_pred1 <- beta1 + beta2*0)
logit(prob_ICE_risk); (lin_pred2 <- beta1 + beta2*1)
# inverse logit of linear predictor
(inv_logit(lin_pred1)) # prob for X1 = 0
(inv_logit(lin_pred2)) # prob for X1 = 1
```

`ocs_exp_covar` *Determine operating characteristics of fits from two-arm trials with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event*

Description

Determine operating characteristics of fits from two-arm trials with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event

Usage

```
ocs_exp_covar(multiple_fits, d_params, m_params)
```

Arguments

`multiple_fits`	List of model fits from `fit_mult_exp_covar`.
`d_params`	List of data parameters as used in `sim_dat_one_trial_exp_covar`.
`m_params`	List of model parameters as used in `fit_single_exp_covar`.

Details

This function is used in `run_sim_exp_covar()`, the output of the two functions is the same.

Value

A list of length 3, containing objects call `ocs`, `d_params`, `m_params`, where `ocs` is a `tibble` containing averaged parameter estimates and operating characteristics, and `d_params` and `m_params` are the objects supplied to the function.

See Also

`ocs_exp_nocovar()` and `run_sim_exp_covar()`.

Examples

```
d_params_covar <- list(
  n = 1000,
  nt = 500,
  prob_X1 = 0.4,
  prob_ice_X1 = 0.5,
  prob_ice_X0 = 0.2,
  fu_max = 48*7,
  prop_cens = 0.15,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
 )
dat_mult_trials <- sim_dat_mult_trials_exp_covar(
  n_iter = 2,
  params = d_params_covar
)
m_params_covar <- list(
  tg = 48,
  p = 2,
  prior_delta = matrix(
    c(0, 5, 0, 5),
    nrow = 2, byrow = TRUE),
  prior_0N = c(1.5, 5),
  prior_1N = c(1.5, 5),
  prior_0T = c(1.5, 5),
  prior_1T = c(1.5, 5),
  t_grid =  seq(7, 7 * 48, 7) / 30,
  chains = 2,
  n_iter = 3000,
  warmup = 1500,
  cores = 2,
  open_progress = FALSE,
  show_messages = TRUE
)

fit_multiple <- fit_mult_exp_covar(
  dat_mult_trials = dat_mult_trials,
  params = m_params_covar,
  seed = 12
)
list_ocs <- ocs_exp_covar(
  multiple_fits = fit_multiple,
  d_params = d_params_covar,
  m_params = m_params_covar
)
print(list_ocs)
```

ocs_exp_nocovar *Determine operating characteristics of fits from two-arm trials with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event*

Description

Determine operating characteristics of fits from two-arm trials with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event

Usage

```
ocs_exp_nocovar(multiple_fits, d_params, m_params)
```

Arguments

multiple_fits	List of model fits from fit_mult_exp_nocovar.
d_params	List of data parameters as used in sim_dat_one_trial_exp_nocovar.
m_params	List of model parameters as used in fit_single_exp_nocovar.

Details

This function is used in run_sim_exp_nocovar(), the output of the two functions is the same.

Value

A list of length 3, containing objects call ocs, d_params, m_params, where ocs is a tibble containing averaged parameter estimates and operating characteristics, and d_params and m_params are the objects supplied to the function.

See Also

ocs_exp_covar() and run_sim_exp_nocovar().

Examples

```
d_params_nocovar <- list(
  n = 500L,
  nt = 250L,
  prob_ice = 0.5,
  fu_max = 336L,
  prop_cens = 0.15,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
)
dat_mult_trials <- sim_dat_mult_trials_exp_nocovar(
```

```
  n_iter = 2,
  params = d_params_nocovar
)
m_params_nocovar <- list(
  tg = 48L,
  prior_piT = c(0.5, 0.5),
  prior_0N = c(1.5, 5),
  prior_1N = c(1.5, 5),
  prior_0T = c(1.5, 5),
  prior_1T = c(1.5, 5),
  t_grid =  seq(7, 7 * 48, 7) / 30,
  chains = 2L,
  n_iter = 3000L,
  warmup = 1500L,
  cores = 2L,
  open_progress = FALSE,
  show_messages = TRUE
)

fit_multiple <- fit_mult_exp_nocovar(
  dat_mult_trials = dat_mult_trials,
  params = m_params_nocovar,
  seed = 12
)
list_ocs <- ocs_exp_nocovar(
  multiple_fits = fit_multiple,
  d_params = d_params_nocovar,
  m_params = m_params_nocovar
)
print(list_ocs)
```

`run_sim_exp_covar` *Run simulation of two-arm trials with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event*

Description

Run simulation of two-arm trials with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event

Usage

