Package 'gscreend'

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Type Package

Title Analysis of pooled genetic screens

Version 1.23.0

Description Package for the analysis of pooled genetic screens (e.g. CRISPR-KO). The analysis of such screens is based on the comparison of gRNA abundances before and after a cell proliferation phase. The gscreend packages takes gRNA counts as input and allows detection of genes whose knockout decreases or increases cell proliferation.

License GPL-3

Encoding UTF-8

LazyData false

Depends R (>= 3.6)

Imports SummarizedExperiment, nloptr, fGarch, methods, BiocParallel, graphics

Suggests knitr, testthat, rmarkdown, BiocStyle

VignetteBuilder knitr

RoxygenNote 7.2.3

biocViews Software, StatisticalMethod, PooledScreens, CRISPR

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BugReports https://github.com/imkeller/gscreend/issues

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Contents

`assignGeneData` *Calculate gene rank*

Description

Calculate gene rank

Usage

```
assignGeneData(object, alpha_cutoff)
```

Arguments

`object`	PoolScreenExp object
`alpha_cutoff`	alpha cutoff for alpha-RRA (default: 0.05)

Value

object

calculateIntervalFits *Calculate fit parameters for every subset of data*

Description

Calculate fit parameters for every subset of data

Usage

```
calculateIntervalFits(object, quant1, quant2)
```

Arguments

object	PoolScreenExp object
quant1	lower quantile for least quantile of squares regression (default: 0.1)
quant2	upper quantile for least quantile of squares regression (default: 0.9)

Value

object

calculateLFC *Calculate log fold changes*

Description

Calculate log fold changes

Usage

```
calculateLFC(object)
```

Arguments

object	PoolScreenExp object

Value

object

calculatePValues *Calculate p-values*

Description

Calculate p-values

Usage

```
calculatePValues(object)
```

Arguments

object	PoolScreenExp object

Value

object

createPoolScreenExp *Create PoolScreenExp Experiment*

Description

Create PoolScreenExp Experiment

Usage

```
createPoolScreenExp(data)
```

Arguments

data	Input data object containing gRNA level data (SummarizedExperiment)

Value

object PoolScreenExp object

Examples

```
raw_counts <- read.table(
                        system.file('extdata', 'simulated_counts.txt',
                        package = 'gscreend'),
                        header=TRUE)

counts_matrix <- cbind(raw_counts$library0, raw_counts$R0_0, raw_counts$R1_0)

rowData <- data.frame(sgRNA_id = raw_counts$sgrna_id,
gene = raw_counts$Gene)

colData <- data.frame(samplename = c('library', 'R1', 'R2'),
```

```
timepoint = c('T0', 'T1', 'T1'))

library(SummarizedExperiment)
se <- SummarizedExperiment(assays=list(counts=counts_matrix),
rowData=rowData, colData=colData)

# create a PoolScreenExp experiment
pse <- createPoolScreenExp(se)
```

`createPoolScreenExpFromSE` *Create PoolScreenExp Experiment from summarized experiment*

Description

Create PoolScreenExp Experiment from summarized experiment

Usage

```
createPoolScreenExpFromSE(data)
```

Arguments

`data`	SummarizedExperiment object containing gRNA level data

Value

object

`defineFittingIntervals` *Calculate interval limits for splitting data into subsets*

Description

Calculate interval limits for splitting data into subsets

Usage

```
defineFittingIntervals(object)
```

Arguments

`object`	PoolScreenExp object

Value

object

fit_least_quantile *Fit paramters for skew normal distribution*

Description

Fit paramters for skew normal distribution

Usage

```
fit_least_quantile(LFC, quant1, quant2)
```

Arguments

LFC	logarithmic fold changes of gRNA counts
quant1	lower quantile for least quantile of squares regression (default: 0.1)
quant2	upper quantile for least quantile of squares regression (default: 0.9)

Value

fit_skewnorm

GeneData *GeneData: set and retrieve GeneData of PoolScreenExp*

Description

GeneData: set and retrieve GeneData of PoolScreenExp

Usage

```
GeneData(x)
```

Arguments

x	PoolScreenExp object

Value

Gene slot of the object

Examples

```
# import a PoolScreenExp object that has been generated using
# RunGscreend()
pse_an <- readRDS(
system.file('extdata', 'gscreend_analysed_experiment.RData',
package = 'gscreend'))

GeneData(pse_an)
```

