Package 'reconsi'

December 16, 2025

Type Package

Title Resampling Collapsed Null Distributions for Simultaneous Inference

Version 1.22.0

Description Improves simultaneous inference under dependence of tests by estimating a collapsed null distribution through resampling. Accounting for the dependence between tests increases the power while reducing the variability of the false discovery proportion. This dependence is common in genomics applications, e.g. when combining flow cytometry measurements with microbiome sequence counts.

License GPL-2

Encoding UTF-8

RoxygenNote 7.2.1

Imports phyloseq, ks, reshape2, ggplot2, stats, methods, graphics, grDevices, matrixStats, Matrix

Suggests knitr, rmarkdown, testthat

VignetteBuilder knitr

biocViews Metagenomics, Microbiome, MultipleComparison, FlowCytometry

BugReports https://github.com/CenterForStatistics-UGent/reconsi/issues

LazyData true

git_url https://git.bioconductor.org/packages/reconsi

git_branch RELEASE_3_22

git_last_commit 68b3cde

git_last_commit_date 2025-10-29

Repository Bioconductor 3.22

Date/Publication 2025-12-15

Author Stijn Hawinkel [cre, aut] (ORCID:

<https://orcid.org/0000-0002-4501-5180>)

Maintainer Stijn Hawinkel <stijn.hawinkel@psb.ugent.be>

2 binStats

Contents

 		٠.	•			•					•			٠		•		•	•		19
 			•																		18
 																					17
 																					14
 																					14
 																					13
 																					13
 																					12
																					10
																					9
																					8
																					7
																					6
																					5
																					4
																					4
																					3
																					3
var	var	var	var	var	var	var	var	var	var	var	var	var	var	var	var	var	var	var	var	var	var

binStats

Bin the test statistic into equally sized bins

Description

Bin the test statistic into equally sized bins

Usage

```
binStats(z, nBins = 82L, binEdges = c(-4.1, 4.1))
```

Arguments

z the matrix of permuted test statistics

nBins an integer, the number of bins

binEdges A vector of length 2 with the outer bin edges

Value

Matrix of binned test statistics

calcWeights 3

calcWeights	Obtain weights as posterior probabilities to calculate the consensus null
carenerghes	

Description

Obtain weights as posterior probabilities to calculate the consensus null

Usage

```
calcWeights(logDensPerm, fdr)
```

Arguments

logDensPerm A matrix with B rows of logged density estimates of the B permutation distribu-

tions, and p columns for the p observed test statistics

fdr A vector of local false discovery rates for the observed tests statistics of length

p

Value

A vector of weights of length B

estNormal	Fast estimation of mean and standard deviation of a normal distrbu-
	tion, optionally with weights

Description

Fast estimation of mean and standard deviation of a normal distribution, optionally with weights

Usage

```
estNormal(y, w = NULL, p = length(y))
```

Arguments

У	vector of observations
W	optional weight vector
р	The number of features

Value

A vector of length 2 with mean and standard deviation

4 getApproxCovar

the fraction of true null hypotheses.
the fraction of true null hypotheses.

Description

Estimate the fraction of true null hypotheses.

Usage

```
estP0(statObs, fitAll, z0quantRange, smooth.df, evalVal, assumeNormal)
```

Arguments

statObs A vector of observed z-values fitAll the estimated normal null

z@quantRange a number of quantiles between 0 and 0.5 smooth.df degrees of freedom for the spline smoother evalVal the value of q at which to evaluate the spline

assumeNormal A boolean, should normality be assumed for the null distribution?

Details

A natural spline is used over a range of intervals. Based on the qvalue::qvalue() function and Storey and Tibshirani, 2003

Value

The estimated null fraction, the value of the spline evaluated at the first element of z0quantRange

getApproxCovar	Obtain a null covariance matrix of binned test statistics

Description

Obtain a null covariance matrix of binned test statistics

Usage

```
getApproxCovar(statsPerm, ...)
```

Arguments

statsPerm The pxB matrix of permutation z-values in the columns
... passed on to binStats

Value

The covariance matrix of binned z-values

getC1prop 5

Note

This is not the covariance matrix of the p test statistic, nor of the data! It is an approximate covariance matrix of binned test statistics for visualization and diagnostic purposes.

