

Package ‘nucleoSim’

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Title Generate synthetic nucleosome maps

Description This package can generate a synthetic map with reads covering the nucleosome regions as well as a synthetic map with forward and reverse reads emulating next-generation sequencing. The synthetic hybridization data of “Tiling Arrays” can also be generated. The user has choice between three different distributions for the read positioning: Normal, Student and Uniform. In addition, a visualization tool is provided to explore the synthetic nucleosome maps.

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Imports stats, IRanges, S4Vectors, graphics, methods

Suggests BiocStyle, BiocGenerics, knitr, rmarkdown, testthat

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URL <https://github.com/arnauddroitlab/nucleoSim>

BugReports <https://github.com/arnauddroitlab/nucleoSim/issues>

VignetteBuilder knitr

NeedsCompilation no

biocViews Genetics, Sequencing, Software, StatisticalMethod, Alignment

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nucleoSim-package	<i>nucleoSim: Generate synthetic nucleosome maps</i>
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Description

This package can generate a synthetic map with reads covering the nucleosome regions as well as a synthetic map with forward and reverse reads emulating next-generation sequencing. The user has choice between three different distributions for the read positioning: Normal, Student and Uniform.

Author(s)

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See Also

- [syntheticNucMapFromDist](#) to generate a synthetic nucleosome map

plot.syntheticNucMap	<i>Generate a graph of a synthetic nucleosome map</i>
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Description

Generate a graph for a list marked as an syntheticNucMap class

Usage

```
## S3 method for class 'syntheticNucMap'
plot(x, ...)
```

Arguments

x	a list marked as an syntheticNucMap class
...	... extra arguments passed to the plot function

Value

a graph of a synthetic nucleosome map

Author(s)

Astrid Deschenes, Rawane Samb

Examples

```
## Generate a synthetic map with 20 well-positioned nucleosomes, 5 fuzzy
## nucleosomes and 10 deleted nucleosomes using a Student distribution
## with a variance of 10 for the well-positioned nucleosomes,
## a variance of 20 for the fuzzy nucleosomes and a seed of 15
syntheticMap <- syntheticNucMapFromDist(wp.num = 30, wp.del = 10,
wp.var = 10, fuz.num = 5, fuz.var = 20, rnd.seed = 15,
distr = "Student")

## Create graph using the synthetic map
plot(syntheticMap, xlab="Position", ylab="Coverage")
```

plot.syntheticNucReads

Generate a graph of a synthetic nucleosome map containing forward and reverse reads

Description

Generate a graph for a list marked as an syntheticNucReads class

Usage

```
## S3 method for class 'syntheticNucReads'
plot(x, ...)
```

Arguments

x a list marked as a syntheticNucReads class
... ... extra arguments passed to the plot function

Value

a graph of a synthetic nucleosome map containing forward and reverse reads

Author(s)

Astrid Deschenes, Rawane Samb

Examples

```
## Generate a synthetic map with 20 well-positioned nucleosomes, 5 fuzzy
## nucleosomes and 10 deleted nucleosomes using a Student distribution
## with a variance of 10 for the well-positioned nucleosomes,
## a variance of 20 for the fuzzy nucleosomes and a seed of 15
syntheticNucSample <- syntheticNucReadsFromDist(wp.num = 30, wp.del = 10,
wp.var = 10, fuz.num = 5, fuz.var = 20, rnd.seed = 15,
distr = "Student", offset = 1000)

## Create graph using the synthetic map
plot(syntheticNucSample, xlab="Position", ylab="Coverage")
```

```
syntheticNucMapFromDist
```

Generate a synthetic nucleosome map containing complete sequences

Description

Generate a synthetic nucleosome map, a map with complete sequences covering the nucleosome regions, using the distribution selected by the user. The distribution is used to assign the start position to the sequences associated with the nucleosomes. The user has choice between three different distributions: Normal, Student and Uniform.

