# Package 'ChromSCape'

November 6, 2025

Title Analysis of single-cell epigenomics datasets with a Shiny App

**Version** 1.21.0

Description ChromSCape - Chromatin landscape profiling for Single Cells - is a ready-to-launch user-friendly Shiny Application for the analysis of single-cell epigenomics datasets (scChIP-seq, scATAC-seq, scCUT&Tag, ...) from aligned data to differential analysis & gene set enrichment analysis. It is highly interactive, enables users to save their analysis and covers a wide range of analytical steps: QC, preprocessing, filtering, batch correction, dimensionality reduction, vizualisation, clustering, differential analysis and gene set analysis.

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biocViews ShinyApps, Software, SingleCell, ChIPSeq, ATACSeq, MethylSeq, Classification, Clustering, Epigenetics, PrincipalComponent, SingleCell, ATACSeq, ChIPSeq, Annotation, BatchEffect, MultipleComparison, Normalization, Pathways, Preprocessing, QualityControl, ReportWriting, Visualization, GeneSetEnrichment, DifferentialPeakCalling

VignetteBuilder knitr

URL https://github.com/vallotlab/ChromSCape

BugReports https://github.com/vallotlab/ChromSCape/issues

Encoding UTF-8 LazyData false

**Suggests** testthat, knitr, markdown, rmarkdown, BiocStyle, Signac, future, igraph, bluster, httr

RoxygenNote 7.3.2

**Roxygen** list(markdown = TRUE)

Imports shiny, colourpicker, shinyjs, rtracklayer, shinyFiles, shinyhelper, shinyWidgets, shinydashboardPlus, shinycssloaders, Matrix, plotly, shinydashboard, colorRamps, kableExtra, viridis, batchelor, BiocParallel, parallel, Rsamtools, ggplot2, ggrepel, gggenes, gridExtra, qualV, stringdist, stringr, fs, qs, DT, scran, scater, ConsensusClusterPlus, Rtsne, dplyr, tidyr, GenomicRanges, IRanges, irlba, rlist, umap, tibble,

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annotation\_from\_merged\_peaks

Find nearest peaks of each gene and return refined annotation

# **Description**

Find nearest peaks of each gene and return refined annotation

# Usage

```
annotation_from_merged_peaks(scExp, odir, merged_peaks, geneTSS_annotation)
```

# **Arguments**

scExp A SingleCellExperiment object odir An output directory where to write the mergedpeaks BED file merged\_peaks A list of GRanges object containing the merged peaks geneTSS\_annotation

A GRanges object with reference genes

### Value

A data.frame with refined annotation

6 annotToCol2

annotToCol2

annotToCol2

# Description

annotToCol2

# Usage

```
annotToCol2(
  annotS = NULL,
  annotT = NULL,
  missing = c("", NA),
  anotype = NULL,
  maxnumcateg = 2,
  categCol = NULL,
  quantitCol = NULL,
  plotLegend = TRUE,
  plotLegendFile = NULL)
```

# **Arguments**

annotS A color matrix A color matrix annotT Convert missing to NA missing anotype Annotation type Maximum number of categories maxnumcateg categCol Categorical columns quantitCol Quantitative columns plotLegend Plot legend? plotLegendFile Which file to plot legend?

#### Value

A matrix of continuous or discrete colors

```
data("scExp")
annotToCol2(SingleCellExperiment::colData(scExp), plotLegend = FALSE)
```

anocol\_binary 7

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anocol	hinary	
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Helper binary column for anocol function

# Description

Helper binary column for anocol function

# Usage

```
anocol_binary(anocol, anotype, plotLegend, annotS)
```

# **Arguments**

anocol The color feature matrix

anotype The feature types plotLegend Plot legend? A color matrix

### Value

A color matrix similar to anocol with binrary columns colored

anocol\_categorical

Helper binary column for anocol function

# **Description**

Helper binary column for anocol function

# Usage

```
anocol_categorical(anocol, categCol, anotype, plotLegend, annotS)
```

# **Arguments**

anocol The color feature matrix

categCol Colors for categorical features

anotype The feature types plotLegend Plot legend?
annotS A color matrix

### Value

A color matrix similar to anocol with binrary columns colored

bams\_to\_matrix\_indexes

Count bam files on interval to create count indexes

# **Description**

Count bam files on interval to create count indexes

### Usage

```
bams_to_matrix_indexes(dir, which, BPPARAM = BiocParallel::bpparam())
```

#### **Arguments**

dir A directory containing single cell BAM files and BAI files

which Genomic Range on which to count

BPPARAM object for multiprocessing. See bpparam for more informations.

Will take the default BPPARAM set in your R session.

### Value

A list containing a "feature index" data.frame and a count vector for non 0 entries, both used to form the sparse matrix

beds\_to\_matrix\_indexes

Count bed files on interval to create count indexes

# Description

Count bed files on interval to create count indexes

# Usage

```
beds_to_matrix_indexes(dir, which, BPPARAM = BiocParallel::bpparam())
```

### **Arguments**

dir A directory containing the single cell BED files

which Genomic Range on which to count

BPPARAM BPPARAM object for multiprocessing. See bpparam for more informations.

Will take the default BPPARAM set in your R session.

### Value

A list containing a "feature index" data.frame and a names of cells as vector both used to form the sparse matrix

calculate\_CNA 9

calculate\_CNA

Estimate copy number alterations in cytobands

# **Description**

Cytobands are considered large enough in order that a variation at the cytoband level is not considered as an epigenetic event but as a genetic event, e.g. Copy Number Alterations. The function successively:

- Calculates the fraction of reads in each cytoband (FrCyto). See calculate\_cyto\_mat
- Calculates the log2-ratio FrCyto of each cell by the average FrCyto in normal cells. See calculate\_logRatio\_CNA
- Estimates if there was a gain or a loss of copy in each cyto band. See calculate\_gain\_or\_loss

The corresponding matrices are accessibles in the reducedDim slots "cytoBands", "logRatio\_cytoBands" and "gainOrLoss\_cytoBands" respectively.

### Usage

```
calculate_CNA(
   scExp,
   control_samples = unique(scExp$sample_id)[1],
   ref_genome = c("hg38", "mm10", "ce11")[1],
   quantiles_to_define_gol = c(0.05, 0.95)
)
```

# **Arguments**

```
A SingleCellExperiment with "logRatio_cytoBand" reducedDim slot filled. See calculate_logRatio_CNA

control_samples

Sample IDs of the normal sample to take as reference.

ref_genome

Reference genome ('hg38' or 'mm10')

quantiles_to_define_gol

Quantiles of normal log2-ratio distribution below/above which cytoband is considered to be a loss/gain. (c(0.05,0.95)). See calculate_gain_or_loss
```

#### Value

The SCE with the fraction of reads, log2-ratio and gain or loss in each cytobands in each cells (of dimension cell x cytoband) in the reducedDim slots.