Examples

```
p = 200; n = 50; B = 5e1
x = rep(c(0,1), each = n/2)
mat = cbind(
matrix(rnorm(n*p/10, mean = 5+x),n,p/10), #DA
matrix(rnorm(n*p*9/10, mean = 5),n,p*9/10) #Non DA
)
mat = mat = mat + rnorm(n, sd = 0.3) #Introduce some dependence
fdrRes = reconsi(mat, x, B = B)
corMat = getApproxCovar(fdrRes$statsPerm)
```

getC1prop

Find the dependence pat C1 of the approximate covariance matrix, and extract the ratio of the first eigenvalue to the sum of all positive eigenvalues

Description

Find the dependence pat C1 of the approximate covariance matrix, and extract the ratio of the first eigenvalue to the sum of all positive eigenvalues

Usage

```
getC1prop(statsPerm, numEig = 1, ...)
```

Arguments

```
statsPerm Matrix of permuted test statistics
numEig An integer, number of first eigenvalues
... passed onto binStats
```

Value

A proportion indicating the ratio of the first eigenvalues to the sum of all eigenvalues

Examples

```
p = 200; n = 50; B = 5e1
x = rep(c(0,1), each = n/2)
mat = cbind(
matrix(rnorm(n*p/10, mean = 5+x),n,p/10), #DA
matrix(rnorm(n*p*9/10, mean = 5),n,p*9/10) #Non DA
)
mat = mat = mat + rnorm(n, sd = 0.3) #Introduce some dependence
fdrRes = reconsi(mat, x, B = B)
getC1prop(fdrRes$statsPerm)
```

6 getFdr

. = 1	
getFdr	Calculate tail-area (Fdr) and local (fdr) false discovery rates, based
	on a certain null distribution

Description

Calculate tail-area (Fdr) and local (fdr) false discovery rates, based on a certain null distribution

Usage

```
getFdr(
   statObs,
   fitAll,
   fdr,
   p,
   p0,
   zValsDensObs,
   smoothObs,
   assumeNormal,
   fitObs,
   ...
)
```

Arguments

stat0bs	Vector of observed z-values
fitAll	The parameters of the estimated random null
fdr	local false discovery rate, already estimated
p	the number of hypotheses
p0	The estimated fraction of null hypotheses
zValsDensObs	estimated densities of observed test statistics
smoothObs	A boolean, should estimated observed densities of the test statistics be used in estimating the Fdr
assumeNormal	A boolean, should normality be assumed for the null distribution?
fitObs	The kernel density estimate object of all test statistics
	more arguments, ignored

Value

A list with components

Fdr Tail are false discovery rate fdr Local false discovery rate getG0 7

getG0 Obtain the consensus null

Description

Obtain the consensus null

Usage

```
getG0(
   statObs,
   statsPerm,
   z0Quant,
   gridsize,
   maxIter,
   tol,
   estP0args,
   testPargs,
   B,
   p,
   pi0,
   assumeNormal,
   resamAssumeNormal
)
```

Arguments

stat0bs A vector of lenght p with observed test statistics statsPerm A pxB matrix with permuation z-values a vector of length of quantiles defining the central part R of the distribution. If a z0Quant single number is supplied, then (z0quant, 1-z0quant) will be used gridsize An integer, the gridsize for the density estimation An integer, the maximum number of iterations in determining R maxIter tol The convergence tolerance. A list of arguments passed on to the estP0args() function estP0args A list of arguments passed on to quantileFun testPargs В an integer, the number of permutations an integer, the number of hypotheses р A known fraction of true null hypotheses pi0 assumeNormal A boolean, should normality be assumed for the null distribution? resamAssumeNormal

A boolean, should normality be assumed for resampling dists

8 getTestStats

Value

A list with following entries

PermDensFits The permutation density fits

zSeq The support of the kernel for density estimation zValsDensObs The estimated densities of the observed z-values

convergence A boolean, has the algorithm converged?

weights Vector of length B with weights for the permutation distributions fdr Estimated local false discovery rate along the support of the kernel

p0 The estimated fraction of true null hypotheses

iter The number of iterations fitAll The consensus null fit

getTestStats A function to calculate observed and permuation z-statistics on a n-

by-p matrix of observations

Description

A function to calculate observed and permuation z-statistics on a n-by-p matrix of observations

Usage

```
getTestStats(
   Y,
   center,
   test = "wilcox.test",
   X,
   B,
   argList,
   tieBreakRan,
   replace,
   scale
)
```

Arguments

Y The nxp data matrix

center a boolean, should data be centered prior to permuation test A function name, possibly user defined. See details.

x A vector defining the groups. Will be coerced to factor.

