

Package ‘GCAI.bias’

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Type Package

Title Guided Correction Approach for Inherited bias (GCAI.bias)

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Description Many inherited biases and effects exists in RNA-seq due to both biological and technical effects. We observed the biological variance of testing target transcripts can influence the yield of sequencing reads, which might indicate a resource competition existing in RNA-seq. We developed this package to capture the bias depending on local sequence and perform the correction of this type of bias by borrowing information from spike-in measurement.

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GCAI.bias-package *Guided Correction Approach for Inherited bias (GCAI.bias)*

Description

Many inherited biases and effects exists in RNA-seq due to both biological and technical effects. We observed the biological variance of testing target transcripts can influence the yield of sequencing reads, which might indicate a resource competition existing in RNA-seq. We developed this package to capture the bias depending on local sequence and perform the correction of this type of bias by borrowing information from spike-in measurement.

Details

Package:	GCAI-bias
Type:	Package
Version:	1.0
Date:	2014-07-14
License:	GPL (>=2)
LazyLoad:	yes

This package is used for correcting bias introduced by the biological variance of sample transcripts sources. Batch effect in measurement of the same biological sample can be corrected by this package directly. However, spike-in are required to correct bias between different biological samples. For strand specific RNA-seq, antisense and sense reads should be corrected separately. Sequencing reads on each base pair are required to be formatted as train.dat.seq, train.dat.counts, test.dat.seq and test.dat.counts objects. Coefficients of local sequence will be estimated in lm.estimate and they will be used to correct bias by correct.guided function. Visualization of coefficients and correcting performance can be achieved by coeplot, corplot and posplot.

Author(s)

Guoshuai Cai

Maintainer: Guoshuai Cai <GCAI.bioinfo@gmail.com>

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

Examples

```
#initialize index matrix
word<-81
word.vec<-c("A","T","C","G")
pos.vec<-c((-word+1)/2):((word-1)/2)

obj.index<-index.mat.generation(word.vec,pos.vec)

#train

data(train.dat.seq)
data(train.dat.counts)

train.index<-index.preprocess(train.dat.seq,word)
obj.train<-counts.preprocess(train.dat.counts)
obj.train[["index"]]<-train.index

coe.lm<-lm.estimate(obj.train,fit.cut.train=5)

coeplot(coe.lm,obj.index)

#test,correct

data(test.dat.seq)
data(test.dat.counts)

test.index<-index.preprocess(test.dat.seq,word)
obj.test<-counts.preprocess(test.dat.counts)
obj.test[["index"]]<-test.index

test.corrected<-correct.guided(coe.lm,obj.test)

corplot(test.corrected)
posplot(test.corrected,obj.test$pos)
```

coeplot

Estimated Coefficients Visualization

Description

To plot estimated coefficients against their corresponding positions

Usage

```
coeplot(coe.lm, obj.index, ylim = c(-1, 1))
```

Arguments

<code>coe.lm</code>	a 1-column matrix of coefficients estimated from lm.estimate
<code>obj.index</code>	an object of index generated from index.mat.generation
<code>ylim</code>	the extremes of the range of y axis

Details

5 end of the mapped reads will be labeled 0 on x axis. Nucleotide A will be colored in red, T will be in yellowgreen, C will be in green and G will be in blue as the baseline.

Value

a plot produced on the current graphic device

Note

index object should match the length of coe.lm

Author(s)

Guoshuai Cai

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

See Also

[lm.estimate](#), [index.mat.generation](#)

Examples

```
word<-81

data(obj.index)
data(train.dat.seq)
data(train.dat.counts)

train.index<-index.preprocessing(train.dat.seq,word)
obj.train<-counts.preprocessing(train.dat.counts)
obj.train[["index"]]<-train.index

coe.lm<-lm.estimate(obj.train,fit.cut.train=5)

coeplot(coe.lm,obj.index)
```

corplot*Pairwise Correlation Calculation and Visualization*

Description

Pairwise comparison matrix of the original and corrected measurements

Usage

```
corplot(mat)
```

Arguments

mat a data matrix from [correct.guided](#) function

Details

Pairwise plots of sequencing reads mapped to each base pair on the log10 scale will be shown in the bottom panel; Pairwise Pearson correlation coefficients calculated on data points larger than 1 will be shown in the top panel.