```
run_sim_exp_covar(n_iter, d_params, m_params, seed)
```

Arguments

`n_iter`	Positive integer value, number of trials to be simulated.
`d_params`	List of data parameters as used in `sim_dat_one_trial_exp_nocovar`.

m_params	List of model parameters as used in fit_single_exp_nocovar.
seed	Numeric value, seed for reproducibility.

Value

A list of length 3, containing objects call ocs, d_params, m_params, where ocs is a tibble containing averaged parameter estimates and operating characteristics, and d_params and m_params are the objects supplied to the function.

See Also

run_sim_exp_nocovar()

Examples

```
d_params_covar <- list(
  n = 1000,
  nt = 500,
  prob_X1 = 0.4,
  prob_ice_X1 = 0.5,
  prob_ice_X0 = 0.2,
  fu_max = 336L,
  prop_cens = 0.15,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
 )
m_params_covar <- list(
  tg = 48,
  p = 2,
  prior_delta = matrix(
    c(0, 5, 0, 5),
    nrow = 2, byrow = TRUE),
  prior_0N = c(1.5, 5),
  prior_1N = c(1.5, 5),
  prior_0T = c(1.5, 5),
  prior_1T = c(1.5, 5),
  t_grid =  seq(7, 7 * 48, 7) / 30,
  chains = 2,
  n_iter = 3000,
  warmup = 1500,
  cores = 2,
  open_progress = FALSE,
  show_messages = TRUE
)

dat_ocs <- run_sim_exp_covar(
  n_iter = 3,
  d_params = d_params_covar,
  m_params = m_params_covar,
  seed = 12
```

```
  )
print(dat_ocs)
```

`run_sim_exp_nocovar` *Run simulation of two-arm trials with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event*

Description

Run simulation of two-arm trials with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event

Usage

```
run_sim_exp_nocovar(n_iter, d_params, m_params, seed)
```

Arguments

`n_iter`	Positive integer value, number of trials to be simulated.
`d_params`	List of data parameters as used in `sim_dat_one_trial_exp_nocovar`.
`m_params`	List of model parameters as used in `fit_single_exp_nocovar`.
`seed`	Numeric value, seed for reproducibility.

Value

A list of length 3, containing objects call `ocs`, `d_params`, `m_params`, where `ocs` is a `tibble` containing averaged parameter estimates and operating characteristics, and `d_params` and `m_params` are the objects supplied to the function.

See Also

`run_sim_exp_covar()`

Examples

```
d_params_nocovar <- list(
  n = 500L,
  nt = 250L,
  prob_ice = 0.5,
  fu_max = 336L,
  prop_cens = 0.15,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
)
```

```
m_params_nocovar <- list(
  tg = 48L,
  prior_piT = c(0.5, 0.5),
  prior_0N = c(1.5, 5),
  prior_1N = c(1.5, 5),
  prior_0T = c(1.5, 5),
  prior_1T = c(1.5, 5),
  t_grid =  seq(7, 7 * 48, 7) / 30,
  chains = 2L,
  n_iter = 3000L,
  warmup = 1500L,
  cores = 2L,
  open_progress = FALSE,
  show_messages = TRUE
)

dat_ocs <- run_sim_exp_nocovar(
  n_iter = 3,
  d_params = d_params_nocovar,
  m_params = m_params_nocovar,
  seed = 12
  )
print(dat_ocs)
```

`sim_dat_mult_trials_exp_covar` *Simulate data from multiple two-arm trials with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event*

Description

Simulate data from multiple two-arm trials with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event

Usage

```
sim_dat_mult_trials_exp_covar(n_iter, params)
```

Arguments

`n_iter`	Positive integer value, number of trials to be simulated.
`params`	List of data parameters as used in `sim_dat_one_trial_exp_covar`.

Value

A list of length `n_iter`, containing objects of class `tibble()`, each containing one simulated trial dataset.

See Also

sim_dat_mult_trials_exp_nocovar()

Examples

