

Package ‘metevalue’

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Type Package

Title E-Value in the Omics Data Association Studies

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Description

In the omics data association studies, it is common to conduct the p-value corrections to control the false significance. Beyond the P-value corrections, E-value is recently studied to facilitate multiple testing correction based on V. Vovk and R. Wang (2021) <[doi:10.1214/20-AOS2020](https://doi.org/10.1214/20-AOS2020)>. This package provides E-value calculation for DNA methylation data and RNA-seq data. Currently, five data formats are supported: DNA methylation levels using DMR detection tools (BiSeq, DMRfinder, MethylKit, Metilene and other DNA methylation tools) and RNA-seq data. The relevant references are listed below: Katja Hebestreit and Hans-Ulrich Klein (2022) <[doi:10.18129/B9.bioc.BiSeq](https://doi.org/10.18129/B9.bioc.BiSeq)>; Al-tuna Akalin et.al (2012) <[doi:10.18129/B9.bioc.methylKit](https://doi.org/10.18129/B9.bioc.methylKit)>.

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demo_biseq_DMR	<i>BiSeq Output Demo Dataset</i>
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Description

The dummy output for BiSeq illustrating purpose. It is dummy.

Details

- seqnames: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- width
- strand: Strand
- median.p
- median.meth.group1
- median.meth.group2
- median.meth.diff

Notice that there are "NaN" within the feature columns.
Please check the vignette "metevale" for details.

demo_biseq_methyrate *BiSeq Methyrate Demo Dataset*

Description

The methyrate for BiSeq illustrating purpose. It is dummy.

Details

The data includes 12 columns.

- chr: string Chromosome
- pos: int Position
- g1~g2: methylation rate data in groups, repeat 5 times. Notice that there are "NaN" within the feature columns.

Please check the vignette "metevale" for details.

demo_desq_out *DESeq Output Dataset*

Description

The output dummy data for "RNA" meythod illustrating purpose.

Details

The data includes 10 columns.

- treated1fb:
- treated2fb:
- treated3fb:
- untreated1fb:
- untreated2fb:
- untreated3fb:
- untreated4fb:

This data contains 8166 rows and 7 columns.

Please check the vignette "metevale" for details.

Examples

```

# library("pasilla")
# pasCts <- system.file("extdata",
#                       "pasilla_gene_counts.tsv",
#                       package="pasilla", mustWork=TRUE)
# pasAnno <- system.file("extdata",
#                        "pasilla_sample_annotation.csv",
#                        package="pasilla", mustWork=TRUE)
# cts <- as.matrix(read.csv(pasCts, sep="\t", row.names="gene_id"))
# coldata <- read.csv(pasAnno, row.names=1)
# coldata <- coldata[,c("condition", "type")]
# coldata$condition <- factor(coldata$condition)
# coldata$type <- factor(coldata$type)
#
# library("DESeq2")
# colnames(cts)=paste0(colnames(cts), 'fb')
# cts = cts[,rownames(coldata)]
# dds <- DESeqDataSetFromMatrix(countData = cts,
#                               colData = coldata,
#                               design = ~ condition)
# dds <- DESeq(dds)
#
#
# dat <- t(t(cts)/(dds$sizeFactor))
# dat.out <- dat[rowSums(dat >5)>=0.8*ncol(dat),]
#
# demo_desq_out <- log(dat.out)

```

demo_DMRfinder_DMRs *DMRfinder Output Demo Dataset*

Description

The output dummy dataset for DMRfinder illustrating purpose.

Details

The data includes 6 columns.

- chr: string Chromosome
- pos: int Position
- g1~g2: methylation rate data in groups, repeat 2 times. Notice that there are "NaN" within the feature columns.

Please check the vignette "metevalue" for details.

demo_DMRfinder_rate_combine

DMRfinder Methyrate Demo Dataset

Description

The methyrate for BiSeq illustrating purpose. It is dummy.