Value

a plot produced on the current graphic device

Author(s)

Guoshuai Cai

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

See Also

[correct.guided](#)

Examples

```
word<-81  
  
data(obj.index)  
data(train.dat.seq)  
data(train.dat.counts)  
data(test.dat.seq)  
data(test.dat.counts)  
  
#train
```

```

train.index<-index.preprocess(train.dat.seq,word)
obj.train<-counts.preprocess(train.dat.counts)
obj.train[["index"]]<-train.index

coe.lm<-lm.estimate(obj.train,fit.cut.train=5)

#test

test.index<-index.preprocess(test.dat.seq,word)
obj.test<-counts.preprocess(test.dat.counts)
obj.test[["index"]]<-test.index

test.corrected<-correct.guided(coe.lm,obj.test)

corplot(test.corrected)

```

correct.guided

*Guided Correction***Description**

Guided by the coefficients estimated from [lm.estimate](#), the bias correction could be performed.

Usage

```
correct.guided(coe.train, obj.test)
```

Arguments

coe.train	a 1-column matrix of coefficients estimated from training data by lm.estimate
obj.test	an object of testing dataset generated from counts.preprocess and index.preprocess

Details

Besides of corrected reads, βx will be also calculated.

Value

a matrix will be returned containing reads counts before and after correction and βx as well.

Author(s)

Guoshuai Cai

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

See Also

[lm.estimate](#), [counts.preprocess](#), [index.preprocess](#)

Examples

```
word<-81

data(obj.index)
data(train.dat.seq)
data(train.dat.counts)
data(test.dat.seq)
data(test.dat.counts)

#train

train.index<-index.preprocess(train.dat.seq,word)
obj.train<-counts.preprocess(train.dat.counts)
obj.train[["index"]]<-train.index

coe.lm<-lm.estimate(obj.train,fit.cut.train=5)

#test

test.index<-index.preprocess(test.dat.seq,word)
obj.test<-counts.preprocess(test.dat.counts)
obj.test[["index"]]<-test.index

test.corrected<-correct.guided(coe.lm,obj.test)
```

counts.preprocess *Data Object Generation*

Description

To extract position and count information from [train.dat.counts](#) or [test.dat.counts](#)

Usage

`counts.preprocess(mat.counts)`

Arguments

`mat.counts` a dataframe of train.dat.counts or test.dat.counts

Value

`counts.preprocess` return a list containing components:

<code>counts</code>	a matrix of counts
<code>pos</code>	a matrix of positions

Author(s)

Guoshuai Cai

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

See Also

[train.dat.counts](#), [test.dat.counts](#)

Examples

```
data(train.dat.counts)
obj.train<-counts.preprocess(train.dat.counts)

data(test.dat.counts)
obj.test<-counts.preprocess(test.dat.counts)
```

index.mat.generation *Index Matrix Generation*

Description

To generate S3 class of index

Usage

```
index.mat.generation(word.vec, pos.vec)
```

Arguments

word.vec	a vector of nucleotides, the default is c("A","T","C","G").
pos.vec	a vector of positions of local sequence, the default is -40 ~ 40.

Value

`index.mat.generation` return a list containing components:

mat	a matrix with nucleotides as row names and positions as column names will be returned, a 4×81 matrix will be returned with default arguments.
-----	--

Author(s)

Guoshuai Cai

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

Examples

```
word<-81  
word.vec<-c("A","T","C","G")  
pos.vec<-c((-word-1)/2):((word-1)/2)  
  
obj.index<-index.mat.generation(word.vec,pos.vec)
```

index.preprocess *Binary Index Matrix Generation*

Description

To initiate the binary variable matrix for linear model

Usage

```
index.preprocess(mat.seq, word)
```

Arguments

mat.seq	a data frame of train.dat.seq or test.dat.seq
word	the number of local sequence positions

Value

a binary index matrix will be returned.

