

Package ‘RobinCar’

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Type Package

Title Robust Estimation and Inference in Covariate-Adaptive Randomization

Version 0.2.0

Description

Performs robust estimation and inference when using covariate adjustment and/or covariate-adaptive randomization in randomized controlled trials.

Ting Ye, Jun Shao, Yanyao Yi, Qinyuan Zhao (2023) <[doi:10.1080/01621459.2022.2049278](https://doi.org/10.1080/01621459.2022.2049278)>.

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Ting Ye, Jun Shao, Yanyao Yi (2023) <[doi:10.1093/biomet/asad045](https://doi.org/10.1093/biomet/asad045)>.

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nick, Jun Shao, Jingyi Liu, Yu Du, Yanyao Yi, Ting Ye (2024) <[doi:10.48550/arXiv.2306.10213](https://doi.org/10.48550/arXiv.2306.10213)>.

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Imports dplyr, magrittr, tidyr, emulator, numDeriv, tidyverse, stats, rlang, survival, fastDummies, data.table, broom, SuperLearner, AIPW, MASS

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Suggests knitr, rmarkdown, ranger, forcats, testthat (>= 3.0.0)

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car_pb	<i>Generate permuted block treatment assignments</i>
--------	--

Description

Generate permuted block treatment assignments

Usage

```
car_pb(z, trt_label, trt_alc, blocksize = 4L)
```

Arguments

z	The car_strata design matrix, as a data frame with factor variables
trt_label	Treatment label
trt_alc	Treatment allocation vector
blocksize	Permuted block blocksize

Value

A vector of treatment assignments with labels from the ‘trt_label’ argument, based on stratified permuted block randomization.

Examples

```
# Create car_strata variables
library(fastDummies)
library(dplyr)

x <- runif(100)
z <- cut(x, breaks=c(0, 0.25, 0.5, 0.75, 1.0))
z <- dummy_cols(z) %>%
  mutate(across(where(is.numeric), as.factor))

car_pb(z[, 2:5], c(0, 1, 2), trt_alc=c(1/4, 1/2, 1/4), blocksize=4L)
```

car_ps

Generate Pocock-Simon minimization treatment assignments

Description

Generate Pocock-Simon minimization treatment assignments

Usage

```
car_ps(z, treat, ratio, imb_measure, p_bc = 0.8)
```

Arguments

z	The car_strata design matrix
treat	A vector of length k (the number of treatment arms), which labels the treatment arms being compared.
ratio	A vector of length k (the number of treatment arms), which indicates the allocation ratio, e.g., c(1,1,1) for equal allocation with three treatment arms.
imb_measure	What measure of imbalance should be minimized during randomization – either "Range" or "SD"
p_bc	The biased probability, i.e., the probability of assigning each patient to the arm that minimizes the imbalance. Default is 0.8

Value

res treatment assignment vector

A vector of treatment assignments with labels from the ‘treat’ argument, based on Pocock-Simon’s minimization.

Author(s)

Ting Ye Yanyao Yi

Examples

```
# Create car_strata variables
library(fastDummies)
library(dplyr)

x <- runif(100)
z <- cut(x, breaks=c(0, 0.25, 0.5, 0.75, 1.0))
z <- dummy_cols(z)
A <- car_ps(
  z=z[, 2:5],
  treat=c(0, 1, 2),
  ratio=c(1, 1, 1),
  imb_measure="Range"
)
```

car_sr

Generate simple randomization treatment assignments

Description

Generate simple randomization treatment assignments

Usage

```
car_sr(n, p_trt)
```

Arguments

n	Number of observations
p_trt	Proportion allotted to treatment

Value

A vector of treatment assignments as 0's and 1's based on simple randomization.

