

Package ‘rstanemax’

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Title Emax Model Analysis with 'Stan'

Description Perform sigmoidal Emax model fit using 'Stan' in a formula notation, without writing 'Stan' model code.

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ByteCompile true

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Suggests testthat, knitr, rmarkdown, spelling

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URL <https://github.com/yoshidk6/rstanemax>,
<https://yoshidk6.github.io/rstanemax/>

BugReports <https://github.com/yoshidk6/rstanemax/issues>

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rstanemax-package *The 'rstanemax' package.*

Description

Perform sigmoidal Emax model fit using Stan without writing Stan model code.

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- Trustees of Columbia University (src/init.cpp, tools/make_cc.R, R/stanmodels.R) [copyright holder]

References

Stan Development Team (2018). RStan: the R interface to Stan. R package version 2.18.2. <http://mc-stan.org>

See Also

Useful links:

- <https://github.com/yoshidk6/rstanemax>
- <https://yoshidk6.github.io/rstanemax/>
- Report bugs at <https://github.com/yoshidk6/rstanemax/issues>

as_draws	<i>Convert stanemax object to a posterior draws object</i>
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Description

Convert stanemax object to a posterior draws object

Usage

```
as_draws(x, ...)

as_draws_list(x, ...)

as_draws_array(x, ...)

as_draws_df(x, ...)

as_draws_matrix(x, ...)

as_draws_rvars(x, ...)

## S3 method for class 'stanemax'
as_draws(x, inc_warmup = FALSE, ...)

## S3 method for class 'stanemaxbin'
as_draws(x, inc_warmup = FALSE, ...)

## S3 method for class 'stanemax'
as_draws_list(x, inc_warmup = FALSE, ...)

## S3 method for class 'stanemaxbin'
as_draws_list(x, inc_warmup = FALSE, ...)

## S3 method for class 'stanemax'
as_draws_array(x, inc_warmup = FALSE, ...)

## S3 method for class 'stanemaxbin'
as_draws_array(x, inc_warmup = FALSE, ...)

## S3 method for class 'stanemax'
as_draws_df(x, inc_warmup = FALSE, ...)

## S3 method for class 'stanemaxbin'
as_draws_df(x, inc_warmup = FALSE, ...)

## S3 method for class 'stanemax'
as_draws_matrix(x, inc_warmup = FALSE, ...)
```

```
## S3 method for class 'stanemaxbin'
as_draws_matrix(x, inc_warmup = FALSE, ...)

## S3 method for class 'stanemax'
as_draws_rvars(x, inc_warmup = FALSE, ...)

## S3 method for class 'stanemaxbin'
as_draws_rvars(x, inc_warmup = FALSE, ...)
```

Arguments

- `x` An object of class stanemax.
- `...` Arguments passed to individual methods (if applicable).'
- `inc_warmup` Should warmup draws be included? Defaults to FALSE.

Value

A draws object of the appropriate subclass

See Also

[draws](#) [subset_draws](#)

Examples

```
## Not run:
data(exposure.response.sample)
fit <- stan_emax(response ~ exposure, exposure.response.sample)
posterior::as_draws_list(fit)
posterior::as_draws_array(fit)
posterior::as_draws_df(fit)
posterior::as_draws_matrix(fit)
posterior::as_draws_rvars(fit)

## End(Not run)
```

exposure.response.sample

Sample simulated data for exposure-response.

Description

Sample simulated data for exposure-response.

Usage

`exposure.response.sample`

Format

A data frame with columns:

dose Dose levels used for simulation of pharmacokinetics

exposure Simulated exposure

response Simulated pharmacodynamic response

Examples

```
exposure.response.sample
```

```
exposure.response.sample.binary
```

Sample simulated data for exposure-response for binary endpoint

Description

Sample simulated data for exposure-response for binary endpoint

Usage

```
exposure.response.sample.binary
```

Format

A data frame with columns:

conc Simulated exposure

y_logit Simulated logit(p)

y_prob Simulated p

y Simulated event (1 or 0)

y_cov_logit Simulated logit(p) for model with covariate

y_cov_prob Simulated p for model with covariate

y_cov Simulated event (1 or 0) for model with covariate

sex_num 1 or 0

sex female or male

Examples

```
exposure.response.sample
```

`exposure.response.sample.with.cov`

Sample simulated data for exposure-response with covariates

Description

Sample simulated data for exposure-response with covariates

Usage

`exposure.response.sample.with.cov`

Format

A data frame with columns:

- dose** Dose levels used for simulation of pharmacokinetics
- conc** Simulated exposure
- resp** Simulated pharmacodynamic response
- cov1** Covariate 1 for e0
- cov2** Covariate 2 for emax
- cov3** Covariate 3 for ec50 (data type factor)
- cov3num** Covariate 3 for ec50 (data type numeric)

Examples

`exposure.response.sample`

`extract_param`

Extract posterior draws of key parameters

Description

Extract posterior draws of key parameters

Usage

`extract_param(object)`

Arguments

<code>object</code>	A stanemax class object
---------------------	-------------------------

Value

A tibble containing posterior draws of key parameters. If covariate(s) are included in the model, posterior draws for different combinations of covariates are supplied in a long format - e.g. if there are posterior draws of 100 samples and 4 levels of the covariates, the returned tibble will have the length of 400

posterior_predict *Outcome prediction from posterior distribution of parameters*

Description

Compute outcome predictions using posterior samples. Exposure data for prediction can be either original data used for model fit or new data.

