# Package 'idr2d'

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Title Irreproducible Discovery Rate for Genomic Interactions Data

Version 1.23.0

**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)

- **Imports** dplyr (>= 0.7.6), futile.logger (>= 1.4.3), GenomeInfoDb (>= 1.14.0), GenomicRanges (>= 1.30), ggplot2 (>= 3.1.1), grDevices, grid, idr (>= 1.2), IRanges (>= 2.18.0), magrittr (>= 1.5), methods, reticulate (>= 1.13), scales (>= 1.0.0), stats, stringr (>= 1.3.1), utils
- Suggests DT (>= 0.4), htmltools (>= 0.3.6), knitr (>= 1.20), rmarkdown (>= 1.10), roxygen2 (>= 6.1.0), testthat (>= 2.1.0)

#### VignetteBuilder knitr

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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

calculate\_midpoint\_distance2d

#### 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_a_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

## 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),
                              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_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

numeric vector; relative overlaps between interaction pairs

#### chiapet

#### 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

#### 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:chrcharacter; genomic location of peak - chromosome (e.g., "chr3")column 2:startinteger; genomic location of peak - start coordinatecolumn 3:endinteger; genomic location of peak - end coordinatecolumn 4:valuenumeric; 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 column 2: start column 3: end		character; genomic location of anchor in replicate 1 - chromosome (e.g., "chr3") integer; genomic location of anchor in replicate 1 - start coordinate integer; genomic location of anchor in replicate 1 - end coordinate
rep2_anchor		data frame with the following columns:
column 1: column 2: column 3:	chr start end	character; genomic location of anchor in replicate 2 - chromosome (e.g., "chr3") integer; genomic location of anchor in replicate 2 - start coordinate integer; genomic location of anchor in replicate 2 - end coordinate
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)

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

#### Examples

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:

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

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	
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	

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

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 rep_rank idr	integer; rank of the peak, established by value column, ascending order integer; rank of corresponding replicate peak. IDR of the peak and the corresponding peak in the other replicate.
remove_na xlab	logical; should NA values be removed? 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	