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# **Examples**

```
data("scExp")
scExp = calculate_CNA(scExp, control_samples = unique(scExp$sample_id)[1],
ref_genome="hg38", quantiles_to_define_gol = c(0.05,0.95))
SingleCellExperiment::reducedDim(scExp, "cytoBand")
SingleCellExperiment::reducedDim(scExp, "logRatio_cytoBand")
SingleCellExperiment::reducedDim(scExp, "gainOrLoss_cytoBand")
```

calculate\_cyto\_mat

Calculate Fraction of reads in each cytobands

# **Description**

Re-Count binned reads onto cytobands and calculate the fraction of reads in each of the cytoband in each cell. For each cell, the fraction of reads in any given cytoband is calculated. Cytobands are considered large enough in order that a variation at the cytoband level is not considered as an epigenetic event but as a genetic event, e.g. Copy Number Alterations.

# Usage

```
calculate_cyto_mat(scExp, ref_genome = c("hg38", "mm10", "ce11")[1])
```

### **Arguments**

SCEXP A SingleCellExperiment with genomic coordinate as features (peaks or bins)

ref\_genome Reference genome ('hg38' or 'mm10')

# Value

The SCE with the fraction of reads in each cytobands in each cells (of dimension cell x cytoband) in the reducedDim slot "cytoBand".

```
data("scExp")
scExp = calculate_cyto_mat(scExp, ref_genome="hg38")
SingleCellExperiment::reducedDim(scExp, "cytoBand")
```

calculate\_gain\_or\_loss 11

```
calculate_gain_or_loss
```

Estimate the copy gains/loss of tumor vs normal based on log2-ratio of fraction of reads

# Description

Given a SingleCellExperiment object with the slot "logRatio\_cytoBand" containing the log2-ratio of the fraction of reads in each cytoband, estimate if the cytoband was lost or acquired a gain in a non-quantitative way. To do so, the quantiles distribution of the normal cells are calculated, and any cytoband below or above will be considered as a loss/gain. The False Discovery Rate is directly proportional to the quantiles.

### Usage

```
calculate_gain_or_loss(scExp, controls, quantiles = c(0.05, 0.95))
```

#### **Arguments**

scExp	$A \ Single Cell Experiment \ with \ "logRatio\_cytoBand" \ reduced Dim \ slot \ filled. \ See \ calculate\_logRatio\_CNA$
controls	Sample IDs or Cell IDs of the normal sample to take as reference.
quantiles	Quantiles of normal log2-ratio distribution below/above which cytoband is considered to be a loss/gain. $(c(0.05,0.95))$

#### Value

The SCE with the gain or loss in each cytobands in each cells (of dimension cell x cytoband) in the reducedDim slot "gainOrLoss\_cytoBand".

```
data("scExp")
scExp = calculate_cyto_mat(scExp, ref_genome="hg38")
scExp = calculate_logRatio_CNA(scExp, controls=unique(scExp$sample_id)[1])
scExp = calculate_gain_or_loss(scExp, controls=unique(scExp$sample_id)[1])
SingleCellExperiment::reducedDim(scExp, "gainOrLoss_cytoBand")
```

```
calculate_logRatio_CNA
```

Calculate the log2-ratio of tumor vs normal fraction of reads in cytobands

# **Description**

Given a SingleCellExperiment object with the slot "cytoBand" containing the fraction of reads in each cytoband, calculates the log2-ratio of tumor vs normal fraction of reads in cytobands, cell by cell. If the average signal in normal sample in a cytoband is 0, set this value to 1 so that the ratio won't affect the fraction of read value.

# Usage

```
calculate_logRatio_CNA(scExp, controls)
```

# **Arguments**

scExp A SingleCellExperiment with "cytoBand" reducedDim slot filled.

• see calculate\_cyto\_mat

controls Sample IDs or Cell IDs of the normal sample to take as reference.

### Value

The SCE with the log2-ratio of fraction of reads in each cytobands in each cells (of dimension cell x cytoband) in the reducedDim slot "logRatio\_cytoBand".

# **Examples**

```
data("scExp")
scExp = calculate_cyto_mat(scExp, ref_genome="hg38")
scExp = calculate_logRatio_CNA(scExp, controls=unique(scExp$sample_id)[1])
SingleCellExperiment::reducedDim(scExp, "logRatio_cytoBand")
```

```
call_macs2_merge_peaks
```

Calling MACS2 peak caller and merging resulting peaks

# Description

Calling MACS2 peak caller and merging resulting peaks

ce11.chromosomes 13

### Usage

```
call_macs2_merge_peaks(
   affectation,
   odir,
   p.value,
   format = c("scBED", "BAM")[1],
   ref,
   peak_distance_to_merge
)
```

# Arguments

affectation Annotation data.frame with cell cluster and cell id information

odir Output directory to write MACS2 output
p.value P value to detect peaks, passed to MACS2
format File format, either "BAM" or "scBED"

ref Reference genome to get chromosome information from.

peak\_distance\_to\_merge

Distance to merge peaks

#### Value

A list of merged GRanges peaks

ce11.chromosomes

Data.frame of chromosome length - cel1

### **Description**

This data frame provides the length of each "canonical" chromosomes of C Elegans genome build cell.

#### **Usage**

```
data("ce11.chromosomes")
```

### **Format**

ce11.chromosomes - a data frame with 7 chromosomes and 3 variables:

```
chr Chromosome - characterstart Start of the chromosome (bp) - integerend End of the chromosome (bp) - integer
```

# Value

ce11.chromosomes - a data frame with 7 chromosomes and 3 variables

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ce11.GeneTSS

Data.frame of gene TSS - cel1

# **Description**

This dataframe was extracted from Gencode v25 and report the Transcription Start Site of each gene in the C Elegans genome build ce11

### Usage

```
data("ce11.GeneTSS")
```

#### **Format**

```
ce11.GeneTSS - a data frame with 20,051 genes and 5 variables:
```

```
chr Chromosome - character
```

start Start of the gene (TSS) - integer

end End of the gene - integer

Gene Gene symbol - character

strand Srand - character

# Value

ce11.GeneTSS - a data frame with 20,051 genes and 5 variables

changeRange

changeRange

# Description

changeRange

# Usage

```
changeRange(v, newmin = 1, newmax = 10)
```

# Arguments

v A numeric vector

newmin New min newmax New max

### Value

A matrix with values scaled between newmin and newmax

CheA3\_TF\_nTargets 15

CheA3\_TF\_nTargets

A data frame with the number of targets of each TF in ChEA3

# **Description**

This data.frame was obtained by downloading datasets from ChEA3 database (https://maayanlab.cloud/chea3/) and merging targets for :