Details

The data includes 6 columns.

- chr: string Chromosome

- pos: int Position

- g1~g2: methylation rate data in groups, repeat 2 times. Notice that there are "NaN" within the feature columns.

Please check the vignette "metevalue" for details.

demo_methylkit_methyrate

Methyrate Dataset

Description

The methyrate dataset samples "myCpG" data from the methylKit (a bioconductor package) for illustrating purpose.

Details

The data includes 6 columns.

- chr: string Chromosome

- pos: int Position

- g1~g2: methylation rate data in groups (4 columns)

Please check the vignette "metevalue" for details.

References

Akalin, Altuna, et al. "methylKit: a comprehensive R package for the analysis of genome-wide DNA methylation profiles." *Genome biology* 13.10 (2012): 1-9. doi: [10.1186/gb20121310r87](https://doi.org/10.1186/gb20121310r87)

demo_methylkit_met_all

Methyrate output dataset from methylKit

Description

The methyrate dataset samples "myCpG" data from the methylKit (a bioconductor package) for illustrating purpose.

Details

The data includes 7 columns:

- chr: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- strand: Strand
- pvalue: The adjusted p-value based on BH method in MWU-test
- qvalue: cutoff for qvalue of differential methylation statistic
- methyl.diff: The difference between the group means of methylation level

Please check the vignette "metevalue" for details.

References

Akalin, Altuna, et al. "methylKit: a comprehensive R package for the analysis of genome-wide DNA methylation profiles." *Genome biology* 13.10 (2012): 1-9. doi: [10.1186/gb20121310r87](https://doi.org/10.1186/gb20121310r87)

demo_metilene_input

Metilene Methyrate Demo Dataset

Description

The methyrate for metilene illustrating purpose. It is dummy.

Details

The data includes 18 columns.

- chr: string Chromosome
- pos: int Position
- g1~g2: methylation rate data in groups.

Notice that there are "NaN" within the feature columns.

Please check the vignette "metevalue" for details.

demo_metilene_out	<i>Metilene Demo Output Dataset</i>
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Description

The output dummy data for "metilene" meymethod illustrating purpose.

Details

The data includes 10 columns.

- V1: string Chromosome
- V2: The positions of the start sites of the corresponding region
- V3: The positions of the end sites of the corresponding region
- V4- V10: data value.

Please check the vignette "metevalue" for details.

evalue_buildin_sql	<i>Build-in data process function</i>
--------------------	---------------------------------------

Description

Build-in data process function

Usage

```
evalue_buildin_sql(a, b, method = "metilene")
```

Arguments

a	data frame of the methylation rate
b	data frame of output data corresponding to the "method" option
method	"metilene" or "biseq", "DMRfinder" or "methylKit"

Value

a data frame combines data frame a and b corresponding to the "method" option

Examples

```
data("demo_metilene_out")
data("demo_metilene_input")
result = evalue_buildin_var_fmt_nm(demo_metilene_input,
                                  demo_metilene_out, method="metilene")
result_sql = evalue_buildin_sql(result$a, result$b, method="metilene")
```

evaluate_buildin_var_fmt_nm

Build-in check file format function Perform the format check and data clean for the "metilene" or "biseq", "DMRfinder" or "methylKit" method correspondingly.

Description

Build-in check file format function Perform the format check and data clean for the "metilene" or "biseq", "DMRfinder" or "methylKit" method correspondingly.

Usage

```
evaluate_buildin_var_fmt_nm(a, b, method = "metilene")
```

Arguments

a	data frame of the methylation rate
b	data frame of output data corresponding to the "method" option
method	"metilene" or "biseq", "DMRfinder" or "methylKit"

Value

list(a, b) which contains the cleaned data correspondingly

Examples

```
data("demo_metilene_out")
data("demo_metilene_input")
evaluate_buildin_var_fmt_nm(demo_metilene_input,
                           demo_metilene_out, method="metilene")
```

metevalue.biseq

Calculate E-value of the BiSeq data format

Description

Please check vignette "metevalue" for details.

