Package 'idr2d'

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Description A tool to measure reproducibility between genomic experiments that produce two-dimensional peaks (interactions between peaks), such as ChIA-PET, HiChIP, and HiC. idr2d is an extension of the original idr package, which is intended for (one-dimensional) ChIP-seq peaks.

Title Irreproducible Discovery Rate for Genomic Interactions Data

```
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URL https://idr2d.mit.edu

Depends R (>= 3.6)

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Suggests DT (>= 0.4), htmltools (>= 0.3.6), knitr (>= 1.20), rmarkdown
```

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```
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```

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Contents

	calculate_midpoint_distance1d
	calculate_midpoint_distance2d
	calculate_relative_overlap1d
	calculate_relative_overlap2d
	chiapet
	chipseq
	determine_anchor_overlap
	draw_hic_contact_map
	draw_idr_distribution_histogram
	draw_rank_idr_scatterplot
	draw_value_idr_scatterplot
	establish_bijection
	establish_bijection1d
	establish_bijection2d
	establish_overlap1d
	establish_overlap2d
	estimate_idr
	estimate_idr1d
	estimate_idr2d
	estimate_idr2d_hic
	hic
	parse_hic_pro_matrix
	parse_juicer_matrix
	preprocess
	remove_nonstandard_chromosomes1d
	remove_nonstandard_chromosomes2d
_	
Index	35

calculate_midpoint_distance1d

Distance between Midpoints of two Peaks

Description

Calculates the distance in nucleotides between the midpoints of two peaks.

Note: peaks must be on the same chromosome; start coordinate is always less than end coordinate

Usage

```
calculate_midpoint_distance1d(peak1_start, peak1_end, peak2_start, peak2_end)
```

Arguments

```
peak1_start integer vector; genomic start coordinate(s) of peak in replicate 1
peak1_end integer vector; genomic end coordinate(s) of peak in replicate 1
peak2_start integer vector; genomic start coordinate(s) of peak in replicate 2
peak2_end integer vector; genomic end coordinate(s) of peak in replicate 2
```

Value

positive integer vector; distances between peak pairs

Examples

 ${\tt calculate_midpoint_distance2d}$

Distance between Anchor Midpoints of two Interactions

Description

Calculates the distance in nucleotides between the anchor midpoints of two interactions, which is the sum of the distance between midpoints of anchor A in interaction 1 and anchor A in interaction 2, and the distance between midpoints of anchor B in interaction 1 and anchor B in interaction 2.

Note: all anchors must be on the same chromosome; start coordinate is always less than end coordinate

Usage

```
calculate_midpoint_distance2d(
  int1_anchor_a_start,
  int1_anchor_a_end,
  int1_anchor_b_start,
  int1_anchor_b_end,
  int2_anchor_a_start,
  int2_anchor_a_end,
  int2_anchor_b_end,
  int2_anchor_b_end
)
```

Arguments

```
int1_anchor_a_start
                  integer vector; genomic start coordinate(s) of anchor A in replicate 1 interaction
int1_anchor_a_end
                  integer vector; genomic end coordinate(s) of anchor A in replicate 1 interaction
int1_anchor_b_start
                  integer vector; genomic start coordinate(s) of anchor B in replicate 1 interaction
int1_anchor_b_end
                  integer vector; genomic end coordinate(s) of anchor B in replicate 1 interaction
int2_anchor_a_start
                  integer vector; genomic start coordinate(s) of anchor A in replicate 2 interaction
int2_anchor_a_end
                  integer vector; genomic end coordinate(s) of anchor A in replicate 2 interaction
int2_anchor_b_start
                  integer vector; genomic start coordinate(s) of anchor B in replicate 2 interaction
int2_anchor_b_end
                  integer vector; genomic end coordinate(s) of anchor B in replicate 2 interaction
```

Value

positive integer vector; distances between interaction pairs

Examples

```
# identical, zero distance
calculate_midpoint_distance2d(100, 120, 240, 260,
                              100, 120, 240, 260)
# centered, zero distance
calculate_midpoint_distance2d(100, 120, 240, 260,
                              90, 130, 230, 270)
# off by 10 per anchor
calculate_midpoint_distance2d(100, 120, 240, 250,
                              110, 130, 230, 240)
# off by 10 (anchor B only)
calculate_midpoint_distance2d(100, 120, 240, 250,
                              90, 130, 250, 260)
# vectorized example
calculate_midpoint_distance2d(c(100, 100, 100, 100),
                              c(120, 120, 120, 120),
                              c(240, 240, 240, 240),
                              c(260, 260, 250, 250),
                              c(100, 90, 110, 90),
                              c(120, 130, 130, 130),
                              c(240, 230, 230, 250),
                              c(260, 270, 240, 260))
```

```
calculate_relative_overlap1d
```

Relative Anchor Overlap of two Peaks

Description

Calculates the overlap between anchor A of interaction 1 and anchor A of interaction 2, as well as anchor B of interaction 1 and anchor B of interaction 2. The overlap (in nucleotides) is then normalized by the length of the anchors.

Usage

```
calculate_relative_overlap1d(peak1_start, peak1_end, peak2_start, peak2_end)
```

Arguments

peak1_start	integer vector; genomic start coordinate(s) of peak in replicate 1
peak1_end	integer vector; genomic end coordinate(s) of peak in replicate 1
peak2_start	integer vector; genomic start coordinate(s) of peak in replicate 2
peak2_end	integer vector; genomic end coordinate(s) of peak in replicate 2

Value

numeric vector; relative overlaps between peak pairs

Examples

```
# 100% overlap
calculate_relative_overlap1d(100, 120,
                        100, 120)
# 50% overlap
calculate_relative_overlap1d(100, 120,
                         100, 110)
# negative overlap
calculate_relative_overlap1d(100, 120,
                         130, 140)
# larger negative overlap
calculate_relative_overlap1d(100, 120,
                         200, 220)
# vectorized example
calculate_relative_overlap1d(c(100, 100, 100, 100),
                         c(120, 120, 120, 120),
                         c(100, 100, 130, 200),
                         c(120, 110, 140, 220))
```

```
calculate_relative_overlap2d
```

Relative Anchor Overlap of two Interactions

Description

Calculates the overlap between anchor A of interaction 1 and anchor A of interaction 2, as well as anchor B of interaction 1 and anchor B of interaction 2. The overlap (in nucleotides) is then normalized by the length of the anchors.

