

Package ‘LinTInd’

September 16, 2024

Title Lineage tracing by indels

Version 1.8.0

Description When we combine gene-editing technology and sequencing technology, we need to reconstruct a lineage tree from alleles generated and calculate the similarity between each pair of groups.

FindIndel() and IndelForm() function will help you align each read to reference sequence and generate scar form strings respectively.

IndelIdents() function will help you to define a scar form for each cell or read.

IndelPlot() function will help you to visualize the distribution of deletion and insertion.

TagProcess() function will help you to extract indels for each cell or read.

TagDist() function will help you to calculate the similarity between each pair of groups across the indwells they contain.

BuildTree() function will help you to reconstruct a tree.

PlotTree() function will help you to visualize the tree.

License MIT + file LICENSE

Encoding UTF-8

LazyData true

VignetteBuilder knitr

Suggests knitr, rmarkdown

Roxygen list(markdown = TRUE)

RoxygenNote 7.3.1

biocViews SingleCell, CRISPR, Alignment

Depends R (>= 4.0), ggplot2, parallel, stats, S4Vectors

Imports data.tree, reshape2, networkD3, stringdist, purrr, ape, cowplot, ggnewscale, stringr, dplyr, rlist, pheatmap, Biostrings, pwalign, IRanges, BiocGenerics(>= 0.36.1), ggtree

git_url <https://git.bioconductor.org/packages/LinTInd>

git_branch RELEASE_3_19

git_last_commit 4f54123

git_last_commit_date 2024-04-30

Repository Bioconductor 3.19

Date/Publication 2024-09-15

Author Luyue Wang [aut, cre],
 Bin Xiang [ctb],
 Hengxin Liu [ctb],
 Wu Wei [ths]

Maintainer Luyue Wang <wly1995310@gmail.com>

Contents

align_to_range	2
BuildTree	3
cellsinfo	4
celltype	4
change_form_stat	4
cutsite	5
data	5
example_data	6
FindIndel	6
IndelForm	7
IndelIdents	8
IndelPlot	9
PlotTree	9
ReadCutsite	10
ReadFasta	11
ref	11
scarform	12
scarinfo	12
scarref	12
scarref_all	13
tag	13
TagDist	13
TagProcess	14
treeinfo	15
Index	16

align_to_range	<i>align_to_range</i>
----------------	-----------------------

Description

Title

Usage

align_to_range(p, s, cut)

Arguments

p	A base sequence in character format
s	A base sequence in character format
cut	The distance between the starting sites of two fragments

Value

A list include two IRanges instances (deletion and insertion)

Examples

```
align_to_range(p="AAGG---AAATTTCGGAATAAGGAATTT",s="AAGGCCCAAATTT-CGGAATAAGGAATTT",cut=0)
```

 BuildTree

BuildTree

Description

Generate an array generant tree of a data.tree data structure and save it

Usage

```
BuildTree(tag)
```

Arguments

tag	List generate from TagProcess, for more see TagProcess
-----	--

Details

Title

Value

list with two elements, a data.tree data structure and a dataframe of array information for each cell barcode

Examples

```
data("example_data",package = "LinTInd")
treeinfo<-BuildTree(tag)
```

cellsinfo	<i>example of cellsinfo</i>
-----------	-----------------------------

Description

This an example of cellsinfo

Usage

```
data("example_data")
```

Format

cellsinfo

celltype	<i>example of celltype</i>
----------	----------------------------

Description

This an example of celltype

Usage

```
data("example_data")
```

Format

celltype

change_form_stat	<i>change_form_stat</i>
------------------	-------------------------

Description

Title

Usage

```
change_form_stat(indel)
```

Arguments

indel	List include two IRanges instances, contain start and end site of deletions and inserstions
-------	---

Value

A scar form string

Examples

```
data("example_data", package = "LinTInd")  
change_form_stat(cellsinfo$indel[[1]])
```

cutsite	<i>example of cutsite</i>
---------	---------------------------

Description

This an example of cutsite

Usage

```
data("example_data")
```

Format

cutsite

data	<i>example of data</i>
------	------------------------

Description

This an example of data

Usage

```
data("example_data")
```

Format

data

example_data	<i>example_data</i>
--------------	---------------------

Description

A set of example data

Usage

```
data("example_data")
```

Format

The format is: chr "example_data"