B an integer, the number of permuations

argList A list of further arguments passed on to the test function

tieBreakRan A boolean, should ties of permutation test statistics be broken randomly? If not,

midranks are used

replace A boolean. If FALSE, samples are permuted (resampled without replacement),

if TRUE the samples are bootstrapped (resampled with replacement)

scale a boolean, should data be scaled prior to resampling

getTstat 9

Details

For test "wilcox.test" and "t.test", fast custom implementations are used. Other functions can be supplied but must accept a y outcome variable, a x as grouping variable, and possibly a list of other arguments. It must return all arguments needed to evaluate its quantile function if z-values are to be used.

Value

A list with components

statObs A vector of length p of observed test statistics

statsPerm A p-by-B matrix of permutation test statistics

resamDesign The resampling design

getTstat

A function to obtain a t-test statistic efficiently. For internal use only

Description

A function to obtain a t-test statistic efficiently. For internal use only

Usage

```
getTstat(y1, y2, mm, nn)
```

Arguments

y1, y2 vectors of observeed values in the two groups

mm, nn number of observations in the corresponding groups

Value

A list with items

tstat The t-test statistic

df The degrees of freedom (Welch approximation)

10 plotApproxCovar

plotApproxCovar

Plot an approximation of the correlation structure of the test statistics

Description

Plot an approximation of the correlation structure of the test statistics

Usage

```
plotApproxCovar(
  reconsiFit,
  col = colorRampPalette(c("yellow", "blue"))(12),
  x = seq(-4.2, 4.2, 0.1),
  y = seq(-4.2, 4.2, 0.1),
  xlab = "Z-values",
  ylab = "Z-values",
  nBins = 82L,
  binEdges = c(-4.1, 4.1),
  ...
)
```

Arguments

```
\label{eq:consifit} \begin{tabular}{ll} \beg
```

Details

By default, yellow indicates negative correlation between bin counts, blue positive correlation

Value

invisible()

Note

This is not the covariance matrix of the p test statistic, nor of the data! It is an approximate covariance matrix of binned test statistics for visualization purposes. See plotCovar for the full covariance matrix.

See Also

```
plotCovar, getApproxCovar
```

plotCovar 11

Examples

```
p = 200; n = 50; B = 5e1 \\ x = rep(c(0,1), each = n/2) \\ mat = cbind( \\ matrix(rnorm(n*p/10, mean = 5+x),n,p/10), #DA \\ matrix(rnorm(n*p*9/10, mean = 5),n,p*9/10) #Non DA ) \\ mat = mat = mat + rnorm(n, sd = 0.3) #Introduce some dependence fdrRes = reconsi(mat, x, B = B) \\ plotApproxCovar(fdrRes)
```

plotCovar

Plot an the corvariance matrix of the test statistics estimated through permutations

Description

Plot an the corvariance matrix of the test statistics estimated through permutations

Usage

```
plotCovar(
  reconsiFit,
  col = colorRampPalette(c("yellow", "blue"))(12),
  xlab = "Test statistic index",
  ylab = xlab,
  ...
)
```

Arguments

```
reconsiFit The reconsi fit col, xlab, ylab, ...
```

A list of arguments for the image() function.

