Package 'swfdr'

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Title Estimation of the science-wise false discovery rate and the false discovery rate conditional on covariates

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Description This package allows users to estimate the science-wise false discovery rate from Jager and Leek, ``Empirical estimates suggest most published medical research is true," 2013, Biostatistics, using an EM approach due to the presence of rounding and censoring. It also allows users to estimate the false discovery rate conditional on covariates, using a regression framework, as per Boca and Leek, ``A direct approach to estimating false discovery rates conditional on covariates," 2018, PeerJ.

Depends R (>= 3.4)

Imports methods, splines, stats4, stats

License GPL (>= 3)

URL https://github.com/leekgroup/swfdr

BugReports https://github.com/leekgroup/swfdr/issues

Encoding UTF-8 LazyData true

RoxygenNote 7.1.1

Suggests dplyr, ggplot2, BiocStyle, knitr, qvalue, reshape2, rmarkdown, testthat

VignetteBuilder knitr

biocViews MultipleComparison, StatisticalMethod, Software

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 ${\tt BMI_GIANT_GWAS_sample} \quad \textit{Subset of SNPs from meta-analysis of BMI GWAS study}.$

Description

A dataset containing 50,000 SNPs and results for their associations with BMI.

Usage

data(BMI_GIANT_GWAS_sample)

Format

A data frame with 50,000 rows and 9 variables:

SNP ID for SNP (single nucleotide polymorphism)

A1 Allele 1 for SNP

A2 Allele 2 for SNP

Freq_MAF_Hapmap Frequency of minor allele (MAF) in Hapmap project

- **b** Estimated beta for association between SNP and BMI
- se Estimated standard error (se) for association between SNP and BMI
- p P-value for association between SNP and BMI
- N Total sample size considered for association of SNP and BMI

Freq_MAF_Int_Hapmap Three approximately equal intervals for the Hapmap MAFs

Value

Object of class tbl_df, tbl, data.frame.

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Source

 $https://www.broadinstitute.org/collaboration/giant/index.php/GIANT_consortium_data_files\#GWAS_Anthropometric_2015_BMI$

calculateSwfdr

Calculate the science-wise FDR (swfdr)

Description

Calculate the science-wise FDR (swfdr)

Usage

```
calculateSwfdr(
  pValues,
  truncated,
  rounded,
  pi0 = 0.5,
  alpha = 1,
  beta = 50,
  numEmIterations = 100
)
```

Arguments

pValues	Numerical	vector of	p-values
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truncated Vector of 0s and 1s with indices corresponding to those in pValues; 1 indicates

that the p-values is truncated, 0 that it is not truncated

rounded Vector of 0s and 1s with indices corresponding to those in pValues; 1 indicates

that the p-values is rounded, 0 that it is not rounded

pi0 Initial prior probability that a hypothesis is null (default is 0.5)

alpha Initial value of parameter alpha from Beta(alpha, beta) true positive distribution

(default is 1)

beta Initial value of parameter beta from Beta(alpha, beta) true positive distribution

(default is 50)

numEmIterations

The number of EM iterations (default is 100)

Value

pi0 Final value of prior probability - estimated from EM - that a hypothesis is null,

i.e. estimated swfdr

alpha Final value of parameter alpha - estimated from EM - from Beta(alpha, beta)

true positive distribution

get_number_decimals

beta	Final value of parameter beta - estimated from EM - from Beta(alpha, beta) true positive distribution
Z	Vector of expected values of the indicator of whether the p-value is null or not - estimated from EM - for the non-rounded p-values (values of NA represent the rounded p-values)
n0	Expected number of rounded null p-values - estimated from EM - between certain cutpoints $(0.005,0.015,0.025,0.035,0.045,0.05)$
n	Number of rounded p-values between certain cutpoints (0.005, 0.015, 0.025, 0.035, 0.045, 0.05)

Examples

```
pVals <- runif(100)
tt <- rr <- rep(0, 100)
resSwfdr <- calculateSwfdr(pValues = pVals, truncated = tt, rounded = rr, numEmIterations=100)</pre>
```

get_number_decimals Get number of decimals (i.e. return total number of digits after decimal point) for any vector of numbers in [0,1) if number of decimals <=6

Description

Get number of decimals (i.e. return total number of digits after decimal point) for any vector of numbers in [0,1) if number of decimals ≤ 6

Usage

```
get_number_decimals(x)
```

Arguments

x Numerical vector where all elements are in [0,1)

Value

Vector giving the number of decimals for each element in x if the number is \leq 6; otherwise return 7 with a warning

Examples

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journals_pVals	P-values from abstracts from articles in 5 biomedical journals (American Journal of Epidemiology, BMJ, JAMA, Lancet, New England Journal of Medicine), over 11 years (2000, 2010)
	nal of Medicine), over 11 years (2000-2010).