Examples

```
car_sr(10, p_trt=0.4)
```

`data_gen`*Data generation function from JRSS-B paper*

Description

Data generation function from JRSS-B paper

Usage

```
data_gen(  
  n,  
  theta,  
  randomization,  
  p_trt,  
  case = c("case1", "case2", "case3", "case4", "case5")  
)
```

Arguments

<code>n</code>	total number of subjects to be generated
<code>theta</code>	true treatment effect
<code>randomization</code>	randomization method in <code>c("SR", "CABC", "permuted_block", "minimization", "urn")</code>
<code>p_trt</code>	proportion of treatment arm
<code>case</code>	simulation case in the paper

Value

A data frame with the following columns:

<code>t</code>	event time
<code>delta</code>	event indicator
<code>I1</code>	assignment to treatment group 1
<code>I0</code>	assignment to treatment group 0
<code>model_z1, model_z2</code>	covariates
<code>car_strata1, ...</code>	strata variables

 data_gen2

Data generation function from covariate adjusted log-rank paper

Description

Data generation function from covariate adjusted log-rank paper

Usage

```
data_gen2(
  n,
  theta,
  randomization,
  p_trt,
  case = c("case1", "case2", "case3", "case4"),
  blocksize = 4
)
```

Arguments

n	total number of subjects to be generated
theta	true treatment effect
randomization	randomization method in c("SR","CABC","permuted_block","minimization","urn")
p_trt	proportion of treatment arm
case	simulation case in the paper
blocksize	block size for permuted block design

Value

A data frame with the following columns:

t	event time
delta	event indicator
I1	assignment to treatment group 1
I0	assignment to treatment group 0
model_w3	covariates
car_strata1, ...	strata variables

```
print.CalibrationResult  
    Print calibration result
```

Description

Print calibration result

Usage

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

Arguments

x	A GLMModel result. If you'd like to calibrate a linear adjustment, use 'robin-car_glm' instead of 'robin-car_linear'.
...	Additional arguments

Value

Prints the treatment mean estimates (and variances) based on a calibration on top of a GLM working model, along with the settings used. See [RobinCar::robin-car_calibrate\(\)](#).

```
print.ContrastResult    Print contrast result
```

Description

Print contrast result

Usage

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

Arguments

x	A ContrastResult object
...	Additional arguments

Value

Prints estimates (and variances) of treatment contrasts based on a linear or GLM working model, along with the settings used. See [RobinCar::robin-car_contrast\(\)](#)

`print.GLMModelResult` *Print glm model result*

Description

Print glm model result

Usage

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

Arguments

<code>x</code>	A GLMModelResult object
<code>...</code>	Additional arguments

Value

Prints the treatment mean estimates (and variances) based on a GLM working model, along with the settings used. See [RobinCar::robincar_glm\(\)](#).

`print.LinModelResult` *Print linear model result*

Description

Print linear model result

Usage

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

Arguments

<code>x</code>	A LinModelResult object
<code>...</code>	Additional arguments

Value

Prints the treatment mean estimates (and variances) based on a linear working model, along with the settings used. See [RobinCar::robincar_linear\(\)](#).

```
print.TTEResult      Print TTE result
```

Description

Print TTE result

Usage

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

Arguments

```
x          A TTEResult object
...        Additional arguments
```

Value

Prints results of time-to-event covariate adjusted analyses including covariate-adjusted (stratified) logrank, robust Cox score, and covariate-adjusted hazard ratio. Prints summary statistics about number of observations and events, possibly by strata, and the test statistics and/or estimates, and p-values. See [RobinCar::robincar_tte\(\)](#) and [RobinCar::robincar_covhr\(\)](#).

```
robincar_calibrate  Perform linear or joint calibration
```

Description

Uses linear or joint calibration to "calibrate" the estimates from a linear or GLM-type adjustment. Linear calibration fits a linear model with treatment (and treatment-by-covariate interactions) and with the predicted $\hat{\mu}(X_i) = (\hat{\mu}_1(X_i), \dots, \hat{\mu}_K(X_i))$ as constructed covariates where K is the number of treatment groups; joint calibration also includes Z_i the strata variables as covariates.

Usage

```
robincar_calibrate(result, joint = FALSE, add_x = NULL)
```

Arguments

```
result      A GLMModelResult
joint       If true, then performs joint calibration with the  $\hat{\mu}(X_i)$  and strata  $Z_i$  to achieve
            universality and efficiency gain rather than just linear calibration that uses  $\hat{\mu}(X_i)$ .
add_x       Additional x to use in the calibration. Must have been in the original dataset that
            robincar_glm was called on.
```

Value

A result object that has the same structure as `RobinCar::robincar_glm()`, with the argument ‘result’ included as "original" in the list.

<code>robincar_contrast</code>	<i>Estimate a treatment contrast</i>
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Description

Estimate a treatment contrast using the result of `RobinCar::robincar_linear()`, `RobinCar::robincar_glm()`, or `RobinCar::robincar_SL()` using the delta method.

