# Package 'GOTHiC'

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Title Binomial test for Hi-C data analysis					
Description This is a Hi-C analysis package using a cumulative binomial test to detect interactions between distal genomic loci that have significantly more reads than expected by chance in Hi-C experiments. It takes mapped paired NGS reads as input and gives back the list of significant interactions for a given bin size in the genome.					
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R topics documented:					
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filtered

A GenomicRangesList object used as an example in the GOTHiC package

## **Description**

filtered is a GenomicRangesList example object used as an example for the binomialHiC package. This GenomicRangesList contains reads from a human lymphoblastoid cell line HiC experiment (Lieberman-Aiden et al. 2009) for chr20, that were mapped to the genome, paired and PCR duplicate-filtered.

## Usage

```
data(lymphoid_chr20_paired_filtered)
```

## **Format**

The format is: GenomicRangesList with 2 slots: \$paired\_reads\_1 contains the coordinates for one end of the paired reads \$paired\_reads\_2 contains the coordinates for the other end of the paired reads

## Author(s)

Borbala Gerle and Robert Sugar

## See Also

 ${\tt mapReadsToRestrictionSites}$ 

## **Examples**

data(lymphoid\_chr20\_paired\_filtered)

GOTHiC

Genome Organisation Through HiC

## **Description**

GOTHiC performs a cumulative binomial test to detect interactions between distal genomic loci that have significantly more reads than expected by chance in Hi-C experiments. It takes mapped paired NGS reads as input and gives back the list of significant interactions for a given bin size in the genome.

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#### Usage

```
GOTHiC(fileName1, fileName2, sampleName, res, BSgenomeName='BSgenome.Hsapiens.UCSC.hg19', genome=BSgenome.Hsapiens.UCSC.hg19, restrictionSite='A^AGCTT', enzyme='HindIII',cistrans='all',filterdist=10000, DUPLICATETHRESHOLD=1, fileType='BAM', parallel=FALSE, cores=NULL)
```

## **Arguments**

fileName1 File containing the mapped reads of the first fragment ends (BAM or Bowtie

format)

fileName2 File containing the mapped reads of the second fragment ends (BAM or Bowtie

format)

sampleName A character string that will be used to name the exported BedGraph file contain-

ing the coverage, R object files with paired and mapped reads, and the final data frame with the results from the binomial test. They will be saved in the current

directory.

res An integer that gives the required bin size or resolution of the contact map e.g.

1000000.

BSgenomeName A character string of the BSgenome package required to make the restriction

fragment file containing information for both the organism the experiment was made in, and the genome version the reads were mapped to. The default is the

current human genome build 'BSgenome. Hsapiens. UCSC. hg19'.

genome The BSgenome package required to make the restriction fragment file containing

information for both the organism the experiment was made in, and the genome version the reads were mapped to. The default is the current human genome

build BSgenome. Hsapiens. UCSC. hg19.

restrictionSite

A character string that specifies the enzymes recognition site, ^ indicating where the enzyme actually cuts. The default is the HindIII restriction site: 'A^AGCTT'.

enzyme A character string containing the name of the enzyme used during the Hi-C

experiment (i.e. "HindIII", "NcoI"). The default is "HindIII".

cistrans A character string with three possibilities. "all" runs the binomial test on all

interactions, "cis" runs the binomial test only on intrachromosomal/cis interactions, "trans" runs the binomial test only on interchromosomal/trans interactions.

filterdist An integer specifying the distance between the midpoint of fragments under

which interactions are filtered out in order to filter for those read-pairs where

the digestion was incomplete. The default is 10000.

DUPLICATETHRESHOLD

An integer specifying the maximum amount of duplicated paired-end reads allowed, over that value it is expected to be PCR bias. The default is 1.

fileType A character string specifying the format of the aligned reads. The default is

'BAM'. Other accepted format is 'Bowtie'.

parallel Logical argument. If TRUE the mapping and the binomial test will be performed

faster using multiple cores. The default is FALSE.

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cores

An integer specifying the number of cores used in the parallel processing if parellel=TRUE. The default is NULL.

#### Value

A data.frame containing elements

chr1 / chr2 chromosome(s) containing interacting regions 1 and 2

locus1 / locus2

start positions of the interacting regions 1 and 2 in the corresponding chromo-

some(s)

relCoverage1 / relCoverage2

relative coverage corresponding to regions 1 and 2

probability expected frequency

expected expected number of reads

readCount observed reads number

pvalue binomial p-value

qvalue binomial p-value corrected for multi-testing with Benjamini-Hochberg

logObservedOverExpected

observed/expected read numbers log ratio

## Author(s)

Borbala Mifsud and Robert Sugar

## See Also

binom.test, pairReads, mapReadsToRestrictionSites

## Examples

```
library(GOTHiC)
dirPath <- system.file("extdata", package="HiCDataLymphoblast")
fileName1 <- list.files(dirPath, full.names=TRUE)[1]
fileName2 <- list.files(dirPath, full.names=TRUE)[2]
binom=GOTHiC(fileName1, fileName2, sampleName='lymphoid_chr20', res=1000000,
BSgenomeName='BSgenome.Hsapiens.UCSC.hg18', genome=BSgenome.Hsapiens.UCSC.hg18,
restrictionSite='A^AGCTT', enzyme='HindIII',cistrans='all', filterdist=10000,
DUPLICATETHRESHOLD=1, fileType='Table', parallel=FALSE, cores=NULL)</pre>
```

mapReadsToRestrictionSites

Function to map aligned and paired reads to the restriction fragments

## **Description**

This function takes mapped paired NGS reads in the format of a GenomicRangesList object where the two end of the reads are in the GenomicRanges paired\_reads\_1 and paired\_reads\_2. It prepares the digestion file from the genome supplied to it with the given restriction enzyme and specificity and maps the reads to the fragments.