The synthetic nucleosome map creation is separated into 3 steps :

1. Adding well-positioned nucleosomes following specified parameters. The nucleosomes are all positioned at equidistance. Assigning sequences of variable length to each nucleosome using a normal distribution and specified variance.
2. Deleting some well-positioned nucleosomes following specified parameters. Each nucleosome has an equal probability to be selected.
3. Adding fuzzy nucleosomes following an uniform distribution ad specified parameters. Assigning sequences of variable length to each nucleosome using the specified distribution and parameters. The sequence length is always following a normal distribution.

This function is a modified version of the syntheticNucMap() function from Bioconductor nucleR package (Flores and Orozco, 2011).

Usage

```
syntheticNucMapFromDist(
  wp.num,
  wp.del,
  wp.var,
  fuz.num,
  fuz.var,
  max.cover = 100,
  nuc.len = 147,
  len.var = 10,
```

```

lin.len = 20,
rnd.seed = NULL,
as.ratio = FALSE,
distr = c("Uniform", "Normal", "Student")
)

```

Arguments

<code>wp.num</code>	a non-negative integer, the number of well-positioned (non-overlapping) nucleosomes.
<code>wp.del</code>	a non-negative integer, the number of well-positioned nucleosomes to remove to create uncovered regions.
<code>wp.var</code>	a non-negative integer, the variance associated with the distribution used to assign the start position to the sequences of the well-positioned nucleosomes. This parameter introduces some variation in the starting positions.
<code>fuz.num</code>	a non-negative integer, the number of fuzzy nucleosomes. Those nucleosomes are distributed accordingly to an uniform distribution all over the region. Those nucleosomes can overlap other well-positioned or fuzzy nucleosomes.
<code>fuz.var</code>	a non-negative integer, the variance associated with the distribution used to assign the start position to the sequences of the fuzzy nucleosomes. This parameter introduces some variation in the starting positions.
<code>max.cover</code>	a positive integer, the maximum coverage for one nucleosome. The final coverage can have a higher value than <code>max.cover</code> since sequences from different nucleosomes can be overlapping. Default = 100.
<code>nuc.len</code>	a non-negative numeric, the nucleosome length. Default = 147.
<code>len.var</code>	a non-negative integer, the variance associated to the normal distribution used to add some variance to the length of each sequence. Default = 10.
<code>lin.len</code>	a non-negative integer, the length of the DNA linker. Default = 20.
<code>rnd.seed</code>	a single value, interpreted as an integer, or NULL. If a integer is given, the value is used to set the seed of the random number generator. By fixing the seed, the generated results can be reproduced. Default = NULL.
<code>as.ratio</code>	a logical, if TRUE, a synthetic naked DNA control map is created and the ratio between it and the nucleosome coverage are calculated. It can be used to simulate hybridization ratio data, like the one in Tiling Arrays. Both control map and obtained ratio are returned. Default = FALSE.
<code>distr</code>	the name of the distribution used to generate the nucleosome map. The choices are : "Uniform", "Normal" and "Student". Default = "Uniform".

Value

an list of class "syntheticNucMap" containing the following elements:

- call the matched call.
- `wp.starts` a vector of integer, the start positions of all well-positioned nucleosome regions. The central position of the nucleosome is calculated as `wp.starts + round(nuc.len/2)`.

- `wp.nreads` a vector of integer, the number of sequences associated to each well-positioned nucleosome.
- `wp.reads` a IRanges containing the well-positioned nucleosome sequences.
- `fuz.starts` a vector of integer, the start position of all the fuzzy nucleosomes.
- `fuz.nreads` a vector of integer, the number of sequences associated to each fuzzy nucleosome.
- `fuz.reads` a IRanges containing the fuzzy nucleosome sequences.
- `syn.reads` a IRanges containing all the synthetic nucleosome sequences (from both fuzzy and well-positioned nucleosomes).
- `nuc.len` a numeric the nucleosome length.

The following elements will be only returned if `as.ratio=TRUE`:

- `ctr.reads` a IRanges containing the naked DNA (control) sequences.
- `syn.ratio` a Rle containing the calculated ratio between the nucleosome coverage and the control coverage.

