Package 'idr2d'

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```
Title Irreproducible Discovery Rate for Genomic Interactions Data Version 1.18.0
```

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.

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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 (>= 1.10), roxygen2 (>= 6.1.0), testthat (>= 2.1.0)
```

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 ${\tt calculate_midpoint_distance1d}$

Distance between Midpoints of two Peaks

Description

Index

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

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_b_end,
  int2_anchor_b_end,
  int2_anchor_b_end,
  int2_anchor_b_start,
  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

```
# 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

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_start,
  int2_anchor_b_start,
  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

Value

numeric vector; relative overlaps between interaction pairs

```
# 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))
```

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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: 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
column 6: end_b integer; genomic location of anchor B - end coordinate
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
            end
  rep2_anchor
                     data frame with the following columns:
column 1:
            chr
                     character; genomic location of anchor in replicate 2 - chromosome (e.g., "chr3")
column 2:
            start
                     integer; genomic location of anchor in replicate 2 - start coordinate
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)
```

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
```

```
start = rep2_df[, 2],
end = rep2_df[, 3])
anchor_a_overlap <- determine_anchor_overlap(rep1_anchor_a, rep2_anchor_a)</pre>
```

```
draw_hic_contact_map
Create Hi-C contact map
```

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
column 5:
            "rep_rank"
                           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
                    character; chromsome name or list of chromosome names to be analyzed, e.g.,
  chromosome
                    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 values_normalized
```

logical; are read counts in value column raw or normalized? Defaults to FALSE

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

Value

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

Examples

```
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 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

df 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

rep_rank 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
)
```

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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

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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_reso	lution_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.
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

See establish_bijection1d or establish_bijection2d, respectively.

```
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*.

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 follow-
                  ing columns (position of columns matter, column names are irrelevant):
   column 1:
                chr
                         character; genomic location of peak - chromosome (e.g., "chr3")
                         integer; genomic location of peak - start coordinate
   column 2:
                start
                         integer; genomic location of peak - end coordinate
   column 3:
                end
                         numeric; p-value, FDR, or heuristic used to rank the interactions
   column 4:
                value
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 two is "overlap" the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate 1 in the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance from

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)

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Value

Data frames rep1_df and rep2_df with the following columns:

```
column 1:
                          character; genomic location of peak - chromosome (e.g., "chr3")
            chr
column 2:
            start
                          integer; genomic location of peak - start coordinate
column 3:
                          integer; genomic location of peak - end coordinate
            end
column 4:
            value
                          numeric; p-value, FDR, or heuristic used to rank the peaks
column 5:
                          numeric; value of corresponding replicate peak. If no corresponding peak was found, rep_value is
            rep_value
column 6:
                          integer; rank of the peak, established by value column, ascending order
column 7:
                          integer; rank of corresponding replicate peak. If no corresponding peak was found, rep_rank is set
            rep_rank
column 8:
            idx
                          integer; peak index, primary key
column 9:
                          integer; specifies the index of the corresponding peak in the other replicate (foreign key). If no corr
            rep_idx
```

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_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

leas			a frame of observations (i.e., genomic interactions) of replicate 1, with at at the following columns (position of columns matter, column names are invant):		
	column 1: column 2: column 3: column 4: column 5: column 6: column 7:	chr_a start_a end_a chr_b start_b end_b value	character; genomic location of anchor A - chromosome (e.g., "chr3") integer; genomic location of anchor A - start coordinate integer; genomic location of anchor A - end coordinate character; genomic location of anchor B - chromosome (e.g., "chr3") integer; genomic location of anchor B - start coordinate integer; genomic location of anchor B - end coordinate 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: column 2: column 3: column 4: column 5: column 6: column 7:	chr_a start_a end_a chr_b start_b end_b value	character; genomic location of anchor A - chromosome (e.g., "chr3") integer; genomic location of anchor A - start coordinate integer; genomic location of anchor A - end coordinate character; genomic location of anchor B - chromosome (e.g., "chr3") integer; genomic location of anchor B - start coordinate integer; genomic location of anchor B - end coordinate numeric; p-value, FDR, or heuristic used to rank the interactions		
	ambiguity _.	defi: laps	on_method nes how ambiguous assignments (when one interaction in replicate 1 overwith multiple interactions in replicate 2 or vice versa) are resolved. Availamethods:		
	"valua"	interaction	os are prioritized by ascending or descending value column (see conting di		

"value" interactions are prioritized by ascending or descending value column (see sorting_direction), e.g., if two is "overlap" the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate 1 in the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance from

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:
            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")
                            integer; genomic location of anchor B - start coordinate
column 5:
            start_b
            end_b
column 6:
                            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 foun
```

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```
column 9: "rank" integer; rank of the interaction, established by value column, ascending order column 10: "rep_rank" integer; rank of corresponding replicate interaction. If no corresponding interaction was found, column 11: "idx" integer; interaction index, primary key column 12: "rep_idx" integer; specifies the index of the corresponding interaction in the other replicate (foreign key).
```

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>
```

establish_overlap1d Establish m:n Mapping Between Peaks 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_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
```

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```
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
    column 3:
                          integer; genomic location of peak - end coordinate
                 end
                          numeric; p-value, FDR, or heuristic used to rank the interactions
    column 4:
                 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 two in
"overlap"
             the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate 1 in
```

