# Package 'PStrata'

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Title Principal Stratification Analysis in R

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Description Estimating causal effects in the presence of post-treatment confounding using principal stratification. 'PStrata' allows for customized monotonicity assumptions and exclusion restriction assumptions, with automatic full Bayesian inference supported by 'Stan'. The main function to use in this package is PStrata(), which provides posterior estimates of principal causal effect with uncertainty quantification. Visualization tools are also provided for diagnosis and interpretation. See Liu and Li (2023) <arXiv:2304.02740> for details.

**Depends** R (>= 3.5.0)

Collate PStrata-package.R PSFormula.R PStrataInfo.R survival.R prior.R make\_standata.R PSObject.R make\_stancode.R PSSample.R PStrata.R PSOutcome.R sim\_data\_normal.R sim\_data\_Cox.R PSContrast.R

**License** GPL (>= 2)

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## **Description**

The **PStrata** package is designed for estimating causal effects in the presense of post-treatment confounding using principal stratification. It provides an interface to fit the Bayesian principal stratification model, which is a complex mixture model, using **Stan**, a C++ package for obtaining full Bayesian inference. The formula syntax is an extended version of the syntax applied in many regression functions and packages, such as **lm**, **glm** and **lme4**, to provide a simple interface. A wide variety of distributions and link functions are supported, allowing users to fit linear, binary or count data, and survival models with principal stratification. Further modeling options include multiple post-treatment confounding variables and cluster random effects. The monotonicity and exclusion restriction assumptions can be easily applied, and prior specifications are flexible and encourage users to reflect their prior belief. In addition, all parameters can be inferred from the posterior distribution, which enables further analysis other than provided by the package. A frequentist weighting-based triply-robust estimator is also implemented for both ordinary outcomes and survival outcomes.

### **Details**

The Bayesian principal stratification analysis relies on two models, the principal stratum model and the outcome model. The main function of **PStrata** is PStrata, which uses formula syntax to specify these models. Based on the supplied formulas, data and additional information allowing users to specify assumptions and prior distributions, it automatically generates the Stan code via make\_stancode and make\_standata, and fits the model using **Stan**.

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The estimated probability for each principal stratum and the estimated mean response are calculated with **Stan** as it is faster and more space-efficient. However, a large number of post-processing methods can also be applied. summary is perfectly suited for an overview of the estimated parameters, and plot provides visualization of the principal stratification and the outcome distribution.

Because **PStrata** heavily relies on **Stan** for posterior sampling, a C++ compiler is required. The program **Rtools** (available on https://cran.r-project.org/bin/windows/Rtools/) comes with a C++ compiler for Windows. On Mac, Xcode is suggested. For further instructions on how to get the compilers running, please refer to the prerequisites section at the **RStan-Getting-Started** page.

#### References

The Stan Development Team. Stan Modeling Language User's Guide and Reference Manual. https://mc-stan.org/users/documentation/

Stan Development Team (2020). RStan: the R interface to Stan. R package version 2.21.2. https://mc-stan.org/

#### See Also

**PStrata** 

make\_stancode

Stan Code for PStrata Models

## Description

Generate the **Stan** code corresponding to the model, which is read by **Stan** to do sampling.

## Usage

```
make_stancode(PSobject, filename = NULL, debug = FALSE)
```

## Arguments

PSobject an object of class PSobject

filename (optional) string. If not NULL, the stan file will be saved via cat in a text file

named after the string supplied.

debug only for testing in development mode. Will be removed in future release.

#### Value

A string, which can be printed on screen using cat.

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make\_standata

Data for **PStrata** Models

## **Description**

Generate data for PStrata models to be passed to Stan

## Usage

```
make_standata(PSobject)
```

## **Arguments**

PSobject

an object of class PSObject

#### Value

a named list of objects containing the required data to fit a PStrata model with Stan.

prior

Prior functions

## Description

Define prior functions used in PStrata.

## Usage

```
prior_flat()
prior_normal(mu = 0, sigma = 1)
prior_t(mu = 0, sigma = 1, df = 1)
prior_cauchy(mu = 0, sigma = 1)
prior_lasso(mu = 0, sigma = 1)
prior_logistic(mu = 0, sigma = 1)
prior_chisq(df = 1)
prior_inv_chisq(df = 1)
prior_exponential(beta = 1)
```

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```
prior_gamma(alpha = 1, beta = 1)
prior_inv_gamma(alpha = 1, beta = 1)
prior_weibull(alpha = 1, sigma = 1)
```

## **Arguments**