GeneData,PoolScreenExp-method

Accessor function for the Gene slot of the PoolScreenExp class

Description

Accessor function for the Gene slot of the PoolScreenExp class

Usage

```
## S4 method for signature 'PoolScreenExp'
GeneData(x)
```

Arguments

x	PoolScreenExp object

Value

Gene slot of the object

Examples

```
# import a PoolScreenExp object that has been generated using
# RunGscreend()
pse_an <- readRDS(
system.file('extdata', 'gscreend_analysed_experiment.RData',
package = 'gscreend'))

GeneData(pse_an)
```

normalizePoolScreenExp

Normalize raw count

Description

Normalize raw count

Usage

```
normalizePoolScreenExp(object)
```

Arguments

object	PoolScreenExp object

Value

object

plotModelParameters *Plot model parameters from the fitting*

Description

Plot model parameters from the fitting

Usage

```
plotModelParameters(object)
```

Arguments

object	PoolScreenExp object

Value

plot

Examples

```
# import a PoolScreenExp object that has been generated using
# RunGscreend()
pse_an <- readRDS(
system.file('extdata', 'gscreend_analysed_experiment.RData',
package = 'gscreend'))
plotModelParameters(pse_an)
```

plotReplicateCorrelation *Plot replicate correlation*

Description

Plot replicate correlation

Usage

```
plotReplicateCorrelation(object, rep1 = "R1", rep2 = "R2")
```

Arguments

object	PoolScreenExp object
rep1	Name of replicate 1
rep2	Name of replicate 2

Value

replicate_plot

Examples

```
# import a PoolScreenExp object that has been generated using RunGscreend()
pse_an <- readRDS(
system.file('extdata', 'gscreend_analysed_experiment.RData',
package = 'gscreend'))
plotReplicateCorrelation(pse_an, rep1 = 'R1', rep2 = 'R2')
```

PoolScreenExp-class *Class to store pooled CRISPR screening experiment*

Description

The poolScreenExp class is an S4 class used to store sgRNA and gene related data as well as parameters necessary for statistical model.

Slots

`sgRNAData` A SummarizedExperiment containing the data related to sgRNAs.

`FittingIntervals` A vector defining the limits of the intervals used for fitting of null model.

`LFCModelParameters` A vector of parameters estimated when fitting the null model.

`GeneData` SummarizedExperiment containing the data related to genes.

`FittingOptions` A named list with options for fitting: IntervalFraction - fraction of sgRNAs used in every fitting interval (default 0.1), alphaCutoff - alpha cutoff for alpha RRA algorithm (default: 0.05).

ResultsTable *Extract a results table*

Description

Extract a results table

Usage

```
ResultsTable(object, direction = "negative")
```

Arguments

`object`	PoolScreenExp object
`direction`	Whether the table should contain information on positive or negative fold changes ['negative'| 'positive']

Value

plot

Examples

```
# import a PoolScreenExp object that has been generated using
# RunGscreend()
pse_an <- readRDS(
system.file('extdata', 'gscreend_analysed_experiment.RData',
package = 'gscreend'))
ResultsTable(pse_an, direction = 'negative')
```

RunGscreend *run gscreend*

Description

run gscreend

Usage

```
RunGscreend(object, quant1 = 0.1, quant2 = 0.9, alphacutoff = 0.05)
```

Arguments

object	PoolScreenExp object
quant1	lower quantile for least quantile of squares regression (default: 0.1)
quant2	upper quantile for least quantile of squares regression (default: 0.9)
alphacutoff	alpha cutoff for alpha-RRA (default: 0.05)

Value

object

Examples

```
raw_counts <- read.table(
                        system.file('extdata', 'simulated_counts.txt',
                        package = 'gscreend'),
                        header=TRUE)

# Create the PoolScreenExp to be analyzed
counts_matrix <- cbind(raw_counts$library0, raw_counts$R0_0, raw_counts$R1_0)

rowData <- data.frame(sgRNA_id = raw_counts$sgrna_id,
gene = raw_counts$Gene)

colData <- data.frame(samplename = c('library', 'R1', 'R2'),
timepoint = c('T0', 'T1', 'T1'))

library(SummarizedExperiment)
se <- SummarizedExperiment(assays=list(counts=counts_matrix),
rowData=rowData, colData=colData)

pse <- createPoolScreenExp(se)
```

```
# Run Analysis
pse_an <- RunGscreend(pse)
```

---

sgRNAData *sgRNAData: set and retrieve sgRNAData of PoolScreenExp*

Description

sgRNAData: set and retrieve sgRNAData of PoolScreenExp

Usage

```
sgRNAData(x)
```

Arguments

x	PoolScreenExp object

Value

sgRNA slot of the object

Examples

```
# import a PoolScreenExp object that has been generated using
# RunGscreend()
pse_an <- readRDS(
system.file('extdata', 'gscreend_analysed_experiment.RData',
package = 'gscreend'))

sgRNAData(pse_an)
```

---

sgRNAData,PoolScreenExp-method
Accessor function for the sgRNA slot of the PoolScreenExp class

---

Description

Accessor function for the sgRNA slot of the PoolScreenExp class

Usage

```
## S4 method for signature 'PoolScreenExp'
sgRNAData(x)
```

Arguments

x	PoolScreenExp object

Value

sgRNA slot of the object

Examples

```
# import a PoolScreenExp object that has been generated using
# RunGscreend()
pse_an <- readRDS(
system.file('extdata', 'gscreend_analysed_experiment.RData',
package = 'gscreend'))

sgRNAData(pse_an)
```

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