Details

By default, yellow indicates negative correlaton between test statistics, blue positive correlation

Value

invisible()

Note

Note the difference with the plotApproxCovar function, where the covariances between binned test statistics are shown to get an idea between covariances between tail and center values of the univariate null distribution. Here the covariance matrix between all test statistics is shown

See Also

```
plotApproxCovar
```

12 plotNull

Examples

```
p = 200; n = 50; B = 5e1 \\ x = rep(c(0,1), each = n/2) \\ mat = cbind( \\ matrix(rnorm(n*p/10, mean = 5+x),n,p/10), #DA \\ matrix(rnorm(n*p*9/10, mean = 5),n,p*9/10) #Non DA ) \\ mat = mat = mat + rnorm(n, sd = 0.3) #Introduce some dependence fdrRes = reconsi(mat, x, B = B) \\ plotCovar(fdrRes)
```

plotNull

Plot the obtained null distribution along with a histogram of observed test statistics

Description

Plot the obtained null distribution along with a histogram of observed test statistics

Usage

```
plotNull(
   fit,
   lowColor = "yellow",
   highColor = "blue",
   idNull = NULL,
   nResampleCurves = length(fit$Weights),
   hSize = 0.5
)
```

Arguments

```
fit an object returned by the reconsi() (or testDAA()) function lowColor, highColor

The low and high ends of the colour scale

idNull indices of known null taxa

nResampleCurves

The number of resampling null distributions to plot

hSize A double, the size of the line of the collapsed null estimate
```

Value

```
a ggplot2 plot object
```

Examples

```
p = 180; n = 50; B = 1e2

#Low number of resamples keeps computation time down

x = rep(c(0,1), each = n/2)

mat = cbind(

matrix(rnorm(n*p/10, mean = 5+x),n,p/10), #DA

matrix(rnorm(n*p*9/10, mean = 5),n,p*9/10) #Non DA
```

ptEdit 13

```
) #Provide just the matrix and grouping factor, and test using the random null fdrRes = reconsi(mat, x, B = B) plotNull(fdrRes)
```

ptEdit

A custom function to calculate the distribution function of the t-test statistic. For internal use only

Description

A custom function to calculate the distribution function of the t-test statistic. For internal use only

Usage

ptEdit(q)

Arguments

q

a vector with t-statistic and degrees of freedom

Value

A value between 0 and 1, the evaluation of the cdf

qtEdit

A custom function to calculate the quantile function of the t-test statistic. For internal use only

Description

A custom function to calculate the quantile function of the t-test statistic. For internal use only

Usage

```
qtEdit(p)
```

Arguments

p

a vector with quantile and degrees of freedom

Value

the corresponding quantile

14 reconsi

quantCorrect

Correct quantiles by not returning 0 or 1

Description

Correct quantiles by not returning 0 or 1

Usage

```
quantCorrect(quants)
```

Arguments

quants

A vector of quantiles

Value

The same vector of quantiles but without 0 or 1 values

reconsi

Perform simultaneous inference through collapsed resampling null distributions

Description

Perform simultaneous inference through collapsed resampling null distributions

Usage

```
reconsi(
 Υ,
 x = NULL
 B = 1000L
 test = "wilcox.test",
 argList = list(),
 distFun = "pnorm",
 zValues = TRUE,
 testPargs = list(),
 z0Quant = 0.25,
 gridsize = 801L,
 maxIter = 100L,
 tol = 1e-06,
 zVals = NULL,
 center = FALSE,
 replace = is.null(x),
 assumeNormal = TRUE,
 estP0args = list(z0quantRange = seq(0.05, 0.45, 0.0125), smooth.df = 3, evalVal = 0.05),
 resamZvals = FALSE,
 smoothObs = TRUE,
 scale = FALSE,
```