Usage

```
robincar_contrast(result, contrast_h, contrast_dh = NULL)
```

Arguments

<code>result</code>	A <code>LinModelResult</code> or <code>GLMModelResult</code>
<code>contrast_h</code>	An optional function to specify a desired contrast
<code>contrast_dh</code>	An optional jacobian function for the contrast

Value

A contrast object which has the following attributes:

<code>result</code>	A <code>dplyr::tibble()</code> with the label of the treatment contrast (e.g., 1 vs. 0), the estimate of the treatment contrast, estimated SE, and p-value based on a z-test with estimate and SE.
<code>varcov</code>	The variance-covariance matrix for the treatment contrast estimates.
<code>settings</code>	List of model settings used for the contrast.

<code>robincar_covhr</code>	<i>Covariate-adjusted estimators for time to event data</i>
-----------------------------	---

Description

Estimate a covariate-adjusted hazard ratio (‘adj_method="CL"’), or a covariate-adjusted stratified hazard ratio (‘adj_method="CSL"’).

Usage

```
robincar_covhr(
  df,
  treat_col,
  response_col,
  event_col,
  car_strata_cols = NULL,
  covariate_cols = NULL,
  p_trt = 0.5,
  ref_arm = NULL,
  car_scheme = "simple",
  adj_method = "CL",
  interval = c(-10, 10)
)
```

Arguments

<code>df</code>	A data.frame with the required columns
<code>treat_col</code>	Name of column in df with treatment variable
<code>response_col</code>	Name of the column in df with response variable
<code>event_col</code>	Name of column in df with event indicator (0/FALSE=no event, 1/TRUE=event)
<code>car_strata_cols</code>	Names of columns in df with car_strata variables
<code>covariate_cols</code>	Names of columns in df with covariate variables
<code>p_trt</code>	Treatment allocation ratio for the reference arm.
<code>ref_arm</code>	Reference arm of the treatment group, defaults to NULL, which results in using the first element of <code>'unique(data[, treat_col])'</code> .
<code>car_scheme</code>	Name of the type of covariate-adaptive randomization scheme. One of: "simple", "pocock-simon", "biased-coin", "permuted-block".
<code>adj_method</code>	Adjustment method (one of "CL", "CSL")
<code>interval</code>	Interval for uniroot function

Value

An object with attribute named "result", which lists:

<code>theta_L</code>	estimate of the hazard ratio
<code>se_theta_L</code>	SE estimate of the hazard ratio
<code>theta_CL</code>	estimate of the covariate-adjusted hazard ratio
<code>se_theta_CL</code>	SE estimate of the covariate-adjusted hazard ratio

Other attributes are the settings used, data attributes, and the original data frame supplied by the user.

robincar_coxscore	<i>Robust cox score adjustment</i>
-------------------	------------------------------------

Description

Robust cox score adjustment

Usage

```
robincar_coxscore(...)
```

Arguments

... Arguments to robincar_tte, other than 'adj_method'

Value

A result object with the following attributes:

result	A list: "statistic" is the robust Cox score test statistic which can be used to obtain p-values; "U" and "se" are the numerator and denominator of the test statistic, respectively.
settings	The covariate adjustment settings used.
original_df	The dataset supplied by the user.

robincar_glm	<i>Covariate adjustment using generalized linear working model</i>
--------------	--

Description

Estimate treatment-group-specific response means and (optionally) treatment group contrasts using a generalized linear working model.

Usage

```
robincar_glm(
  df,
  treat_col,
  response_col,
  car_strata_cols = NULL,
  covariate_cols = NULL,
  car_scheme = "simple",
  adj_method = "heterogeneous",
  covariate_to_include_strata = NULL,
  g_family = stats::gaussian,
```

```

g_accuracy = 7,
formula = NULL,
contrast_h = NULL,
contrast_dh = NULL
)

