FraseR

July 8, 2017

Type Package

Title Find RAre Splicing Events in RNA-Seq Data

Version 0.99.1 **Date** 2017-05-27

Author Christian Mertes <mertes@in.tum.de>, Prof. Julien Gagneur

<gagneur@in.tum.de>

Maintainer Christian Mertes <mertes@in.tum.de>

Description Detection of rare aberrant splicing events in transcriptome profiles. The workflow aims to assist the diagnostics in the field of rare diseases where RNA-seq is performed to identify splicing defects.

License Artistic-2.0

Depends R (>= 3.4), BiocParallel, data.table, Rsamtools, SummarizedExperiment (>= 1.6.3)

Imports BBmisc, biomaRt, DelayedArray, DT, GenomicAlignments, GenomicRanges, IRanges, graphics, gplots, htmlwidgets, HDF5Array, methods, parallel, plotly (>= 4.6.0), rhdf5, Rsubread, R.utils, shiny, stats, tidyr, tools, VGAM

biocViews RNASeq, AlternativeSplicing, Sequencing, Software, Genetics, Coverage

URL https://i12g-gagneurweb.in.tum.de/gitlab/mertes/FraseR/

RoxygenNote 6.0.1

Collate 'FraseR-package.R'

'FraseRDataSet-class.R'

'AllGenerics.R'

'Fraser-pipeline.R'

'accessor-methods.R'

'annotationOfRanges.R'

'beta-binomial-testing.R'

'calculatePSIValue.R'

'calculateStats.R'

'countRNAseqData.R'

'example_functions.R'

'getURLs.R'

'helper-functions.R'

'plotResults.R'

'saveHDF5Objects.R'

'shiny.R'

VignetteBuilder knitr

Index

Suggests BiocStyle, knitr, testthat, covr, RSelenium

${\sf R}$ topics documented:

annotateRanges
assayNames,FraseRDataSet-method
assays,FraseRDataSet-method
bamFile,FraseRDataSet-method
calculatePSIValues
calculatePValues
calculateZScores
cleanCache
condition,FraseRDataSet-method
countRNAData
counts,FraseRDataSet-method
createFullLinkTable
createTestFraseRDataSet
createTestFraseRSettings
FraseR
FraseRDataSet
FraseRDataSet-class
FraseRShinyApp
length,FraseRDataSet-method
loadFraseRDataSet
mcols,FraseRDataSet-method
method,FraseRDataSet-method
na2false
name,FraseRDataSet-method
nonSplicedReads,FraseRDataSet-method
parallel,FraseRDataSet-method
plotCountsAtSite
plotSampleResults
results
samples,FraseRDataSet-method
saveFraseRDataSet
scanBamParam,FraseRDataSet-method
strandSpecific,FraseRDataSet-method
subsetFraseR
workingDir,FraseRDataSet-method

20

annotateRanges 3

annotateRanges	Annotates the given biomaRt	FraseRDataSet with	the HGNC symbol with
	O to mark		

Description

Annotates the given FraseRDataSet with the HGNC symbol with biomaRt

Usage

```
annotateRanges(fds, feature = "hgnc_symbol",
  biotype = list("protein_coding"), ensembl = NULL)
```

```
assayNames,FraseRDataSet-method
```

Returns the assayNames of FraseR

Description

Returns the assayNames of FraseR

Usage

```
## S4 method for signature 'FraseRDataSet'
assayNames(x)
```

```
assays,FraseRDataSet-method
```

Returns the assay corrensonding to the given name/index of the FraseRDataSet

Description

Returns the assay corrensonding to the given name/index of the FraseRDataSet

```
## S4 method for signature 'FraseRDataSet'
assays(x, ..., type = NULL, withDimnames = TRUE)
```

4 calculatePSIValues

```
bamFile,FraseRDataSet-method
```

Get/Set the bamFile

Description

Get/Set the bamFile

Usage

```
## S4 method for signature 'FraseRDataSet'
bamFile(object)

## S4 replacement method for signature 'FraseRDataSet'
bamFile(object) <- value</pre>
```

Arguments

object AF1

A FraseRDataSet object.