```
d_params_covar <- list(
  n = 1000,
  nt = 500,
  prob_X1 = 0.4,
  prob_ice_X1 = 0.5,
  prob_ice_X0 = 0.2,
  fu_max = 336L,
  prop_cens = 0.15,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
 )
dat_mult_trials <- sim_dat_mult_trials_exp_covar(
  n_iter = 3L,
  params = d_params_covar
)
lapply(dat_mult_trials, dim)
head(dat_mult_trials[[1]])
```

---

sim_dat_mult_trials_exp_nocovar

Simulate data from multiple two-arm trials with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event

---

Description

Simulate data from multiple two-arm trials with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event

Usage

```
sim_dat_mult_trials_exp_nocovar(n_iter, params)
```

Arguments

n_iter	Positive integer value, number of trials to be simulated.
params	List of data parameters as used in sim_dat_one_trial_exp_nocovar.

Value

A list of length n_iter, containing objects of class tibble(), each containing one simulated trial dataset.

See Also

sim_dat_mult_trials_exp_covar()

Examples

```
d_params_nocovar <- list(
  n = 500L,
  nt = 250L,
  prob_ice = 0.5,
  fu_max = 336L,
  prop_cens = 0.15,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
)
dat_mult_trials <- sim_dat_mult_trials_exp_nocovar(
  n_iter = 3L,
  params = d_params_nocovar
)
lapply(dat_mult_trials, dim)
head(dat_mult_trials[[1]])
```

---

sim_dat_one_trial_exp_covar

Simulate data from a single two-arm trial with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event

---

Description

Simulate data from a single two-arm trial with an exponentially distributed time-to-event endpoint and one predictor of the intercurrent event

Usage

```
sim_dat_one_trial_exp_covar(
  n,
  nt,
  prob_X1,
  prob_ice_X1,
  prob_ice_X0,
  fu_max,
  prop_cens = 0,
  T0T_rate,
  T0N_rate,
  T1T_rate,
  T1N_rate
)
```

Arguments

n	Positive integer value, number of subjects in the trial.
nt	Positive integer value, number of treated subjects.
prob_X1	Numeric value on the interval $(0, 1)$, probability of being at high risk of experiencing the intercurrent event of interest when treated (i.e. the event that determines the principal stratum membership).
prob_ice_X1	Numeric value on the interval $(0, 1)$, probability of the intercurrent event of interest if treated and at high risk of the intercurrent event.
prob_ice_X0	Numeric value on the interval $(0, 1)$, probability of the intercurrent event of interest if treated and not at high risk of the intercurrent event.
fu_max	Positive integer value, maximum follow-up time in days (administrative censoring assumed afterwards).
prop_cens	Numeric value on the interval $[0, 1)$, proportion of uniformly censored patients (default is 0).
T0T_rate	Positive numeric value, monthly event rate in control subjects that would develop the intercurrent event if treated.
T0N_rate	Positive numeric value, monthly event rate in control subjects that never develop the intercurrent event.
T1T_rate	Positive numeric value, monthly event rate in treated subjects that develop the intercurrent event.
T1N_rate	Positive numeric value, monthly event rate in treated subjects that never develop the intercurrent event.

Value

...

See Also

sim_dat_one_trial_exp_nocovar()

Examples

```
d_params_covar <- list(
  n = 1000,
  nt = 500,
  prob_X1 = 0.4,
  prob_ice_X1 = 0.5,
  prob_ice_X0 = 0.2,
  fu_max = 48*7,
  prop_cens = 0.15,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
 )
dat_single_trial <- sim_dat_one_trial_exp_covar(
```

```
  n = d_params_covar[["n"]],
  nt = d_params_covar[["nt"]],
  prob_X1 = d_params_covar[["prob_X1"]],
  prob_ice_X1 = d_params_covar[["prob_ice_X1"]],
  prob_ice_X0 = d_params_covar[["prob_ice_X0"]],
  fu_max = d_params_covar[["fu_max"]],
  prop_cens = d_params_covar[["prop_cens"]],
  T0T_rate = d_params_covar[["T0T_rate"]],
  T0N_rate = d_params_covar[["T0N_rate"]],
  T1T_rate = d_params_covar[["T1T_rate"]],
  T1N_rate = d_params_covar[["T1N_rate"]]
)
dim(dat_single_trial)
head(dat_single_trial)
```

`sim_dat_one_trial_exp_nocovar` *Simulate data from a single two-arm trial with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event*

Description

Simulate data from a single two-arm trial with an exponentially distributed time-to-event endpoint and no predictor of the intercurrent event

Usage