Note: anchors A and B of the same interaction have to be on the same chromosome; start coordinate is always less than end coordinate

Usage

```
calculate_relative_overlap2d(
  int1_anchor_a_start,
  int1_anchor_a_end,
  int1_anchor_b_start,
  int1_anchor_b_end,
  int2_anchor_a_start,
  int2_anchor_a_end,
  int2_anchor_b_end,
  int2_anchor_b_end)
```

Arguments

```
int1_anchor_a_start
                  integer vector; genomic start coordinate(s) of anchor A in replicate 1 interaction
int1_anchor_a_end
                  integer vector; genomic end coordinate(s) of anchor A in replicate 1 interaction
int1_anchor_b_start
                  integer vector; genomic start coordinate(s) of anchor B in replicate 1 interaction
int1_anchor_b_end
                  integer vector; genomic end coordinate(s) of anchor B in replicate 1 interaction
int2_anchor_a_start
                  integer vector; genomic start coordinate(s) of anchor A in replicate 2 interaction
int2_anchor_a_end
                  integer vector; genomic end coordinate(s) of anchor A in replicate 2 interaction
int2_anchor_b_start
                  integer vector; genomic start coordinate(s) of anchor B in replicate 2 interaction
int2_anchor_b_end
                  integer vector; genomic end coordinate(s) of anchor B in replicate 2 interaction
```

Value

numeric vector; relative overlaps between interaction pairs

chiapet 7

Examples

```
# 100% overlap
calculate_relative_overlap2d(100, 120, 240, 260,
                             100, 120, 240, 260)
# 50% overlap
calculate_relative_overlap2d(100, 120, 240, 250,
                             100, 110, 240, 260)
# negative overlap
calculate_relative_overlap2d(100, 120, 240, 250,
                             130, 140, 260, 280)
# larger negative overlap
calculate_relative_overlap2d(100, 120, 240, 250,
                             200, 220, 340, 350)
# vectorized example
calculate_relative_overlap2d(c(100, 100, 100, 100),
                             c(120, 120, 120, 120),
                             c(240, 240, 240, 240),
                             c(260, 250, 250, 250),
                              c(100, 100, 130, 200),
                             c(120, 110, 140, 220),
                              c(240, 240, 260, 340),
                              c(260, 260, 280, 350))
```

chiapet

Example Genomic Interaction Data Set

Description

This object contains genomic interactions on chromosomes 1 to 5, which could be the results of Hi-C or ChIA-PET experiments, done in duplicates.

Usage

chiapet

Format

A list with two components, the data frames rep1_df and rep2_df, which have the following seven columns:

```
column 1:
                       character; genomic location of anchor A - chromosome (e.g., "chr3")
            chr_a
column 2:
           start_a
                       integer; genomic location of anchor A - start coordinate
column 3:
            end_a
                       integer; genomic location of anchor A - end coordinate
column 4:
           chr_b
                       character; genomic location of anchor B - chromosome (e.g., "chr3")
column 5:
           start_b
                       integer; genomic location of anchor B - start coordinate
                       integer; genomic location of anchor B - end coordinate
column 6:
            end_b
column 7:
            fdr
                       numeric; False Discovery Rate - significance of interaction
```

chipseq	Example Genomic Peak Data Set	

Description

This object contains genomic peaks from two replicate ChIP-seq experiments.

Usage

chipseq

Format

A list with two components, the data frames rep1_df and rep2_df, which have the following four columns:

```
column 1: chr character; genomic location of peak - chromosome (e.g., "chr3")
column 2: start integer; genomic location of peak - start coordinate
column 3: end integer; genomic location of peak - end coordinate
column 4: value numeric; heuristic used to rank the peaks
```

```
determine_anchor_overlap
```

Identifies Overlapping Anchors

Description

Identifies all overlapping anchor pairs (m:n mapping).

Usage

```
determine_anchor_overlap(rep1_anchor, rep2_anchor, max_gap = -1L)
```

Arguments

```
rep1_anchor
                     data frame with the following columns:
column 1:
            chr
                     character; genomic location of anchor in replicate 1 - chromosome (e.g., "chr3")
column 2:
            start
                     integer; genomic location of anchor in replicate 1 - start coordinate
column 3:
                      integer; genomic location of anchor in replicate 1 - end coordinate
  rep2_anchor
                     data frame with the following columns:
                      character; genomic location of anchor in replicate 2 - chromosome (e.g., "chr3")
column 1:
            chr
column 2:
                     integer; genomic location of anchor in replicate 2 - start coordinate
            start
column 3:
            end
                      integer; genomic location of anchor in replicate 2 - end coordinate
                     integer; maximum gap in nucleotides allowed between two anchors for them to
  max_gap
                     be considered as overlapping (defaults to -1, i.e., overlapping anchors)
```

draw_hic_contact_map

Value

A data frame containing overlapping anchor pairs with the following columns:

```
column 1: rep1_idx anchor index in data frame rep1_anchor column 2: rep2_idx anchor index in data frame rep2_anchor
```

9

Examples

Description

Creates Hi-C contact maps to visualize the results of estimate_idr2d_hic.

Usage

```
draw_hic_contact_map(
   df,
   idr_cutoff = NULL,
   chromosome = NULL,
   start_coordinate = NULL,
   end_coordinate = NULL,
   title = NULL,
   values_normalized = FALSE,
   log_values = TRUE
)
```

Arguments

df output of estimate_idr2d_hic, a data frame with the following columns:

```
column 1:
            interaction
                             character; genomic location of interaction block (e.g., "chr1:204940000-204940000")
column 2:
            value
                             numeric; p-value, FDR, or heuristic used to rank the interactions
column 3:
            "rep_value"
                             numeric; value of corresponding replicate interaction
column 4:
            "rank"
                             integer; rank of the interaction, established by value column, ascending order
             "rep_rank"
column 5:
                             integer; rank of corresponding replicate interaction
column 6:
            "idr"
                             integer; IDR of the block and the corresponding block in the other replicate
```

idr_cutoff numeric; only show blocks with IDR < idr_cutoff, shows all blocks by default chromosome character; chromsome name or list of chromosome names to be analyzed, e.g., UCSC chromosome 1, "chr1", defaults to all chromosomes (chromosome = NULL) start coordinate integer; only show contact map window between "start_coordinate" and "end_coordinate", by default shows entire chromosome end_coordinate integer; only show contact map window between "start_coordinate" and "end_coordinate", by default shows entire chromosome title character; plot title logical; are read counts in value column raw or normalized? Defaults to FALSE

values_normalized

log_values logical; log-transform value column? Defaults to TRUE

Value

```
ggplot2 object; Hi-C contact map
```

Examples

```
idr_results_df <- estimate_idr2d_hic(idr2d:::hic$rep1_df,</pre>
                                       idr2d:::hic$rep2_df)
draw_hic_contact_map(idr_results_df, idr_cutoff = 0.05, chromosome = "chr1")
```

```
draw_idr_distribution_histogram
```

Create histogram of IDR values

Description

Creates diagnostic plots to visualize the results of estimate_idr.

Usage

```
draw_idr_distribution_histogram(
 df,
  remove_na = TRUE,
 xlab = "IDR",
 ylab = "density",
  title = "IDR value distribution"
)
```

Arguments

df part of output of estimate_idr, a data frame with at least the following named columns:

IDR of the peak and the corresponding peak in the other replicate.

```
remove_na logical; should NA values be removed?

xlab character; x axis label

ylab character; y axis label

title character; plot title
```

Value

```
ggplot2 object; IDR distribution histogram
```

Examples

```
draw_rank_idr_scatterplot
```

Create scatterplot of IDR values

Description

Creates diagnostic plots to visualize the results of estimate_idr.

