

Package ‘DChIPRep’

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Title DChIPRep - Analysis of chromatin modification ChIP-Seq data with replication

Version 1.0.4

Description The DChIPRep package implements a methodology to assess differences between chromatin modification profiles in replicated ChIP-Seq studies as described in Chabbert et. al - <http://www.dx.doi.org/10.15252/msb.20145776>.

Depends R (>= 3.2.0)

Imports methods, ggplot2, DESeq2, fdrtool, reshape2, GenomicRanges, smoothmest, plyr, tidyr, assertthat, S4Vectors

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VignetteBuilder knitr

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DChIPRep	<i>DChIPRep: A package for differential analysis of histone modification ChIP-Seq profiles</i>
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Description

The DChIPRep package provides functions to perform a differential analysis of histone modification profiles at base-pair resolution

DChIPRep functions

The DChIPRep packages provides functions for data import `importData` and performing position wise tests. After data import, a `DChIPRepResults` object on which the function `runTesting` is run to perform the tests and add the result to the object. Then, plots can be created from this object. See the vignette for additional details: `vignette("DChIPRepVignette")`

DChIPRepResults-class	<i>DChIPRepResults object and constructor</i>
-----------------------	---

Description

The `DChIPRepResults` contains a `DESeqDataSet` as obtained after the initial import.

Usage

```
DChIPRepResults(object)
```

Arguments

object	A <code>DESeqDataSet</code>
--------	-----------------------------

Value

A DChIPRepResult object.

Examples

```
data(testData)
dcr <- DChIPRepResults(testData)
```

DESeq2Data

Accessors for the 'DESeq2Data' slot of a DChIPRepResults object.

Description

The slot contains the DESeqDataSet as it is obtained after the initial data import. The DESeqDataSet contains the counts per position and the normalization factors as computed using the input counts.

Usage

```
## S4 method for signature 'DChIPRepResults'
DESeq2Data(object)

## S4 replacement method for signature 'DChIPRepResults,DESeqDataSet'
DESeq2Data(object) <- value
```

Arguments

object	a DChIPRepResults object
value	A DESeqDataSet object

Value

the DESeq2Data object contained in the DChIPRepResults object

Examples

```
data(testData)
dcr <- DChIPRepResults(testData)
DESeq2Data(dcr)
```

exampleChipData *An example ChIP data.*

Description

An example Chip data set that can be used with [importDataFromMatrices](#). Its associated sample table can be accessed via `data(data(exampleSampleTable))`.

Usage

```
data(exampleChipData)
```

Format

a matrix

Value

a matrix

See Also

[exampleSampleTable](#) [exampleInputData](#)

exampleInputData *An example input data.*

Description

An example input data set that can be used with [importDataFromMatrices](#). Its associated sample table can be accessed via `data(data(exampleSampleTable))`.

Usage

```
data(exampleInputData)
```

Format

a matrix

Value

a matrix

See Also

[exampleSampleTable](#) [exampleChipData](#)

exampleSampleTable	<i>An example sample table data.frame</i>
--------------------	---

Description

An example sample table

Usage

```
data(exampleSampleTable)
```

Format

a data.frame

Value

a data.frame

See Also

[exampleChipData](#) [exampleInputData](#)

FDRresults	<i>Accessor and setter for the 'FDRresults' slot of a DChIPRepResults object.</i>
------------	---

Description

The slot contains the results of the FDR estimation as performed within the function [runTesting](#). It is the complete output of the [fdrtool](#) function.

Usage

```
## S4 method for signature 'DChIPRepResults'  
FDRresults(object)  
  
## S4 replacement method for signature 'DChIPRepResults,list'  
FDRresults(object) <- value
```

Arguments

object	a DChIPRepResults object
value	A DESeqDataSet object

Value

a list containing the estimated false discovery rates

Examples

```
data(testData)
dcr <- DChIPRepResults(testData)
dcr <- runTesting(dcr)
str(FDRresults(dcr))
```

getMATfromDataFrame	<i>Helper function to turn a data.frame into a matrix and remove the ID column.</i>
---------------------	---

Description

This function takes a data.frame, with the genomic features (e.g. transcripts or genes) in the rows and the positions upstream and downstream of the TSS in the columns as well as a column ID containing a genomic feature ID and returns the data.frame with the ID column removed. The input for this function are tables obtained after running the Python import script.