Author(s)

Guoshuai Cai

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

See Also

[train.dat.seq](#), [test.dat.seq](#)

Examples

```
word<-81

data(train.dat.seq)
train.index<-index preprocess(train.dat.seq,word)

data(test.dat.seq)
test.index<-index preprocess(test.dat.seq,word)
```

lm.estimate

Linear Model Fitting

Description

To estimating the coefficients by fitting the linear model

Usage

```
lm.estimate(obj.train, fit.cut.train = 5)
```

Arguments

- | | |
|---------------|--|
| obj.train | an object of training data generated from index.preprocess and counts.preprocess |
| fit.cut.train | the minimum counts of the data points used for model fitting, the default for spike-in training dataset is 5. |

Details

It models the influence on the local sequence from the dissimilarity of spike-in transcripts measurement. The region of positions around hexamer primers can be defined by the variable "word".

Value

a 1-column matrix of coefficients will be returned.

Author(s)

Guoshuai Cai

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

See Also

[index.preprocess](#),[counts.preprocess](#)

Examples

```
word<-81  
  
data(train.dat.seq)  
data(train.dat.counts)  
  
train.index<-index.preprocess(train.dat.seq,word)  
obj.train<-counts.preprocess(train.dat.counts)  
obj.train[["index"]]<-train.index  
  
coe.lm<-lm.estimate(obj.train,fit.cut.train=5)
```

*obj.index**Index S3 Class*

Description

Index S3 class contains two objects: a 2-dimension matrix with nucleotides as row names and positions as column names and a flattened one with 1-dimension combinations of nucleotides and positions.

Usage

```
data(obj.index)
```

Format

The format is:

List of 2

```
$ mat : num [1:4, 1:81] 0 0 0 0 0 0 0 0 0 ...  
..- attr(*, "dimnames")=List of 2  
.. ..$ : chr [1:4] "A" "T" "C" "G"  
.. ..$ : chr [1:81] "-40" "-39" "-38" "-37" ...  
$ flat: chr [1:324, 1:3] "1" "2" "3" "4" ...
```

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

Examples

```
data(obj.index)
```

panel.cor

*Pearson Correlation Calculation***Description**

To calculate the Pearson correlation coefficient, a part of [corplot](#)

Usage

```
panel.cor(x, y, digits = 2, prefix = "", cex.cor, cor.cut = 1)
```

Arguments

```
x  
y  
digits  
prefix  
cex.cor  
cor.cut
```

Author(s)

Guoshuai Cai

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

See Also

[corplot](#)

panel.smooth.r

*Scatter Plot Generation***Description**

To generate a base pair x-y plot, a part of [corplot](#)

Usage

```
panel.smooth.r(x, y, col = par("col"), bg = NA, pch = par("pch"),
cex = par("cex"), col.smooth = "red", span = 2/3, iter = 3, ...)
```

Arguments

```
x  
y  
col  
bg  
pch  
cex  
col.smooth  
span  
iter  
...
```

Author(s)

Guoshuai Cai

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

See Also

[corplot](#)

posplot

Correction Performance Visualization

Description

Pairwise comparison matrix of the original and corrected measurements

Usage

```
posplot(test.corrected, pos.test, n.lim = 1000, fit.cut.lr = 50)
```

Arguments

test.corrected	a data matrix from correct.guided function
pos.test	a data matrix of test transcripts' position information
n.lim	the maximum number of log ratios points will be plotted
fit.cut.lr	the minimum number of sequencing reads points will be used for log ratio plotting

Details

Distributions of sequencing reads of samples measured before and after correction will be plotted with the positions on the x-axis and the number of sequencing reads as bars. Also patterns of the fluctuation factor, log ratio before the correction (black line) and log ratio after the correction (blue line) will be plotted.

Value

a plot produced on the current graphic device

Author(s)

Guoshuai Cai

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

See Also

[correct.guided](#), [counts.preprocess](#)

Examples

```
word<-81

data(obj.index)
data(train.dat.seq)
data(train.dat.counts)
data(test.dat.seq)
data(test.dat.counts)

#train

train.index<-index.preprocess(train.dat.seq,word)
obj.train<-counts.preprocess(train.dat.counts)
obj.train[["index"]]<-train.index

coe.lm<-lm.estimate(obj.train,fit.cut.train=5)

#test

test.index<-index.preprocess(test.dat.seq,word)
obj.test<-counts.preprocess(test.dat.counts)
obj.test[["index"]]<-test.index

test.corrected<-correct.guided(coe.lm,obj.test)

posplot(test.corrected,obj.test$pos)
```