Value

A vector with the bamFile for each sample

Author(s)

Christian Mertes < mertes@in.tum.de>

Examples

```
settings <- createTestFraseRSettings()
bamFile(settings)
bamFile(settings) <- file.path("bamfiles", samples(settings), "rna-seq.bam")</pre>
```

calculatePSIValues This function calculates the PSI values for each junction and splice site based on the FraseRDataSet object

Description

This function calculates the PSI values for each junction and splice site based on the FraseRDataSet object

Usage

```
calculatePSIValues(fds)
```

```
fds <- countRNAData(createTestFraseRSettings())
fds <- calculatePSIValues(fds)</pre>
```

calculatePValues 5

calculatePValues

calculates the P-Value for the given FraseR dataset object The P-Value calculation is based on the given method in the FraseRSettings object

Description

calculates the P-Value for the given FraseR dataset object The P-Value calculation is based on the given method in the FraseRSettings object

Usage

```
calculatePValues(fds, internBPPARAM = SerialParam(), ...)
```

Examples

```
fds <- countRNAData(createTestFraseRSettings())
fds <- calculatePSIValues(fds)
fds <- calculatePValues(fds)</pre>
```

calculateZScores

Calculate the zscore for each PSI value.

Description

Calculate the zscore for each PSI value.

Usage

```
calculateZScores(fds)
```

Examples

```
fds <- countRNAData(createTestFraseRSettings())
fds <- calculatePSIValues(fds)
fds <- calculateZScores(fds)</pre>
```

cleanCache

clear the files in the cache to start fresh

Description

clear the files in the cache to start fresh

```
cleanCache(fds, all = FALSE, cache = TRUE, assays = FALSE,
  results = FALSE)
```

```
fds <- createTestFraseRSettings()
cleanCache(fds)</pre>
```

```
{\it Condition, FraseRDataSet-method} \\ {\it Get the condition for each sample}
```

Description

Get the condition for each sample

Usage

```
## S4 method for signature 'FraseRDataSet'
condition(object)

## S4 replacement method for signature 'FraseRDataSet'
condition(object) <- value</pre>
```

Arguments

```
object A FraseRDataSet object.
```

Value

A vector with the condition per sample If the condition column is not set sequence of numbers is returned.

Author(s)

```
Christian Mertes <mertes@in.tum.de>
```

```
fds <- createTestFraseRSettings()
condition(fds)
condition(fds) <- 1:dim(fds)[2]</pre>
```

countRNAData 7

countRNAData

Count RNA-seq data

Description

This method extracts and counts the split reads and non spliced reads from a RNA bam file

Usage

```
countRNAData(fds, NcpuPerSample = 1, junctionMap = NULL)
```

Arguments

fds A FraseRDataSet object with all the information how and what to count NcpuPerSample

A BiocParallel param object or a positive integer to configure the parallel backend of the internal loop per sample

junctionMap A object or file containing a map of all junctions of interest across all samples

Details

TODO

Value

FraseRDataSet

Examples

```
countRNAData(createTestFraseRSettings())
countRNAData(createTestFraseRSettings(), 5)
```

Description

getter for count data

```
## S4 method for signature 'FraseRDataSet'
counts(object, type = NULL, side = c("ofInterest",
    "otherSide"))
```

createFullLinkTable

get links

Description

get links

Usage

createFullLinkTable(data, addHRef = TRUE)

createTestFraseRDataSet

Create a test case dataset based on the test sample annotation filled with counts to be used in the vignette and to explore the functionallity of the FraseR package.

Description

Create a test case dataset based on the test sample annotation filled with counts to be used in the vignette and to explore the functionallity of the FraseR package.

Usage

```
createTestFraseRDataSet(BPPARAM = NULL)
```

Value

a FraseRDataSet object which contains a test case

Examples

createTestFraseRDataSet()

createTestFraseRSettings

Create a test case dataset (sample information only) to be used in the vignette and to explore the functionallity of the FraseR package.