Usage

```
draw_rank_idr_scatterplot(
   df,
   remove_na = TRUE,
   xlab = "rank in replicate 1",
   ylab = "rank in replicate 2",
   log_idr = FALSE,
   title = "rank - IDR dependence",
   color_gradient = c("rainbow", "default"),
   alpha = 1,
   max_points_shown = 2500
)
```

Arguments

```
part of output of estimate_idr, a data frame with at least the following named columns:

rank integer; rank of the peak, established by value column, ascending order integer; rank of corresponding replicate peak.

idr IDR of the peak and the corresponding peak in the other replicate.

remove_na logical; should NA values be removed?

xlab character; x axis label
```

```
ylab character; y axis label

log_idr logical; use logarithmized IDRs for colors to better distinguish highly significant IDRs

title character; plot title

color_gradient character; either "rainbow" or "default"

alpha numeric; transparency of dots, from 0.0 - 1.0, where 1.0 is completely opaque; default is 1.0

max_points_shown integer; default is 2500
```

Value

```
ggplot2 object; IDR rank scatterplot
```

Examples

```
draw_value_idr_scatterplot

Create scatterplot of IDR values
```

Description

Creates diagnostic plots to visualize the results of estimate_idr.

Usage

```
draw_value_idr_scatterplot(
    df,
    remove_na = TRUE,
    remove_outliers = TRUE,
    xlab = "transformed value in replicate 1",
    ylab = "transformed value in replicate 2",
    log_axes = FALSE,
    log_idr = FALSE,
    title = "value - IDR dependence",
    color_gradient = c("rainbow", "default"),
    alpha = 1,
    max_points_shown = 2500
)
```

establish_bijection 13

Arguments

df part of output of estimate_idr, a data frame with at least the following named

columns:

value numeric; p-value, FDR, or heuristic used to rank the peaks

rep_value numeric; value of corresponding replicate peak

idr IDR of the peak and the corresponding peak in the other replicate.

remove_na logical; should NA values be removed?

remove_outliers

logical; removes extreme data points

xlab character; x axis label ylab character; y axis label

log_axes logical; show logarithmized values from replicate 1 and 2 (default value is

FALSE)

log_idr logical; use logarithmized IDRs for colors to better distinguish highly significant

IDRs (default value is FALSE)

title character; plot title

color_gradient character; either "rainbow" or "default"

alpha numeric; transparency of dots, from 0.0 - 1.0, where 1.0 is completely opaque;

default is 1.0

max_points_shown

integer; default is 2500

Value

```
ggplot2 object; IDR value scatterplot
```

Examples

establish_bijection Finds One-to-One Correspondence between Peaks or interactions from Replicate 1 and 2

Description

This method establishes a bijective assignment between observations (genomic peaks in case of ChIP-seq, genomic interactions in case of ChIA-PET, HiChIP, and Hi-C) from replicate 1 and 2. An observation in replicate 1 is assigned to an observation in replicate 2 if and only if (1) the observation loci in both replicates overlap (or the gap between them is less than or equal to max_gap), and (2) there is no other observation in replicate 2 that overlaps with the observation in replicate 1 and has a lower *ambiguity resolution value*.

Usage

```
establish_bijection(
  rep1_df,
  rep2_df,
  analysis_type = c("IDR1D", "IDR2D"),
  ambiguity_resolution_method = c("overlap", "midpoint", "value"),
  max_gap = -1L
)
```

Arguments

rep1_df data frame of observations (i.e., genomic peaks or genomic interactions) of replicate 1. If analysis_type is IDR1D, the columns of rep1_df are described in establish_bijection1d, otherwise in establish_bijection2d rep2_df data frame of observations (i.e., genomic peaks or genomic interactions) of replicate 2. Same columns as rep1_df. analysis_type "IDR2D" for genomic interaction data sets, "IDR1D" for genomic peak data sets ambiguity_resolution_method defines how ambiguous assignments (when one interaction or peak in replicate 1 overlaps with multiple interactions or peaks in replicate 2 or vice versa) are resolved. For available methods, see establish_overlap1d or establish_overlap2d, respectively. integer; maximum gap in nucleotides allowed between two anchors for them to max_gap be considered as overlapping (defaults to -1, i.e., overlapping anchors)

Value

See establish_bijection1d or establish_bijection2d, respectively.

Examples

```
rep1_df <- idr2d:::chipseq$rep1_df
rep1_df$value <- preprocess(rep1_df$value, "log")

rep2_df <- idr2d:::chipseq$rep2_df
rep2_df$value <- preprocess(rep2_df$value, "log")

mapping <- establish_bijection(rep1_df, rep2_df, analysis_type = "IDR1D")</pre>
```

establish_bijection1d Finds One-to-One Correspondence between Peaks from Replicate 1 and 2

Description

This method establishes a bijective assignment between peaks from replicate 1 and 2. A peak in replicate 1 is assigned to a peak in replicate 2 if and only if (1) they overlap (or the gap between the peaks is less than or equal to max_gap), and (2) there is no other peak in replicate 2 that overlaps with the peak in replicate 1 and has a lower *ambiguity resolution value*.

establish_bijection1d 15

Usage

```
establish_bijection1d(
  rep1_df,
  rep2_df,
  ambiguity_resolution_method = c("overlap", "midpoint", "value"),
  max_gap = -1L
)
```

Arguments

rep1_df data frame of observations (i.e., genomic peaks) of replicate 1, with at least the following columns (position of columns matter, column names are irrelevant):

column 1: chr character; genomic location of peak - chromosome (e.g., "chr3") column 2: start integer; genomic location of peak - start coordinate

column 3: end integer; genomic location of peak - end coordinate

column 4: value numeric; p-value, FDR, or heuristic used to rank the interactions

rep2_df data frame of observations (i.e., genomic peaks) of replicate 2, with the following columns (position of columns matter, column names are irrelevant):

column 1: chr character; genomic location of peak - chromosome (e.g., "chr3")

column 2: start integer; genomic location of peak - start coordinate column 3: end integer; genomic location of peak - end coordinate

column 4: value numeric; p-value, FDR, or heuristic used to rank the interactions

ambiguity_resolution_method

defines how ambiguous assignments (when one interaction in replicate 1 overlaps with multiple interactions in replicate 2 or vice versa) are resolved. Available methods:

the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate

the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance

able methods:

"value" interactions are prioritized by ascending or descending value column (see sorting_direction), e.g., if to

max_gap integer; maximum gap in nucleotides allowed between two anchors for them to

be considered as overlapping (defaults to -1, i.e., overlapping anchors)

Value

"overlap"

"midpoint"

Data frames rep1_df and rep2_df with the following columns:

```
column 1:
            chr
                          character; genomic location of peak - chromosome (e.g., "chr3")
column 2:
            start
                          integer; genomic location of peak - start coordinate
column 3:
                          integer; genomic location of peak - end coordinate
            end
column 4:
                          numeric; p-value, FDR, or heuristic used to rank the peaks
            value
column 5:
            rep_value
                          numeric; value of corresponding replicate peak. If no corresponding peak was found, rep_val
column 6:
            rank
                          integer; rank of the peak, established by value column, ascending order
                          integer; rank of corresponding replicate peak. If no corresponding peak was found, rep_rank i
column 7:
            rep_rank
column 8:
                          integer; peak index, primary key
            idx
```

column 9: rep_idx integer; specifies the index of the corresponding peak in the other replicate (foreign key). If no

establish_bijection2d

Examples

16

```
rep1_df <- idr2d:::chipseq$rep1_df
rep1_df$value <- preprocess(rep1_df$value, "log")
rep2_df <- idr2d:::chipseq$rep2_df
rep2_df$value <- preprocess(rep2_df$value, "log")
mapping <- establish_bijection1d(rep1_df, rep2_df)</pre>
```

establish_bijection2d Finds One-to-One Correspondence between Interactions from Replicate 1 and 2

Description

This method establishes a bijective assignment between interactions from replicate 1 and 2. An interaction in replicate 1 is assigned to an interaction in replicate 2 if and only if (1) both anchors of the interactions overlap (or the gap between anchor A/B in replicate 1 and 2 is less than or equal to max_gap), and (2) there is no other interaction in replicate 2 that overlaps with the interaction in replicate 1 and has a lower *ambiguity resolution value*.