Usage

```
metevaluate.biseq(
  methyrate,
  BiSeq.output,
  adjust.methods = "BH",
  sep = "\t",
  bheader = FALSE
)
```

Arguments

`methyrate` is the methyrate file. For example:

```
chr  pos  g1  ...  g1  g2  ...  g2
chr1  1  0.1  ...  0.1  0.2  ...  0.2
```

The columns are (in order):

- chr: Chromosome
- pos: int Position
- g1~g2: methylation rate data in groups

`BiSeq.output` is the output file of BiSeq. The columns are (in order):

- seqnames: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- width: The number of CpG sites within the corresponding region
- strand: Strand
- median.p: The median p-value among CpG sites within the corresponding region
- median.meth.group1: The median methylation rate in the first group among CpG sites within the corresponding region
- median.meth.group2: The median methylation rate in the second group among CpG sites within the corresponding region
- median.meth.diff: The median methylation difference between groups among CpG sites within the corresponding region

`adjust.methods` is the adjust methods of e-value. It can be 'bonferroni', 'hochberg', 'holm', 'hommel', 'BH', 'BY'

`sep` seperator, default is the TAB key.

`bheader` a logical value indicating whether the BiSeq.output file contains the names of the variables as its first line. By default, `bheader = FALSE`.

Value

a dataframe, the columns are (in order):

- chr: Chromosome

- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- q-value: The adjusted p-value based on BH method in MWU-test
- methyl.diff: The difference between the group means of methylation level
- CpGs: The number of CpG sites within the corresponding region
- p : p-value based on MWU-test
- p2: p-value based on 2D KS-test
- m1: The absolute mean methylation level for the corresponding segment of group 1
- m2: The absolute mean methylation level for the corresponding segment of group 2
- e_value: The e-value of the corresponding region

Examples

```

#\donttest{
#data("demo_biseq_methyrate")
#data("demo_biseq_DMR")
#example_tempfiles = tempfile(c("demo_biseq_methyrate", "demo_biseq_DMR"))
#tempdir()
#### write to temp file ####
#write.table(demo_biseq_methyrate, file=example_tempfiles[1],row.names=FALSE,
#           col.names=TRUE, quote=FALSE, sep='\t')
#write.table(demo_biseq_DMR, file=example_tempfiles[2],
#           sep ="\t", row.names =FALSE, col.names =TRUE, quote =FALSE)
#### compute e-value and its adjustment ####
#result = metevalue.biseq(example_tempfiles[1],
#                          example_tempfiles[2], bheader = TRUE)
#}

```

metevalue.biseq.chk *Check the BiSeq data format*

Description

Check the BiSeq data format

Usage

```

metevalue.biseq.chk(
  input_filename_a,
  input_filename_b,
  sep = "\t",
  bheader = FALSE
)

```

metevalue.DMRfinder *Calculate E-value of the DMRfinder data format*

Description

Calculate E-value of the DMRfinder data format

Usage

```
metevalue.DMRfinder(  
  methyrate,  
  DMRfinder.output,  
  adjust.methods = "BH",  
  sep = "\t",  
  bheader = FALSE  
)
```

Arguments

methyrate is the methyrate file. For example:

```
chr  pos  g1  ...  g1  g2  ...  g2  
chr1  1  0.1  ...  0.1  0.2  ...  0.2
```

The columns are (in order):

- chr: Chromosome
- pos: int Position
- g1~g2: methylation rate data in groups

DMRfinder.output

is the output file of DMRfinder.