```
mu, sigma, df, alpha, beta parameters for the prior distribution
```

## Value

A list, including the following items.

name name of the distribution

type type of the distribution, one character string of "real" or "positive"

args a named list of all the input parameters

call a function call object of the prior distribution on the parameters

**PSContrast** 

Contrast of potential outcome for principal stratification analysis

## **Description**

Create an object that represents contrast of potential outcomes by treatment arms, strata or time points.

## Usage

```
PSContrast(
  outcome,
  S = NULL,
  Z = NULL,
  T = NULL,
  type = c("all", "sequential", "cycle")
)
```

## Arguments

outcome	an object of class PSoutcome or PSContrast
S	a vector denoting which strata to take contrasts. Default is NULL indicating no contrasts are taken. Set to 'TRUE' to take contrasts between all strata.
Z	a vector denoting which treatment arms to take contrasts. Default is NULL indicating no contrasts are taken. Set to 'TRUE' to take contrasts between all treatment arms.

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T a vector denoting which time points to take contrasts. Default is NULL indicating

no contrasts are taken. Set to 'TRUE' to take contrasts between all time points.

This is used only when 'object' is obtained under survival outcome.

type Either "all" (default), "sequential" or "cycle". If "all", every pairwise

contrasts are taken. If "sequential", contrasts are taken over every consecutive pairs. If "cycle", contrasts are taken over every consecutive pairs and also

between the first and the last levels.

#### Value

An S3 object of class PSContrast and PSOutcome, containing

outcome\_array A num\_strata \* num\_treatment \* num\_iter array of contrast if the outcome type

is non-survival or a num\_strata \* num\_treatment \* num\_time\_points \* num\_iter

array of contrast if the outcome type is survival.

is.survival A boolean value, whether the outcome type is survival.

time\_points The time points at which the outcome is evaluated, if the outcome type is sur-

vival.

The S3 method summary and plot can be applied to the returned object.

PSFormula	Set up a model formula for use in <b>PStrata</b>
-----------	--

## Description

Set up a model formula for use in **PStrata** package allowing users to specify the treatment indicator, the post-randomization confounding variables, the outcome variable, and possibly the covariates. For survival outcome, a censoring indicator is also specified. Users can also define (potentially non-linear) transforms of the covariates and include random effects for clusters.

## Usage

PSFormula(formula, data)

## Arguments

formula an object of class formula (or one that can be coerced to that class): a symbolic

description of the model to be fitted. The details of model specification are given

in 'Details'.

data a data frame containing the variables named in formula.

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#### **Details**

Two models are required for the principal stratification analysis: the principal stratum model and the outcome model.

**General formula structure:** For the principal stratum model, the formula argument accepts formulas of the following syntax:

treatment + postrand ~ terms

The treatment variable refers to the name of the binary treatment indicator. The postrand variable refers to the name of the binary post-randomization confounding variable. The terms part includes all of the predictors used for the principal stratum model.

For the outcome model, the formula argument accepts formulas of the similar syntax:

response [+ observed] ~ terms

The response variable refers to the name of the outcome variable. The terms part includes all of the predictors used for the outcome model. The observed variable shall not be used for ordinary response. When the true response is subject to right censoring (also called survival outcome in relevant literature), the response variable should refer to the observed or censored response, and the observed variable should be an indicator of whether the true response is observed. For example, suppose the true time for an event is T and the time of censoring is T, Then, the response variable should refer to T0, the actual time of the event or censoring, whichever comes earlier, and the indicator observed is 1 if T1 of T2 and 0 otherwise.

The terms specified in the principal stratum model and the outcome model can be different.

**Multiple post-randomization confounding variables:** If multiple post-randomization confounding variables exist, one can specify all of them using the following syntax:

treatment + postrand\_1 + postrand\_2 + ... + postrand\_n ~ terms

The post-randomization confounding variables are provided in place of postrand\_1 to postrand\_n. Up to this version, all of these variables should be binary indicators. Note that the order of these post-randomization confounding variables will not affect the result of the estimation of the parameters, but it will be important in specifying other parameters, such as strata and ER (see PStrata).

**Non-linear transformation of the predictors:** The syntax for the predictors follow the conventions as used in link{formula}. The part terms consists of a series of terms concatenated by +, each term being the name of a variable, or the interaction of several variables separated by :.

Arithmetic expressions such as  $a + \log(b)$  are also legal. However, arithmetic expressions may contain special symbols that are defined for other use, such as +, \*, ^ and -. To avoid confusion, the function I() can be used to bracket portions where the operators should be interpreted in arithmetic sense. For example, in x + I(y + z), the term y + z is interpreted as the sum of y and z.

**Group level random effect:** When effects assumed to vary across grouping variables are considered, one can specify such effects by adding terms in the form of gterms | group, where group

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refers to the group indicator (usually a factor), and gterms specifies the terms whose coefficients are group-specific, drawn from a population normal distribution.

The most common situation for group level random effect is to include group-specific intercepts to account for unmeasured confounding. For example,  $x + y + (1 \mid g)$  specifies a model with population predictors x and y, as well as random intercept for each level of g.

For more complex random effect structures, refer to lme4::lmer. However, structures other than simple random intercepts and slopes may lead to unexpected behaviors.

#### Value

PSFormula returns an object of class PSFormula, which is a list containing for following components.

full\_formula input formula as is

data input data frame

fixed\_eff\_formula input formula with only fixed effects

response\_names character vector with names of variables that appear on the left hand side of input formula

has\_random\_effect logical indicating whether random effects are specified in the input formula

has\_intercept logical indicating whether the input formula has an intercept

fixed\_eff\_names character vector with names of all variables included as fixed effects

fixed\_eff\_count integer indicating the number of variables (factors are converted to and counted as dummy variables)

fixed\_eff\_matrix fixed-effect design matrix

random\_eff\_list a list containing information for each random effect. Such information is a list with the corresponding design matrix, the term names and the factor levels.

## See Also

formula, 1mer.

#### **Examples**