Usage

```
establish_bijection2d(
  rep1_df,
  rep2_df,
  ambiguity_resolution_method = c("overlap", "midpoint", "value"),
  max_gap = -1L
)
```

Arguments

column 3:

end_a

rep1_df data frame of observations (i.e., genomic interactions) of replicate 1, with at least the following columns (position of columns matter, column names are irrelevant):

```
column 1:
            chr_a
                       character; genomic location of anchor A - chromosome (e.g., "chr3")
column 2:
                       integer; genomic location of anchor A - start coordinate
            start_a
column 3:
            end_a
                       integer; genomic location of anchor A - end coordinate
                       character; genomic location of anchor B - chromosome (e.g., "chr3")
column 4:
            chr b
column 5:
                       integer; genomic location of anchor B - start coordinate
            start_b
column 6:
            end_b
                        integer; genomic location of anchor B - end coordinate
column 7:
            value
                        numeric; p-value, FDR, or heuristic used to rank the interactions
                  data frame of observations (i.e., genomic interactions) of replicate 2, with the
rep2_df
                  following columns (position of columns matter, column names are irrelevant):
column 1:
                        character; genomic location of anchor A - chromosome (e.g., "chr3")
            chr_a
                       integer; genomic location of anchor A - start coordinate
column 2:
            start_a
```

integer; genomic location of anchor A - end coordinate

establish_overlap1d 17

```
column 4:
                                                  chr_b
                                                                                                  character; genomic location of anchor B - chromosome (e.g., "chr3")
column 5:
                                                   start_b
                                                                                                 integer; genomic location of anchor B - start coordinate
column 6:
                                                   end_b
                                                                                                  integer; genomic location of anchor B - end coordinate
column 7:
                                                                                                  numeric; p-value, FDR, or heuristic used to rank the interactions
                                                  value
ambiguity_resolution_method
                                                                             defines how ambiguous assignments (when one interaction in replicate 1 over-
                                                                             laps with multiple interactions in replicate 2 or vice versa) are resolved. Avail-
                                                                            able methods:
     "value"
                                                   interactions are prioritized by ascending or descending value column (see sorting_direction), e.g., if the solution of the sol
```

"overlap" the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance

max_gap integer; maximum gap in nucleotides allowed between two anchors for them to be considered as overlapping (defaults to -1, i.e., overlapping anchors)

Value

Data frames rep1_df and rep2_df with the following columns:

```
column 1:
                              character; genomic location of anchor A - chromosome (e.g., "chr3")
             chr a
 column 2:
                              integer; genomic location of anchor A - start coordinate
             start_a
 column 3:
             end_a
                              integer; genomic location of anchor A - end coordinate
 column 4:
             chr_b
                              character; genomic location of anchor B - chromosome (e.g., "chr3")
 column 5: start_b
                              integer; genomic location of anchor B - start coordinate
 column 6: end_b
                              integer; genomic location of anchor B - end coordinate
 column 7:
             value
                              numeric; p-value, FDR, or heuristic used to rank the interactions
 column 8:
             "rep_value"
                              numeric; value of corresponding replicate interaction. If no corresponding interaction was
 column 9:
             "rank"
                              integer; rank of the interaction, established by value column, ascending order
              "rep_rank"
column 10:
                              integer; rank of corresponding replicate interaction. If no corresponding interaction was for
              "idx"
column 11:
                              integer; interaction index, primary key
column 12:
                              integer; specifies the index of the corresponding interaction in the other replicate (foreign k
              "rep_idx"
```

Examples

```
rep1_df <- idr2d:::chiapet$rep1_df
rep1_df$fdr <- preprocess(rep1_df$fdr, "log_additive_inverse")
rep2_df <- idr2d:::chiapet$rep2_df
rep2_df$fdr <- preprocess(rep2_df$fdr, "log_additive_inverse")
mapping <- establish_bijection2d(rep1_df, rep2_df)</pre>
```

18 establish_overlap1d

Description

This method returns all overlapping interactions between two replicates. For each pair of overlapping interactions, the *ambiguity resolution value* (ARV) is calculated, which helps to reduce the m:n mapping to a 1:1 mapping. The semantics of the ARV depend on the specified ambiguity_resolution_method, but in general interaction pairs with lower ARVs have priority over interaction pairs with higher ARVs when the bijective mapping is established.

Usage

```
establish_overlap1d(
  rep1_df,
  rep2_df,
  ambiguity_resolution_method = c("overlap", "midpoint", "value"),
  max_gap = -1L
)
```

Arguments

```
rep1_df
                  data frame of observations (i.e., genomic peaks) of replicate 1, with at least the
                  following columns (position of columns matter, column names are irrelevant):
   column 1:
                chr
                         character; genomic location of peak - chromosome (e.g., "chr3")
   column 2:
                start
                         integer; genomic location of peak - start coordinate
   column 3:
                end
                         integer; genomic location of peak - end coordinate
   column 4:
                value
                         numeric; p-value, FDR, or heuristic used to rank the interactions
rep2_df
                  data frame of observations (i.e., genomic peaks) of replicate 2, with the follow-
                  ing columns (position of columns matter, column names are irrelevant):
   column 1:
                chr
                         character; genomic location of peak - chromosome (e.g., "chr3")
   column 2:
                start
                         integer; genomic location of peak - start coordinate
```

integer; genomic location of peak - end coordinate

ambiguity_resolution_method

end

value

column 3:

column 4:

defines how ambiguous assignments (when one interaction in replicate 1 overlaps with multiple interactions in replicate 2 or vice versa) are resolved. Available methods:

numeric; p-value, FDR, or heuristic used to rank the interactions

"value" interactions are prioritized by ascending or descending value column (see sorting_direction), e.g., if to the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate "midpoint" the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance

max_gap integer; maximum gap in nucleotides allowed between two anchors for them to be considered as overlapping (defaults to -1, i.e., overlapping anchors)

Value

data frame with the following columns:

```
column 1: rep1_idx index of interaction in replicate 1 (i.e., row index in rep1_df) column 2: rep2_idx index of interaction in replicate 2 (i.e., row index in rep2_df)
```

establish_overlap2d 19

column 3: arv

ambiguity resolution value used turn m:n mapping into 1:1 mapping. Interaction pairs with lower

Examples

```
rep1_df <- idr2d:::chipseq$rep1_df
rep1_df$value <- preprocess(rep1_df$value, "log_additive_inverse")
rep2_df <- idr2d:::chipseq$rep2_df
rep2_df$value <- preprocess(rep2_df$value, "log_additive_inverse")
# shuffle to break preexisting order
rep1_df <- rep1_df[sample.int(nrow(rep1_df)), ]
rep2_df <- rep2_df[sample.int(nrow(rep2_df)), ]
# sort by value column
rep1_df <- dplyr::arrange(rep1_df, value)
rep2_df <- dplyr::arrange(rep2_df, value)
pairs_df <- establish_overlap1d(rep1_df, rep2_df)</pre>
```

establish_overlap2d

Establish m:n mapping between interactions from replicate 1 and 2

Description

This method returns all overlapping interactions between two replicates. For each pair of overlapping interactions, the *ambiguity resolution value* (ARV) is calculated, which helps to reduce the m:n mapping to a 1:1 mapping. The semantics of the ARV depend on the specified ambiguity_resolution_method, but in general interaction pairs with lower ARVs have priority over interaction pairs with higher ARVs when the bijective mapping is established.