Examples

```
data("example_data", package = "LinTInd")
```

FindIndel	<i>FindIndel</i>
-----------	------------------

Description

This function can identify indels for each read in input data, and create IRanges instances for deletion and insertion

Usage

```
FindIndel(  
  data,  
  scarfull,  
  scar,  
  align_score = NULL,  
  type = NULL,  
  indel.coverage = NULL,  
  cln  
)
```

Arguments

data	data frame, include cell barcode, UMI and reads.
scarfull	DNASTring of reference sequence
scar	The cutsite data frame
align_score	The minimum alignment score that matched sequence should get, default in this parameter is the score that the reads which all of the target set were cutted got
type	Group name for this data ("None" in default)
indel.coverage	Choose indels selected scope: "Accurate" (default) means only the indles happened in target site will be idented; "All" means each indel will be detected even they locate on the anchors
cln	The number of threads

Details

Title

Value

list include IRanges instances (deletion and insertion), a data frame of reads' informations, reference sequenc, dataframe of cut sites

Examples

```
data("example_data", package = "LinTInd")
scarinfo<-FindIndel(data=data, scarfull=ref, scar=cutsite, indel.coverage="All", type="test", cln=1)
```

IndelForm

IndelForm

Description

Generate scar form strings from scarinfo list for each reads

Usage

```
IndelForm(scarinfo, cln)
```

Arguments

scarinfo	List generate from FindIndel, for more see FindIndel
cln	The number of threads

Details

Title

Value

A new list of scarinfo, the scarform of each reads will add in the data frame of reads' informations

Examples

```
data("example_data", package = "LinTInd")
IndelForm(scarinfo, cln=1)
```

IndelIdents

IndelIdents

Description

Function to define a scarform for each cell(single cell) or each reads(bulk seq, generate 'cell barcode' for each reads)

Usage

```
IndelIdents(scarinfo, method.use = NULL, cln)
```

Arguments

scarinfo	List generate from IndelForm, for more see IndelForm
method.use	Select how to determine a scar form string for each cell: "reads.num" (default):find the scar with the most reads in the cell; "umi.num":find the scar with the most UMIs in the cell; "consensus":find the consistent sequences in each cell, and then generate scar form strings from the new reads
cln	The number of threads

Details

Title

Value

The list generate from FindIndel, but in 'Scar' element a new column contain scar form strings

Examples

```
data("example_data", package = "LinTInd")
IndelIdents(scarinfo, method.use="umi.num", cln=1)
```

IndelPlot	<i>IndelPlot</i>
-----------	------------------

Description

Return 2 line charts, show the probability of insertion and deletion at each site

Usage

```
IndelPlot(cellsinfo)
```

Arguments

cellsinfo List generate from IndelIdents, for more see [IndelIdents](#)

Details

Title

Value

2 line charts

Examples

```
data("example_data", package = "LinTInd")
IndelPlot(cellsinfo = cellsinfo)
```

PlotTree	<i>PlotTree</i>
----------	-----------------

Description

Function to visualise the array generate tree

Usage

```
PlotTree(treeinfo, data.extract = NULL, annotation = NULL, prefix = NULL)
```

Arguments

treeinfo List generate from BuildTree, for more see [BuildTree](#)

data.extract (optional) If "FALSE" (default), will not return the indel's information, if it's "TRUE", the opposite

annotation (optional) If "TRUE" (default), the annotation of each cell barcodes have to be provided before, and a heatmap of cells' distribution for each array will be return

prefix (optional) Indels' prefix

Details

Title

Value

A list include a ggplot object, a dataframe show the distribution of each array contained in each group of cells (optional), and a dataframe to create the ggplot object

Examples

```
data("example_data", package = "LinTInd")
plotinfo<-PlotTree(treeinfo = treeinfo, data.extract = "TRUE", annotation = "TRUE")
plotinfo<-PlotTree(treeinfo = treeinfo, data.extract = "TRUE", annotation = "FALSE")
```

ReadCutsite

ReadCutsite

Description

Function to create a reference dataframe include each position and its' group

Usage

```
ReadCutsite(segref, reftype=NULL)
```

Arguments

segref	The cutsite file
reftype	Choose the reference type you want, if reftype="Accurate" (default), there will only the target sites be generated; if reftype="All", each site will be generated