- chr: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- CpG: The number of CpG sites within the corresponding region
- Control.mu: The average methylation rate in control group
- Expt1.mu: The average methylation rate in experiment group
- Control.Expt1.diff: The methylation difference between control and experiment groups
- Control.Expt1.pval: P-value based on Wald-test.

adjust.methods is the adjust methods of e-value. It can be 'bonferroni', 'hochberg', 'holm', 'hommel', 'BH', 'BY'

sep seperator, default is the TAB key.

bheader a logical value indicating whether the DMRfinder.output file contains the names of the variables as its first line. By default, bheader = FALSE.

Value

a dataframe, the columns are (in order):

- chr: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- q-value: The adjusted p-value based on BH method in MWU-test
- methyl.diff: The difference between the group means of methylation level
- CpGs: The number of CpG sites within the corresponding region
- p : p-value based on MWU-test
- p2: p-value based on 2D KS-test
- m1: The absolute mean methylation level for the corresponding segment of group 1
- m2: The absolute mean methylation level for the corresponding segment of group 2
- e_value: The e-value of the corresponding region

Examples

```
data(demo_DMRfinder_rate_combine)
data(demo_DMRfinder_DMRs)
#example_tempfiles = tempfile(c("rate_combine", "DMRfinder_out"))
#tempdir()
#write.table(demo_DMRfinder_rate_combine, file=example_tempfiles[1],
#           row.names=FALSE, col.names=TRUE, quote=FALSE, sep='\t')
#write.table(demo_DMRfinder_DMRs, file=example_tempfiles[2],
#           sep = "\t", row.names =FALSE, col.names =TRUE, quote =FALSE)
#result = metevalue._DMRfinder(example_tempfiles[1], example_tempfiles[2],
#           bheader = TRUE)
#head(result)
```

metevalue.DMRfinder.chk

Check the DMRfinder data format

Description

Check the DMRfinder data format

Usage

```
metevalue.DMRfinder.chk(
  input_filename_a,
  input_filename_b,
  sep = "\t",
  bheader = FALSE
)
```

Arguments

`input_filename_a`
the combined data of methylation rate file. This file is a sep (e.g. TAB) separated file with two key columns and several value columns. For example:

```
chr  pos  g1  ...  g1  g2  ...  g2
chr1  1  0.1  ...  0.1  0.2  ...  0.2
```

The columns are (in order):

- chr and pos are keys;
- g1~g2: methylation rate data in groups.

`input_filename_b`
the output file of DMRfinder. The columns are (in order):

- chr: Chromosome
- start: The position of the start sites of the corresponding region
- end: The position of the end sites of the corresponding region
- CpG: The number of CpG sites within the corresponding region
- 'Control:mu': The absolute mean methylation level for the corresponding segment of the control group
- 'Exptl:mu': The absolute mean methylation level for the corresponding segment of the experimental group
- 'Control->Exptl:diff': The difference between the group means of methylation level
- p: p-value

`sep`
separator, default is the TAB key.

`bheader`
a logical value indicating whether the `input_filename_b` file contains the names of the variables as its first line. By default, `bheader = FALSE`.

Value

`list(file_a, file_b, file_a_b)` returns a list with three pre-handled data.frames corresponding to the `input_filename_a`, `input_filename_b` file and a A JOIN B file.

Examples

```
data(demo_DMRfinder_rate_combine)
data(demo_DMRfinder_DMRs)
#example_tempfiles = tempfile(c("rate_combine", "DMRfinder_out"))
#tempdir()
#write.table(demo_DMRfinder_rate_combine, file=example_tempfiles[1],
#  row.names=FALSE, col.names=TRUE, quote=FALSE, sep='\t')
#write.table(demo_DMRfinder_DMRs, file=example_tempfiles[2],
#  sep ="\t", row.names =FALSE, col.names =TRUE, quote =FALSE)
#result = metevalue.DMRfinder.chk(example_tempfiles[1], example_tempfiles[2],
#  bheader = TRUE)
```

metevalue.methylKit *Calculate E-value of the methylKit data format*

Description

Calculate E-value of the methylKit data format

Usage

```
metevalue.methylKit(
  methyrate,
  methylKit.output,
  adjust.methods = "BH",
  sep = "\t",
  bheader = FALSE
)
```