```
df <- data.frame(
    X = 1:10,
    Z = c(0,0,0,0,0,1,1,1,1,1),
    D = c(0,0,0,1,1,1,0,0,1,1),
    R = c(1,1,1,1,2,2,2,3,3,3)
)
PSFormula(Z + D ~ X + I(X^2) + (1 | R), df)</pre>
```

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PSObject

Create a Principal Stratification Object

#### **Description**

Create an object containing essential information to create the Stan file and data for Stan to draw posterior samples. Such information includes the specified model for principal stratum and outcome, the type of outcome, assumptions, and prior specification, etc.

#### Usage

```
PSObject(
    S.formula,
    Y.formula,
    Y.family,
    data = NULL,
    strata = NULL,
    ER = NULL,
    prior_intercept = prior_flat(),
    prior_coefficient = prior_normal(),
    prior_sigma = prior_inv_gamma(),
    prior_alpha = prior_inv_gamma(),
    prior_lambda = prior_inv_gamma(),
    prior_theta = prior_normal(),
    survival.time.points = 50
)
```

#### **Arguments**

S.formula, Y.formula

an object of class "PSFormula" (or an object of class "formula" that can be coerced to that class with data provided) specifying the model for principal stratum and outcome respectively. See PSFormula for details.

Y.family

an object of class "family": specifying the parametric family of the model for the response and the link function. See the documentation for glm for details on how such model fitting takes place. Supported families and corresponding link functions are presented in 'Details' below.

data

(optional) a data frame object. This is required when either S. formula or Y. formula is a formula object, to coerce it into a PSFormula object. When this happens, the data frame should contain all of the variables with names given in S. formula or Y. formula.

strata, ER

arguments to define the principal strata. See PStrataInfo for details.

Alternatively, one can pass an object of class PStrataInfo to strata, and ER will be ignored.

prior\_intercept, prior\_coefficient, prior\_sigma, prior\_alpha, prior\_lambda, prior\_theta prior distribution for corresponding parameters in the model.

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survival.time.points

a vector of time points at which the estimated survival probability is evaluated (only used when the type of outcome is survival), or an integer specifying the number of time points to be chosen. By default, the time points are chosen with equal distance from 0 to the 90% quantile of the observed outcome.

#### **Details**

The supported family objects include two types: native families for ordinary outcome and survival family for survival outcome.

For ordinary outcome, the below families and links are supported. See family for more details.

family	link				
binomial	logit, probit, cauchit, log, cloglog				
gaussian	identity, log, inverse				
Gamma	inverse, identity, log				
poisson	log, identity, log				
inverse.gamma	1/mu^2, inverse, identity, log				

The quasi family is not supported for the current version of the package.

For survival outcome, the family object is created by survival (method = "Cox", link = "identity"), where method can be either "Cox" for Weibull-Cox model or "AFT" for accelerated failure time model. See survival for more details. For the current version, only "identity" is used as the link function.

The gaussian family and the survival family with method = "AFT" introduce an additional parameter sigma for the standard deviation, whose prior distribution is specified by prior\_sigma. Similarly, prior\_alpha specifies the prior distribution of alpha for Gamma family, prior\_lambda specifies the prior distribution of theta for inverse.gaussian family, and prior\_theta specifies the prior distribution of theta for survival family with method = "Cox".

The models for principal stratum S. formula and response Y. formula also involve a linear combination of terms, where the prior distribution of the intercept and coefficients are specified by prior\_intercept and prior\_coefficient respectively.

#### Value

A list, containing important information describing the principal stratification model.

S.formula, Y.formula

A PSFormula object converted from the input S. formula and Y. formula

Y. family Same as input.

is.survival A boolean value. TRUE if Y.family is survival\_Cox or survival\_AFT.

strata\_info A PStrataInfo object converted from the input strata and ER.

survival.time.points

A list of time points at which the estimated survival probability is evaluated.

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SZDG\_table A matrix. Each row corresponds to a valid (stratum, treatment, confounder, group) combination.

Z\_names A character vector. The names of the levels of the treatment.

## **Examples**