```

Arguments

df	A data.frame with the required columns
treat_col	Name of column in df with treatment variable
response_col	Name of the column in df with response variable
car_strata_cols	Names of columns in df with car_strata variables
covariate_cols	Names of columns in df with covariate variables
car_scheme	Name of the type of covariate-adaptive randomization scheme. One of: "simple", "pocock-simon", "biased-coin", "permuted-block".
adj_method	Name of adjustment method to use, one of "heterogeneous" (interaction model) or "homogeneous"
covariate_to_include_strata	Whether to include car_strata variables in covariate adjustment. Defaults to F for homogeneous; defaults to T for heterogeneous. User may override by passing in this argument.
g_family	Family that would be supplied to <code>glm()</code> , e.g., binomial. If no link specified, will use default link, like behavior in <code>glm</code> . If you wish to use a negative binomial working model with an unknown dispersion parameter, then use <code>'g_family="nb"</code> .
g_accuracy	Level of accuracy to check prediction un-biasedness.
formula	An optional formula to use for adjustment specified using <code>as.formula("...")</code> . This overrides <code>car_strata_cols</code> and <code>covariate_cols</code> .
contrast_h	An optional function to specify a desired contrast
contrast_dh	An optional jacobian function for the contrast (otherwise use numerical derivative)

Details

The output is the AIPW estimator given by (for each treatment group a):

$$\frac{1}{n} \sum_{i=1}^n \hat{\mu}_a(X_i) + \frac{1}{n_a} \sum_{i:A_i=a} \{Y_i - \hat{\mu}(X_i)\}$$

where Y_i is the outcome, A_i is the treatment assignment, X_i are the covariates, $n_a = \sum_{i=1}^n A_i = a$, and $\hat{\mu}_a$ is the estimated conditional mean function based on the GLM working model. This working model has treatment a -specific coefficients if `'adj_method'` is "heterogeneous". Otherwise, they are shared across the treatment arms. Alternatively, if `'formula'` is used, the working model can be specified according to the user.

Importantly, the estimated variance accounts for misspecification of the working model, and for covariate-adaptive randomization.

Value

If ‘contrast_h’ argument is used, outputs a ‘main’ and a ‘contrast’ object. The ‘main’ object has the following structure:

result	A <code>dplyr::tibble()</code> with the treatment label, treatment mean estimate using AIPW, estimated SE, and p-value based on a z-test with estimate and SE.
varcov	The variance-covariance matrix for the treatment mean estimates.
settings	List of model settings used in covariate adjustment.
original_df	The original dataset provided by the user.
mod	The fit from the <code>glm()</code> working model used for covariate adjustment.
mu_a	Predicted potential outcomes for each treatment category (columns) and individual (rows). These are the $\hat{\mu}_a$
.	.
g.estimate	The G-computation estimate based only on $\frac{1}{n} \sum_{i=1}^n \hat{\mu}_a(X_i)$. This is equivalent to the AIPW estimate when a canonical link function is used.
data	Attributes about the dataset.

The ‘contrast’ object has a structure that is documented in `RobinCar::robincar_contrast()`.

robincar_glm2	<i>Covariate adjustment using generalized linear working model, with simplified interface.</i>
---------------	--

Description

Covariate adjustment using generalized linear working model, with simplified interface.

Usage

```
robincar_glm2(
  df,
  treat_col,
  response_col,
  formula = NULL,
  car_strata_cols = NULL,
  car_scheme = "simple",
  g_family = stats::gaussian,
  g_accuracy = 7,
  contrast_h = NULL,
  contrast_dh = NULL
)
```

Arguments

df	A data.frame with the required columns
treat_col	Name of column in df with treatment variable
response_col	Name of the column in df with response variable
formula	The formula to use for adjustment specified using <code>as.formula("...")</code> . This overrides <code>car_strata_cols</code> and <code>covariate_cols</code> .
car_strata_cols	Names of columns in df with <code>car_strata</code> variables
car_scheme	Name of the type of covariate-adaptive randomization scheme. One of: "simple", "pocock-simon", "biased-coin", "permuted-block".
g_family	Family that would be supplied to <code>glm(...)</code> , e.g., binomial. If no link specified, will use default link, like behavior in <code>glm</code> . If you wish to use a negative binomial working model with an unknown dispersion parameter, then use <code>'g_family="nb"'</code> .
g_accuracy	Level of accuracy to check prediction un-biasedness.
contrast_h	An optional function to specify a desired contrast
contrast_dh	An optional jacobian function for the contrast (otherwise use numerical derivative)

Value

See value of [RobinCar::robincar_glm\(\)](#). This function is a wrapper for [RobinCar::robincar_glm\(\)](#).

robincar_linear	<i>Covariate adjustment using linear working model</i>
-----------------	--

Description

Estimate treatment-group-specific response means and (optionally) treatment group contrasts using a linear working model for continuous outcomes.