Usage

```
posterior_predict(object, ...)

## S3 method for class 'stanemax'
posterior_predict(
  object,
  newdata = NULL,
  returnType = c("matrix", "dataframe", "tibble"),
  newDataType = c("raw", "modelframe"),
  ...
)

## S3 method for class 'stanemaxbin'
posterior_predict(
  object,
  newdata = NULL,
  returnType = c("matrix", "dataframe", "tibble"),
  newDataType = c("raw", "modelframe"),
  ...
)

posterior_predict_quantile(
  object,
  newdata = NULL,
  ci = 0.9,
  pi = 0.9,
  newDataType = c("raw", "modelframe")
)
```

Arguments

<code>object</code>	A <code>stanemax</code> class object
<code>...</code>	Additional arguments passed to methods.
<code>newdata</code>	An optional data frame that contains columns needed for model to run (exposure and covariates). If the model does not have any covariate, this can be a numeric vector corresponding to the exposure metric.
<code>returnType</code>	An optional string specifying the type of return object.
<code>newDataType</code>	An optional string specifying the type of newdata input, whether in the format of an original data frame or a processed model frame. Mostly used for internal purposes and users can usually leave at default.
<code>ci</code>	Credible interval of the response without residual variability.
<code>pi</code>	Prediction interval of the response with residual variability.

Details

Run `vignette("emaxmodel", package = "rstanemax")` to see how you can use the posterior prediction for plotting estimated Emax curve.

Value

An object that contain predicted response with posterior distribution of parameters. The default is a matrix containing predicted response for `stan_emax()` and `.epred` for `stan_emax_binary()`. Each row of the matrix is a vector of predictions generated using a single draw of the model parameters from the posterior distribution.

If either `dataframe` or `tibble` is specified, the function returns a data frame or tibble object in a long format - each row is a prediction generated using a single draw of the model parameters and a corresponding exposure.

Several types of predictions are generated with this function.

For continuous endpoint model (`stan_emax()`),

- `.linpred & .epred`: prediction without considering residual variability and is intended to provide credible interval of "mean" response.
- `.prediction`: include residual variability in its calculation, therefore the range represents prediction interval of observed response.
- (deprecated) `respHat`: replaced by `.linpred & .epred`
- (deprecated) `response`: replaced by `.prediction`

For binary endpoint model (`stan_emax_binary()`),

- `.linpred`: predicted probability on logit scale
- `.epred`: predicted probability on probability scale
- `.prediction`: predicted event (1) or non-event (0)

The return object also contains exposure and parameter values used for calculation.

With `posterior_predict_quantile()` function, you can obtain quantiles of `respHat` and `response` as specified by `ci` and `pi`.

Description

Methods for stanemax objects

Usage

```
## S3 method for class 'stanemax'  
print(x, digits_summary = 2, ...)  
  
## S3 method for class 'stanemaxbin'  
print(x, digits_summary = 2, ...)  
  
extract_stanfit(x)  
  
extract_obs_mod_frame(x)  
  
## S3 method for class 'stanemax'  
plot(x, show.ci = TRUE, show.pi = FALSE, ci = 0.9, pi = 0.9, ...)  
  
## S3 method for class 'stanemaxbin'  
plot(x, show.ci = TRUE, show.pi = FALSE, ci = 0.9, pi = 0.9, ...)
```

Arguments

- x An object of class `stanemax` or `stanemaxbin`
- digits_summary The number of significant digits to use when printing the summary, defaulting to 2. Applies to the quantities other than the effective sample size, which is always rounded to the nearest integer.
- ... Additional arguments passed to methods.
- show.ci An logical specifying if the output figure include credible interval of posterior prediction. Default TRUE.
- show.pi An logical specifying if the output figure include prediction interval. Default FALSE.
- ci Credible interval range.
- pi Prediction interval range.

stan_emax*Bayesian Emax model fit with Stan*

Description

Run sigmoidal Emax model fit with formula notation

Usage

```
stan_emax(
  formula,
  data,
  gamma.fix = 1,
  e0.fix = NULL,
  emax.fix = NULL,
  priors = NULL,
  param.cov = NULL,
  ...
)
```