```
df <- data.frame(</pre>
  Z = rbinom(10, 1, 0.5),
  D = rbinom(10, 1, 0.5),
  Y = rnorm(10),
  X = 1:10
)
PSObject(
  S.formula = Z + D \sim X,
  Y.formula = Y \sim X,
  Y.family = gaussian("identity"),
  data = df,
  strata = c(n = "00*", c = "01", a = "11*")
   _____
PSObject(
  S.formula = Z + D \sim 1,
  Y.formula = Y \sim 1,
  Y.family = gaussian("identity"),
  data = sim_data_normal,
  strata = c(n = "00*", c = "01", a = "11*")
)
```

PSOutcome

Estimated potential outcome for principal stratification analysis

## Description

Create an object useful to present the potential outcomes under each treatment arm for each principal stratum. Contrasts between treatment arms or principal strata are easy to obtain from this object.

#### Usage

```
PSOutcome(PStrataObj, type = c("probability", "RACE"))
```

#### **Arguments**

PStrataObj an object of class PStrata or PStrata\_survival

type whether the causal estimand is survival probability or RACE, ignored for non-

survival outcomes.

#### Value

An S3 object of type PSOutcome, containing

outcome\_array A num\_strata \* num\_treatment \* num\_iter array of mean outcome if the outcome type is non-survival or a num\_strata \* num\_treatment \* num\_time\_points \* num\_iter array of mean outcome if the outcome type is survival.

is.survival A boolean value, whether the outcome type is survival.

time\_points The time points at which the outcome is evaluated, if the outcome type is survival.

The S3 method summary and plot can be applied to the returned object.

**PSSample** 

Sample with Stan

## **Description**

Sample from the posterior distribution by calling stan. Check stan for details of the arguments.

## Usage