Usage

```
robincar_linear(
  df,
  treat_col,
  response_col,
  car_strata_cols = NULL,
  covariate_cols = NULL,
  car_scheme = "simple",
  adj_method = "ANOVA",
  covariate_to_include_strata = NULL,
  contrast_h = NULL,
  contrast_dh = NULL
)
```

Arguments

df	A data.frame with the required columns
treat_col	Name of column in df with treatment variable
response_col	Name of the column in df with response variable
car_strata_cols	Names of columns in df with car_strata variables
covariate_cols	Names of columns in df with covariate variables
car_scheme	Name of the type of covariate-adaptive randomization scheme. One of: "simple", "pocock-simon", "biased-coin", "permuted-block".
adj_method	Name of linear adjustment method to use. One of: "ANOVA", "ANCOVA", "ANHECOVA".
covariate_to_include_strata	Whether to include car_strata variables in covariate adjustment. Defaults to F for ANOVA and ANCOVA; defaults to T for ANHECOVA. User may override by passing in this argument.
contrast_h	An optional function to specify a desired contrast
contrast_dh	An optional jacobian function for the contrast (otherwise use numerical derivative)

Value

See value of [RobinCar::robincar_glm\(\)](#), this function is a wrapper using an identity link for the glm.

robincar_linear2	<i>Covariate adjustment using linear working model, with simplified interface.</i>
------------------	--

Description

Estimate treatment-group-specific response means and (optionally) treatment group contrasts using a linear working model for continuous outcomes.

Usage

```
robincar_linear2(
  df,
  treat_col,
  response_col,
  car_strata_cols = NULL,
  covariate_cols = NULL,
  car_scheme = "simple",
  adj_method = "ANOVA",
  contrast_h = NULL,
  contrast_dh = NULL
)
```


Arguments

df	A data.frame with the required columns
treat_col	Name of column in df with treatment variable
response_col	Name of the column in df with response variable
car_strata_cols	Names of columns in df with car_strata variables
covariate_cols	Names of columns in df with covariate variables. **If you want to include the strata variables as covariates also, add them here.**
car_scheme	Name of the type of covariate-adaptive randomization scheme. One of: "simple", "pocock-simon", "biased-coin", "permuted-block".
adj_method	Name of linear adjustment method to use. One of: "ANOVA", "ANCOVA", "ANHECOVA".
contrast_h	An optional function to specify a desired contrast
contrast_dh	An optional jacobian function for the contrast (otherwise use numerical derivative)

Details

Note the difference between [RobinCar::robincar_linear\(\)](#) and [RobinCar::robincar_linear2\(\)](#) with regards to their 'covariate_cols' argument.

Value

See value of [RobinCar::robincar_linear\(\)](#), this function is a wrapper.

robincar_logrank	<i>Robust (potentially stratified) logrank adjustment</i>
------------------	---

Description

Perform a robust covariate-adjusted logrank test ("CL") that can be stratified ("CSL") if desired.

Usage

```
robincar_logrank(adj_method, ...)
```

Arguments

adj_method	Adjustment method, one of "CL", "CSL"
...	Additional arguments to 'robincar_tte'

Value

A result object with the following attributes:

result	A list: "statistic" is the adjusted logrank test statistic which can be used to obtain p-values; "U" and "se" are the numerator and denominator of the test statistic, respectively.
settings	The covariate adjustment settings used.
original_df	The dataset supplied by the user.

Examples

```
library(magrittr)
library(dplyr)
library(forcats)
set.seed(0)
n=100
data.simu0=data_gen(n=n,
                    theta=0,
                    randomization="permuted_block",
                    p_trt=0.5,
                    case="case2") %>% mutate(strata1=sample(letters[1:3],n,replace=TRUE),
                                             strata2=sample(LETTERS[4:5],n,replace=TRUE))

out <- robincar_logrank(df=data.simu0,
                       treat_col="I1",
                       p_trt=0.5,
                       ref_arm=0,
                       response_col="t",
                       event_col="delta",
                       covariate_cols=c("model_z1", "model_z2"),
                       car_scheme="simple",
                       adj_method=c("CL"))

set.seed(0)
n=100
data.simu0=data_gen(n=n,
                    theta=0,
                    randomization="permuted_block",
                    p_trt=0.5,
                    case="case1")

data.simu <- data.simu0 %>%
  tidyr::pivot_longer(cols=starts_with("car_strata"),
                     names_prefix="car_strata",
                     names_to="strt") %>%
  filter(value==1) %>% select(-value) %>%
  mutate(strt=forcats::as_factor(strt)) %>%
  select(t,strt) %>%
  left_join(data.simu0, .)

out1 <- robincar_logrank(df=data.simu,
```

```

    treat_col="I1",
    p_trt=0.5,
    ref_arm=0,
    response_col="t",
    event_col="delta",
    car_strata_cols="strt",
    covariate_cols=NULL,
    car_scheme=c("permuted-block"),
    adj_method=c("CSL")
)