- ARCHS4\_Coexpression
- ENCODE\_ChIP-seq
- Enrichr\_Queries
- GTEx\_Coexpression
- Literature\_ChIP-seq
- ReMap\_ChIP-seq

# Usage

```
data("CheA3_TF_nTargets")
```

#### **Format**

CheA3\_TF\_nTargets - a data.frame with 1632 rows (unique TFs) and 2 columns

### Value

A CheA3\_TF\_nTargets - a data.frame with 1632 rows (unique TFs) and 2 columns

# References

Keenan AB, Torre D, Lachmann A, Leong AK, Wojciechowicz M, Utti V, Jagodnik K, Kropiwnicki E, Wang Z, Ma'ayan A (2019) ChEA3: transcription factor enrichment analysis by orthogonal omics integration. Nucleic Acids Research. doi: 10.1093/nar/gkz446

The data.frame is composed of two columns:

- TF column containing the TF gene names (human)
- nTargets\_TF containing the number of targets for this TF in the combined database.

```
data("CheA3_TF_nTargets")
head(CheA3_TF_nTargets)
```

```
check_correct_datamatrix
```

Check if matrix rownames are well formated and correct if needed

# Description

Throws warnings / error if matrix is in the wrong format

# Usage

```
check_correct_datamatrix(datamatrix_single, sample_name = "")
```

# Arguments

### Value

A sparseMatrix in the right rownames format

```
choose_cluster_scExp Choose a number of clusters
```

#### **Description**

This functions takes as input a SingleCellExperiment object and a number of cluster to select. It outputs a SingleCellExperiment object with each cell assigned to a correlation cluster in colData. Also calculates a hierarchical clustering of the consensus associations calculated by Consensus-ClusterPlus.

# Usage

```
choose_cluster_scExp(
   scExp,
   nclust = 3,
   consensus = FALSE,
   hc_linkage = "ward.D"
)
```

# **Arguments**

scExp A SingleCellExperiment object containing consclust in metadata.

nclust Number of cluster to pick (3)

consensus Use consensus clustering results instead of simple hierarchical clustering? (FALSE)

hc\_linkage A linkage method for hierarchical clustering. See cor. ('ward.D')

choose\_perplexity 17

# Value

Returns a SingleCellExperiment object with each cell assigned to a correlation cluster in colData.

# **Examples**

```
data("scExp")
scExp_cf = correlation_and_hierarchical_clust_scExp(scExp)
scExp_cf = choose_cluster_scExp(scExp_cf,nclust=3,consensus=FALSE)
table(scExp_cf$cell_cluster)
scExp_cf = consensus_clustering_scExp(scExp)
scExp_cf_consensus = choose_cluster_scExp(scExp_cf,nclust=3,consensus=TRUE)
table(scExp_cf_consensus$cell_cluster)
```

choose\_perplexity

Choose perplexity depending on number of cells for Tsne

# **Description**

Choose perplexity depending on number of cells for Tsne

### Usage

```
choose_perplexity(dataset)
```

# Arguments

dataset

A matrix of features x cells (rows x columns)

#### Value

A number between 5 and 30 to use in Rtsne function

col2hex

Col2Hex

# Description

Transform character color to hexadecimal color code.

# Usage

```
col2hex(cname)
```

18 colors\_scExp

# **Arguments**

cname

Color name

#### Value

The HEX color code of a particular color

colors\_scExp

Adding colors to cells & features

# **Description**

Adding colors to cells & features

### Usage

```
colors_scExp(
  scExp,
  annotCol = "sample_id",
  color_by = "sample_id",
  color_df = NULL
)
```

# Arguments

scExp A SingleCellExperiment Object

annotCol Column names to color

color\_by If specifying color\_df, column names to color

color\_df Color data.frame to specify which color for which condition

# Value

A SingleCellExperiment with additionnal "color" columns in colData

```
data("scExp")
scExp = colors_scExp(scExp,annotCol = c("sample_id",
   "total_counts"),
   color_by = c("sample_id","total_counts"))

#Specific colors using a manually created data.frame :
color_df = data.frame(sample_id=unique(scExp$sample_id),
   sample_id_color=c("red","blue","green","yellow"))
scExp = colors_scExp(scExp,annotCol="sample_id",
   color_by="sample_id",color_df=color_df)
```

combine\_datamatrix 19

combine\_datamatrix

Combine two matrices and emit warning if no regions are in common

# **Description**

Combine two matrices and emit warning if no regions are in common

# Usage

```
combine_datamatrix(datamatrix, datamatrix_single, file_names, i)
```

# **Arguments**

### Value

A combined sparse matrix

```
combine_enrichmentTests
```

Run enrichment tests and combine into list

# **Description**

Run enrichment tests and combine into list

# Usage

```
combine_enrichmentTests(
   diff,
   enrichment_qval,
   qval.th,
   logFC.th,
   min.percent,
   annotFeat_long,
   peak_distance,
   refined_annotation,
   GeneSets,
   GeneSetsDf,
   GenePool,
   progress = NULL
)
```

### **Arguments**

diff Differential list

enrichment\_qval

Adusted p-value threshold above which a pathway is considered significative list

qval.th Differential analysis adjusted p.value threshold logFC.th Differential analysis log-fold change threshold

min.percent Minimum fraction of cells having the feature active to consider it as significantly

differential. (0.01)

annotFeat\_long Long annotation

peak\_distance Maximum gene to peak distance

refined\_annotation

Refined annotation data.frame if peak calling is done

GeneSets List of pathways

GeneSetsDf Data.frame of pathways

GenePool Pool of possible genes for testing

progress A shiny Progress instance to display progress bar.

#### Value

A list of list of pathway enrichment data.frames for Both / Over / Under and for each cluster

comparable\_variables Find comparable variable scExp

# Description

Find comparable variable scExp

# Usage

```
comparable_variables(scExp, allExp = TRUE)
```

# Arguments

scExp A SingleCellExperiment

allExp A logical indicating wether alternative experiments comparable variables should

also be fetch.