```
PSSample(
  file,
  model_name = "anon_model",
 model_code = "",
  fit = NA,
  data = list(),
  pars = NA,
  chains = 4,
  iter = 2000,
 warmup = floor(iter/2),
  thin = 1,
  init = "random",
  seed = sample.int(.Machine$integer.max, 1),
  algorithm = c("NUTS", "HMC", "Fixed_param"),
  control = NULL,
  sample_file = NULL,
  diagnostic_file = NULL,
  save\_dso = TRUE,
  verbose = FALSE,
  include = TRUE,
  cores = getOption("mc.cores", 1L),
 open_progress = interactive() && !isatty(stdout()) && !identical(Sys.getenv("RSTUDIO"),
    "1"),
 boost_lib = NULL,
  eigen_lib = NULL
)
```

#### **Arguments**

file

The path to the Stan program to use. file should be a character string file name or a connection that R supports containing the text of a model specification in the Stan modeling language.

A model may also be specified directly as a character string using the model\_code argument, but we recommend always putting Stan programs in separate files with a .stan extension.

The stan function can also use the Stan program from an existing stanfit object via the fit argument. When fit is specified, the file argument is ignored.

model\_name

A string to use as the name of the model; defaults to "anon\_model". However, the model name will be derived from file or model\_code (if model\_code is the name of a character string object) if model\_name is not specified. This is not a particularly important argument, although since it affects the name used in printed messages, developers of other packages that use **rstan** to fit models may want to use informative names.

model\_code

A character string either containing the model definition or the name of a character string object in the workspace. This argument is used only if arguments file and fit are not specified.

fit

An instance of S4 class stanfit derived from a previous fit; defaults to NA. If fit is not NA, the compiled model associated with the fitted result is re-used; thus the time that would otherwise be spent recompiling the C++ code for the model can be saved.

data

A named list or environment providing the data for the model, or a character vector for all the names of objects to use as data. See the **Passing data to Stan** section below.

pars

A character vector specifying parameters of interest to be saved. The default is to save all parameters from the model. If include = TRUE, only samples for parameters named in pars are stored in the fitted results. Conversely, if include = FALSE, samples for all parameters *except* those named in pars are stored in the fitted results.

chains

A positive integer specifying the number of Markov chains. The default is 4.

iter

A positive integer specifying the number of iterations for each chain (including warmup). The default is 2000.

warmup

A positive integer specifying the number of warmup (aka burnin) iterations per chain. If step-size adaptation is on (which it is by default), this also controls the number of iterations for which adaptation is run (and hence these warmup samples should not be used for inference). The number of warmup iterations should be smaller than iter and the default is iter/2.

thin

A positive integer specifying the period for saving samples. The default is 1, which is usually the recommended value. Unless your posterior distribution takes up too much memory we do *not* recommend thinning as it throws away information. The tradition of thinning when running MCMC stems primarily from the use of samplers that require a large number of iterations to achieve the desired effective sample size. Because of the efficiency (effective samples per second) of Hamiltonian Monte Carlo, rarely should this be necessary when using Stan.

init

Specification of initial values for all or some parameters. Can be the digit 0, the strings "0" or "random", a function that returns a named list, or a list of named lists:

init="random" (default): Let Stan generate random initial values for all parameters. The seed of the random number generator used by Stan can be specified via the seed argument. If the seed for Stan is fixed, the same initial values are used. The default is to randomly generate initial values between -2 and 2 on the unconstrained support. The optional additional parameter init\_r can be set to some value other than 2 to change the range of the randomly generated inits.

init="0", init=0: Initialize all parameters to zero on the unconstrained support.

**inits via list:** Set inital values by providing a list equal in length to the number of chains. The elements of this list should themselves be named lists, where each of these named lists has the name of a parameter and is used to specify the initial values for that parameter for the corresponding chain.

inits via function: Set initial values by providing a function that returns a list for specifying the initial values of parameters for a chain. The function can take an optional parameter chain\_id through which the chain\_id (if specified) or the integers from 1 to chains will be supplied to the function for generating initial values. See the **Examples** section below for examples of defining such functions and using a list of lists for specifying initial values.

When specifying initial values via a list or function, any parameters for which values are not specified will receive initial values generated as described in the init="random" description above.

The seed for random number generation. The default is generated from 1 to the maximum integer supported by R on the machine. Even if multiple chains are used, only one seed is needed, with other chains having seeds derived from that of the first chain to avoid dependent samples. When a seed is specified by a number, as.integer will be applied to it. If as.integer produces NA, the seed is generated randomly. The seed can also be specified as a character string of digits, such as "12345", which is converted to integer.

Using R's set. seed function to set the seed for Stan will not work.

algorithm

One of the sampling algorithms that are implemented in Stan. The default and preferred algorithm is "NUTS", which is the No-U-Turn sampler variant of Hamiltonian Monte Carlo (Hoffman and Gelman 2011, Betancourt 2017). Currently the other options are "HMC" (Hamiltonian Monte Carlo), and "Fixed\_param". When "Fixed\_param" is used no MCMC sampling is performed (e.g., for simulating with in the generated quantities block).

control

A named list of parameters to control the sampler's behavior. It defaults to NULL so all the default values are used. First, the following are adaptation parameters for sampling algorithms. These are parameters used in Stan with similar names here.

- adapt\_engaged (logical)
- adapt\_gamma (double, positive, defaults to 0.05)
- adapt\_delta (double, between 0 and 1, defaults to 0.8)

seed

- adapt\_kappa (double, positive, defaults to 0.75)
- adapt\_t0 (double, positive, defaults to 10)
- adapt\_init\_buffer (integer, positive, defaults to 75)
- adapt\_term\_buffer (integer, positive, defaults to 50)
- adapt\_window (integer, positive, defaults to 25)

In addition, algorithm HMC (called 'static HMC' in Stan) and NUTS share the following parameters:

- stepsize (double, positive, defaults to 1) Note: this controls the *initial* stepsize only, unless adapt\_engaged=FALSE.
- stepsize\_jitter (double, [0,1], defaults to 0)
- metric (string, one of "unit\_e", "diag\_e", "dense\_e", defaults to "diag\_e")

For algorithm NUTS, we can also set:

• max\_treedepth (integer, positive, defaults to 10)

For algorithm HMC, we can also set:

• int\_time (double, positive)

For test\_grad mode, the following parameters can be set:

- epsilon (double, defaults to 1e-6)
- error (double, defaults to 1e-6)

sample\_file

An optional character string providing the name of a file. If specified the draws for *all* parameters and other saved quantities will be written to the file. If not provided, files are not created. When the folder specified is not writable, tempdir() is used. When there are multiple chains, an underscore and chain number are appended to the file name.

#### diagnostic\_file

An optional character string providing the name of a file. If specified the diagnostics data for *all* parameters will be written to the file. If not provided, files are not created. When the folder specified is not writable, tempdir() is used. When there are multiple chains, an underscore and chain number are appended to the file name.

save\_dso

Logical, with default TRUE, indicating whether the dynamic shared object (DSO) compiled from the C++ code for the model will be saved or not. If TRUE, we can draw samples from the same model in another R session using the saved DSO (i.e., without compiling the C++ code again). This parameter only takes effect if fit is not used; with fit defined, the DSO from the previous run is used. When save\_dso=TRUE, the fitted object can be loaded from what is saved previously and used for sampling, if the compiling is done on the same platform, that is, same operating system and same architecture (32bits or 64bits).

verbose

TRUE or FALSE: flag indicating whether to print intermediate output from Stan on the console, which might be helpful for model debugging.

include

Logical scalar defaulting to TRUE indicating whether to include or exclude the parameters given by the pars argument. If FALSE, only entire multidimensional parameters can be excluded, rather than particular elements of them.

cores

The number of cores to use when executing the Markov chains in parallel. The default is to use the value of the "mc.cores" option if it has been set and otherwise to default to 1 core. However, we recommend setting it to be as many processors as the hardware and RAM allow (up to the number of chains). See detectCores if you don't know this number for your system.

open\_progress

Logical scalar that only takes effect if cores > 1 but is recommended to be TRUE in interactive use so that the progress of the chains will be redirected to a file that is automatically opened for inspection. For very short runs, the user might prefer FALSE.

.. Other optional parameters:

- chain\_id (integer)
- init\_r (double, positive)
- test\_grad (logical)
- append\_samples (logical)
- refresh(integer)
- save\_warmup(logical)
- deprecated: enable\_random\_init(logical)

chain\_id can be a vector to specify the chain\_id for all chains or an integer. For the former case, they should be unique. For the latter, the sequence of integers starting from the given chain\_id are used for all chains.

init\_r is used only for generating random initial values, specifically when init="random" or not all parameters are initialized in the user-supplied list or function. If specified, the initial values are simulated uniformly from interval [-init\_r, init\_r] rather than using the default interval (see the manual of (cmd)Stan).

test\_grad (logical). If test\_grad=TRUE, Stan will not do any sampling. Instead, the gradient calculation is tested and printed out and the fitted stanfit object is in test gradient mode. By default, it is FALSE.

append\_samples (logical). Only relevant if sample\_file is specified *and* is an existing file. In that case, setting append\_samples=TRUE will append the samples to the existing file rather than overwriting the contents of the file.

refresh (integer) can be used to control how often the progress of the sampling is reported (i.e. show the progress every refresh iterations). By default, refresh = max(iter/10, 1). The progress indicator is turned off if refresh  $\le 0$ 

Deprecated: enable\_random\_init (logical) being TRUE enables specifying initial values randomly when the initial values are not fully specified from the user.

save\_warmup (logical) indicates whether to save draws during the warmup phase and defaults to TRUE. Some memory related problems can be avoided by setting it to FALSE, but some diagnostics are more limited if the warmup draws are not stored.

boost\_lib The path for an alternative version of the Boost C++ to use instead of the one in the **BH** package.

eigen\_lib The path for an alternative version of the Eigen C++ library to the one in **RcppEigen**.

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#### Value

An object of S4 class rstan::stanfit.

**PStrata** 

Principal Stratification Analysis for Data with Post-Randomization Intervention

## **Description**

Perform pincipal stratification analysis when there are confounding variables after randomization

#### Usage