Arguments

`methyrate` is the data of methylation rates of each sites and group. For example:

```
chr  pos  g1  ...  g1  g2  ...  g2
chr1  1  0.1  ...  0.1  0.2  ...  0.2
```

The columns are (in order):

- chr: Chromosome
- pos: int Position
- g1~g2: methylation rate data in groups

`methylKit.output`

is the output data with e-value of each region

- chr: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- strand: Strand
- pvalue: The adjusted p-value based on BH method in MWU-test
- qvalue: cutoff for qvalue of differential methylation statistic
- methyl.diff: The difference between the group means of methylation level

`adjust.methods` is the adjust methods of e-value. It can be 'bonferroni', 'hochberg', 'holm', 'hommel', 'BH', 'BY'

`sep` separator, default is the TAB key.

`bheader` a logical value indicating whether the `input_filename_b` file contains the names of the variables as its first line. By default, `bheader = FALSE`.

Value

a dataframe, the columns are (in order):

- chr: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- q-value: The adjusted p-value based on BH method in MWU-test
- methyl.diff: The difference between the group means of methylation level
- CpGs: The number of CpG sites within the corresponding region
- p : p-value based on MWU-test
- p2: p-value based on 2D KS-test
- m1: The absolute mean methylation level for the corresponding segment of group 1
- m2: The absolute mean methylation level for the corresponding segment of group 2
- e_value: The e-value of the corresponding region

Examples

```
data(demo_methylkit_methyrate)
data(demo_methylkit_met_all)
## example_tempfiles = tempfile(c("rate_combine", "methylKit_DMR_raw"))
## tempdir()
## write.table(demo_methylkit_methyrate, file=example_tempfiles[1],
##             row.names=FALSE, col.names=TRUE, quote=FALSE, sep='\t')
## write.table(demo_methylkit_met_all, file=example_tempfiles[2],
##             sep="\t", row.names =FALSE, col.names =TRUE, quote =FALSE)
## result = metevalue.methylKit(example_tempfiles[1], example_tempfiles[2],
##                               bheader = TRUE)
## str(result)
```

metevalue.methylKit.chk

Check the methylKit data format

Description

Check the methylKit data format

Usage

```
metevalue.methylKit.chk(
  input_filename_a,
  input_filename_b,
  sep = "\t",
  bheader = FALSE
)
```

Arguments

- `input_filename_a`
the combined data of methylation rate file. This file is a sep (e.g. TAB) separated file with two key columns and several value columns: For example:
- ```
chr pos g1 ... g1 g2 ... g2
chr1 1 0.1 ... 0.1 0.2 ... 0.2
```
- chr and pos are keys;
  - g1~g2: methylation rate data in groups.
- `input_filename_b`  
the output file of methylKit. a methylDiff or methylDiffDB object containing the differential methylated locations satisfying the criteria. The columns are (in order):
- chr: Chromosome
  - start: The position of the start sites of the corresponding region
  - end: The position of the end sites of the corresponding region
  - strand: Strand
  - p: p-value
  - qvalue: The adjusted p-value based on BH method
  - meth.diff : The difference between the group means of methylation level
- `sep`  
separator, default is the TAB key.
- `bheader`  
a logical value indicating whether the `input_filename_b` file contains the names of the variables as its first line. By default, `bheader = FALSE`.

**Value**

`list(file_a, file_b, file_a_b)` returns a list with three pre-handled data.frames corresponding to the `input_filename_a`, `input_filename_b` file and a A JOIN B file.