## Usage

mapReadsToRestrictionSites
(pairedReadsFile, sampleName,
BSgenomeName, genome, restrictionSite, enzyme, parallel=F, cores=1)

step 2

Arguments

pairedReadsFile

这步会产生的文件: coverage\_lymphoid\_chr20.bed, Digest\_BSgenome.Hsapiens.UCSC.hg18\_HindIII, interactingLoci\_lymphoid\_chr20

R object of GenomicRangesList containing paired\_reads\_1 and paired\_reads\_2 GenomicRanges with the paired mapped reads from a Hi-C experiment.

sampleName A character string that will be used to name the exported R object file with the mapped reads containing a GenomicRangesList with slots locus1 and locus2. It

will be saved in the current directory.

BSgenomeName A character string of the BSgenome package required to make the restriction

fragment file containing information for both the organism the experiment was made in, and the genome version the reads were mapped to. The default is the

current human genome build 'BSgenome. Hsapiens. UCSC. hg19'.

genome The BSgenome package required to make the restriction fragment file containing

information for both the organism the experiment was made in, and the genome version the reads were mapped to. The default is the current human genome

build BSgenome. Hsapiens. UCSC. hg19.

restrictionSite

enzyme

A character string that specifies the enzymes recognition site, ^ indicating where the enzyme actually cuts. The default is the HindIII restriction site: 'A^AGCTT'.

A character string containing the name of the enzyme used during the Hi-C

experiment (i.e. "HindIII", "NcoI"). The default is "HindIII".

parallel Logical argument. If TRUE the mapping will be performed faster using multiple

cores. The default is FALSE.

cores An integer specifying the number of cores used in the parallel processing if

parellel=TRUE. The default is 1.

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#### Value

## A GenomicRangesList

locus1 GenomicRanges with the coordinates of the start of the fragment where one end

of the read mapped

locus2 GenomicRanges with the coordinates of the start of the fragment where the other

end of the read mapped

#### Author(s)

Borbala Mifsud and Robert Sugar

#### See Also

```
pairReads, GOTHiC
```

## **Examples**

```
library(GOTHiC)
data(lymphoid_chr20_paired_filtered)
mapped=mapReadsToRestrictionSites(filtered, sampleName='lymphoid_chr20',
BSgenomeName='BSgenome.Hsapiens.UCSC.hg18', genome=BSgenome.Hsapiens.UCSC.hg18,
restrictionSite='A^AGCTT', enzyme='HindIII', parallel=FALSE, cores=1)
```

pairReads

Function pairs aligned paired NGS reads

## **Description**

This function takes bowtie output files, pairs the reads, only keeps those where both ends mapped, filters for perfect duplicates to avoid PCR bias, and saves and returns a GenomicRangesList object that contains the paired\_reads\_1 and paired\_reads\_2 GenomicRanges with the paired reads

## Usage

```
pairReads(fileName1, fileName2, sampleName, DUPLICATETHRESHOLD = 1,
fileType='BAM')
```

#### **Arguments**

fileNamel	File containing the	mapped reads of	the first fragment	ends (BAM or Bowtie
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format)

fileName2 File containing the mapped reads of the second fragment ends (BAM or Bowtie

format)

sampleName A character string that will be used to name the exported BedGraph file contain-

ing the coverage, and the R object file with paired reads. They will be saved in

the current directory.

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#### DUPLICATETHRESHOLD

An integer specifying the maximum amount of duplicated paired-end reads allowed, over that value it is expected to be PCR bias. The default is 1.

fileType

A character string specifying the format of the aligned reads. The default is 'BAM'. Other accepted format is 'Bowtie'.

## Value

A GenomicRangesList called filtered

paired\_reads\_1 GenomicRanges with the coordinates of where one end of the read mapped paired\_reads\_2 GenomicRanges with the coordinates of where the other end of the read mapped

## Author(s)

Borbala Mifsud and Robert Sugar

#### See Also

mapReadsToRestrictionSites, GOTHiC

## **Examples**

```
library(GOTHiC)
dirPath <- system.file("extdata", package="HiCDataLymphoblast")
fileName1 <- list.files(dirPath, full.names=TRUE)[1]
fileName2 <- list.files(dirPath, full.names=TRUE)[2]
paired <- pairReads(fileName1, fileName2, sampleName='lymphoid_chr20',
DUPLICATETHRESHOLD = 1, fileType='Table')</pre>
```

step one

这一步是为了让读入的mapping好的文件转为 GRanges 文件格式,会产生文件 lymphoid\_chr20\_paired\_filtered

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