```
PStrata(
  PSobject = NULL,
  S.formula,
  Y.formula,
  Y.family,
  data = NULL,
  strata = NULL,
  ER = NULL
  prior_intercept = prior_flat(),
  prior_coefficient = prior_normal(),
  prior_sigma = prior_inv_gamma(),
  prior_alpha = prior_inv_gamma(),
  prior_lambda = prior_inv_gamma(),
  prior_theta = prior_normal(),
  survival.time.points = 50,
  filename = NULL,
)
```

## Arguments

**PSobject** 

an object of class PSObject. If left blank, the object is constructed using the following arguments. See PSObject for details.

S.formula, Y.formula

an object of class "PSFormula" (or an object of class "formula" that can be coerced to that class with data provided) specifying the model for principal stratum and outcome respectively. See PSFormula for details.

Y.family

an object of class "family": specifying the parametric family of the model for the response and the link function. See the documentation for glm for details on how such model fitting takes place. Supported families and corresponding link functions are presented in 'Details' below.

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data (optional) a data frame object. This is required when either S.formula or

Y. formula is a formula object, to coerce it into a PSFormula object. When this happens, the data frame should contain all of the variables with names given in

S. formula or Y. formula.

strata, ER arguments to define the principal strata. See PStrataInfo for details.

Alternatively, one can pass an object of class PStrataInfo to strata, and ER will be ignored.

prior\_intercept, prior\_coefficient, prior\_sigma, prior\_alpha, prior\_lambda, prior\_theta prior distribution for corresponding parameters in the model.

survival.time.points

a vector of time points at which the estimated survival probability is evaluated (only used when the type of outcome is survival), or an integer specifying the number of time points to be chosen. By default, the time points are chosen with equal distance from 0 to the 90% quantile of the observed outcome.

filename (optional) string. If not NULL, the stan file will be saved via cat in a text file

named after the string supplied.

additional parameters to be passed into PSSample.

#### Value

An object of class PStrata or PStrata\_survival, which is a list containing

PSobject An object of PSObject.

post\_samples An object of class rstan::stanfit returned by **Stan**.

## **Examples**