**Examples**

```
data(demo_methylkit_methyrate)
data(demo_methylkit_met_all)
example_tempfiles = tempfile(c("rate_combine", "methylKit_DMR_raw"))
tempdir()
write.table(demo_methylkit_methyrate, file=example_tempfiles[1],
row.names=FALSE, col.names=TRUE, quote=FALSE, sep='\t')
write.table(demo_methylkit_met_all, file=example_tempfiles[2],
sep="\t", row.names =FALSE, col.names =TRUE, quote =FALSE)
result = metevalue.methylKit.chk(example_tempfiles[1], example_tempfiles[2],
bheader = TRUE)
```

---

metevalue.metilene      *Calculate E-value of the Metilene data format*

---

## Description

Calculate E-value of the Metilene data format

## Usage

```
metevalue.metilene(
 methyrate,
 metilene.output,
 adjust.methods = "BH",
 sep = "\t",
 bheader = FALSE
)
```

## Arguments

**methyrate**      metilene input file path. This file is a sep (e.g. TAB) separated file with two key columns and several value columns. For example:

```
chr pos g1 ... g1 g2 ... g2
chr1 1 0.1 ... 0.1 0.2 ... 0.2
```

The columns are (in order):

- chr and pos are keys;
- g1~g2: methylation rate data in groups.

**metilene.output**

metilene input file path. This file should be stored as a sep (e.g. TAB) separated file with two key columns and several value columns: The columns are (in order):

- chr: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- q-value: The adjusted p-value based on BH method in MWU-test
- methyl.diff: The difference between the group means of methylation level
- CpGs: The number of CpG sites within the corresponding region
- p : p-value based on MWU-test
- p2: p-value based on 2D KS-test
- m1: The absolute mean methylation level for the corresponding segment of group 1
- m2: The absolute mean methylation level for the corresponding segment of group 2

adjust.methods is the adjust methods of e-value. It can be 'bonferroni', 'hochberg', 'holm', 'hommel', 'BH', 'BY'

sep separator, default is the TAB key.

bheader a logical value indicating whether the metilene.output file contains the names of the variables as its first line. By default, bheader = FALSE.

### Value

a dataframe, the columns are (in order):

- chr: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- q-value: The adjusted p-value based on BH method in MWU-test
- methyl.diff: The difference between the group means of methylation level
- CpGs: The number of CpG sites within the corresponding region
- p : p-value based on MWU-test
- p2: p-value based on 2D KS-test
- m1: The absolute mean methylation level for the corresponding segment of group 1
- m2: The absolute mean methylation level for the corresponding segment of group 2
- e\_value: The e-value of the corresponding region

### Examples

```
metilene example ####'
data(demo_metilene_input)
data(demo_metilene_out)
#example_tempfiles = tempfile(c("metilene_input", "metilene_out"))
#tempdir()
#write.table(demo_metilene_input, file=example_tempfiles[1],
row.names=FALSE, col.names=TRUE, quote=FALSE, sep='\t')
#write.table(demo_metilene_out, file=example_tempfiles[2],
sep = "\t", row.names =FALSE, col.names =TRUE, quote =FALSE)
#result = metevaluate.metilene(example_tempfiles[1], example_tempfiles[2],
bheader = TRUE)
#head(result)
```

---

metevaluate.metilene.chk

*Check the Metilene data format*

---

### Description

Check the Metilene data format

**Usage**

```
metevalue.metilene.chk(
 input_filename_a,
 input_filename_b,
 sep = "\t",
 bheader = FALSE
)
```

**Arguments**

`input_filename_a`  
metilene input file path. This file is a sep (e.g. TAB) separated file with two key columns and several value columns. For example:

```
chr pos g1 ... g1 g2 ... g2
chr1 1 0.1 ... 0.1 0.2 ... 0.2
```

The columns are (in order):

- chr and pos are keys;
- g1~g2: methylation rate data in groups.