Usage

```
getMATfromDataFrame(df, ID = "name")
```

Arguments

df	the input data frame with positions in the columns and the genomic features in the rows.
ID	the name of the ID column to be removed.

Value

a matrix with the ID column removed

Examples

```
data(exampleSampleTable)
directory <- file.path(system.file("extdata", package="DChIPRep"))
df <- lapply(file.path(directory, exampleSampleTable$Input), read.delim)[[1]]
mat <- getMATfromDataFrame(df)
```

importData	<i>Import the data after running the Python script</i>
------------	--

Description

This function imports the data from the count table files as returned by the accompanying Python script.

Usage

```
importData(sampleTable, directory = "", ID = "name", ...)
```

Arguments

sampleTable	a data.frame that has to contain the columns ChiP, Input, sampleID, upstream, downstream and condition. Each row of the table describes one experimental sample. Each row of the table describes one experimental sample. See <code>data(exampleSampleTable)</code> for an example table. and the vignette for further information.
directory	the directory relative to which the filenames are specified given as a character.
ID	character giving the name of the feature identifier column in the count tables. Defaults to "name"
...	parameters passed to summarizeCountsPerPosition

Value

a DChIPRepResults object containing the imported data as a [DESeqDataSet](#).

Examples

```
data(exampleSampleTable)
directory <- file.path(system.file("extdata", package="DChIPRep"))
importedData <- importData(exampleSampleTable, directory)
```

importDataFromMatrices	<i>Import the data from ChiP and input matrices</i>
------------------------	---

Description

This function imports the data from two matrices that contain counts summarized per position. It computes the normalization factors from the input (one per position) and creates a DChIPRepResults object.

Usage

```
importDataFromMatrices(inputData, chipData, sampleTable)
```

Arguments

inputData	a matrix containing the counts for the input per position.
chipData	a matrix containing the counts for the ChIP per position.
sampleTable	a data.frame that has to contain the columns sampleID, upstream, downstream and condition. Each row of the table describes one experimental sample. See <code>data(exampleSampleTable)</code> for an example table. and the vignette for further information.

Details

The normalization factors are computed as $t(t(inputData) * (covC/covI))$, Where covC and covI contain the total sum of the ChIP and the input samples. Zero normalization factors can arise if the input has zero counts for certain positions. That's why input values equal to zero are set to 1 in order to always obtain valid normalizationFactors.

Value

a DChIPRepResults object containing the imported data as a [DESeqDataSet](#).

Examples

```
data(exampleSampleTable)
data(exampleInputData)
data(exampleChipData)
imDataFromMatrices <- importDataFromMatrices(inputData = exampleInputData,
chipData = exampleChipData,
sampleTable = exampleSampleTable)
```

plotProfiles

Produce a TSS plot of the two conditions in the data

Description

This function plots the positionwise mean of the log₂ of the normalized counts of the two conditions after [runTesting](#) has been run on a DChIPRepResults object.

Usage

```
## S4 method for signature 'DChIPRepResults'
plotProfiles(object, meanFunction = function(x) {
  smhuber(x)$mu }, ...)
```


Arguments

object a DChIPRepResults object after [runTesting](#)

meanFunction a function to compute the positionwise mean per group, defaults to a Huber estimator of the mean.

... additional parameters for plotting (NOT YET IMPLEMENTED)

Value

a ggplot2 object

Examples

```
if (requireNamespace("mgcv", quietly=TRUE)) {
  data(testData)
  dcr <- DChIPRepResults(testData)
  dcr <- runTesting(dcr)
  plotProfiles(dcr)
}
```

plotSignificance *Produce a plot that colors the positions identified as significant*

Description

This function plots the positionwise mean of the two conditions after [runTesting](#) has been run on a DChIPRepResults object. The points corresponding to significant positions are colored black in both of the conditions. The function returns the plot as a ggplot2 object that can be modified afterwards.