```

robincar_SL	<i>BETA: Covariate adjustment using working models from the super learner libraries through the AIPW package with cross-fitting.</i>
-------------	--

Description

Estimate treatment-group-specific response means and (optionally) treatment group contrasts using a generalized linear working model.

Usage

```

robincar_SL(
  df,
  treat_col,
  response_col,
  car_strata_cols = NULL,
  covariate_cols = NULL,
  car_scheme = "simple",
  covariate_to_include_strata = NULL,
  SL_libraries = c(),
  SL_learners = c(),
  k_split = 2,
  g_accuracy = 7,
  contrast_h = NULL,
  contrast_dh = NULL
)

```

Arguments

<code>df</code>	A data.frame with the required columns
<code>treat_col</code>	Name of column in df with treatment variable
<code>response_col</code>	Name of the column in df with response variable
<code>car_strata_cols</code>	Names of columns in df with car_strata variables
<code>covariate_cols</code>	Names of columns in df with covariate variables

car_scheme	Name of the type of covariate-adaptive randomization scheme. One of: "simple", "pocock-simon", "biased-coin", "permuted-block".
covariate_to_include_strata	Whether to include car_strata variables in covariate adjustment. Defaults to F for ANOVA and ANCOVA; defaults to T for ANHECOVA. User may override by passing in this argument.
SL_libraries	Vector of super-learner libraries to use for the covariate adjustment (see SuperLearner::listWrappers)
SL_learners	Optional list of super-learner "learners" to use for the covariate adjustment (see SuperLearner::create.Learner())
k_split	Number of splits to use in cross-fitting
g_accuracy	Level of accuracy to check prediction un-biasedness (in digits).
contrast_h	An optional function to specify a desired contrast
contrast_dh	An optional jacobian function for the contrast (otherwise use numerical derivative)

Details

WARNING: This function is still under development and has not been extensively tested. This function currently only works for two treatment groups. Before using this function, you must load the [SuperLearner](#) library with 'library(SuperLearner)', otherwise the function call will fail.

Value

See value of [RobinCar::robincar_glm](#), but the working model for $\hat{\mu}(X_i)$ is based on the [AIPW](#) package that uses specified SuperLearner libraries and cross-fitting. Also, 'mod' attribute is an object of class [AIPW::AIPW](#).

Examples

```
library(SuperLearner)
library(ranger)
n <- 1000
set.seed(10)
DATA2 <- data.frame(A=rbinom(n, size=1, prob=0.5),
                    y=rbinom(n, size=1, prob=0.2),
                    x1=rnorm(n),
                    x2=rnorm(n),
                    x3=as.factor(rbinom(n, size=1, prob=0.5)),
                    z1=rbinom(n, size=1, prob=0.5),
                    z2=rbinom(n, size=1, prob=0.5))

DATA2[, "y"] <- NA
As <- DATA2$A == 1
DATA2[DATA2$A == 1, "y"] <- rbinom(
  sum(As),
  size=1,
  prob=exp(DATA2[As,]$x1)/(1+exp(DATA2[As,]$x1)))
DATA2[DATA2$A == 0, "y"] <- rbinom(
```

```

n=sum(As),
size=1,
prob=exp(1 +
  5*DATA2[!As,]$x1 + DATA2[!As,]$x2)/
  (1+exp(1 + 5*DATA2[!As,]$x1 + DATA2[!As,]$x2)))
DATA2$A <- as.factor(DATA2$A)

sl.mod <- robincar_SL(
  df=DATA2,
  response_col="y",
  treat_col="A",
  car_strata_cols=c("z1"),
  covariate_cols=c("x1"),
  SL_libraries=c("SL.ranger"),
  car_scheme="permuted-block",
  covariate_to_include_strata=TRUE
)

sl.mod$result

```

robincar_SL_median *BETA: Covariate adjustment using working models from the super learner libraries through the AIPW package with cross-fitting, with median adjustment.*

Description

Estimate treatment-group-specific response means and (optionally) treatment group contrasts using a generalized linear working model. Perform median adjustment to limit randomness induced from cross-fitting.