```
sim_dat_one_trial_exp_nocovar(
  n,
  nt,
  prob_ice,
  fu_max,
  prop_cens = 0,
  T0T_rate,
  T0N_rate,
  T1T_rate,
  T1N_rate
)
```

Arguments

`n`	Positive integer value, number of subjects in the trial.
`nt`	Positive integer value, number of treated subjects.
`prob_ice`	Numeric value on the interval $(0, 1)$, probability of the intercurrent event of interest (i.e. the event that determines the principal stratum membership).

fu_max	Positive integer value, maximum follow-up time in days (administrative censoring assumed afterwards).
prop_cens	Numeric value on the interval $[0, 1)$, proportion of uniformly censored patients (default is 0).
T0T_rate	Positive numeric value, monthly event rate in control subjects that would develop the intercurrent event if treated.
T0N_rate	Positive numeric value, monthly event rate in control subjects that never develop the intercurrent event.
T1T_rate	Positive numeric value, monthly event rate in treated subjects that develop the intercurrent event.
T1N_rate	Positive numeric value, monthly event rate in treated subjects that never develop the intercurrent event.

Value

A `tibble()`containing the trial data for analysis.

See Also

`sim_dat_one_trial_exp_covar()`

Examples

```
d_params_nocovar <- list(
  n = 500L,
  nt = 250L,
  prob_ice = 0.5,
  fu_max = 336L,
  prop_cens = 0.15,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
)
dat_single_trial <- sim_dat_one_trial_exp_nocovar(
  n = d_params_nocovar[["n"]],
  nt = d_params_nocovar[["nt"]],
  prob_ice = d_params_nocovar[["prob_ice"]],
  fu_max = d_params_nocovar[["fu_max"]],
  prop_cens = d_params_nocovar[["prop_cens"]],
  T0T_rate = d_params_nocovar[["T0T_rate"]],
  T0N_rate = d_params_nocovar[["T0N_rate"]],
  T1T_rate = d_params_nocovar[["T1T_rate"]],
  T1N_rate = d_params_nocovar[["T1N_rate"]]
)
dim(dat_single_trial)
head(dat_single_trial)
```

`true_vals_exp_covar` *Adding true values to estimates for models with an exponential endpoint and consideration of predictors of the intercurrent event*

Description

Adding true values to estimates for models with an exponential endpoint and consideration of predictors of the intercurrent event

Usage

```
true_vals_exp_covar(x, d_params, m_params)
```

Arguments

`x`	Model object as returned by `fit_single_exp_covar()`.
`d_params`	List of data parameters as used in `fit_single_exp_covar()`.
`m_params`	List of model parameters as used in `fit_single_exp_covar()`.

Value

A summary table with parameter estimates, true values and differences.

See Also

`true_vals_exp_nocovar()`

Examples

```
d_params_covar <- list(
  n = 1000,
  nt = 500,
  prob_X1 = 0.4,
  prob_ice_X1 = 0.5,
  prob_ice_X0 = 0.2,
  fu_max = 48*7,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
 )
dat_single_trial <- sim_dat_one_trial_exp_covar(
  n = d_params_covar[["n"]],
  nt = d_params_covar[["nt"]],
  prob_X1 = d_params_covar[["prob_X1"]],
  prob_ice_X1 = d_params_covar[["prob_ice_X1"]],
  prob_ice_X0 = d_params_covar[["prob_ice_X0"]],
  fu_max = d_params_covar[["fu_max"]],
  T0T_rate = d_params_covar[["T0T_rate"]],
```

```
  T0N_rate = d_params_covar[["T0N_rate"]],
  T1T_rate = d_params_covar[["T1T_rate"]],
  T1N_rate = d_params_covar[["T1N_rate"]]
)
m_params_covar <- list(
  tg = 48,
  p = 2,
  prior_delta = matrix(
    c(0, 5, 0, 5),
    nrow = 2, byrow = TRUE),
  prior_0N = c(1.5, 5),
  prior_1N = c(1.5, 5),
  prior_0T = c(1.5, 5),
  prior_1T = c(1.5, 5),
  t_grid =  seq(7, 7 * 48, 7) / 30,
  chains = 2,
  n_iter = 3000,
  warmup = 1500,
  cores = 2,
  open_progress = FALSE,
  show_messages = TRUE
)

fit_single <- fit_single_exp_covar(
  data = dat_single_trial,
  params = m_params_covar,
  summarize_fit = TRUE
)
print(fit_single)
tab_obs_truth <- true_vals_exp_covar(
  x = fit_single,
  d_params = d_params_covar,
  m_params = m_params_covar
)
print(tab_obs_truth)
```

`true_vals_exp_nocovar` *Adding true values to estimates for models with an exponential endpoint and no consideration of predictors of the intercurrent event*

Description

Adding true values to estimates for models with an exponential endpoint and no consideration of predictors of the intercurrent event

Usage