# Value

A character vector with the comparable variable names

CompareedgeRGLM 21

CompareedgeRGLM	Creates a summary table with the number of genes under- or overex- pressed in each group and outputs several graphical representations
	pressed in each group and outputs several graphical representations

# **Description**

Creates a summary table with the number of genes under- or overexpressed in each group and outputs several graphical representations

# Usage

```
CompareedgeRGLM(
  dataMat = NULL,
  annot = NULL,
  ref_group = NULL,
  groups = NULL,
  featureTab = NULL,
  norm_method = "TMMwsp"
)
```

# **Arguments**

dataMat	reads matrix
annot	selected annotation of interest
ref_group	List containing one or more vectors of reference samples. Name of the vectors will be used in the results table. The length of this list should be 1 or the same length as the groups list
groups	List containing the IDs of groups to be compared with the reference samples. Names of the vectors will be used in the results table
featureTab	Feature annotations to be added to the results table
norm_method	Which method to use for normalizing ('upperquantile')

# Value

A dataframe containing the foldchange and p.value of each feature

# Author(s)

Eric Letouze & Celine Vallot

```
data("scExp")
scExp_cf = correlation_and_hierarchical_clust_scExp(scExp)
scExp_cf = choose_cluster_scExp(scExp_cf,nclust=2,consensus=FALSE)
featureTab = as.data.frame(SummarizedExperiment::rowRanges(scExp_cf))
```

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```
rownames(featureTab) = featureTab$ID
ref_group = list("C1"=scExp_cf$cell_id[which(scExp_cf$cell_cluster=="C1")])
groups = list("C2"=scExp_cf$cell_id[which(scExp_cf$cell_cluster=="C2")])
myres = CompareedgeRGLM(as.matrix(SingleCellExperiment::counts(scExp_cf)),
annot=as.data.frame(SingleCellExperiment::colData(scExp_cf)),
    ref_group=ref_group,groups=groups, featureTab=featureTab)
```

CompareWilcox

**CompareWilcox** 

# **Description**

CompareWilcox

# Usage

```
CompareWilcox(
  dataMat = NULL,
  annot = NULL,
  ref_group = NULL,
  groups = NULL,
  featureTab = NULL,
  block = NULL,
  BPPARAM = BiocParallel::bpparam()
)
```

### **Arguments**

dataMat A raw count matrix

annot A cell annotation data.frame

ref\_group List with cells in reference group(s)
groups List with cells in group(s) to test
featureTab data.frame with feature annotation

block Use a blocking factor to conteract batch effect?

BPPARAM object for multiprocessing. See bpparam for more informations.

Will take the default BPPARAM set in your R session.

# Value

A dataframe containing the foldchange and p.value of each feature

# Author(s)

Eric Letouze & Celine Vallot & Pacome Prompsy

# **Examples**

```
data("scExp")
scExp_cf = correlation_and_hierarchical_clust_scExp(scExp)
scExp_cf = choose_cluster_scExp(scExp_cf,nclust=2,consensus=FALSE)
featureTab = as.data.frame(SummarizedExperiment::rowRanges(scExp_cf))
rownames(featureTab) = featureTab$ID
ref_group = list("C1"=scExp_cf$cell_id[which(scExp_cf$cell_cluster=="C1")])
groups = list("C2"=scExp_cf$cell_id[which(scExp_cf$cell_cluster=="C2")])
myres = CompareWilcox(as.matrix(SingleCellExperiment::normcounts(scExp_cf)),
annot=as.data.frame(SingleCellExperiment::colData(scExp_cf)),
    ref_group=ref_group,groups=groups, featureTab=featureTab)
```

```
concatenate_scBed_into_clusters
```

Concatenate single-cell BED into clusters

# **Description**

Concatenate single-cell BED into clusters

# Usage

```
concatenate_scBed_into_clusters(affectation, files_list, odir)
```

### **Arguments**

affectation	Annotation data frame containing cluster information
files_list	Named list of scBED file paths to concatenate. List Names must match affectation\$sample_id and basenames must match affectation\$barcode.
odir	Output directory to write concatenate pseudo-bulk BEDs.

### Value

Merge single-cell BED files into cluster BED files. Ungzip file if BED is gzipped.

```
consensus_clustering_scExp
```

Wrapper to apply ConsensusClusterPlus to scExp object

# **Description**

Runs consensus hierarchical clustering on PCA feature space of scExp object. Plot consensus scores for each number of clusters. See ConsensusClusterPlus - Wilkerson, M.D., Hayes, D.N. (2010). ConsensusClusterPlus: a class discovery tool with confidence assessments and item tracking. Bioinformatics, 2010 Jun 15;26(12):1572-3.

# Usage

```
consensus_clustering_scExp(
   scExp,
   prefix = NULL,
   maxK = 10,
   reps = 100,
   pItem = 0.8,
   pFeature = 1,
   distance = "pearson",
   clusterAlg = "hc",
   innerLinkage = "ward.D",
   finalLinkage = "ward.D",
   plot_consclust = "pdf",
   plot_icl = "png"
)
```

# Arguments

scExp	A SingleCellExperiment object containing 'PCA' in reducedDims.
prefix	character value for output directory. Directory is created only if plot_consclust is not NULL. This title can be an abosulte or relative path.
maxK	integer value. maximum cluster number to evaluate. (10)
reps	integer value. number of subsamples. (100)
pItem	numerical value. proportion of items to sample. (0.8)
pFeature	numerical value. proportion of features to sample. (1)
distance	character value. 'pearson': (1 - Pearson correlation), 'spearman' (1 - Spearman correlation), 'euclidean', 'binary', 'maximum', 'canberra', 'minkowski' or custom distance function. ('pearson')
clusterAlg	character value. cluster algorithm. 'he' heirarchical (hclust), 'pam' for paritioning around medoids, 'km' for k-means upon data matrix, 'kmdist' ('he') for k-means upon distance matrices (former km option), or a function that returns a clustering. ('he')

innerLinkage hierarchical linkage method for subsampling. ('ward.D')

finalLinkage hierarchical linkage method for consensus matrix. ('ward.D')

plot\_consclust character value. NULL - print to screen, 'pdf', 'png', 'pngBMP' for bitmap png, helpful for large datasets. ('pdf')

plot\_icl same as above for item consensus plot. ('png')

#### **Details**

This functions takes as input a SingleCellExperiment object that must have 'PCA' in reducedDims and outputs a SingleCellExperiment object containing consclust list calculated cluster consensus and item consensus scores in metadata.

#### Value

Returns a SingleCellExperiment object containing consclust list, calculated cluster consensus and item consensus scores in metadata.

#### References

ConsensusClusterPlus package by Wilkerson, M.D., Hayes, D.N. (2010). ConsensusClusterPlus: a class discovery tool with confidence assessments and item tracking. Bioinformatics, 2010 Jun 15;26(12):1572-3.

# **Examples**

```
data("scExp")
scExp_cf = correlation_and_hierarchical_clust_scExp(scExp)
scExp_cf = consensus_clustering_scExp(scExp)
```