Usage

```
establish_overlap2d(
  rep1_df,
  rep2_df,
  ambiguity_resolution_method = c("overlap", "midpoint", "value"),
  max_gap = -1L
)
```

Arguments

```
rep1_df data frame of observations (i.e., genomic interactions) of replicate 1, with at least the following columns (position of columns matter, column names are irrelevant):
```

```
column 1: chr_a character; genomic location of anchor A - chromosome (e.g., "chr3")
column 2: start_a column 3: end_a integer; genomic location of anchor A - end coordinate
column 4: chr_b character; genomic location of anchor B - chromosome (e.g., "chr3")
column 5: start_b integer; genomic location of anchor B - start coordinate
```

20 establish_overlap2d

integer; genomic location of anchor B - end coordinate

```
column 7:
              value
                          numeric; p-value, FDR, or heuristic used to rank the interactions
  rep2_df
                     data frame of observations (i.e., genomic interactions) of replicate 2, with the
                     following columns (position of columns matter, column names are irrelevant):
  column 1:
              chr_a
                          character; genomic location of anchor A - chromosome (e.g., "chr3")
  column 2:
                          integer; genomic location of anchor A - start coordinate
              start_a
  column 3:
              end_a
                          integer; genomic location of anchor A - end coordinate
  column 4:
              chr_b
                          character; genomic location of anchor B - chromosome (e.g., "chr3")
  column 5:
               start_b
                          integer; genomic location of anchor B - start coordinate
  column 6:
               end_b
                          integer; genomic location of anchor B - end coordinate
  column 7:
              value
                          numeric; p-value, FDR, or heuristic used to rank the interactions
  ambiguity_resolution_method
                     defines how ambiguous assignments (when one interaction in replicate 1 over-
                     laps with multiple interactions in replicate 2 or vice versa) are resolved. Avail-
                     able methods:
   "value"
               interactions are prioritized by ascending or descending value column (see sorting_direction), e.g., if to
 "overlap"
               the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate
"midpoint"
              the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance
```

Value

max_gap

column 6:

end b

data frame with the following columns:

```
column 1: rep1_idx index of interaction in replicate 1 (i.e., row index in rep1_df)
column 2: rep2_idx index of interaction in replicate 2 (i.e., row index in rep2_df)
column 3: arv ambiguity resolution value used turn m:n mapping into 1:1 mapping. Interaction pairs with lower
```

integer; maximum gap in nucleotides allowed between two anchors for them to

be considered as overlapping (defaults to -1, i.e., overlapping anchors)

Examples

```
rep1_df <- idr2d:::chiapet$rep1_df
rep1_df$fdr <- preprocess(rep1_df$fdr, "log_additive_inverse")
rep2_df <- idr2d:::chiapet$rep2_df
rep2_df$fdr <- preprocess(rep2_df$fdr, "log_additive_inverse")
# shuffle to break preexisting order
rep1_df <- rep1_df[sample.int(nrow(rep1_df)), ]
rep2_df <- rep2_df[sample.int(nrow(rep2_df)), ]
# sort by value column
rep1_df <- dplyr::arrange(rep1_df, rep1_df$fdr)
rep2_df <- dplyr::arrange(rep2_df, rep2_df$fdr)
pairs_df <- establish_overlap2d(rep1_df, rep2_df)</pre>
```

estimate_idr 21

estimate_idr

Estimates IDR for Genomic Peaks or Genomic Interactions

Description

Estimates IDR for Genomic Peaks or Genomic Interactions

Usage

```
estimate_idr(
  rep1_df,
  rep2_df,
  analysis_type = "IDR2D",
 value_transformation = c("identity", "additive_inverse", "multiplicative_inverse",
    "log", "log_additive_inverse"),
  ambiguity_resolution_method = c("overlap", "midpoint", "value"),
  remove_nonstandard_chromosomes = TRUE,
 max_factor = 1.5,
  jitter_factor = 1e-04,
 max_gap = -1L,
 mu = 0.1,
  sigma = 1,
  rho = 0.2,
  p = 0.5,
  eps = 0.001,
 max_iteration = 30,
  local_idr = TRUE
```

Arguments

rep1_df

data frame of observations (i.e., genomic peaks or genomic interactions) of replicate 1. If analysis_type is IDR1D, the columns of rep1_df are described in establish_bijection1d, otherwise in establish_bijection2d

rep2_df

data frame of observations (i.e., genomic peaks or genomic interactions) of replicate 2. Same columns as rep1_df.

 $analysis_type \quad "IDR2D" \ for genomic interaction \ data \ sets, "IDR1D" \ for genomic \ peak \ data \ sets \\ value_transformation$

the values in x have to be transformed in a way such that when ordered in descending order, more significant interactions end up on top of the list. If the values in x are p-values, "log_additive_inverse" is recommended. The following transformations are supported:

either "ascending" (more significant interactions have lower value in value column) or "descending" (more significant interactions have higher value in value column)

22 estimate_idr1d

ambiguity_resolution_method

defines how ambiguous assignments (when one interaction or peak in replicate 1 overlaps with multiple interactions or peaks in replicate 2 or vice versa) are resolved. For available methods, see establish_overlap1d or establish_overlap2d, respectively.

remove_nonstandard_chromosomes

removes peaks and interactions containing genomic locations on non-standard

chromosomes using keepStandardChromosomes (default is TRUE)

max_factor numeric; controls the replacement values for Inf and -Inf. Inf are replaced by

 $max(x) * max_factor and -Inf are replaced by min(x) / max_factor.$

jitter_factor numeric; controls the magnitude of the noise that is added to x. This is done to

break ties in x. Set jitter_factor = NULL for no jitter.

max_gap integer; maximum gap in nucleotides allowed between two anchors for them to

be considered as overlapping (defaults to -1, i.e., overlapping anchors)

mu a starting value for the mean of the reproducible component.

sigma a starting value for the standard deviation of the reproducible component.

rho a starting value for the correlation coefficient of the reproducible component.

p a starting value for the proportion of reproducible component.

eps Stopping criterion. Iterations stop when the increment of log-likelihood is <

eps*log-likelihood, Default=0.001.

max_iteration integer; maximum number of iterations for IDR estimation (defaults to 30)

local_idr see est.IDR

Value

See estimate_idr1d or estimate_idr2d, respectively.

References

Q. Li, J. B. Brown, H. Huang and P. J. Bickel. (2011) Measuring reproducibility of high-throughput experiments. Annals of Applied Statistics, Vol. 5, No. 3, 1752-1779.