Description

Create a test case dataset (sample information only) to be used in the vignette and to explore the functionallity of the FraseR package.

```
createTestFraseRSettings()
```

FraseR 9

Value

a FraseRSettings object which contains a test case

Examples

```
createTestFraseRSettings()
```

FraseR

FraseR: A package providing a workflow to detect aberrant splicing events in RNA sequencing data in a rare disease cohort

Description

FraseR: A package providing a workflow to detect aberrant splicing events in RNA sequencing data in a rare disease cohort

This function performs a default analysis of RNA-seq data

Usage

```
FraseR(settings = createTestFraseRSettings(), NcpuPerSample = 1)
```

Arguments

settings A FraseRDataSet object with all the information how and what to count NcpuPerSample

A BiocParallel param object to configure the parallel backend of the internal loop for counting

Value

FraseRDataSet

```
fds <- FraseR(createTestFraseRSettings())
plotSampleResults(fds)</pre>
```

10 FraseRDataSet-class

FraseRDataSet

The constructor function for FraseRSettings

Description

The constructor function for FraseRSettings

Usage

```
FraseRDataSet(colData = NULL, ...)
```

Arguments

Any parameters corresponding to the slots and their possible values. See FraseR-DataSet

Value

A FraseRDataSet object.

Author(s)

Christian Mertes <mertes@in.tum.de>

Examples

```
fraser <- FraseRDataSet()
fraser <- countRNAData(createTestFraseRSettings())</pre>
```

```
FraseRDataSet-class
```

FraseRDataSet

Description

This class is designed to store the whole FraseR data set needed for an analysis of a disease cohort

Author(s)

```
Christian Mertes <mertes@in.tum.de>
```

FraseRShinyApp 11

FraseRShinyApp

Present the FraseR results as shiny app

Description

Present the FraseR results as shiny app

Usage

```
FraseRShinyApp(fds, fdsres = NULL, server = !interactive(), ...)
```

Examples

```
fds <- createTestFraseRDataSet()

# for interactive sessions
myShinyApp <- FraseRShinyApp(fds)

# for running a shiny application as server
myShinyAppObj <- FraseRShinyApp(fds, server=TRUE)</pre>
```

```
length, FraseRDataSet-method
```

retrive the length of the object (aka number of junctions)

Description

retrive the length of the object (aka number of junctions)

Usage

```
\#\# S4 method for signature 'FraseRDataSet' length(x)
```

loadFraseRDataSet Load a saved FraseR object into memory

Description

Load a saved FraseR object into memory

Usage

```
loadFraseRDataSet(dir, name = NULL)
```

Arguments

dir

a path to the working directory of FraseR

```
loadFraseRDataSet(file.path(Sys.getenv("HOME"), "FraseR"))
```

Description

getter and setter for mcols

Usage

```
## S4 method for signature 'FraseRDataSet'
mcols(x, type = NULL, ...)
```

method, FraseRDataSet-method

Get/Set the statistical method to use for P-value calculation from the FraseRDataSet object

Description

Get/Set the statistical method to use for P-value calculation from the FraseRDataSet object

Usage

```
## S4 method for signature 'FraseRDataSet'
method(object)

## S4 replacement method for signature 'FraseRDataSet'
method(object) <- value</pre>
```

Arguments

object A FraseRDataSet object.

Value

A character string representing the statistical method

Author(s)

Christian Mertes <mertes@in.tum.de>

```
settings <- createTestFraseRSettings()
method(settings)
method(settings) <- "betaBin"</pre>
```

na2false 13

na2false

convert all NA's of a input vector or of a single dimension matrix/data.table to FALSE

Description

convert all NA's of a input vector or of a single dimension matrix/data.table to FALSE

Usage

```
na2false(x)
```

Examples

```
a <- c(TRUE, FALSE, NA, TRUE, NA)
na2false(a)
dt <- data.table(a)
na2false(dt)</pre>
```

```
name, FraseRDataSet-method
```

Get/Set the name of the analysis

Description

Get/Set the name of the analysis

Usage

```
## S4 method for signature 'FraseRDataSet'
name(object)
## S4 replacement method for signature 'FraseRDataSet'
name(object) <- value</pre>
```