```
correlation_and_hierarchical_clust_scExp

Correlation and hierarchical clustering
```

# Description

Calculates cell to cell correlation matrix based on the PCA feature space and runs hierarchical clustering taking 1 - correlation scores as distance.

### Usage

```
correlation_and_hierarchical_clust_scExp(scExp, hc_linkage = "ward.D")
```

# **Arguments**

```
scExp A SingleCellExperiment object, containing 'PCA' in reducedDims.
hc_linkage A linkage method for hierarchical clustering. See cor. ('ward.D')
```

26 count\_coverage

# **Details**

This functions takes as input a SingleCellExperiment object that must have PCA calculated and outputs a SingleCellExperiment object with correlation matrix and hierarchical clustering.

#### Value

Return a SingleCellExperiment object with correlation matrix & hiearchical clustering.

# **Examples**

```
data("scExp")
scExp_cf = correlation_and_hierarchical_clust_scExp(scExp)
```

count\_coverage

Create a smoothed and normalized coverage track from a BAM file and given a bin GenomicRanges object (same as deepTools bamCoverage)

# **Description**

Normalization is CPM, smoothing is done by averaging on n smoothBin regions left and right of any given region.

### Usage

```
count_coverage(
  input,
  format = "BAM",
 bins,
  canonical_chr,
  norm_factor,
  n_{smoothBin} = 5,
  ref = "hg38",
  read_size = 101,
  original_bins = NULL
)
```

## **Arguments**

input

Either a named list of character vector of path towards single-cell BED files or a sparse raw matrix of small bins («500bp). If a named list specifying scBEDn the names MUST correspond to the 'sample\_id' column in your SingleCellExperiment object. The single-cell BED files names MUST match the barcode names in your SingleCellExperiment (column 'barcode'). The scBED files can be gzipped or not.

File format, either "BAM" or "BED"

bins A GenomicRanges object of binned genome

format

create\_project\_folder 27

canonical\_chr GenomicRanges of the chromosomes to read the BAM file.

norm\_factor Then number of cells or total number of reads in the given sample, for normal-

ization.

n\_smoothBin Number of bins left and right to smooth the signal.

ref Genomic reference read\_size Length of the reads

original\_bins Original bins GenomicRanges in case the format is raw

matrix.

# Value

A binned GenomicRanges that can be readily exported into bigwig file.

```
create_project_folder Create ChromSCape project folder
```

### **Description**

Creates a project folder that will be recognizable by ChromSCape Shiny application.

# Usage

```
create_project_folder(
  output_directory,
  analysis_name = "Analysis_1",
  ref_genome = c("hg38", "mm10", "ce11")[1]
)
```

#### **Arguments**

output\_directory

Path towards the directory to create the 'ChromSCape\_Analyses' folder and the analysis subfolder. If this path already contains the 'ChromSCape\_Analyses'

folder, will only create the analysis subfolder.

analysis\_name Name of the analysis. Must only contain alphanumerical characters or '\_'.

ref\_genome Reference genome, either 'hg38' or 'mm10'.

#### Value

Creates the project folder and returns the root of the project.

```
dir = tempdir()
create_project_folder(output_directory = dir,
    analysis_name = "Analysis_1")
list.dirs(file.path(dir))
```

28 create\_scDataset\_raw

```
create_sample_name_mat
```

Create a sample name matrix

# Description

Create a sample name matrix

# Usage

```
create_sample_name_mat(nb_samples, samples_names)
```

# Arguments

```
nb_samples Number of samples
samples_names Character vector of sample names
```

#### Value

A matrix

# Description

Create a simulated single cell datamatrix & cell annotation

# Usage

```
create_scDataset_raw(
  cells = 300,
  features = 600,
  featureType = c("window", "peak", "gene"),
  sparse = TRUE,
  nsamp = 4,
  ref = "hg38",
  batch_id = factor(rep(1, nsamp))
)
```

create\_scExp 29

# Arguments

cells Number of cells (300)

features Number of features (600)

featureType Type of feature (window)

sparse Is matrix sparse ? (TRUE)

nsamp Number of samples (4)

ref Reference genome ('hg38')

batch\_id Batch\_origin (factor((1,1,1,1)))

### Value

A list composed of \* mat : a sparse matrix following an approximation of the negative binomial law (adapted to scChIPseq) \* annot : a data.frame of cell annotation \* batches : an integer vector with the batch number for each cell

### **Examples**

```
# Creating a basic sparse 600 genomic bins x 300 cells matrix and annotation
1 = create_scDataset_raw()
head(1$mat)
head(1$annot)
head(1$batches)
# Specifying number of cells, features and samples
12 = create_scDataset_raw(cells = 500, features = 500, nsamp=2)
# Specifying species
mouse_1 = create_scDataset_raw(ref="mm10")
# Specifying batches
batch_1 = create_scDataset_raw(nsamp=4, batch_id = factor(c(1,1,2,2)))
# Peaks of different size as features
peak_1 = create_scDataset_raw(featureType="peak")
head(peak_1$mat)
# Genes as features
gene_l = create_scDataset_raw(featureType="gene")
head(gene_1$mat)
```

create\_scExp Wrapper to create the single cell experiment from count matrix and

feature dataframe

30 create\_scExp

# **Description**

Create the single cell experiment from (sparse) datamatrix and feature dataframe containing feature names and location. Also optionally removes zero count Features, zero count Cells, non canconical chromosomes, and chromosome M. Calculates QC Metrics (scran).

# Usage

```
create_scExp(
  datamatrix,
  annot,
  remove_zero_cells = TRUE,
  remove_zero_features = TRUE,
  remove_non_canonical = TRUE,
  remove_chr_M = TRUE,
  mainExpName = "main",
  verbose = TRUE
)
```

### **Arguments**

```
A matrix or sparseMatrix of raw counts. Features x Cells (rows x columns).
datamatrix
annot
                 A data frame containing informations on cells. Should have the same number of
                 rows as the number of columns in datamatrix.
remove_zero_cells
                 remove cells with zero counts? (TRUE)
remove_zero_features
                 remove cells with zero counts? (TRUE)
remove_non_canonical
                 remove non canonical chromosomes ?(TRUE)
remove_chr_M
                 remove chromosomes M? (TRUE)
mainExpName
                 Name of the mainExpName e.g. 'bins', 'peaks'... ("default")
verbose
                 (TRUE)
```

### Value

Returns a SingleCellExperiment object.

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp</pre>
```

DA\_custom 31

DA_custom $Di$	ferential Analysis Custom in 'One vs One' mode
----------------	--

# Description

Differential Analysis Custom in 'One vs One' mode

# Usage

```
DA_custom(
   affectation,
   by,
   counts,
   method,
   feature,
   block,
   ref,
   group,
   progress = NULL,
   BPPARAM = BiocParallel::bpparam()
)
```

# **Arguments**

affectation	An annotation data.frame with cell_id and
by	= A character specifying the column of the object containing the groups of cells to compare.
counts	Count matrix
method	DA method: Wilcoxon or EdgeR
feature	Feature tables
block	Blocking feature
ref	If de_type is custom, the reference to compare (data.frame), must be a one-column data.frame with cell_clusters or sample_id as character in rows
group	If de_type is custom, the group to compare (data.frame), must be a one-column data.frame with cell_clusters or sample_id as character in rows
progress	A shiny Progress instance to display progress bar.
BPPARAM	BPPARAM object for multiprocessing. See bpparam for more informations.