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

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

```
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>
```

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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):

```
character; genomic location of anchor A - chromosome (e.g., "chr3")
column 1:
            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
                       character; genomic location of anchor B - chromosome (e.g., "chr3")
column 4:
            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
```

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
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
```

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```
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 two is "overlap" the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate 1 in the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance from

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)
column 3: arv ambiguity resolution value used turn m:n mapping into 1:1 mapping. Interaction pairs with lower ar
```

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

Estimates IDR for Genomic Peaks or Genomic Interactions

Description

Estimates IDR for Genomic Peaks or Genomic Interactions

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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 repl_df.

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 or peak in replicate 1 overlaps with multiple interactions or peaks in replicate 2 or vice versa) are re-

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solved. 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.

estimate_idr1d 25

estimate_idr1d

Estimates IDR for Genomic Peak Data

Description

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

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 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
   column 3:
                end
   column 4:
               value
                         numeric; p-value, FDR, or heuristic used to rank the interactions
```

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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\$doubl

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 two in "overlap" the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate 1 in "midpoint" the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance from

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.

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)

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

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

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

a starting value for the proportion of reproducible component.

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

eps*log-likelihood, Default=0.001.

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

local_idr see est. IDR estimate_idr2d 27

Value

column 1:

chr

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 2:
             start
column 3:
             end
                           integer; genomic location of peak - end coordinate
                           numeric; p-value, FDR, or heuristic used to rank the peaks
 column 4:
             value
             rep_value
column 5:
                           numeric; value of corresponding replicate peak. If no corresponding peak was found, rep_value
                           integer; rank of the peak, established by value column, ascending order
 column 6:
             rank
                           integer; rank of corresponding replicate peak. If no corresponding peak was found, rep_rank is s
 column 7:
             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 co
                           IDR of the peak and the corresponding peak in the other replicate. If no corresponding peak was t
column 10:
             idr
```

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

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) 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,
```

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```
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:
            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:
                        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
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:
                        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
                        character; genomic location of anchor B - chromosome (e.g., "chr3")
column 4:
           chr b
```

integer; genomic location of anchor B - start coordinate

integer; genomic location of anchor B - end coordinate

value_transformation

column 5: start_b

end b

value

column 6:

column 7:

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:

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

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

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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 two is "overlap" the interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate 1 in the interaction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance from

remove_nonstandard_chromosomes

see est. IDR

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_factor $max(x) * max_factor and -Inf are replaced by min(x) / max_factor.$ 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. 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) a starting value for the mean of the reproducible component. mu a starting value for the standard deviation of the reproducible component. sigma a starting value for the correlation coefficient of the reproducible component. rho a starting value for the proportion of reproducible component. Stopping criterion. Iterations stop when the increment of log-likelihood is < eps eps*log-likelihood, Default=0.001. max_iteration integer; maximum number of iterations for IDR estimation (defaults to 30)

Value

local idr

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 1:
            chr_a
 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"
column 11:
            "idx"
column 12:
            "rep_idx"
```

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

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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_idr2d_hic

Estimates IDR for Genomic Interactions measured by Hi-C experiments

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,
 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
)
```

estimate_idr2d_hic 31

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_v	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_v	ralue
	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 \neg Inf. Inf are replaced by $\max(x) * \max_{f} \arctan \neg F$ are replaced by $\min(x) / \max_{f} F$.
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.
р	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

Data frame with the following columns:

```
column 1:
            interaction
                            character; genomic location of interaction block (e.g., "chr1:204940000-204940000")
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
column 6:
            "idr"
                            integer; IDR of the block and the corresponding block in the other replicate
```

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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
```

parse_hic_pro_matrix Parse .matrix and .bed files from HiC-Pro for IDR2D analysis

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")
```

parse_juicer_matrix 33

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 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.

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
)
```

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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, 100000, 50000, 25000, 10000, 5000) normalization normalization step performed by Python package hic-straw, one of the following: "NONE", "VC", "VC_SQRT", "KR". chromsome name to be analyzed, defaults to UCSC chromosome 1 ("chr1") chromosome if Python is not on PATH, specify path to Python binary here (see use_python) use_python if Python package hic-straw is not in base virtualenv environment, specify use_virtualenv environment here (see use_virtualenv) use_condaenv if Python package hic-straw is not in base conda environment, specify environment here (see use_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

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 Analysis

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.

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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.

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

```
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

```
rep1_df <- remove_nonstandard_chromosomes1d(idr2d:::chipseq$rep1_df)</pre>
```

remove_nonstandard_chromosomes2d

Removes Interactions on Non-standard Chromosomes

Description

Removes Interactions on Non-standard Chromosomes

Usage

remove_nonstandard_chromosomes2d(x)

Arguments

X	aata	a frame of genomic interactions, with the following columns (position of
	colı	umns matter, column names are irrelevant):
column 1:	chr a	character; genomic location of anchor A chromosome (e.g. "chr2")

```
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
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

x without non-standard chromosomes.

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

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