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
)
```

## Arguments

df	part of output of estimate_idr, a data frame with at least the following named columns:
value rep_value idr	numeric; value of corresponding replicate peak
remove_na	logical; should NA values be removed?
remove_outliers	3
	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) $% \left( {{\left( {{{\left( {{{\left( {{{\left( {{{\left( {{{\left( {{{{\left( {{{\left( {{{\left( {{{\left( {{{{\left( {{{\left( {{{{\left( {{{{}}}}}} \right)}}}}\right.}$
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 $1.0$
<pre>max_points_show</pre>	<i>i</i> n
	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 repli- cate 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 repli- cate 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 re- solved. 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.

#### 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

## Usage

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

## Arguments

		frame of observations (i.e., genomic peaks) of replicate 1, with at least the wing columns (position of columns matter, column names are irrelevant):
column 1: column 2:	chr start	character; genomic location of peak - chromosome (e.g., "chr3") 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 c	olumns (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_re	esolutio	n_method
0 9-		es how ambiguous assignments (when one interaction in replicate 1 over-
	laps	with multiple interactions in replicate 2 or vice versa) are resolved. Avail- methods:
"value" in	nteractions	s are prioritized by ascending or descending value column (see sorting_direction), e.g., if t
"overlap" th	ne interact	ion pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate
"midpoint" th	ne interact	ion 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:	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
column 5:	rep_value	numeric; value of corresponding replicate peak. If no corresponding peak was found, rep_value
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 i
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

#### 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

rep1_df	leas	frame of observations (i.e., genomic interactions) of replicate 1, with at t the following columns (position of columns matter, column names are ir- vant):
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
rep2_df		frame of observations (i.e., genomic interactions) of replicate 2, with the owing 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: column 5: column 6: column 7:	chr_b start_b end_b value	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	
"value" "overlap" "midpoint"	the interac	as are prioritized by ascending or descending value column (see sorting_direction), e.g., if the stion pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate extion pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance
max_gap		ger; maximum gap in nucleotides allowed between two anchors for them to considered as overlapping (defaults to -1, i.e., overlapping anchors)

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")
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 i
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 for
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 k

## 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 column 2 column 3 column 4	2: start 3: end	character; genomic location of peak - chromosome (e.g., "chr3") integer; genomic location of peak - start coordinate integer; genomic location of peak - end coordinate numeric; p-value, FDR, or heuristic used to rank the interactions	
rep2_df		frame of observations (i.e., genomic peaks) of replicate 2, with the follow- olumns (position of columns matter, column names are irrelevant):	
column 1	1: chr	character; genomic location of peak - chromosome (e.g., "chr3")	
column 2	2: start	integer; genomic location of peak - start coordinate	
column 3		integer; genomic location of peak - end coordinate	
column 4	4: value	numeric; p-value, FDR, or heuristic used to rank the interactions	
ambiguity_	define laps v	n_method es how ambiguous assignments (when one interaction in replicate 1 over- with multiple interactions in replicate 2 or vice versa) are resolved. Avail- methods:	
"value" "overlap" "midpoint"	the interaction	s are prioritized by ascending or descending value column (see sorting_direction), e.g., if to ion pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate ion pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance	
max_gap	U	er; maximum gap in nucleotides allowed between two anchors for them to onsidered 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

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 <- 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 a least the following columns (position of columns matter, column names are in relevant):		
column 2: column 3: column 4:	end_a chr_b	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")	
column 5:	start_b	integer; genomic location of anchor B - start coordinate	

column 6: column 7:	end_b value	integer; genomic location of anchor B - end coordinate numeric; p-value, FDR, or heuristic used to rank the interactions
rep2_df		a frame of observations (i.e., genomic interactions) of replicate 2, with the owing columns (position of columns matter, column names are irrelevant):
column 1: column 2:	chr_a start_a	character; genomic location of anchor A - chromosome (e.g., "chr3") integer; genomic location of anchor A - start coordinate
column 3: column 4: column 5:	end_a chr_b start_b	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
column 6: column 7:	end_b value	integer; genomic location of anchor B - end coordinate numeric; p-value, FDR, or heuristic used to rank the interactions
ambiguity_	laps	on_method nes how ambiguous assignments (when one interaction in replicate 1 over- s with multiple interactions in replicate 2 or vice versa) are resolved. Avail- e methods:
"value" "overlap" "midpoint"	the interac	ns are prioritized by ascending or descending value column (see sorting_direction), e.g., if to ction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate ction pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance
max_gap		ger; maximum gap in nucleotides allowed between two anchors for them to considered as overlapping (defaults to -1, i.e., overlapping anchors)

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 lowe

#### 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

## 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 repli- cate 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 repli- cate 2. Same columns as rep1_df.		
analysis_type value_transform	"IDR2D" for genomic interaction data sets, "IDR1D" for genomic peak data sets nation		
	the values in x have to be transformed in a way such that when ordered in de- scending 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 fol- lowing transformations are supported:		
additive_: 	<pre>dentity" no transformation is performed on x inverse" x. = -x inverse" x. = 1 / x "log" x. = log(x). Note: zeros are replaced by .Machine\$double.xmin inverse" x. = -log(x), recommended if x are p-values. Note: zeros are replaced by .Machine\$do</pre>		
	either "ascending" (more significant interactions have lower value in value column) or "descending" (more significant interactions have higher value in value column)		

ambiguity_resol	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 re- solved. For available methods, see <pre>establish_overlap1d</pre> or <pre>establish_overlap2d</pre> respectively.
remove_nonstand	dard_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.
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.
<pre>max_iteration</pre>	integer; maximum number of iterations for IDR estimation (defaults to 30)
local_idr	see est.IDR

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