```
require(abind)
PSobj <- PSObject(</pre>
  S.formula = Z + D \sim 1,
  Y.formula = Y \sim 1,
  Y.family = gaussian("identity"),
  data = sim_data_normal,
  strata = c(n = "00*", c = "01", a = "11*")
)
PStrata(PSobj, cores = 2, chains = 2, iter = 200)
# Another example for survival data
PSobj <- PSObject(</pre>
  S.formula = Z + D \sim 1,
  Y.formula = Y + delta \sim 1,
  Y.family = survival("Cox"),
  data = sim_data_Cox,
  strata = c('never-taker' = "00*", complier = "01", 'always-taker' = "11*")
)
PStrata(PSobj, cores = 2, chains = 2, iter = 200)
```

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PStrataInfo	Create an object that defines the principal strata

#### **Description**

PStrataInfo is a class of object that defines all principal strata to be considered, by specifying the potential value of each post-randomization confounding variable under each treatment arm.

## Usage

```
PStrataInfo(strata, ER = NULL)
```

#### **Arguments**

strata a list or a vector defining all principal strata. Details of the syntax are given in

'Details' below.

ER a vector indicating on which strata exclusion restriction is assumed. Details are

given in 'Details' below.

#### Details

Since definition of the principal strata appears fundamental and essential in principal stratification analyses, the creation of such an object is designed to be user-friendly - various ways are accommodated to create a PStrataInfo object, some possibly preferable over others under different settings.

There are mainly two ways to easily create a PStrataInfo object.

**By string:** To define the principal strata by strings, the strata argument should receive a named vector, each component being the description of one strata with the name of that strata. The naming does not affect the actual inference, but informative names can be helpful for users to distinguish among strata.

Each stratum is defined by the potential values of the post-randomization confounding variable D under each treatment arm. By convention, assume that the K treatment arms are numbered from 0 to K-1. Then, each stratum is defined by the tuple  $(D(0),\ldots,D(K-1))$ , which can be written compactly as a string. For example, under binary treatment, the never-takers (i.e. D(0) = D(1) = 0) can be represented by string "00" and the compliers (i.e. D(0) = 0, D(1) = 1) can be represented by string "01". Note that the value that the post-randomization confounding variable can take is limited between 0 to 9 for the string to be parsed correctly. This should be more than enough in most of the applications, and in cases where a number above 10 is needed, please create the PStrataInfo object by matrix (see below).

When multiple post-randomization confounding variables exist, the string for each confounding variable is concatenated with the symbol "|". For example, if  $D_0$  and  $D_1$  are both binary post-randomization confounding variables, the stratum defined by  $D_0(0) = D_0(1) = 0$ ,  $D_1(0) = 0$ ,  $D_1(1) = 1$  can be represented by string "00|11". The order of these confounding variables should be the same as they appear in the S. formula parameter in PSObject.

A common assumption in practice is the exclusion restriction (ER) assumption, which assumes that the causal effect of the treatment on the outcome is totally realized through the post-randomization

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confounding variables. For example, the ER assumption on the stratum of never-takers can be interpreted as the outcome is identically distributed across the treated and control group, because all causal effect of the treatment is realized through the post-randomization variable, which is the same (0) under both treatment arms. To assume ER for some stratum, simply put an asterisk "\*" at the end of the string, such as "00\*" for the never-taker stratum. Note that under the context of multiple post-randomization variables, the package treats all such variables as a unity. The outcome is assumed to be identical under different treatment arms only when all post-randomization variables remain the same under these treatment arms.

Another way to specify the stratum where ER is assumed is to use the ER argument. It either takes a logical vector of the same length of strata with TRUE indicating ER is assumed and FALSE otherwise, or takes a character vector with the names of all strata where ER is to be assumed upon. When names to the strata are not provided in strata, the strata can be referred to by their canonical name, which is the string used to define the stratum with asterisks removed. For example, the strata "00|11\*" can be referred to with name "00|11".

**By matrix:** To define the principal strata by matrices, the strata argument should receive a named list, each component being a matrix. The number of rows matches the number of post-randomization variables, and the number of columns matches that of possible treatment arms. For any fixed row i, column j stores the potential value of the i-th post-randomization variable under treatment arm j.

When this approach is used, there is no shorthand to specify ER assumption. The ER argument is required to do this.

**Warning:** When ER assumption is specified in both strata and ER argument, the shorthand notation for ER in strata is ignored, and a warning is given regardless of whether the specification given by strata and ER actually match.

#### Value

an object of class PSStrataInfo, which is a list of the following components.

num\_strata number of principal strata defined

num\_treatment number of treatment arms

num\_postrand\_var number of post-randomization variables

max\_postrand\_level integer vector, the biggest number used by each post-randomization variable strata\_matrix integer matrix, each row corresponding to one stratum and each column corresponding to one treatment arm. The matrix is designed only for internal use.

**ER\_list** logical vector, each component corresponding to one stratum, indicating whether ER is assumed for the specific stratum

strata\_names character vector, the names of all strata

## **Examples**