Examples

estimate_idr1d

Estimates IDR for Genomic Peak Data

Description

This method estimates Irreproducible Discovery Rates (IDR) for peaks in replicated ChIP-seq experiments.

estimate_idr1d 23

Usage

```
estimate_idr1d(
  rep1_df,
  rep2_df,
 value_transformation = c("identity", "additive_inverse", "multiplicative_inverse",
    "log", "log_additive_inverse"),
  ambiguity_resolution_method = c("overlap", "midpoint", "value"),
  remove_nonstandard_chromosomes = TRUE,
 max_factor = 1.5,
  jitter_factor = 1e-04,
 max_gap = -1L,
 mu = 0.1,
 sigma = 1,
  rho = 0.2,
  p = 0.5,
  eps = 0.001,
 max_iteration = 30,
  local_idr = TRUE
)
```

Arguments

rep1_df data frame of observations (i.e., genomic peaks) of replicate 1, with at least the following columns (position of columns matter, column names are irrelevant):

```
column 1: chr character; genomic location of peak - chromosome (e.g., "chr3")
column 2: start integer; genomic location of peak - start coordinate
column 3: end integer; genomic location of peak - end coordinate
column 4: value numeric; p-value, FDR, or heuristic used to rank the interactions
```

rep2_df data frame of observations (i.e., genomic peaks) of replicate 2, with the following columns (position of columns matter, column names are irrelevant):

```
column 1: chr character; genomic location of peak - chromosome (e.g., "chr3")
column 2: start integer; genomic location of peak - start coordinate
column 3: end integer; genomic location of peak - end coordinate
column 4: value numeric; p-value, FDR, or heuristic used to rank the interactions
```

value_transformation

the values in x have to be transformed in a way such that when ordered in descending order, more significant interactions end up on top of the list. If the values in x are p-values, "log_additive_inverse" is recommended. The following transformations are supported:

```
"identity" no transformation is performed on x

"additive_inverse" x. = -x

"multiplicative_inverse" x. = 1 / x

"log" x. = \log(x). Note: zeros are replaced by .Machine$double.xmin

"log_additive_inverse" x. = -\log(x), recommended if x are p-values. Note: zeros are replaced by .Machine$double.xmin
```

either "ascending" (more significant interactions have lower value in value column) or "descending" (more significant interactions have higher value in value column)

24 estimate_idr1d

ambiguity_resolution_method

defines how ambiguous assignments (when one interaction in replicate 1 overlaps with multiple interactions in replicate 2 or vice versa) are resolved. Available methods:

"value" interactions are prioritized by ascending or descending value column (see sorting_direction), e.g., if to the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance

remove_nonstandard_chromosomes

removes peaks containing genomic locations on non-standard chromosomes us-

ing keepStandardChromosomes (default is TRUE)

max_factor numeric; controls the replacement values for Inf and -Inf. Inf are replaced by

 $max(x) * max_factor and -Inf are replaced by min(x) / max_factor.$

jitter_factor numeric; controls the magnitude of the noise that is added to x. This is done to

break ties in x. Set jitter_factor = NULL for no jitter.

max_gap integer; maximum gap in nucleotides allowed between two anchors for them to

be considered as overlapping (defaults to -1, i.e., overlapping anchors)

mu a starting value for the mean of the reproducible component.

sigma a starting value for the standard deviation of the reproducible component.

rho a starting value for the correlation coefficient of the reproducible component.

p a starting value for the proportion of reproducible component.

eps Stopping criterion. Iterations stop when the increment of log-likelihood is <

eps*log-likelihood, Default=0.001.

max_iteration integer; maximum number of iterations for IDR estimation (defaults to 30)

local_idr see est.IDR

chr

idr

start

Value

column 1:

column 2:

List with three components, (rep1_df, rep2_df, and analysis_type) containing the interactions from input data frames rep1_df and rep2_df with the following additional columns:

integer; genomic location of peak - start coordinate

column 3:	end	integer; genomic location of peak - end coordinate
column 4:	value	numeric; p-value, FDR, or heuristic used to rank the peaks
column 5:	rep_value	numeric; value of corresponding replicate peak. If no corresponding peak was found, rep_va
column 6:	rank	integer; rank of the peak, established by value column, ascending order
column 7:	rep_rank	integer; rank of corresponding replicate peak. If no corresponding peak was found, rep_rank
column 8:	idx	integer; peak index, primary key
column 9:	rep_idx	integer; specifies the index of the corresponding peak in the other replicate (foreign key). If no

IDR of the peak and the corresponding peak in the other replicate. If no corresponding peak v

character; genomic location of peak - chromosome (e.g., "chr3")

References

column 10:

Q. Li, J. B. Brown, H. Huang and P. J. Bickel. (2011) Measuring reproducibility of high-throughput experiments. Annals of Applied Statistics, Vol. 5, No. 3, 1752-1779.

estimate_idr2d 25

Examples

estimate_idr2d

Estimates IDR for Genomic Interaction Data

Description

This method estimates Irreproducible Discovery Rates (IDR) between two replicates of experiments identifying genomic interactions, such as Hi-C, ChIA-PET, and HiChIP.

Usage

```
estimate_idr2d(
  rep1_df,
  rep2_df,
 value_transformation = c("identity", "additive_inverse", "multiplicative_inverse",
    "log", "log_additive_inverse"),
  ambiguity_resolution_method = c("overlap", "midpoint", "value"),
  remove_nonstandard_chromosomes = TRUE,
 max_factor = 1.5,
  jitter_factor = 1e-04,
 max_gap = -1L,
 mu = 0.1,
 sigma = 1,
  rho = 0.2,
 p = 0.5,
 eps = 0.001,
 max_iteration = 30,
 local_idr = TRUE
)
```

Arguments

rep1_df data frame of observations (i.e., genomic interactions) of replicate 1, with at least the following columns (position of columns matter, column names are irrelevant):

```
column 1:
                       character; genomic location of anchor A - chromosome (e.g., "chr3")
            chr_a
column 2:
            start_a
                       integer; genomic location of anchor A - start coordinate
column 3:
                        integer; genomic location of anchor A - end coordinate
            end_a
column 4:
                       character; genomic location of anchor B - chromosome (e.g., "chr3")
            chr_b
column 5:
            start_b
                       integer; genomic location of anchor B - start coordinate
                        integer; genomic location of anchor B - end coordinate
column 6:
            end_b
column 7:
            value
                        numeric; p-value, FDR, or heuristic used to rank the interactions
```

26 estimate_idr2d

rep2_df data frame of observations (i.e., genomic interactions) of replicate 2, with the following columns (position of columns matter, column names are irrelevant): column 1: chr_a character; genomic location of anchor A - chromosome (e.g., "chr3") column 2: start a integer; genomic location of anchor A - start coordinate integer; genomic location of anchor A - end coordinate column 3: end_a column 4: chr_b character; genomic location of anchor B - chromosome (e.g., "chr3") column 5: integer; genomic location of anchor B - start coordinate start_b column 6: end_b integer; genomic location of anchor B - end coordinate column 7: value numeric; p-value, FDR, or heuristic used to rank the interactions

value_transformation

the values in x have to be transformed in a way such that when ordered in descending order, more significant interactions end up on top of the list. If the values in x are p-values, "log_additive_inverse" is recommended. The following transformations are supported:

either "ascending" (more significant interactions have lower value in value column) or "descending" (more significant interactions have higher value in value column)

ambiguity_resolution_method

defines how ambiguous assignments (when one interaction in replicate 1 overlaps with multiple interactions in replicate 2 or vice versa) are resolved. Available methods:

"value" interactions are prioritized by ascending or descending value column (see sorting_direction), e.g., if to "overlap" the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate "midpoint" the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance

remove_nonstandard_chromosomes

max_factor

max_gap

mu

rho

sigma

jitter_factor

removes interactions containing genomic locations on non-standard chromosomes using keepStandardChromosomes (default is TRUE)

numeric; controls the replacement values for Inf and -Inf. Inf are replaced by max(x) * max_factor and -Inf are replaced by min(x) / max_factor.

numeric; controls the magnitude of the noise that is added to x. This is done to break ties in x. Set jitter_factor = NULL for no jitter.

integer; maximum gap in nucleotides allowed between two anchors for them to be considered as overlapping (defaults to -1, i.e., overlapping anchors)

a starting value for the mean of the reproducible component.

a starting value for the correlation coefficient of the reproducible component.

a starting value for the proportion of reproducible component.

p a starting value for the proportion of reproducible component.

eps Stopping criterion. Iterations stop when the increment of log-likelihood is < eps*log-likelihood, Default=0.001.

max_iteration integer; maximum number of iterations for IDR estimation (defaults to 30)

local_idr see est.IDR

estimate_idr2d_hic 27

Value

column 1:

chr_a

List with three components, (rep1_df, rep2_df, and analysis_type) containing the interactions from input data frames rep1_df and rep2_df with the following additional columns:

```
column 2:
           start_a
 column 3: end a
 column 4: chr_b
 column 5: start_b
 column 6:
           end_b
 column 7: value
 column 8:
           "rep_value"
 column 9:
           "rank"
column 10:
           "rep_rank"
           "idx"
column 11:
            "rep_idx"
column 12:
```

idr IDR of the interaction and the corresponding interaction in the other replicate. If no corresponding interaction

References

Q. Li, J. B. Brown, H. Huang and P. J. Bickel. (2011) Measuring reproducibility of high-throughput experiments. Annals of Applied Statistics, Vol. 5, No. 3, 1752-1779.

Examples

Description

This method estimates Irreproducible Discovery Rates (IDR) of genomic interactions between two replicates of Hi-C experiments.

Before calling this method, call Juicer .hic contact matrix c

The contact matrix is subdivided into blocks, where the block size is determined by resolution. The reads per block are used to rank blocks and replicate blocks are easily matched by genomic location.

Usage

```
estimate_idr2d_hic(
  rep1_df,
  rep2_df,
  combined_min_value = 30,
  combined_max_value = Inf,
```

28 estimate_idr2d_hic

```
min_value = -Inf,
max_value = Inf,
max_factor = 1.5,
jitter_factor = 1e-04,
mu = 0.1,
sigma = 1,
rho = 0.2,
p = 0.5,
eps = 0.001,
max_iteration = 30,
local_idr = TRUE
)
```

Arguments

rep1_df data frame of either parsed .hic file from Juicer (output of parse_juicer_matrix)

or parsed .matrix and .bed files from HiC-Pro (output of parse_hic_pro_matrix)

for replicate 1

rep2_df data frame of either parsed .hic file from Juicer (output of parse_juicer_matrix)

or parsed .matrix and .bed files from HiC-Pro (output of parse_hic_pro_matrix)

for replicate 2

combined_min_value

exclude blocks with a combined (replicate 1 + replicate 2) read count or normalized read count of less than combined_min_value (default is 20 reads, set

combined_min_value = -Inf to disable)

combined_max_value

exclude blocks with a combined (replicate 1 + replicate 2) read count or normalized read count of more than combined_max_value (disabled by default, set

combined_max_value = Inf to disable)

min_value exclude blocks with a read count or normalized read count of less than min_value

in one replicate (disabled by default, set min_value = -Inf to disable)

max_value exclude blocks with a read count or normalized read count of more than max_value

in one replicate (disabled by default, set max_value = Inf to disable)

max_factor numeric; controls the replacement values for Inf and -Inf. Inf are replaced by

 $max(x) * max_factor and -Inf are replaced by min(x) / max_factor.$

jitter_factor numeric; controls the magnitude of the noise that is added to x. This is done to

break ties in x. Set jitter_factor = NULL for no jitter.

mu a starting value for the mean of the reproducible component.

sigma a starting value for the standard deviation of the reproducible component.

rho a starting value for the correlation coefficient of the reproducible component.

p a starting value for the proportion of reproducible component.

eps Stopping criterion. Iterations stop when the increment of log-likelihood is <

eps*log-likelihood, Default=0.001.

max_iteration integer; maximum number of iterations for IDR estimation (defaults to 30)

local_idr see est.IDR

hic 29

Value

Data frame with the following columns:

```
character; genomic location of interaction block (e.g., "chr1:204940000-204940000")
column 1:
            interaction
column 2:
                             numeric; p-value, FDR, or heuristic used to rank the interactions
            value
column 3:
            "rep_value"
                             numeric; value of corresponding replicate interaction
column 4:
            "rank"
                             integer; rank of the interaction, established by value column, ascending order
column 5:
            "rep_rank"
                             integer; rank of corresponding replicate interaction
            "idr"
column 6:
                             integer; IDR of the block and the corresponding block in the other replicate
```

References

Q. Li, J. B. Brown, H. Huang and P. J. Bickel. (2011) Measuring reproducibility of high-throughput experiments. Annals of Applied Statistics, Vol. 5, No. 3, 1752-1779.

Examples

hic

Example Hi-C data set

Description

This object contains data from a Hi-C contact map of human chromosome 1 and a resolution of 2.5 * 10^6, extracted from GEO series GSE71831.

Usage

hic

Format

A list with two components, the data frames rep1_df and rep2_df, which have the following four columns:

```
column 1: chr character; genomic location of block - chromosome (e.g., "chr3")
column 2: region1 integer; genomic location of block - coordinate A
column 3: region2 integer; genomic location of block - coordinate B
column 4: value numeric; heuristic used to rank blocks, in this case: number of reads
```

30 parse_juicer_matrix

Description

This function is used to convert the contact matrix from a HiC-Pro pipeline analysis run into an IDR2D compatible format. It takes one .matrix and one .bed file per replicate from HiC-Pro and returns the contact matrix for a specific chromosome for IDR2D analysis (see estimate_idr2d_hic)

Usage

```
parse_hic_pro_matrix(matrix_file, bed_file, chromosome = "chr1")
```

Arguments

matrix_file path to .matrix file from HiC-Pro analysis run

bed_file path to .bed file from HiC-Pro analysis run

chromosome chromosome name to be analyzed, defaults to UCSC chromosome 1 ("chr1")

Value

Data frame with the following columns:

```
column 1: chr character; chromosome of block (e.g., "chr3")
column 2: region1 integer; genomic location of side A of block in Hi-C contact matrix
column 3: region2 integer; genomic location of side B of block in Hi-C contact matrix
column 4: value numeric; (normalized) read count in block
```

References

Servant, N., Varoquaux, N., Lajoie, B.R. et al. HiC-Pro: an optimized and flexible pipeline for Hi-C data processing. Genome Biol 16, 259 (2015) doi:10.1186/s13059-015-0831-x

Description

parse_juicer_matrix uses the Python package hic-straw internally to read .hic contact matrix files (see hic-straw on PyPI or the Aiden lab GitHub repository for more information).