Arguments

```
object A FraseRDataSet object.
```

Value

A character string representing the name of the analysis

Author(s)

```
Christian Mertes <mertes@in.tum.de>
```

```
settings <- createTestFraseRSettings()
name(settings)
name(settings) <- "My Analysis"</pre>
```

nonSplicedReads,FraseRDataSet-method

accessor for the non spliced reads object within the FraseRDataSet object

Description

accessor for the non spliced reads object within the FraseRDataSet object setter for the non spliced reads object within the FraseRDataSet object

Usage

```
## S4 method for signature 'FraseRDataSet'
nonSplicedReads(object)

## S4 replacement method for signature 'FraseRDataSet'
nonSplicedReads(object) <- value</pre>
```

parallel, FraseRDataSet-method

Get/Set the parallel param object from the FraseRDataSet object

Description

Get/Set the parallel param object from the FraseRDataSet object

Usage

```
## S4 method for signature 'FraseRDataSet'
parallel(object)

## S4 replacement method for signature 'FraseRDataSet'
parallel(object) <- value</pre>
```

Arguments

object A FraseRDataSet object.

Value

A parallel param object

Author(s)

Christian Mertes <mertes@in.tum.de>

plotCountsAtSite 15

Examples

```
settings <- createTestFraseRSettings()
parallel(settings)
parallel(settings) <- SerialParam()</pre>
```

```
plotCountsAtSite plot count distribution
```

Description

plot count distribution

Usage

```
plotCountsAtSite(gr, fds, type, sample = NULL, plotLog = TRUE)
```

```
plotSampleResults Plot the results of the FraseR analysis pipeline All three types are plotted into one HTML file based on plotly.
```

Description

Plot the results of the FraseR analysis pipeline All three types are plotted into one HTML file based on plotly.

Usage

```
plotSampleResults(fds, sampleID = NULL, file = NULL, dir = NULL,
    browseIt = FALSE)
```

Examples

```
fds <- FraseR()
plotSampleResults(fds, "sample1")
plotSampleResults(fds, "sample1", "result.html")</pre>
```

results

obtain the results for the given analysis pipeline

Description

obtain the results for the given analysis pipeline

```
results(fds, sampleIDs = samples(fds), pvalueCut = 1e-05, zscoreCut = 2,
    psiType = c("psi3", "psi5", "psiSite"), redo = FALSE)
```

16 saveFraseRDataSet

```
samples, FraseRDataSet-method {\it Getter/Setter for the \ sampleIDs}
```

Description

Getter/Setter for the sampleIDs

Usage

```
## S4 method for signature 'FraseRDataSet'
samples(object)

## S4 replacement method for signature 'FraseRDataSet'
samples(object) <- value</pre>
```

Arguments

object A FraseRDataSet object.

Value

A vector with all sample IDs

Author(s)

Christian Mertes <mertes@in.tum.de>

Examples

```
fds <- createTestFraseRSettings()
samples(fds)
samples(fds) <- 1:dim(fds)[2]</pre>
```

saveFraseRDataSet Saves the FraseRDataSet object on disk under the given working dir.

It furthermore uses HDF5 to save all internal assays

Description

After saving this can be loaded again with the corresponding load function

Usage

```
saveFraseRDataSet(fds, dir = NULL)
```

Arguments

fds	A FraseRDataSet object ot be saved
dir	a directory name where to save the objects (replaces the working directory)

```
fds <- countRNAData(createTestFraseRSettings())
saveFraseRDataSet(fds)</pre>
```

scanBamParam,FraseRDataSet-method

Get/Set the ScanBamParam object from the FraseRDataSet object

Description

Get/Set the ScanBamParam object from the FraseRDataSet object

Usage

```
## S4 method for signature 'FraseRDataSet'
scanBamParam(object)

## S4 replacement method for signature 'FraseRDataSet'
scanBamParam(object) <- value</pre>
```

Arguments

object A FraseRDataSet object.

Value

A ScanBamParam object

Author(s)

Christian Mertes <mertes@in.tum.de>

```
settings <- createTestFraseRSettings()
scanBamParam(settings)
scanBamParam(settings) <- ScanBamParam(mapqFilter=30)</pre>
```

18 subsetFraseR

```
strandSpecific, FraseRDataSet-method

Get/Set if the analysis is strand specific or not
```

Description

Get/Set if the analysis is strand specific or not

Usage

```
## S4 method for signature 'FraseRDataSet'
strandSpecific(object)

## S4 replacement method for signature 'FraseRDataSet'
strandSpecific(object) <- value</pre>
```

Arguments

object A FraseRDataSet object.