Description

A dataset containing 15,653 p-values.

Usage

```
journals_pVals
```

Format

A tbl data frame with 15,653 rows and 5 variables:

```
pvalue P-value
pvalueTruncated Equals to 1 if the p-value is truncated, 0 otherwise
pubmedID Pubmed ID of the article
year Year of publication
journal Journal
```

Value

Object of class tbl_df, tbl, data.frame.

Source

Code for extracting p-values at: inst/script/getPvalues.R

lm_pi0	Estimation of pi0, proportion of p-values consistent with a null hypothesis
	· ·

Description

Estimation of pi0, proportion of p-values consistent with a null hypothesis

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Usage

```
lm_pi0(
   p,
   lambda = seq(0.05, 0.95, 0.05),
   X,
   type = c("logistic", "linear"),
   smooth.df = 3,
   threshold = TRUE,
   smoothing = c("unit.spline", "smooth.spline")
)
```

Arguments

p numeric vector, p-values

lambda numeric vector, thresholds used to bin pvalues, must be in [0,1).

X numeric matrix, covariates that might be related to p values (one test per row,

one variable per column).

type character, type of regression used to fit features to pvalues

smooth. df integer, degrees of freedom when estimating pi0(x) with a smoother.

threshold logical, if TRUE, all estimates are thresholded into unit interval; if FALSE, all

estimates are left as they are computed

smoothing character, type of smoothing used to fit pi0

Value

object of class 'lm pi0', which is a list with several components

call matched function call

lambda numeric vector of thresholds used in calculating pi0.lambda

X. names character vector of covariates used in modeling

pi0.1ambda numeric matrix of estimated pi0(x) for each value of lambda. The number of

columns is the number of tests, the number of rows is the length of lambda.

pi0 numerical vector of smoothed estimate of pi0(x). The length is the number of

rows in X.

pi0. smooth (only output with smoothing="smooth.spline") Matrix of fitted values from the

smoother fit to the pi0(x) estimates at each value of lambda (same number of

rows and columns as pi0.lambda)

Examples

```
# define a covariate
X <- seq(-1,2,length=1000)
# set probability of being null
pi0 <- 1/4*X + 1/2
# generate null/alternative p-values
nullI <- rbinom(1000,prob=pi0,size=1)> 0
```

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```
# vector of p-values
pValues <- rep(NA,1000)
pValues[nullI] <- runif(sum(nullI)) # from U(0,1)
pValues[!nullI] <- rbeta(sum(!nullI),1,2) # from Beta
pi0x <- lm_pi0(pValues, X=X)</pre>
```

lm_qvalue

Estimation of qvalues conditioned on covariates

Description

The recipe for turning pvalues into qvalues is adapted from package 'qvalue' and articles by Storey, Tibshirani, Taylor, Siegmund.

Usage

```
lm_qvalue(
   p,
   X,
   pfdr = FALSE,
   pi0 = NULL,
   smoothing = c("unit.spline", "smooth.spline"),
   ...
)
```

Arguments

p	numeric vector of p-values
X	matrix of covariates (can be missing if pi0 is specified instead)
pfdr	logical, making estimates robust for small p-values and a small sample size
pi0	list with pi0 estimates from lm_pi0. If this is not provided, pi0 is estimated using function lm_pi0.
smoothing	character, type of smoothing used to fit pi0. Note the default in this function is different than in lm_pi0 .
	other parameters (passed on to lm_pi0 if pi0 is not provided)

Value

object of class 'lm_qvalue', which is a list with several components

call matched function call
pvalues numeric vector of original p-values
qvalues numeric vector of q-values

other list elements transferred from pi0

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Examples

```
# define a covariate X \leftarrow rep(c(0, 1), each=1000) # generate p-values, randomly for group 0 and with low values for group 1 pVal \leftarrow c(runif(1000), rbeta(1000, 0.2, 1)) # compute qvalues, using the covariate qVal \leftarrow lm_qvalue(pVal, X=X)
```

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