Will take the default BPPARAM set in your R session.

# Value

A list of results, groups compared and references

DA\_one\_vs\_rest

DA\_one\_vs\_rest

Differential Analysis in 'One vs Rest' mode

# **Description**

Differential Analysis in 'One vs Rest' mode

# Usage

```
DA_one_vs_rest(
   affectation,
   by,
   counts,
   method,
   feature,
   block,
   progress = NULL,
   BPPARAM = BiocParallel::bpparam()
)
```

# **Arguments**

affectation An annotation data.frame with cell\_id and cell\_cluster columns

by = A character specifying the column of the object containing the groups of cells

to compare.

counts Count matrix

method DA method: Wilcoxon or EdgeR

feature Feature tables

block Blocking feature

progress A shiny Progress instance to display progress bar.

BPPARAM object for multiprocessing. See bpparam for more informations.

Will take the default BPPARAM set in your R session.

### Value

A list of results, groups compared and references

DA\_pairwise 33

DA_pairwise	Run differential analysis in Pairwise mode

# **Description**

Run differential analysis in Pairwise mode

# Usage

```
DA_pairwise(
   affectation,
   by,
   counts,
   method,
   feature,
   block,
   progress = NULL,
   BPPARAM = BiocParallel::bpparam()
)
```

# **Arguments**

affectation An annotation data.frame with cell\_cluster and cell\_id columns

by = A character specifying the column of the object containing the groups of cells

to compare.

counts Count matrix

method DA method, Wilcoxon or edgeR

feature Feature data.frame block Blocking feature

progress A shiny Progress instance to display progress bar.

BPPARAM object for multiprocessing. See bpparam for more informations.

Will take the default BPPARAM set in your R session.

### Value

A list of results, groups compared and references

34 detect\_samples

define\_feature

Define the features on which reads will be counted

### **Description**

Define the features on which reads will be counted

### Usage

```
define_feature(ref = c("hg38","mm10", "ce11")[1],
  peak_file = NULL,
  bin_width = NULL,
  genebody = FALSE,
  extendPromoter = 2500)
```

# **Arguments**

ref Reference genome

peak\_file A bed file if counting on peaks

bin\_width A number of bins if divinding genome into fixed width bins

genebody A logical indicating if feature should be counted in genebodies and promoter.

extendPromoter Extension length before TSS (2500).

# Value

A GRanges object

# **Examples**

```
gr_bins = define_feature("hg38", bin_width = 50000)
gr_genes = define_feature("hg38", genebody = TRUE, extendPromoter = 5000)
```

detect\_samples

Heuristic discovery of samples based on cell labels

# Description

Identify a fixed number of common string (samples) in a set of varying strings (cells). E.g. in the set "Sample1\_cell1", "Sample1\_cell2", "Sample2\_cell1", "Sample2\_cell2" and with nb\_samples=2, the function returns "Sample1", "Sample1", "Sample2", "Sample2".

# Usage

```
detect_samples(barcodes, nb_samples = 1)
```

differential\_activation 35

# Arguments

barcodes Vector of cell barcode names (e.g. Sample1\_cell1, Sample1\_cell2...)

nb\_samples Number of samples to find

### Value

character vector of sample names the same length as cell labels

# **Examples**

```
barcodes = c(paste0("HBCx22_BC_",seq_len(100)),
paste0("mouse_sample_XX",208:397))
samples = detect_samples(barcodes, nb_samples=2)
```

differential\_activation

Find Differentialy Activated Features (One vs All)

# **Description**

Based on the statement that single-cell epigenomic dataset are very sparse, specifically when analysis small bins or peaks, we can define each feature as being 'active' or not simply by the presence or the absence of reads in this feature. This is the equivalent of binarize the data. When trying to find differences in signal for a feature between multiple cell groups, this function simply compare the percentage of cells 'activating' the feature in each of the group. The p.values are then calculated using a Pearson's Chi-squared Test for Count Data (comparing the number of active cells in one group vs the other) and corrected using Benjamini-Hochberg correction for multiple testing.

# Usage

```
differential_activation(
   scExp,
   by = c("cell_cluster", "sample_id")[1],
   verbose = TRUE,
   progress = NULL
)
```

#### **Arguments**

scExp A SingleCellExperiment object containing consclust with selected number of

cluster.

by Which grouping to run the marker enrichment?

verbose Print?

progress A shiny Progress instance to display progress bar.

#### **Details**

To calculate the logFC, the percentage of activation of the features are corrected for total number of reads to correct for library size bias. For each cluster ('group') the function consider the rest of the cells as the reference.

#### Value

Returns a dataframe of differential activation results that contains the rowData of the SingleCellExperiment with additional logFC, q.value, group activation (fraction of cells active for each feature in the group cells), reference activation (fraction of cells active for each feature in the reference cells).

# See Also

For Pearson's Chi-squared Test for Count Data chisq.test. For other differential analysis see differential\_analysis\_scExp.

# **Examples**

```
data("scExp")
res = differential_activation(scExp, by = "cell_cluster")
res = differential_activation(scExp, by = "sample_id")

differential_analysis_scExp
```

Runs differential analysis between cell clusters

# **Description**

Based on clusters of cell defined previously, runs non-parametric Wilcoxon Rank Sum test to find significantly depleted or enriched features, in 'one\_vs\_rest' mode or 'pairwise' mode. In pairwise mode, each cluster is compared to all other cluster individually, and then pairwise comparisons between clusters are combined to find overall differential features using combineMarkers function from scran.