Usage

```
## S4 method for signature 'DChIPRepResults'
plotSignificance(object,
  meanFunction = function(x) { smhuber(x)$mu }, lfdThreshold = 0.2, ...)
```

Arguments

object a DChIPRepResults object after [runTesting](#)

meanFunction a function to compute the positionwise mean per group, defaults to a Huber estimator of the mean.

lfdThreshold Threshold for the local FDR

... additional parameters for plotting (NOT YET IMPLEMENTED)

Value

a ggplot2 object

Examples

```
data(testData)
dcr <- DChIPRepResults(testData)
dcr <- runTesting(dcr)
plotSignificance(dcr)
```

resultsDChIPRep	<i>Accessors and setter for the 'results' slot of a DChIPRepResults object.</i>
-----------------	---

Description

The slot contains the results of the position wise tests in a data.frame after running the function [runTesting](#). It is a modified output of the [results](#) function of the DESeq2 package.

Usage

```
## S4 method for signature 'DChIPRepResults'
resultsDChIPRep(object)

## S4 replacement method for signature 'DChIPRepResults,list'
resultsDChIPRep(object) <- value
```

Arguments

object	a DChIPRepResults object
value	A DESeqDataSet object

Value

a data.frame containing the results of the position wise tests

Examples

```
data(testData)
dcr <- DChIPRepResults(testData)
dcr <- runTesting(dcr)
head(resultsDChIPRep(dcr))
```

runTesting	<i>Run the tests on a DChIPRepResults object.</i>
------------	---

Description

This function runs the testing on a DChIPRepResults object. It adds the FDR calculations and the result table to the DChIPRepResults object.

Usage

```
## S4 method for signature 'DChIPRepResults'  
runTesting(object, lfcThreshold = 0.05,  
           plotFDR = FALSE, ...)
```

Arguments

object	A DChIPRepResults object.
lfcThreshold	A non-negative threshold value, which determines the null hypothesis. The null hypothesis is $H_0 : \log_2(FC) > \text{lfcThreshold}$
plotFDR	If set to TRUE a plot showing the estimated FDRs will be displayed
...	not used currently

Value

a modified [DChIPRepResults](#) object containing the testing results

See Also

[resultsDChIPRep](#)

Examples

```
data(testData)  
dcr <- DChIPRepResults(testData)  
dcr <- runTesting(dcr)
```

show *prints the DESeq2Data slot of the DChIPRepResults object*

Description

prints the data

Usage

```
## S4 method for signature 'DChIPRepResults'  
show(object)
```

Arguments

object A DChIPRepResults object

Value

A compact representation of the DChIPRepResults object

Examples

```
data(testData)  
dcr <- DChIPRepResults(testData)  
dcr  
dcr <- runTesting(dcr)  
dcr
```

summarizeCountsPerPosition

Helper function to summarize the counts per position

Description

This function takes a matrix of counts, with the genomic features (e.g. transcripts or genes) in the rows and the positions upstream and downstream of the TSS in the columns and returns a vector with the summarized counts per position.

Usage

```
summarizeCountsPerPosition(mat, ct = 0, trim = 0.15)
```

Arguments

<code>mat</code>	the input matrix with positions in the columns and the genomic features in the rows.
<code>ct</code>	the count threshold to use.
<code>trim</code>	the trimming percentage for the trimmed mean.

Details

The summary per condition is computed as a trimmed mean per position. First, counts greater than `ct` are removed and then a trimmed mean with a trimming percentage of `trim` is computed on the log scale. This mean is then exponentiated again and multiplied by the total number of features passing the threshold `ct` per position. If a position contains only zero counts, its mean is returned as zero.

Value

a vector containing the summarized counts per condition

Examples

```
data(exampleSampleTable)
directory <- file.path(system.file("extdata", package="DChIPRep"))
df <- lapply(file.path(directory, exampleSampleTable$Input), read.delim)[[1]]
mat <- getMATfromDataFrame(df)
summaryPerPos <- summarizeCountsPerPosition(mat)
```

<code>testData</code>	<i>A test DESeqDataSet</i>
-----------------------	----------------------------

Description

test data to check the functions

Usage

```
data(testData)
```

Format

a DESeqDataSet

Value

a DESeqDataSet

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