

# Package ‘DChIPRep’

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

**Version** 1.0.0

**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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**Suggests** testthat, BiocStyle, knitr, rmarkdown, knitcitations

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**VignetteBuilder** knitr

**biocViews** Sequencing, ChIPSeq

**SystemRequirements** Python 2.7, HTSeq (>= 0.6.1), numpy, argparse, sys

**NeedsCompilation** no

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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")`

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

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

### Usage

```
DChIPRepResults(object)
```

### Arguments

object	A DESeqDataSet
--------	----------------

### Value

A DChIPRepResult object.

### Examples

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

---

DESeq2Data	<i>Accessors for the 'DESeq2Data' slot of a DChIPRepResults object.</i>
------------	-------------------------------------------------------------------------

---

**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	<i>An example ChIP data.</i>
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---

**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      *An example sample table data.frame*

---

**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>
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---

### 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

---

*Import the data after running the Python script*


---

**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 <a href="#">summarizeCountsPerPosition</a>

**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`*Import the data from ChiP and input matrices*

---

## 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

<code>inputData</code>	a matrix containing the counts for the input per position.
<code>chipData</code>	a matrix containing the counts for the ChIP per position.
<code>sampleTable</code>	a data.frame that has to contain the columns <code>sampleID</code> , <code>upstream</code> , <code>downstream</code> and <code>condition</code> . 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	<i>Produce a TSS plot of the two conditions in the data</i>
--------------	-------------------------------------------------------------

---

### 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

<code>object</code>	a <code>DChIPRepResults</code> object after <code>runTesting</code>
<code>meanFunction</code>	a function to compute the positionwise mean per group, defaults to a Huber estimator of the mean.
<code>...</code>	additional parameters for plotting (NOT YET IMPLEMENTED)

### Value

a `ggplot2` object

### Examples

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

---

plotSignificance	<i>Produce a plot that colors the positions identified as significant</i>
------------------	---------------------------------------------------------------------------

---

### 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 }, lfdrThresh = 0.2, ...)
```

**Arguments**

object	a DChIPRepResults object after <a href="#">runTesting</a>
meanFunction	a function to compute the positionwise mean per group, defaults to a Huber estimator of the mean.
lfdrThresh	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 <a href="#">DChIPRepResults</a> 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

[results](#)

### Examples

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

---

show	<i>prints the DESeq2Data slot of the DChIPRepResults object</i>
------	-----------------------------------------------------------------

---

### 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**

mat            the input matrix with positions in the columns and the genomic features in the rows.

ct            the count threshold to use.

trim          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)
```

---

testData

*A test DESeqDataSet*

---

**Description**

test data to check the functions

**Usage**

```
data(testData)
```

**Format**

a DESeqDataSet

**Value**

a DESeqDataSet

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