`input_filename_b`  
metilene input file path. This file should be stored as a sep (e.g. TAB) separated file with two key columns and several value columns: The columns are (in order):

- chr: Chromosome
- start: The position of the start sites of the corresponding region
- end: The position of the end sites of the corresponding region
- q-value: The adjusted p-value based on BH method in MWU-test
- methyl.diff: The difference between the group means of methylation level
- CpGs: The number of CpG sites within the corresponding region
- p : p-value based on MWU-test
- p2: p-value based on 2D KS-test
- m1: The absolute mean methylation level for the corresponding segment of group 1
- m2: The absolute mean methylation level for the corresponding segment of group 2

`sep` separator, default is the TAB key.

`bheader` a logical value indicating whether the `input_filename_b` file contains the names of the variables as its first line. By default, `bheader = FALSE`.

**Value**

`list(file_a, file_b, file_a_b)` returns a list with three pre-handled data.frames corresponding to the `input_filename_a`, `input_filename_b` file and a A JOIN B file.

**Examples**

```
#data(demo_metilene_input)
#data(demo_metilene_out)
#example_tempfiles = tempfile(c("metilene_input", "metilene_out"))
#tempdir()
#write.table(demo_metilene_input, file=example_tempfiles[1],
row.names=FALSE, col.names=TRUE, quote=FALSE, sep='\t')
#write.table(demo_metilene_out, file=example_tempfiles[2],
sep ="\t", row.names =FALSE, col.names =TRUE, quote =FALSE)
#result = metevaluate.metilene.chk(example_tempfiles[1], example_tempfiles[2],
bheader = TRUE)
```

---

metevaluate.RNA\_general *A general method to calculate the e-value for RNA-seq data.*

---

**Description**

A general method to calculate the e-value for RNA-seq data.

**Usage**

```
metevaluate.RNA_general(rna, group1_name, group2_name)
```

**Arguments**

rna                    data.frame: A data.frame object of RNAseq data. For example:

|      | TAG | treated1fb | treated2fb | untreated1fb | untreated2fb |
|------|-----|------------|------------|--------------|--------------|
| TAG1 |     | 4.449648   | 4.750104   | 4.392285     | 4.497514     |
| TAG2 |     | 8.241116   | 8.302852   | 8.318125     | 8.488796     |
|      | ... | ...        | ...        | ...          | ...          |

Row names (TAG1 and TAG2 in the above example) is also suggested.

group1\_name          character: The name (pattern) of the first group. For example, "treated" in the above example. For example 'treated\_abc' and 'treated' will be considered as the same group if 'group1\_name = "treated"'. Use this with care in practice.

group2\_name          character: The name (pattern) of the second group. For example, "untreated" in the above example. For example 'untreated\_abc' and 'untreated' will be considered as the same group if 'group2\_name = "untreated"'. Use this with care in practice.

**Value**

evaluate

**Examples**

```
data("demo_desq_out")

evaluate = metevaluate.RNA_general(demo_desq_out, 'treated', 'untreated')
```

---

varevalue.metilene      *Calculate E-value of the Metilene data*

---

**Description**

The data file could be pre-handled by the evaluate.metilene.chk function.

**Usage**

```
varevalue.metilene(
 a,
 b,
 a_b,
 group1_name = "g1",
 group2_name = "g2",
 adjust.methods = "BH"
)
```

**Arguments**

a                    A data.frame object:

```
chr pos g1 ... g1 g2 ... g2
chr1 1 0.1 ... 0.1 0.2 ... 0.2
```

i.e two key columns (chrom, pos) with several value columns in groups.

b                    A data.frame object stores the data, the columns are (in order):