Arguments

<code>formula</code>	a symbolic description of variables for Emax model fit.
<code>data</code>	an optional data frame containing the variables in the model.
<code>gamma.fix</code>	a (positive) numeric or NULL to specify gamma (Hill coefficient) in the sigmoidal Emax model. If NULL, gamma will be estimated from the data. If numeric, gamma is fixed at the number provided. Default = 1 (normal Emax model).
<code>e0.fix</code>	a numeric or NULL to specify E0 in the Emax model. If NULL, E0 will be estimated from the data. If numeric, E0 is fixed at the number provided. Default = NULL (estimate from the data).
<code>emax.fix</code>	a numeric or NULL to specify Emax in the Emax model. If NULL, Emax will be estimated from the data. If numeric, Emax is fixed at the number provided. Default = NULL (estimate from the data).
<code>priors</code>	a named list specifying priors of parameters (ec50, emax, e0, gamma, sigma). Each list item should be length 2 numeric vector, one corresponding to mean and another corresponding to standard deviation. Currently only supports normal distribution for priors.
<code>param.cov</code>	a named list specifying categorical covariates on parameters (ec50, emax, e0). Convert a column into factor if specific order of covariates are needed.
<code>...</code>	Arguments passed to rstan::sampling (e.g. iter, chains).

Details

The following structure is used for the Emax model:

$$\text{Response} = e_0 + e_{\max} \times \text{exposure}^{\gamma} / (\text{ec50}^{\gamma} + \text{exposure}^{\gamma}) + \epsilon$$

$$\epsilon \sim N(0, \sigma^2)$$

Value

An object of class `stanemax`

Examples

```
## Not run:
data(exposure.response.sample)
fit1 <- stan_emax(response ~ exposure,
  data = exposure.response.sample,
  # the next line is only to make the example go fast enough
  chains = 1, iter = 500, seed = 12345
)
print(fit1)

# Set priors manually, also estimate gamma instead of the default of fix to 1
fit2 <- stan_emax(response ~ exposure,
  data = exposure.response.sample, gamma.fix = NULL,
  priors = list(
    ec50 = c(100, 30), emax = c(100, 30), e0 = c(10, 5),
    gamma = c(0, 3), sigma = c(0, 30)
  ),
  # the next line is only to make the example go fast enough
  chains = 1, iter = 500, seed = 12345
)
print(fit2)

data(exposure.response.sample.with.cov)
# Specify covariates
fit3 <- stan_emax(
  formula = resp ~ conc, data = exposure.response.sample.with.cov,
  param.cov = list(emax = "cov2", ec50 = "cov3", e0 = "cov1"),
  # the next line is only to make the example go fast enough
  chains = 1, iter = 500, seed = 12345
)
print(fit3)

## End(Not run)
```

`stan_emax_binary`*Bayesian Emax model fit with Stan for binary endpoint*

Description

Bayesian Emax model fit with Stan for binary endpoint

Usage

```
stan_emax_binary(
  formula,
  data,
  gamma.fix = 1,
  e0.fix = NULL,
  emax.fix = NULL,
  priors = NULL,
  param.cov = NULL,
  ...
)
```

Arguments

<code>formula</code>	a symbolic description of variables for Emax model fit.
<code>data</code>	an optional data frame containing the variables in the model.
<code>gamma.fix</code>	a (positive) numeric or NULL to specify gamma (Hill coefficient) in the sigmoidal Emax model. If NULL, gamma will be estimated from the data. If numeric, gamma is fixed at the number provided. Default = 1 (normal Emax model).
<code>e0.fix</code>	a numeric or NULL to specify E0 in the Emax model. If NULL, E0 will be estimated from the data. If numeric, E0 is fixed at the number provided. Default = NULL (estimate from the data).
<code>emax.fix</code>	a numeric or NULL to specify Emax in the Emax model. If NULL, Emax will be estimated from the data. If numeric, Emax is fixed at the number provided. Default = NULL (estimate from the data).
<code>priors</code>	a named list specifying priors of parameters (ec50, emax, e0, gamma, sigma). Each list item should be length 2 numeric vector, one corresponding to mean and another corresponding to standard deviation. Currently only supports normal distribution for priors.
<code>param.cov</code>	a named list specifying categorical covariates on parameters (ec50, emax, e0). Convert a column into factor if specific order of covariates are needed.
<code>...</code>	Arguments passed to rstan::sampling (e.g. iter, chains).

Examples

```
## Not run:  
data(exposure.response.sample.binary)  
fit1 <- stan_emax_binary(  
  y ~ conc,  
  data = exposure.response.sample.binary,  
  # the next line is only to make the example go fast enough  
  chains = 2, iter = 500, seed = 12345  
)  
print(fit1)  
  
# Specify covariates  
fit2 <- stan_emax_binary(  
  formula = y ~ conc, data = exposure.response.sample.binary,  
  param.cov = list(emax = "sex"),  
  # the next line is only to make the example go fast enough  
  chains = 2, iter = 500, seed = 12345  
)  
print(fit2)  
  
## End(Not run)
```

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