15 reconsi

```
tieBreakRan = FALSE,
pi0 = NULL,
resamAssumeNormal = TRUE
```

Arguments

Υ	the matrix of sequencing counts
х	a grouping factor. If provided, this grouping factor is permuted. Otherwise a bootstrap procedure is performed
В	the number of resampling instances
test	Character string, giving the name of the function to test for differential absolute abundance. Must accept the formula interface. Features with tests resulting in observed NA test statistics will be discarded
argList	A list of arguments, passed on to the testing function
distFun	the distribution function of the test statistic, or its name. Must at least accept an argument named \dot{q} , \dot{p} and \dot{x} respectively.
zValues	A boolean, should test statistics be converted to z-values. See details
testPargs	A list of arguments passed on to distFun
z0Quant	A vector of length 2 of quantiles of the null distribution, in between which only null values are expected
gridsize	The number of bins for the kernel density estimates
maxIter	An integer, the maximum number of iterations in the estimation of the null distribution
tol	The tolerance for the infinity norm of the central borders in the iterative procedure
zVals	An optional list of observed (statObs) and resampling (statsPerm) z-values. If supplied, the calculation of the observed and resampling test statistics is skipped and the algorithm proceeds with calculation of the consensus null distribution
center	A boolean, should observations be centered in each group prior to permuations? See details.
replace	A boolean. Should resampling occur with replacement (boostrap) or without replacement (permutation)
assumeNormal	A boolean, should normality be assumed for the null distribution?
estP0args	A list of arguments passed on to the estP0 function
resamZvals	$\label{lem:condition} A \ boolean, should \ resampling \ rather \ than \ theoretical \ null \ distributions \ be \ used?$
smoothObs	A boolean, should the fitted rather than estimated observed distribution be used in the Fdr calculation?
scale	a boolean, should data be scaled prior to resampling
tieBreakRan	A boolean, should ties of resampling test statistics be broken randomly? If not, midranks are used
pi0	A known fraction of true null hypotheses. If provided, the fraction of true null hypotheses will not be estimated. Mainly for oracle purposes.
resamAssumeNorm	nal

A boolean, should normality be assumed for resampling dists

16 reconsi

Details

Efron (2007) centers the observations in each group prior to permutation. As permutations will remove any genuine group differences anyway, we skip this step by default. If zValues = FALSE, the density is fitted on the original test statistics rather than converted to z-values. This unlocks the procedure for test statistics with unknown distributions, but may be numerically less stable.

Value

A list with entries

statsPerm Resampling Z-values statObs Observed Z-values

distFun Density, distribution and quantile function as given

testPargs Same as given

zValues A boolean, were z-values used?

resamZvals A boolean, were the resampling null distribution used?

cdfValObs Cumulative distribution function evaluation of observed test statistics

p@estimated A boolean, was the fraction of true null hypotheses estimated from the data?

Fdr, fdr Estimates of tail-area and local false discovery rates

p0 Estimated or supplied fraction of true null hypotheses

iter Number of iterations executed

fitAll Mean and standard deviation estimated collapsed null

PermDensFits Mean and standard deviations of resamples

convergence A boolean, did the iterative algorithm converge?

zSeq Basis for the evaluation of the densities weights weights of the resampling distributions

zValsDensObs Estimated overall densities, evaluated in zSeq

Note

Ideally, it would be better to only use unique resamples, to avoid unnecesarry replicated calculations of the same test statistics. Yet this issue is almost always ignored in practice; as the sample size grows it also becomes irrelevant. Notice also that this would require to place weights in case of the bootstrap, as some bootstrap samples are more likely than others.

Examples

```
#Important notice: low number of resamples B necessary to keep # computation time down, but not recommended. Pray set B at 200 or higher. p = 60; n = 20; B = 5e1 x = rep(c(0,1), each = n/2) mat = cbind( matrix(rnorm(n*p/10, mean = 5+x), n, p/10), #DA matrix(rnorm(n*p*9/10, mean = 5), n, p*9/10) #Non DA ) fdrRes = reconsi(mat, x, B = B) fdrRes$p0 #Indeed close to 0.9 estFdr = fdrRes$Fdr
```

rowMultiply 17

```
#The estimated tail-area false discovery rates.
#With another type of test. Need to supply quantile function in this case
fdrResLm = reconsi(mat, x, B = B,
test = function(x, y){
fit = lm(y^x)
c(summary(fit)$coef["x","t value"], fit$df.residual)},
distFun = function(q)\{pt(q = q[1], df = q[2])\}
#With a test statistic without known null distribution(for small samples)
fdrResKruskal = reconsi(mat, x, B = B,
test = function(x, y){
kruskal.test(y~x)$statistic}, zValues = FALSE)
#Provide an additional covariate through the 'argList' argument
z = rpois(n, lambda = 2)
fdrResLmZ = reconsi(mat, x, B = B,
test = function(x, y, z){
fit = lm(y\sim x+z)
c(summary(fit)$coef["x","t value"], fit$df.residual)},
distFun = function(q)\{pt(q = q[1], df = q[2])\},
argList = list(z = z))
#When nog grouping variable is provided, a bootstrap is performed
matBoot = cbind(
matrix(rnorm(n*p/10, mean = 1), n, p/10), #DA
matrix(rnorm(n*p*9/10, mean = 0), n, p*9/10) #Non DA
fdrResBoot = reconsi(matBoot, B = B,
test = function(y, x){testRes = t.test(y, mu = 0, var.equal = TRUE);
c(testRes$statistic, testRes$parameter)},
distFun = function(q)\{pt(q = q[1], df = q[2])\},
center = TRUE, replace = TRUE)
```