# Usage

```
differential_analysis_scExp(
    scExp,
    de_type = c("one_vs_rest_fast", "one_vs_rest", "pairwise", "custom")[1],
    by = "cell_cluster",
    method = "wilcox",
    block = NULL,
    group = NULL,
    ref = NULL,
    prioritize_genes = nrow(scExp) > 20000,
    max_distanceToTSS = 1000,
    progress = NULL,
    BPPARAM = BiocParallel::bpparam()
)
```

### **Arguments**

scExp	A SingleCellExperiment object containing consclust with selected number of cluster.	
de_type	Type of comparisons. Either 'one_vs_rest', to compare each cluster against all others, or 'pairwise' to make 1 to 1 comparisons. ('one_vs_rest')	
by	= A character specifying the column of the object containing the groups of cells to compare. Exclusive with de_type == custom	
method	Differential testing method, either 'wilcox' for Wilcoxon non- parametric testing or 'neg.binomial' for edgerGLM based testing. ('wilcox')	
block	Use batches as blocking factors? If TRUE, block will be taken as the column "batch_id" from the SCE. Cells will be compared only within samples belonging to the same batch.	
group	If de_type = "custom", the sample / cluster of interest as a one- column data.frame. The name of the column is the group name and the values are character either cluster ("C1", "C2",) or sample_id.	
ref	If de_type = "custom", the sample / cluster of reference as a one- column data.frame. The name of the column is the group name and the values are character either cluster ("C1", "C2",) or sample_id.	
prioritize_genes		
	First filter by loci being close to genes ? E.g. for differential analysis, it is more relevant to keep features close to genes	
max_distanceToTSS		
	If prioritize_genes is TRUE, the maximum distance to consider a feature close to a gene.	

# Details

progress

**BPPARAM** 

This functions takes as input a SingleCellExperiment object with consclust, the type of comparison, either 'one\_vs\_rest' or 'pairwise', the adjusted p-value threshold (qval.th) and the fold-change threshold (logFC.th). It outputs a SingleCellExperiment object containing a differential list.

BPPARAM object for multiprocessing. See bpparam for more informations.

A shiny Progress instance to display progress bar.

Will take the default BPPARAM set in your R session.

# Value

Returns a SingleCellExperiment object containing a differential list.

```
data("scExp")
scExp_cf = differential_analysis_scExp(scExp)
```

38 enrichmentTest

distPearson

distPearson

# Description

distPearson

# Usage

distPearson(m)

# Arguments

m

A matrix

# Value

A dist object

enrichmentTest

enrichment Test

# Description

enrichmentTest

# Usage

```
enrichmentTest(gene.sets, mylist, possibleIds, sep = ";", silent = FALSE)
```

# **Arguments**

gene.sets A list of reference gene sets

mylist A list of genes to test possibleIds All existing genes

sep Separator used to collapse genes

silent Silent mode?

### Value

A dataframe with the gene sets and their enrichment p.value

enrich\_TF\_ChEA3\_genes Find the TF that are enriched in the differential genes using ChEA3 API

# **Description**

Find the TF that are enriched in the differential genes using ChEA3 API

### Usage

```
enrich_TF_ChEA3_genes(genes)
```

#### **Arguments**

genes

A character vector with the name of genes to enrich for TF.

#### Value

Returns a SingleCellExperiment object containing list of enriched Gene Sets for each cluster, either in depleted features, enriched features or simply differential features (both).

#### References

Keenan AB, Torre D, Lachmann A, Leong AK, Wojciechowicz M, Utti V, Jagodnik K, Kropiwnicki E, Wang Z, Ma'ayan A (2019) ChEA3: transcription factor enrichment analysis by orthogonal omics integration. Nucleic Acids Research. doi: 10.1093/nar/gkz446 +

### **Examples**

```
data(scExp)
enrich_TF_ChEA3_genes(head(unlist(strsplit(SummarizedExperiment::rowData(scExp)$Gene, split = ",", fixed = TRUE)
```

enrich\_TF\_ChEA3\_scExp Find the TF that are enriched in the differential genes using ChEA3 database

### **Description**

Find the TF that are enriched in the differential genes using ChEA3 database

# Usage

```
enrich_TF_ChEA3_scExp(
    scExp,
    qval.th = 0.01,
    logFC.th = 1,
    min.percent = 0.01,
    peak_distance = 1000,
    use_peaks = FALSE,
    progress = NULL,
    verbose = TRUE
)
```

# **Arguments**

scExp	A SingleCellExperiment object containing list of differential features.
SCLXP	A Single Centraperiment object containing list of differential readures.
qval.th	Adjusted p-value threshold to define differential features. (0.01)
logFC.th	Fold change threshold to define differential features. (1)
min.percent	Minimum fraction of cells having the feature active to consider it as significantly differential. $(0.01)$
peak_distance	Maximum distance ToTSS of feature to gene TSS to consider associated, in bp. $(1000)$
use_peaks	Use peak calling method (must be calculated beforehand). (FALSE)
progress	A shiny Progress instance to display progress bar.
verbose	A logical to print message or not. (TRUE)

# Value

Returns a SingleCellExperiment object containing list of enriched Gene Sets for each cluster, either in depleted features, enriched features or simply differential features (both).

```
data("scExp")
scExp = enrich_TF_ChEA3_scExp(
    scExp,
    qval.th = 0.01,
    logFC.th = 1,
    min.percent = 0.01)
```

```
exclude_features_scExp
```

Remove specific features (CNA, repeats)

# **Description**

Remove specific features (CNA, repeats)

# Usage

```
exclude_features_scExp(
   scExp,
   features_to_exclude,
   by = "region",
   verbose = TRUE
)
```

### **Arguments**

```
scExp A SingleCellExperiment object.

features_to_exclude

A GenomicRanges object or data.frame containing genomic regions or features to exclude or path towards a BED file containing the features to exclude.

by Type of features. Either 'region' or 'feature_name'. If 'region', will look for genomic coordinates in columns 1-3 (chr,start,stop). If 'feature_name', will look for a genes in first column. ('region')

verbose (TRUE)
```

#### Value

A SingleCellExperiment object without features to exclude.

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
features_to_exclude = data.frame(chr=c("chr4","chr7","chr17"),
start=c(50000,8000000,2000000),
end=c(100000,16000000,2500000))
features_to_exclude = as(features_to_exclude,"GRanges")
scExp = exclude_features_scExp(scExp,features_to_exclude)
scExp</pre>
```

```
feature_annotation_scExp
```

Add gene annotations to features

# **Description**

Add gene annotations to features

### Usage

```
feature_annotation_scExp(scExp, ref = "hg38", reference_annotation = NULL)
```

### **Arguments**

scExp A SingleCellExperiment object.

ref Reference genome. Either 'hg38', 'mm10' or 'ce11'. ('hg38')

reference\_annotation

A data frame containing gene (or else) annotation with genomic coordinates.

### Value

A SingleCellExperiment object with annotated rowData.

### **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp = feature_annotation_scExp(scExp)
head(SummarizedExperiment::rowRanges(scExp))

# Mouse
raw = create_scDataset_raw(ref = "mm10")
scExp = create_scExp(raw$mat, raw$annot)
scExp = feature_annotation_scExp(scExp,ref="mm10")
head(SummarizedExperiment::rowRanges(scExp))</pre>
```

```
filter_correlated_cell_scExp
```

Filter lowly correlated cells

# **Description**

Remove cells that have a correlation score lower than what would be expected by chance with other cells.