```
true_vals_exp_nocovar(x, d_params, m_params)
```

Arguments

x	Model object as returned by `fit_single_exp_nocovar()`.
d_params	List of data parameters as used in `fit_single_exp_nocovar()`.
m_params	List of model parameters as used in `fit_single_exp_nocovar()`.

Value

A summary table with parameter estimates, true values and differences.

See Also

`true_vals_exp_covar()`

Examples

```
d_params_nocovar <- list(
  n = 500L,
  nt = 250L,
  prob_ice = 0.5,
  fu_max = 336L,
  T0T_rate = 0.2,
  T0N_rate = 0.2,
  T1T_rate = 0.15,
  T1N_rate = 0.1
)
dat_single_trial <- sim_dat_one_trial_exp_nocovar(
  n = d_params_nocovar[["n"]],
  nt = d_params_nocovar[["nt"]],
  prob_ice = d_params_nocovar[["prob_ice"]],
  fu_max = d_params_nocovar[["fu_max"]],
  T0T_rate = d_params_nocovar[["T0T_rate"]],
  T0N_rate = d_params_nocovar[["T0N_rate"]],
  T1T_rate = d_params_nocovar[["T1T_rate"]],
  T1N_rate = d_params_nocovar[["T1N_rate"]]
)
m_params_nocovar <- list(
  tg = 48L,
  prior_piT = c(0.5, 0.5),
  prior_0N = c(1.5, 5),
  prior_1N = c(1.5, 5),
  prior_0T = c(1.5, 5),
  prior_1T = c(1.5, 5),
  t_grid =  seq(7, 7 * 48, 7) / 30,
  chains = 2L,
  n_iter = 3000L,
  warmup = 1500L,
  cores = 2L,
  open_progress = FALSE,
  show_messages = TRUE
)
```

```
fit_single <- fit_single_exp_nocovar(
  data = dat_single_trial,
  params = m_params_nocovar,
  summarize_fit = TRUE
)
print(fit_single)
tab_obs_truth <- true_vals_exp_nocovar(
  x = fit_single,
  d_params = d_params_nocovar,
  m_params = m_params_nocovar
)
print(tab_obs_truth)
```

Index

BPrinStratTTE *(*BPrinStratTTE-package*)*, 2
BPrinStratTTE-package, 2

fit_mult_exp_covar, 3
fit_mult_exp_covar(), *5*
fit_mult_exp_nocovar, 4
fit_mult_exp_nocovar(), *3*
fit_single_exp_covar, 6
fit_single_exp_covar(), *3*, *9*
fit_single_exp_nocovar, 8
fit_single_exp_nocovar(), *5*, *7*

inv_logit, 10
inv_logit(), *11*

logit, 11
logit(), *11*

ocs_exp_covar, 12
ocs_exp_covar(), *14*
ocs_exp_nocovar, 14
ocs_exp_nocovar(), *12*

rstan::sampling(), *7*, *9*
run_sim_exp_covar, 15
run_sim_exp_covar(), *12*, *17*
run_sim_exp_nocovar, 17
run_sim_exp_nocovar(), *14*, *16*

sim_dat_mult_trials_exp_covar, 18
sim_dat_mult_trials_exp_covar(), *3*, *20*
sim_dat_mult_trials_exp_nocovar, 19
sim_dat_mult_trials_exp_nocovar(), *5*, *19*
sim_dat_one_trial_exp_covar, 20
sim_dat_one_trial_exp_covar(), *23*
sim_dat_one_trial_exp_nocovar, 22
sim_dat_one_trial_exp_nocovar(), *21*

true_vals_exp_covar, 24
true_vals_exp_covar(), *26*
true_vals_exp_nocovar, 25
true_vals_exp_nocovar(), *24*