Value

A logical value if the analysis is strand specific

Author(s)

Christian Mertes <mertes@in.tum.de>

Examples

```
settings <- createTestFraseRSettings()
strandSpecific(settings)
strandSpecific(settings) <- TRUE</pre>
```

subsetFraseR

Subsetting by indices for junctions

Description

Providing subsetting by indices through the single-bracket operator

Usage

```
subsetFraseR(x, i, j)
```

Arguments

- x A FraseRDataSet object
- i A integer vector to subset the rows/ranges
- j A integer vector to subset the columns/samples

Value

 $A \ subsetted \ {\tt FraseRDataSet} \ object$

Examples

```
fds <- countRNAData(createTestFraseRSettings())
fds[1:10,1:10]
fds[,samples(fds) %in% c("sample1", "sample2")]</pre>
```

workingDir,FraseRDataSet-method

Get/Set the working directory from the FraseRDataSet object

Description

Get/Set the working directory from the FraseRDataSet object

Usage

```
## S4 method for signature 'FraseRDataSet'
workingDir(object)

## S4 replacement method for signature 'FraseRDataSet'
workingDir(object) <- value</pre>
```

Arguments

object A FraseRDataSet object.

Value

A path

Author(s)

Christian Mertes <mertes@in.tum.de>

```
settings <- createTestFraseRSettings()
workingDir(settings)
workingDir(settings) <- tempdir()</pre>
```

Index

```
annotateRanges, 3
                                        nonSplicedReads, FraseRDataSet-method,
assayNames, FraseRDataSet-method,
       3
                                        nonSplicedReads<-,FraseRDataSet-method
assays, FraseRDataSet-method, 3
                                                (nonSplicedReads, FraseRDataSet-method),
bamFile, FraseRDataSet-method, 4
                                        parallel, FraseRDataSet-method, 14
bamFile<-,FraseRDataSet-method
       (bamFile, FraseRDataSet-method), parallel<-, FraseRDataSet-method
                                                (parallel, FraseRDataSet-method),
                                                14
                                        plotCountsAtSite, 15
calculatePSIValues, 4
                                        plotSampleResults, 15
calculatePValues, 5
calculateZScores, 5
                                        results, 15
cleanCache. 5
condition, FraseRDataSet-method,
                                        samples, FraseRDataSet-method, 16
                                        samples<-,FraseRDataSet-method</pre>
condition<-,FraseRDataSet-method
                                               (samples, FraseRDataSet-method),
       (condition, FraseRDataSet-method),
       6
                                        saveFraseRDataSet, 16
countRNAData, 7
                                        scanBamParam, FraseRDataSet-method,
counts, FraseRDataSet-method, 7
                                                17
createFullLinkTable, 8
                                        scanBamParam<-,FraseRDataSet-method</pre>
createTestFraseRDataSet, 8
                                                (scanBamParam, FraseRDataSet-method),
createTestFraseRSettings, 8
                                                17
                                        strandSpecific,FraseRDataSet-method,
FraseR, 9
FraseRDataSet, 10, 10
                                        strandSpecific<-,FraseRDataSet-method
FraseRDataSet-class, 10
                                               (strandSpecific, FraseRDataSet-method),
FraseRShinyApp, 11
                                        subsetFraseR, 18
length, FraseRDataSet-method, 11
loadFraseRDataSet, 11
                                        workingDir, FraseRDataSet-method,
mcols,FraseRDataSet-method,12
                                        workingDir<-,FraseRDataSet-method
method, FraseRDataSet-method, 12
                                                (workingDir, FraseRDataSet-method),
method<-,FraseRDataSet-method
       (method, FraseRDataSet-method),
       12
na2false, 13
name, FraseRDataSet-method, 13
name<-,FraseRDataSet-method
       (name, FraseRDataSet-method),
       13
```