- chr: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- q-value: The adjusted p-value based on BH method in MWU-test
- methyl.diff: The difference between the group means of methylation level
- CpGs: The number of CpG sites within the corresponding region
- p : p-value based on MWU-test
- p2: p-value based on 2D KS-test
- m1: The absolute mean methylation level for the corresponding segment of group 1
- m2: The absolute mean methylation level for the corresponding segment of group 2

|                |                                                                                                                                     |
|----------------|-------------------------------------------------------------------------------------------------------------------------------------|
| a_b            | A data.frame object of a join b with particular data clean processes. Check the function [evalue.methylKit.chk()] for more details. |
| group1_name    | character: The name of the first group. For example, "g1" in the above example.                                                     |
| group2_name    | character: The name of the second group. For example, "g2" in the above example.                                                    |
| adjust.methods | is the adjust methods of e-value. It can be 'bonferroni', 'hochberg', 'holm', 'hommel', 'BH', 'BY'. The default value is 'BH'.      |

### Value

a dataframe, the columns are (in order):

- chr: Chromosome
- start: The positions of the start sites of the corresponding region
- end: The positions of the end sites of the corresponding region
- q-value: The adjusted p-value based on BH method in MWU-test
- methyl.diff: The difference between the group means of methylation level
- CpGs: The number of CpG sites within the corresponding region
- p : p-value based on MWU-test
- p2: p-value based on 2D KS-test
- m1: The absolute mean methylation level for the corresponding segment of group 1
- m2: The absolute mean methylation level for the corresponding segment of group 2
- e\_value: The e-value of the corresponding region

### Examples

```
#data(demo_metilene_input)
#data(demo_metilene_out)
#result = evalue_buildin_var_fmt_nm(demo_metilene_input, demo_metilene_out, method="metilene")
#result = list(a = result$a,
b = result$b,
a_b = evalue_buildin_sql(result$a, result$b, method="metilene"))
#result = varevalue.metilene(result$a, result$b, result$a_b)
```

---

varevalue.single\_general

*A general method to calculate the e-value for other DNA methylation tools not described above. The input data is the DNA methylation rates using the similar format with Metilene.*

---

### Description

The input data file is just the DNA methylation rates using the similar format above, with no need for another data file output by different tools. The Chromosome name, start and end sites should be specified in the function.

**Usage**

```
varevalue.single_general(
 methyrate,
 group1_name = "g1",
 group2_name = "g2",
 chr,
 start,
 end
)
```

**Arguments**

**methyrate**      **data.frame:** A data.frame object of methylation rates, the columns should be (name of groups can be self-defined)

```
chr pos g1 ... g1 g2 ... g2
chr1 1 0.1 ... 0.1 0.2 ... 0.2
```

**group1\_name**      **character:** The name (pattern) of the first group. For example, "g1" in the above example. For example 'g1\_abc' and 'g1' will be considered as the same group if 'group1\_name = "g1"'. Use this with care in practice.

**group2\_name**      **character:** The name (pattern) of the second group. For example, "g2" in the above example. For example 'g2\_abc' and 'g2' will be considered as the same group if 'group2\_name = "g2"'. Use this with care in practice.

**chr**              **character:** The Chromosome name. Typically, it is a string like "chr21" and so on.

**start**            **integer:** The position of the start site of the corresponding region

**end**              **integer:** The position of the end site of the corresponding region

**Value**

evaluate

**Examples**

```
#data("demo_metilene_input")
#varevalue.single_general(demo_metilene_input, chr = "chr21", start = 9437432, end = 9437540)
[1] 2.626126e+43

Compare to `metevaluate.metilene`
data(demo_metilene_out)
#example_tempfiles = tempfile(c("metilene_input", "metilene_out"))
#tempdir()
#write.table(demo_metilene_input, file=example_tempfiles[1],
row.names=FALSE, col.names=TRUE, quote=FALSE, sep='\t')
#write.table(demo_metilene_out, file=example_tempfiles[2],
sep = "\t", row.names =FALSE, col.names =TRUE, quote =FALSE)
```

```
#result = metevalue.metilene(example_tempfiles[1], example_tempfiles[2],
bheader = TRUE)
result[with(result, chr == 'chr21' & start == '9437432' & end == '9437540'), ncol(result)]
[1] 2.626126e+43
```

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