test.dat.counts *Testing Counts Dataset*

Description

An example, a data frame of reads counts of testing samples

Usage

```
data(test.dat.counts)
```

Format

A data frame with 1527 observations on the following 6 variables.

```
gene a factor with levels ERCC-00002 ERCC-00003 ERCC-00007 ERCC-00009 ERCC-00012 ERCC-00013  
ERCC-00014 ERCC-00016 ERCC-00017 ERCC-00018 ERCC-00019 ERCC-00022 ERCC-00023  
ERCC-00024 ERCC-00025 ERCC-00028 ERCC-00031 ERCC-00033 ERCC-00034 ERCC-00035  
ERCC-00039 ERCC-00040 ERCC-00041 ERCC-00042 ERCC-00044 ERCC-00048 ERCC-00054  
ERCC-00057 ERCC-00058 ERCC-00059 ERCC-00061 ERCC-00079 ERCC-00084 ERCC-00170  
pos a numeric vector  
code_strand a factor with levels +  
seq_strand a factor with levels S  
GSM516589_run32_s_2_ERCC_map_Pos a numeric vector  
GSM517059_run29_s_1_ERCC_map_Pos a numeric vector
```

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

Examples

```
data(test.dat.counts)
```

test.dat.seq*Testing Sequence Index Dataset***Description**

An example, a data frame of sequence index of testing samples

Usage

```
data(test.dat.seq)
```

Format

A data frame with 1527 observations on the following 85 variables.

gene	a factor with levels ERCC-00002 ERCC-00003 ERCC-00007 ERCC-00009 ERCC-00012 ERCC-00013 ERCC-00014 ERCC-00016 ERCC-00017 ERCC-00018 ERCC-00019 ERCC-00022 ERCC-00023 ERCC-00024 ERCC-00025 ERCC-00028 ERCC-00031 ERCC-00033 ERCC-00034 ERCC-00035 ERCC-00039 ERCC-00040 ERCC-00041 ERCC-00042 ERCC-00044 ERCC-00048 ERCC-00054 ERCC-00057 ERCC-00058 ERCC-00059 ERCC-00061 ERCC-00079 ERCC-00084 ERCC-00170
pos	a numeric vector
code_strand	a factor with levels +
seq_strand	a factor with levels S
X1	a numeric vector
X2	a numeric vector
X3	a numeric vector
X4	a numeric vector
X5	a numeric vector
X6	a numeric vector
X7	a numeric vector
X8	a numeric vector
X9	a numeric vector
X10	a numeric vector
X11	a numeric vector
X12	a numeric vector
X13	a numeric vector
X14	a numeric vector
X15	a numeric vector
X16	a numeric vector
X17	a numeric vector
X18	a numeric vector

X19 a numeric vector
X20 a numeric vector
X21 a numeric vector
X22 a numeric vector
X23 a numeric vector
X24 a numeric vector
X25 a numeric vector
X26 a numeric vector
X27 a numeric vector
X28 a numeric vector
X29 a numeric vector
X30 a numeric vector
X31 a numeric vector
X32 a numeric vector
X33 a numeric vector
X34 a numeric vector
X35 a numeric vector
X36 a numeric vector
X37 a numeric vector
X38 a numeric vector
X39 a numeric vector
X40 a numeric vector
X41 a numeric vector
X42 a numeric vector
X43 a numeric vector
X44 a numeric vector
X45 a numeric vector
X46 a numeric vector
X47 a numeric vector
X48 a numeric vector
X49 a numeric vector
X50 a numeric vector
X51 a numeric vector
X52 a numeric vector
X53 a numeric vector
X54 a numeric vector
X55 a numeric vector

X56 a numeric vector
X57 a numeric vector
X58 a numeric vector
X59 a numeric vector
X60 a numeric vector
X61 a numeric vector
X62 a numeric vector
X63 a numeric vector
X64 a numeric vector
X65 a numeric vector
X66 a numeric vector
X67 a numeric vector
X68 a numeric vector
X69 a numeric vector
X70 a numeric vector
X71 a numeric vector
X72 a numeric vector
X73 a numeric vector
X74 a numeric vector
X75 a numeric vector
X76 a numeric vector
X77 a numeric vector
X78 a numeric vector
X79 a numeric vector
X80 a numeric vector
X81 a numeric vector