Author(s)

Rawane Samb, Astrid Deschenes

Examples

```
## Generate a synthetic map with 20 well-positioned nucleosomes and 10 fuzzy
## nucleosomes using a Normal distribution with a variance of 30 for the
## well-positioned nucleosomes, a variance of 40 for the fuzzy nucleosomes
## and a seed of 15.
syntheticNucMapFromDist(wp.num = 20, wp.del = 0, wp.var = 30,
  fuz.num = 10, fuz.var = 40, rnd.seed = 15,
  distr = "Normal")

## Same output but with ratio
syntheticNucMapFromDist(wp.num = 20, wp.del = 0, wp.var = 30,
  fuz.num = 10, fuz.var = 40,
  rnd.seed = 15, as.ratio = TRUE, distr = "Normal")
```

syntheticNucReadsFromDist

Generate a synthetic nucleosome map containing forward and reverse reads (paired-end reads)

Description

Generate a synthetic nucleosome map, a map with forward and reverse reads (paired-end reads) covering the nucleosome regions, using the distribution selected by the user. The distribution is used to assign the start position to the forward reads associated with the nucleosomes. The user has choice between three different distributions: Normal, Student and Uniform. The final map is composed of paired-end reads.

#' The synthetic nucleosome map creation is separated into 3 steps :

1. Adding well-positioned nucleosomes following specified parameters. The nucleosomes are all positioned at equidistance. Assigning the starting positions of forward reads using the specified distribution and parameters. The distance between starting positions of paired-end reads is assigned using a normal distribution and specified variance.
2. Deleting some well-positioned nucleosomes following specified parameters. Each nucleosome has an equal probability to be selected.
3. Adding fuzzy nucleosomes following an uniform distribution and specified parameters. Assigning the starting positions of forward reads using the specified distribution and parameters. The distance between starting positions of paired-end reads is assigned using a normal distribution and specified variance.

This function has been largely inspired by the Generating synthetic maps section of the nucleR package (Flores et Orozco, 2011).

Usage

```
syntheticNucReadsFromDist(
  wp.num,
  wp.del,
  wp.var,
  fuz.num,
  fuz.var,
  max.cover = 100,
  nuc.len = 147,
  len.var = 10,
  lin.len = 20,
  read.len = 40,
  rnd.seed = NULL,
  distr = c("Uniform", "Normal", "Student"),
  offset
)
```

Arguments

wp.num	a non-negative integer, the number of well-positioned (non-overlapping) nucleosomes.
wp.del	a non-negative integer, the number of well-positioned nucleosomes to remove to create uncovered regions.
wp.var	a non-negative integer, the variance associated with the distribution used to assign the start position to the forward reads of the well-positioned nucleosomes. This parameter introduces some variation in the starting positions.

<code>fuz.num</code>	a non-negative numeric, the number of fuzzy nucleosomes. Those nucleosomes are distributed accordingly to an uniform distribution all over the region. Those nucleosomes can overlap other well-positioned or fuzzy nucleosomes.
<code>fuz.var</code>	a non-negative numeric, the maximum variance of the fuzzy nucleosomes. This variance can be different than the one used for the well-positioned nucleosome reads.
<code>max.cover</code>	a positive numeric, the maximum coverage for one nucleosome. The final coverage can have a higher value than <code>max.cover</code> since reads from different nucleosomes can be overlapping. Default = 100.
<code>nuc.len</code>	a positive integer, the nucleosome length. Default = 147.
<code>len.var</code>	a positive numeric, the variance of the distance between a forward read and its paired reverse read. Default = 10.
<code>lin.len</code>	a non-negative integer, the length of the DNA linker. Default = 20.
<code>read.len</code>	a positive integer, the length of each of the paired-end reads. Default = 40.
<code>rnd.seed</code>	a single value, interpreted as an integer, or NULL. If an integer is given, the value is used to set the seed of the random number generator. By fixing the seed, the generated results can be reproduced. Default = NULL.
<code>distr</code>	the name of the distribution used to generate the nucleosome map. The choices are : "Uniform", "Normal" and "Student". Default = "Uniform".
<code>offset</code>	a non-negative integer, the number of bases used to offset all nucleosomes and reads. This is done to ensure that all nucleosome positions and read alignment are of positive values.