```
PStrataInfo(strata = c(n = "00*", c = "01", a = "11"))
PStrataInfo(
    strata = list(n = c(0, 0), c = c(0, 1), a = c(1, 1)),
    ER = c(TRUE, FALSE, FALSE)
)
```

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```
PStrataInfo(
   strata = list(n = c(0, 0), c = c(0, 1), a = c(1, 1)),
   ER = c("n")
)
```

sim\_data\_Cox

Simulated Dataset for Survival Outcome (Cox Model)

## Description

A dataset generated for illustration of the principal stratification analysis. This dataset represents the common case of non-compliance.

## Usage

```
sim_data_Cox
```

#### **Format**

## 'sim\_data\_Cox' A data frame with 1,000 rows and 7 columns:

S Principal Strata: "never taker", "complier" or "always taker"

**Z** Randomized treatment arm: 0 = control, 1 = treatment

**D** Actual treatment arm: 0 = control, 1 = treatment

T True outcome: event time

C Censor time

delta Event indicator. 1 means true outcome is observed; 0 means otherwise

Y The observed event time or censor time

#### **Details**

The dataset represents the scenario where actual treatment might not be in compliance with the randomized (assigned) treatment. Defiers and always-takers are ruled out, leaving two strata, "nevertaker" and "complier" randomly sampled with probability  $0.3,\,0.7$  respectively. The assigned treatment Z is randomized with 0.5 probability for either arm. The true event time T is given by the following Weibull-Cox distribution

```
never-taker Y \sim Weibull - Cox(theta = 1, mu = 0.3)
complier Y \sim Weibull - Cox(theta = 1, mu = 2 - 0.6 * Z)
```

and the censor time C is uniformly drawn between 0.5 and 2.

The exclusion restriction assumption holds for never-takers in this generated dataset.

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sim\_data\_normal

Simulated Dataset for Normal Outcome

## **Description**

A dataset generated for illustration of the principal stratification analysis. This dataset represents the common case of non-compliance.

## Usage

```
sim_data_normal
```

#### **Format**

## 'sim\_data\_normal' A data frame with 1,000 rows and 4 columns:

S Principal Strata: "never taker", "complier" or "always taker"

**Z** Randomized treatment arm: 0 = control, 1 = treatment

**D** Actual treatment arm: 0 = control, 1 = treatment

Y Outcome

#### **Details**

The dataset represents the scenario where actual treatment might not be in compliance with the randomized (assigned) treatment. Defiers are ruled out, leaving three strata, "never taker", "complier" and "always taker" randomly sampled with probability 0.3, 0.2 and 0.5 respectively. The assigned treatment Z is randomized with 0.5 probability for either arm. The outcome Y is given by the following.

```
never taker Y \sim N(3,1) complier Y \sim N(-1-Z,0.5) always taker Y \sim N(1,2)
```

The exclusion restriction assumption holds for never takers and always takers in this generated dataset.

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The family function for survival data

## Description

Construct a family object for survival data

## Usage

```
survival(method = "Cox", link = "identity")
```

## Arguments

method the parametric method used for survival data. Can be Cox or AFT.

link a link function, currently only identity is implemented and used

## Value

A family object

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