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

Examples

```
data(test.dat.seq)
```

train.dat.counts *Training Counts Dataset*

Description

An example, a data frame of reads counts of training samples

Usage

```
data(train.dat.counts)
```

Format

A data frame with 5203 observations on the following 6 variables.

```
gene a factor with levels ERCC-00002 ERCC-00003 ERCC-00007 ERCC-00009 ERCC-00012 ERCC-00013  
ERCC-00014 ERCC-00016 ERCC-00017 ERCC-00018 ERCC-00019 ERCC-00022 ERCC-00023  
ERCC-00024 ERCC-00025 ERCC-00028 ERCC-00031 ERCC-00033 ERCC-00034 ERCC-00035  
ERCC-00039 ERCC-00040 ERCC-00041 ERCC-00042 ERCC-00044 ERCC-00048 ERCC-00054  
ERCC-00058 ERCC-00061 ERCC-00079 ERCC-00171  
pos a numeric vector  
code_strand a factor with levels +  
seq_strand a factor with levels S  
GSM516589_run32_s_2_ERCC_map_Pos a numeric vector  
GSM517059_run29_s_1_ERCC_map_Pos a numeric vector
```

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

Examples

```
data(train.dat.counts)
```

train.dat.seq

*Training Sequence Index Dataset***Description**

An example, a data frame of sequence index of training samples

Usage

```
data(train.dat.seq)
```

Format

A data frame with 5203 observations on the following 85 variables.

gene	a factor with levels ERCC-00002 ERCC-00003 ERCC-00007 ERCC-00009 ERCC-00012 ERCC-00013 ERCC-00014 ERCC-00016 ERCC-00017 ERCC-00018 ERCC-00019 ERCC-00022 ERCC-00023 ERCC-00024 ERCC-00025 ERCC-00028 ERCC-00031 ERCC-00033 ERCC-00034 ERCC-00035 ERCC-00039 ERCC-00040 ERCC-00041 ERCC-00042 ERCC-00044 ERCC-00048 ERCC-00054 ERCC-00058 ERCC-00061 ERCC-00079 ERCC-00171
pos	a numeric vector
code_strand	a factor with levels +
seq_strand	a factor with levels S
X1	a numeric vector
X2	a numeric vector
X3	a numeric vector
X4	a numeric vector
X5	a numeric vector
X6	a numeric vector
X7	a numeric vector
X8	a numeric vector
X9	a numeric vector
X10	a numeric vector
X11	a numeric vector
X12	a numeric vector
X13	a numeric vector
X14	a numeric vector
X15	a numeric vector
X16	a numeric vector
X17	a numeric vector
X18	a numeric vector

X19 a numeric vector
X20 a numeric vector
X21 a numeric vector
X22 a numeric vector
X23 a numeric vector
X24 a numeric vector
X25 a numeric vector
X26 a numeric vector
X27 a numeric vector
X28 a numeric vector
X29 a numeric vector
X30 a numeric vector
X31 a numeric vector
X32 a numeric vector
X33 a numeric vector
X34 a numeric vector
X35 a numeric vector
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X38 a numeric vector
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X43 a numeric vector
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X52 a numeric vector
X53 a numeric vector
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X55 a numeric vector

X56 a numeric vector
X57 a numeric vector
X58 a numeric vector
X59 a numeric vector
X60 a numeric vector
X61 a numeric vector
X62 a numeric vector
X63 a numeric vector
X64 a numeric vector
X65 a numeric vector
X66 a numeric vector
X67 a numeric vector
X68 a numeric vector
X69 a numeric vector
X70 a numeric vector
X71 a numeric vector
X72 a numeric vector
X73 a numeric vector
X74 a numeric vector
X75 a numeric vector
X76 a numeric vector
X77 a numeric vector
X78 a numeric vector
X79 a numeric vector
X80 a numeric vector
X81 a numeric vector

References

Cai G, RNA-SEQUENCING APPLICATIONS: GENE EXPRESSION QUANTIFICATION AND METHYLATOR PHENOTYPE IDENTIFICATION, Ph.D. Thesis, 2013

Examples

```
data(train.dat.seq)
```

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