Value

an list of class "syntheticNucReads" containing the following elements:

- `call` the matched call.
- `dataIP` a `data.frame` with the chromosome name, the starting and ending positions and the direction of all forward and reverse reads for all well-positioned and fuzzy nucleosomes. Paired-end reads are identified with an unique id.
- `wp` a `data.frame` with the positions of all the well-positioned nucleosomes, as well as the number of paired-reads associated to each one.
- `fuz` a `data.frame` with the positions of all the fuzzy nucleosomes, as well as the number of paired-reads associated to each one.
- `paired` a `data.frame` with the starting and ending positions of the reads used to generate the paired-end reads. Paired-end reads are identified with an unique id.

Author(s)

Pascal Belleau, Rawane Samb, Astrid Deschenes

Examples

```
## Generate a synthetic map with 20 well-positioned + 10 fuzzy nucleosomes
## using a Normal distribution with a variance of 30 for the well-positioned
## nucleosomes, a variance of 40 for the fuzzy nucleosomes and a seed of 15.
## Because of the fixed seed, each time is going to be run, the results
## are going to be the seed.
res <- syntheticNucReadsFromDist(wp.num = 20, wp.del = 0, wp.var = 30,
fuz.num = 10, fuz.var = 40, rnd.seed = 15, distr = "Normal",
offset = 1000)
```

```
syntheticNucReadsFromMap
```

Generate a synthetic nucleosome map containing forward and reverse reads

Description

Generate a synthetic nucleosome map using a synthetic nucleosome map.

This function is using a modified version of the `syntheticNucMap()` function from Bioconductor `nucleR` package (Flores and Orozco, 2011).

Usage

```
syntheticNucReadsFromMap(syntheticNucMap, read.len = 40, offset)
```

Arguments

<code>syntheticNucMap</code>	a list of class "syntheticNucMap"
<code>read.len</code>	a positive integer, the length of each of the paired-end reads. Default = 40.
<code>offset</code>	a non-negative integer, the number of bases used to offset all nucleosomes and reads. This is done to ensure that all nucleosome positions and read alignments are of positive values.

Value

a list of class "syntheticNucReads" containing the following elements:

- `call` the matched call.
- `dataIP` a `data.frame` with the chromosome name, the starting and ending positions and the direction of all forward and reverse reads for all well-positioned and fuzzy nucleosomes. Paired-end reads are identified with a unique id.
- `wp` a `data.frame` with the positions of all the well-positioned nucleosomes, as well as the number of paired-reads associated to each one.
- `fuz` a `data.frame` with the positions of all the fuzzy nucleosomes, as well as the number of paired-reads associated to each one.

- paired a `data.frame` with the starting and ending positions of the reads used to generate the paired-end reads. Paired-end reads are identified with a unique id.

Author(s)

Pascal Belleau, Rawane Samb, Astrid Deschenes

Examples

```
## Generate a synthetic map with 20 well-positioned + 10 fuzzy nucleosomes
## using a Normal distribution with a variance of 30 for the well-positioned
## nucleosomes, a variance of 40 for the fuzzy nucleosomes and a seed of 15
## Because of the fixed seed, each time is going to be run, the results
## are going to be the seed
syntheticMap <- syntheticNucMapFromDist(wp.num = 20, wp.del = 0,
  wp.var = 30, fuz.num = 10, fuz.var = 40,
  rnd.seed = 335, as.ratio = FALSE, distr = "Uniform")

res <- nucleoSim::syntheticNucReadsFromMap(syntheticMap, read.len = 45,
  offset = 1000)
```

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