Usage

```

robincar_SL_median(
  n_times,
  seed,
  df,
  treat_col,
  response_col,
  car_strata_cols = NULL,
  covariate_cols = NULL,
  car_scheme = "simple",
  covariate_to_include_strata = NULL,
  SL_libraries = c(),
  SL_learners = c(),
  k_split = 2,
  g_accuracy = 7,
  contrast_h = NULL,

```

```

    contrast_dh = NULL
  )

```

Arguments

n_times	Number of times to run the robincar_SL function
seed	Seed to set before running the set of functions
df	A data.frame with the required columns
treat_col	Name of column in df with treatment variable
response_col	Name of the column in df with response variable
car_strata_cols	Names of columns in df with car_strata variables
covariate_cols	Names of columns in df with covariate variables
car_scheme	Name of the type of covariate-adaptive randomization scheme. One of: "simple", "pocock-simon", "biased-coin", "permuted-block".
covariate_to_include_strata	Whether to include car_strata variables in covariate adjustment. Defaults to F for ANOVA and ANCOVA; defaults to T for ANHECOVA. User may override by passing in this argument.
SL_libraries	Vector of super-learner libraries to use for the covariate adjustment (see SuperLearner::listWrappers)
SL_learners	Optional list of super-learner "learners" to use for the covariate adjustment (see SuperLearner::create.Learner())
k_split	Number of splits to use in cross-fitting
g_accuracy	Level of accuracy to check prediction un-biasedness (in digits).
contrast_h	An optional function to specify a desired contrast
contrast_dh	An optional jacobian function for the contrast (otherwise use numerical derivative)

Details

WARNING: This function is still under development and has not been extensively tested. This function currently only works for two treatment groups. Before using this function, you must load the SuperLearner library with `library(SuperLearner)`, otherwise the function call will fail.

Value

See value of [RobinCar::robincar_SL](#). Attributes `'mods'` and `'mu_as'` are lists of `'mod'` and `'mu_a'` attributes, respectively, for each replicate of `'robincar_SL'` used in the median.

robincar_tte	<i>Covariate adjustment for time to event data</i>
--------------	--

Description

Perform a covariate-adjusted logrank test ('adj_method="CL"'), covariate-adjusted stratified logrank test ('adj_method="CSL"'), or a covariate-adjusted robust Cox score test ('adj_method="coxscore").

Usage

```
robincar_tte(
  df,
  treat_col,
  response_col,
  event_col,
  adj_method,
  car_strata_cols = NULL,
  covariate_cols = NULL,
  p_trt = 0.5,
  ref_arm = NULL,
  sparse_remove = TRUE,
  car_scheme = "simple"
)
```

Arguments

<code>df</code>	A data.frame with the required columns
<code>treat_col</code>	Name of column in df with treatment variable
<code>response_col</code>	Name of the column in df with response variable
<code>event_col</code>	Name of column in df with event indicator (0/FALSE=no event, 1/TRUE=event)
<code>adj_method</code>	Adjustment method (one of "CL", "CSL", or "coxscore")
<code>car_strata_cols</code>	Names of columns in df with car_strata variables
<code>covariate_cols</code>	Names of columns in df with covariate variables
<code>p_trt</code>	Treatment allocation ratio for the reference arm.
<code>ref_arm</code>	Reference arm of the treatment group, defaults to NULL, which results in using the first element of 'unique(data[, treat_col])'.
<code>sparse_remove</code>	Remove sparse car_strata from calculation
<code>car_scheme</code>	Name of the type of covariate-adaptive randomization scheme. One of: "simple", "pocock-simon", "biased-coin", "permuted-block".

Details

'robincar_coxscore' and 'robincar_logrank' are wrapper functions around 'robincar_tte'.

Value

For adjustment method "CL" or "CSL", see value of [RobinCar::robincar_logrank\(\)](#); for adjustment method "coxscore" see value of [RobinCar::robincar_coxscore\(\)](#).

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