The contact matrix is subdivided into blocks, where the block size is determined by resolution. The reads per block are used to rank blocks and replicate blocks are easily matched by genomic location.

preprocess 31

Usage

```
parse_juicer_matrix(
  hic_file,
  resolution = 1e+06,
  normalization = c("NONE", "VC", "VC_SQRT", "KR"),
  chromosome = "chr1",
  use_python = NULL,
  use_virtualenv = NULL,
  use_condaenv = NULL
)
```

Arguments

hic_file path to .hic file (either local file path or URL). resolution block resolution of Hi-C contact matrix in base pairs, defaults to 1,000,000 bp (usually one of the following: 2500000, 1000000, 500000, 250000, 1000000, 50000, 25000, 10000, 5000) normalization step performed by Python package hic-straw, one of the follownormalization ing: "NONE", "VC", "VC_SQRT", "KR". chromosome chromsome name to be analyzed, defaults to UCSC chromosome 1 ("chr1") use_python if Python is not on PATH, specify path to Python binary here (see use_python) use_virtualenv if Python package hic-straw is not in base virtualenv environment, specify environment here (see use_virtualenv) if Python package hic-straw is not in base conda environment, specify enviuse_condaenv

Value

Data frame with the following columns:

```
column 1: chr character; chromosome of block (e.g., "chr3")
column 2: region1 integer; genomic location of side A of block in Hi-C contact matrix
column 3: region2 integer; genomic location of side B of block in Hi-C contact matrix
column 4: value numeric; (normalized) read count in block
```

ronment here (see use_condaenv)

References

Neva C. Durand, James T. Robinson, Muhammad S. Shamim, Ido Machol, Jill P. Mesirov, Eric S. Lander, and Erez Lieberman Aiden. "Juicebox provides a visualization system for Hi-C contact maps with unlimited zoom." Cell Systems 3(1), 2016.

preprocess	Prepares Data for IDR Analysi	is

32 preprocess

Description

This method removes invalid values, establishes the correct ranking, and breaks ties prior to IDR analysis.

Inf and -Inf are replaced by $\max(x) * \max_{factor}$ and $\min(x) / \max_{factor}$, respectively.

NA values in x are replaced by mean(x).

All values in x are transformed using the transformation specified in value_transformation.

Lastly, a small amount of noise is added to x to break ties. The magnitude of the noise is controlled by jitter_factor.

Usage

```
preprocess(
    x,
    value_transformation = c("identity", "additive_inverse", "multiplicative_inverse",
        "log", "log_additive_inverse"),
    max_factor = 1.5,
    jitter_factor = 1e-04
)
```

Arguments

x numeric vector of values value_transformation

the values in x have to be transformed in a way such that when ordered in descending order, more significant interactions end up on top of the list. If the values in x are p-values, "log_additive_inverse" is recommended. The following transformations are supported:

```
"identity" no transformation is performed on x
    "additive_inverse" x. = -x
"multiplicative_inverse" x. = 1 / x
    "log" x. = log(x). Note: zeros are replaced by .Machine$double.xmin
    "log_additive_inverse" x. = -log(x), recommended if x are p-values. Note: zeros are replaced by .Machine$double.xmin
    either "ascending" (more significant interactions have lower value in value)
```

column) or "descending" (more significant interactions have higher value in value column)

max_factor numeric; controls the replacement values for Inf and -Inf. Inf are replaced by $max(x) * max_factor$ and -Inf are replaced by $min(x) / max_factor$.

jitter_factor numeric; controls the magnitude of the noise that is added to x. This is done to break ties in x. Set jitter_factor = NULL for no jitter.

Value

numeric vector; transformed and stripped values of x, ready for IDR analysis

Examples

```
rep1_df <- idr2d:::chiapet$rep1_df
rep1_df$fdr <- preprocess(rep1_df$fdr, "log_additive_inverse")</pre>
```

remove_nonstandard_chromosomes1d

Removes Peaks on Non-standard Chromosomes

Description

Removes Peaks on Non-standard Chromosomes

Usage

```
remove_nonstandard_chromosomes1d(x)
```

Arguments

x data frame of genomic peaks, with the following columns (position of columns matter, column names are irrelevant):

```
column 1: chr character; genomic location of peak - chromosome (e.g., "chr3")
column 2: start integer; genomic location of peak - start coordinate
column 3: end integer; genomic location of peak - end coordinate
column 4: value numeric; p-value, FDR, or heuristic used to rank the peaks
```

Value

x without non-standard chromosomes.

Examples

```
\label{lem:constandard_chromosomes1d} rep1\_df <- \ remove\_nonstandard\_chromosomes1d(idr2d:::chipseq\$rep1\_df)
```

```
remove_nonstandard_chromosomes2d
```

Removes Interactions on Non-standard Chromosomes

Description

Removes Interactions on Non-standard Chromosomes

Usage

 $remove_nonstandard_chromosomes2d(x)$

Arguments

x data frame of genomic interactions, with the following columns (position of columns matter, column names are irrelevant):

```
column 1: chr_a
                       character; genomic location of anchor A - chromosome (e.g., "chr3")
column 2: start_a
                       integer; genomic location of anchor A - start coordinate
column 3: end a
                       integer; genomic location of anchor A - end coordinate
column 4: chr_b
                       character; genomic location of anchor B - chromosome (e.g., "chr3")
column 5: start_b
                       integer; genomic location of anchor B - start coordinate
                       integer; genomic location of anchor B - end coordinate
column 6:
           end_b
column 7:
           value
                       numeric; p-value, FDR, or heuristic used to rank the interactions
```

Value

x without non-standard chromosomes.

Examples

```
rep1_df <- remove_nonstandard_chromosomes2d(idr2d:::chiapet$rep1_df)</pre>
```

Index

```
* datasets
    chiapet, 7
    chipseq, 8
    hic, 29
calculate_midpoint_distance1d, 2
calculate_midpoint_distance2d, 3
calculate_relative_overlap1d, 5
calculate_relative_overlap2d, 6
chiapet, 7
chipseq, 8
determine_anchor_overlap, 8
draw_hic_contact_map, 9
{\tt draw\_idr\_distribution\_histogram,}\ 10
draw_rank_idr_scatterplot, 11
draw_value_idr_scatterplot, 12
est.IDR, 22, 24, 26, 28
establish_bijection, 13
establish_bijection1d, 14, 14, 21
establish_bijection2d, 14, 16, 21
establish_overlap1d, 14, 17, 22
establish_overlap2d, 14, 19, 22
estimate_idr, 10-13, 21
estimate_idr1d, 22, 22
estimate_idr2d, 22, 25
estimate_idr2d_hic, 9, 27, 30
hic, 29
keepStandardChromosomes, 22, 24, 26
parse_hic_pro_matrix, 28, 30
parse_juicer_matrix, 28, 30
preprocess, 31
remove_nonstandard_chromosomes1d, 33
remove_nonstandard_chromosomes2d, 33
use_condaenv, 31
use_python, 31
use_virtualenv, 31
```