Details

Title

Value

reference dataframe

Examples

```
data("example_data", package = "LinTInd")
ReadCutsite(cutsite)
ReadCutsite(cutsite, reftype="All")
```

ReadFasta

ReadFasta

Description

Function to read fasta file to DNString object

Usage

```
ReadFasta(filename)
```

Arguments

filename The input fasta file name

Details

Title

Value

A DNString object

Examples

```
fafilename=paste0(system.file("extdata",package = 'LinTInd'),"/V3.fasta")
ReadFasta(fafilename)
```

ref

example of ref

Description

This an example of ref

Usage

```
data("example_data")
```

Format

ref

scarform	<i>example of scarform</i>
----------	----------------------------

Description

This an example of scarform

Usage

```
data("example_data")
```

Format

scarform

scarinfo	<i>example of scarinfo</i>
----------	----------------------------

Description

This an example of scarinfo

Usage

```
data("example_data")
```

Format

scarinfo

scarref	<i>example of scarref</i>
---------	---------------------------

Description

This an example of scarref

Usage

```
data("example_data")
```

Format

scarref

scarref_all	<i>example of scarref_all</i>
-------------	-------------------------------

Description

This an example of scarref_all

Usage

```
data("example_data")
```

Format

```
scarref_all
```

tag	<i>example of tag</i>
-----	-----------------------

Description

This an example of tag

Usage

```
data("example_data")
```

Format

```
tag
```

TagDist	<i>TagDist</i>
---------	----------------

Description

If the cell barcode and the anntation of each cell are provided, this function can calculate the relationship between each cell type in three way

Usage

```
TagDist(tag, method = NULL)
```

Arguments

tag	List generate from TagProcess, for more see TagProcess
method	Denote which method to use: <ul style="list-style-type: none"> • "Jaccard"(default): calculate the weighted jaccard similarity of indels between each pair of groups; • "P": right-tailed test, compare the Indels intersection level with the hypothetical result generated from random editing, and the former is expected to be significantly higher than the latter; • "spearman": Spearman correlation of indels between each pair of groups

Details

Title

Value

2 figures are saved to show the distribution of INDEL and the relationship between groups respectively, the matrix of the relationship between groups is returned

Examples

```
tag_dist=TagDist(tag,method = "spearman")
tag_dist=TagDist(tag)
tag_dist=TagDist(tag,method = "Jaccard")
tag_dist=TagDist(tag,method = "P")
tag_dist=TagDist(tag,method = "spearman")
```

TagProcess

TagProcess

Description

Split each indel from scar form strings and map indel information to cell barcodes

Usage

```
TagProcess(data, Cells = NULL, prefix = NULL)
```

Arguments

data	List generate from IndelIdents, for more see IndelIdents
Cells	(optional) Dataframe of cells' annotation, with two columns: "Cell.BC" and "Cell.type"
prefix	(optional) Indels' prefix

Details

Title

Value

List with two dataframes: Indels for each cell barcode and cells' annotation

Examples

```
data("example_data", package = "LinTInd")  
TagProcess(cellsinfo$info, Cells=celltype)
```

treeinfo	<i>example of treeinfo</i>
----------	----------------------------

Description

This an example of treeinfo

Usage

```
data("example_data")
```

Format

treeinfo

Index

* datasets

- cellsinfo, [4](#)
- celltype, [4](#)
- cutsite, [5](#)
- data, [5](#)
- example_data, [6](#)
- ref, [11](#)
- scarform, [12](#)
- scarinfo, [12](#)
- scarref, [12](#)
- scarref_all, [13](#)
- tag, [13](#)
- treeinfo, [15](#)

[align_to_range, 2](#)

[BuildTree, 3, 9](#)

[cellsinfo, 4](#)

[celltype, 4](#)

[change_form_stat, 4](#)

[cutsite, 5](#)

[data, 5](#)

[example_data, 6](#)

[FindIndel, 6, 7](#)

[IndelForm, 7, 8](#)

[IndelIdents, 8, 9, 14](#)

[IndelPlot, 9](#)

[PlotTree, 9](#)

[ReadCutsite, 10](#)

[ReadFasta, 11](#)

[ref, 11](#)

[scarform, 12](#)

[scarinfo, 12](#)

[scarref, 12](#)

[scarref_all, 13](#)

[tag, 13](#)

[TagDist, 13](#)

[TagProcess, 3, 14, 14](#)

[treeinfo, 15](#)