rowMultiply

A function to efficiently row multiply a a-by-b matrix by a vector of length b. More memory intensive but that does not matter with given matrix sizes

Description

A function to efficiently row multiply a a-by-b matrix by a vector of length b. More memory intensive but that does not matter with given matrix sizes

Usage

```
rowMultiply(matrix, vector)
```

Arguments

matrix a numeric matrix of dimension a-by-b vector a numeric vector of length b

18 testDAA

Details

```
t(t(matrix)*vector) but then faster
```

Value

a matrix, row multplied by the vector

stabExp

A function to numerically stabilize an exponentiation. For internal use only

Description

A function to numerically stabilize an exponentiation. For internal use only

Usage

```
stabExp(exps)
```

Arguments

exps

the vector to be exponentiated

Value

the vector with the maximum subtracted

testDAA

A function to test for differential absolute abundance on a phyloseq object

Description

A function to test for differential absolute abundance on a phyloseq object

Usage

```
testDAA(Y, ...)
## S4 method for signature 'phyloseq'
testDAA(Y, groupName, FCname, ...)
## S4 method for signature 'matrix'
testDAA(Y, FC, x, S = rowSums(Y), tieBreakRan = TRUE, assumeNormal = TRUE, ...)
```

Vandeputte 19

Arguments

Y A phyloseq object, or a data matrix with samples in the rows and OTUs in the

columns

... passed on to the reconsi() function

groupName A character string, the name of a variable in physeq indicating the grouping

factor

FCname A character string, the name of a variable in physeq containing the total cell

count

FC a vector of length n with total flow cytometry cell counts

x a grouping factor of length n

S a vector of library sizes. Will be calculated if not provided

tieBreakRan A boolean, should ties be broken at random.

assumeNormal A boolean, should normality be assumed for the null distribution?

Value

See the reconsi() function

Examples

```
#Test for phyloseq object
library(phyloseq)
data("VandeputteData")
VandeputtePruned = prune_samples(Vandeputte,
samples = sample_names(Vandeputte)[20:40])
testVanDePutte = testDAA(VandeputtePruned, "Health.status", "absCountFrozen",
B = 15)
#Test for matrix
testMat = testDAA(as(otu_table(VandeputtePruned), "matrix"),
get_variable(VandeputtePruned, "Health.status"),
get_variable(VandeputtePruned, "absCountFrozen"), B = 15)
```

Vandeputte

Microbiomes of Crohn's disease patients and healthy controls

Description

Microbiome sequencing data of Crohn's disease patients, and healthy controls, together with other baseline covariates. Both sequencing and flow cytometry data are available.

Usage

Vandeputte

Format

A phyloseq object with an OTU-table and sample data

otu_table Count data matrix of 234 taxa in 135 samplessample_data Data frame of patient covariates

20 Vandeputte

Source

https://www.ncbi.nlm.nih.gov/pubmed/29143816

Index

```
* datasets
    Vandeputte, 19
binStats, 2
calcWeights, 3
estNormal, 3
estP0,4
getApproxCovar, 4, 10
getC1prop, 5
getFdr, 6
getG0, 7
getTestStats, 8
getTstat, 9
plotApproxCovar, 10, 11
plotCovar, 10, 11
plotNull, \\ \textcolor{red}{12}
ptEdit, 13
\mathsf{qtEdit}, \textcolor{red}{13}
quantCorrect, 14
reconsi, 14
rowMultiply, 17
stabExp, 18
testDAA, 18
testDAA, matrix-method (testDAA), 18
testDAA, phyloseq-method (testDAA), 18
Vandeputte, 19
```