```
idr_results <- estimate_idr(idr2d:::chiapet$rep1_df,</pre>
                             idr2d:::chiapet$rep2_df,
                             analysis_type = "IDR2D",
                             value_transformation = "log_additive_inverse")
```

summary(idr\_results)

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

#### 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		frame of observations (i.e., genomic peaks) of replicate 1, with at least the wing columns (position of columns matter, column names are irrelevant):
column 1 column 2 column 3 column 4	: start : end	character; genomic location of peak - chromosome (e.g., "chr3") integer; genomic location of peak - start coordinate integer; genomic location of peak - end coordinate numeric; p-value, FDR, or heuristic used to rank the interactions
rep2_df		frame of observations (i.e., genomic peaks) of replicate 2, with the follow- columns (position of columns matter, column names are irrelevant):
column 1 column 2 column 3 column 4	: start : end	character; genomic location of peak - chromosome (e.g., "chr3") integer; genomic location of peak - start coordinate integer; genomic location of peak - end coordinate numeric; p-value, FDR, or heuristic used to rank the interactions
value_trans	the v scene value	n ralues in x have to be transformed in a way such that when ordered in de- ding order, more significant interactions end up on top of the list. If the es in x are p-values, "log_additive_inverse" is recommended. The fol- ng transformations are supported:
"addit: "multiplicat: "log_addit:	"1	se" x. = -x se" x. = 1 / x og" x. = log(x). Note: zeros are replaced by .Machine\$double.xmin
		r "ascending" (more significant interactions have lower value in value mn) or "descending" (more significant interactions have higher value in

value column)

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:	
"overlap" th	teractions are prioritized by ascending or descending value column (see sorting_direction), e.g., if to e interaction pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate e 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_facto	<pre>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.</pre>	
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.	
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.	
<pre>max_iteratio</pre>	n integer; maximum number of iterations for IDR estimation (defaults to 30)	
local_idr	see est.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	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
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
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
column 10:	idr	IDR of the peak and the corresponding peak in the other replicate. If no corresponding peak v

## 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\_idr2d

#### 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:	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

rep2_df		frame of observations (i.e., genomic interactions) of replicate 2, with the owing 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		
value_tran				
		values in x have to be transformed in a way such that when ordered in de- nding order, more significant interactions end up on top of the list. If the		
		tes in x are p-values, "log_additive_inverse" is recommended. The fol-		
		ing transformations are supported:		
	"ident:	ty" no transformation is performed on x		
	ive_inve			
"multiplicat				
"lag addit		log" x. = log(x). Note: zeros are replaced by .Machine\$double.xmin		
"log_addit	1Ve_1nver	rse" x. = -log(x), recommended if x are p-values. Note: zeros are replaced by .Machine\$do		
		er "ascending" (more significant interactions have lower value in value		
		umn) or "descending" (more significant interactions have higher value in		
		ue column)		
ambiguity_				
		nes how ambiguous assignments (when one interaction in replicate 1 over- with multiple interactions in replicate 2 or vice versa) are resolved. Avail-		
	-	e methods:		
"value"	interaction	ns are prioritized by ascending or descending value column (see sorting_direction), e.g., if t		
"overlap"		tion pair is chosen which has the highest relative overlap, i.e., overlap in nucleotides of replicate		
"midpoint"	the interac	tion pair is chosen which has the smallest distance between their anchor midpoints, i.e., distance		
remove_non	standard_	chromosomes		
		oves interactions containing genomic locations on non-standard chromo-		
		es using keepStandardChromosomes (default is TRUE)		
<pre>max_factor</pre>		neric; controls the replacement values for Inf and -Inf. Inf are replaced by		
		(x) * max_factor and -Inf are replaced by min(x) / max_factor.		
jitter_fac		heric; controls the magnitude of the noise that is added to x. This is done to a k ties in x. Set jitter_factor = NULL for no jitter.		
max_gap		ger; maximum gap in nucleotides allowed between two anchors for them to		
ιιαν_βαμ		onsidered as overlapping (defaults to -1, i.e., overlapping anchors)		
mu	a sta	arting value for the mean of the reproducible component.		
sigma	a sta	arting value for the standard deviation of the reproducible component.		
rho	a sta	arting value for the correlation coefficient of the reproducible component.		
р	a sta	arting 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.		
<pre>max_iterat</pre>	ion inte	ger; maximum number of iterations for IDR estimation (defaults to 30)		
local_idr		est.IDR		

#### estimate\_idr2d\_hic

#### Value

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:

1 1 .	alua a
column 1:	cnr_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"
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

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
)
```

## 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_matri or parsed .matrix and .bed files from HiC-Pro (output of parse_hic_pro_matri for replicate 2	
combined_min_va	lue	
	exclude blocks with a combined (replicate 1 + replicate 2) read count or nor- malized read count of less than combined_min_value (default is 20 reads, set combined_min_value = -Inf to disable)	
combined_max_va	lue	
	exclude blocks with a combined (replicate 1 + replicate 2) read count or nor- malized 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.	
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	

hic

## Value

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

## 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:chrcharacter; genomic location of block - chromosome (e.g., "chr3")column 2:region1integer; genomic location of block - coordinate Acolumn 3:region2integer; genomic location of block - coordinate Bcolumn 4:valuenumeric; 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 <a href="mailto:estimate\_idr2d\_hic">estimate\_idr2d\_hic</a>)

#### 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	chromsome 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

parse\_juicer\_matrix Parse .hic files from Juicer for IDR2D analysis

## 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

#### 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: 250000, 1000000, 500000, 250000, 100000, 50000, 250000, 100000, 50000)
normalization	normalization step performed by Python package hic-straw, one of the follow- 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)
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

## 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

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:

"ic	dentity"	no transformation is performed on x
"additive_inverse"		x. = -x
"multiplicative_i	inverse"	x. = 1 / x
	"log"	<pre>x. = log(x). Note: zeros are replaced by .Machine\$double.xmin</pre>
"log_additive_inverse"		x. = -log(x), recommended if x are p-values. Note: zeros are replaced by .Machine\$do
max_factor	column) c value col	<i>,</i>
max_ractor	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		controls the magnitude of the noise that is added to x. This is done to 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

х	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		a frame of genomic interactions, with the following columns (position of imms 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

## Value

x without non-standard chromosomes.

## Examples

rep1\_df <- remove\_nonstandard\_chromosomes2d(idr2d:::chiapet\$rep1\_df)</pre>

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