### **Usage**

```
filter_correlated_cell_scExp(scExp, random_iter = 5,
corr_threshold = 99, percent_correlation = 1,
downsample = 2500, verbose = TRUE, n_process = 250,
BPPARAM = BiocParallel::bpparam())
```

### **Arguments**

A SingleCellExperiment object containing 'Cor', a correlation matrix, in rescExp ducedDims. Number of random matrices to create to calculate random correlation scores. random\_iter (50)corr\_threshold Quantile of random correlation score above which a cell is considered to be 'correlated' with another cell. (99) percent\_correlation Percentage of the cells that any cell must be 'correlated' to in order to not be filtered. (1) downsample Number of cells to calculate correlation filtering threshold? (2500) Print messages ? (TRUE) verbose Number of cell to proceed at a time. Increase this number to increase speed at n\_process memory cost BPPARAM object for multiprocessing. See bpparam for more informations. **BPPARAM** 

#### **Details**

This functions takes as input a SingleCellExperiment object that must have correlation matrix calculated and outputs a SingleCellExperiment object without lowly correlated cells. TSNE is recalculated.

Will take the default BPPARAM set in your R session.

#### Value

Returns a SingleCellExperiment object without lowly correlated cells. The calculated correlation score limit threshold is saved in metadata.

```
data("scExp")
dim(scExp)
scExp_cf = filter_correlated_cell_scExp(scExp,
corr_threshold = 99, percent_correlation = 1)
dim(scExp_cf)
```

filter\_scExp

```
filter_genes_with_refined_peak_annotation

Filter genes based on peak calling refined annotation
```

# **Description**

Filter genes based on peak calling refined annotation

# Usage

```
filter_genes_with_refined_peak_annotation(
  refined_annotation,
  peak_distance,
  signific,
  over,
  under
)
```

# **Arguments**

refined\_annotation

A data.frame containing each gene distance to real peak

peak\_distance Minimum distance to an existing peak to accept a given gene

signific Indexes of all significantly differential genes
over Indexes of all significantly overexpressed genes
under Indexes of all significantly underexpressed genes

#### Value

List of significantly differential, overexpressed and underexpressed genes close enough to existing peaks

filter\_scExp Filter cells and features

# **Description**

Function to filter out cells & features from SingleCellExperiment based on total count per cell, number of cells 'ON' in features and top covered cells that might be doublets.

### Usage

```
filter_scExp(
   scExp,
   min_cov_cell = 1600,
   quant_removal = 95,
   min_count_per_feature = 10,
   verbose = TRUE
)
```

### **Arguments**

#### Value

Returns a filtered SingleCellExperiment object.

### **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp. = filter_scExp(scExp)

# No feature filtering (all features are valuable)
scExp. = filter_scExp(scExp,min_count_per_feature=30)

# No cell filtering (all features are valuable)
scExp. = filter_scExp(scExp,min_cov_cell=0,quant_removal=100)</pre>
```

```
find_clusters_louvain_scExp
```

Build SNN graph and find cluster using Louvain Algorithm

### **Description**

Build SNN graph and find cluster using Louvain Algorithm

46 find\_top\_features

### Usage

```
find_clusters_louvain_scExp(
    scExp,
    k = 10,
    resolution = 1,
    use.dimred = "PCA",
    type = c("rank", "number", "jaccard")[3],
    BPPARAM = BiocParallel::bpparam()
)
```

# **Arguments**

A SingleCellExperiment with PCA calculated

An integer scalar specifying the number of nearest neighbors to consider during graph construction.

A numeric specifying the resolution of clustering to pass to igraph::cluster\_louvain function.

A string specifying the dimensionality reduction to use.

A string specifying the type of weighting scheme to use for shared neighbors.

BPPARAM BPPARAM object for multiprocessing. See bpparam for more informations.

Will take the default BPPARAM set in your R session.

#### Value

A SingleCellExperiment containing the vector of clusters (named C1, C2 ....)

### **Examples**

```
data('scExp')
scExp = find_clusters_louvain_scExp(scExp, k = 10)
```

find\_top\_features
Find most covered features

# **Description**

Find the top most covered features that will be used for dimensionality reduction. Optionally remove non-top features.

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# Usage

```
find_top_features(
   scExp,
   n = 20000,
   keep_others = FALSE,
   prioritize_genes = FALSE,
   max_distanceToTSS = 10000,
   verbose = TRUE
)
```

### Arguments

scExp A SingleCellExperiment.

n Either an integer indicating the number of top covered regions to find or a char-

acter vector of the top percentile of features to keep (e.g. 'q20' to keep top 20%

features).

keep\_others Logical indicating if non-top regions are to be removed from the SCE or not

(FALSE).

prioritize\_genes

First filter by loci being close to genes? E.g. for differential analysis, it is more

relevant to keep features close to genes

max\_distanceToTSS

If prioritize\_genes is TRUE, the maximum distance to consider a feature close

to a gene.

verbose Print?

### Value

A SCE with top features

# **Examples**

```
data(scExp)
scExp_top = find_top_features(scExp, n = 4000, keep_others = FALSE)
```

generate\_analysis

Generate a complete ChromSCape analysis

### Description

Generate a complete ChromSCape analysis

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#### Usage

```
generate_analysis(input_data_folder,
   analysis_name = "Analysis_1",
   output_directory = "./",
    input_data_type = c("scBED", "DenseMatrix", "SparseMatrix", "scBAM")[1],
    feature_count_on = c("bins", "genebody", "peaks")[1],
    feature_count_parameter = 50000,
    rebin_sparse_matrix = FALSE,
   ref_genome = c("hg38","mm10", "ce11")[1],
    run = c("filter", "CNA","cluster", "consensus", "coverage",
    "DA", "GSA", "report")[c(1,3,5,6,7,8)],
   min_reads_per_cell = 1000,
   max_quantile_read_per_cell = 99,
    n_{top_features} = 40000,
   norm_type = "CPM",
    subsample_n = NULL,
   exclude_regions = NULL,
   n_{clust} = NULL,
   corr_threshold = 99,
   percent_correlation = 1,
   maxK = 10,
   qval.th = 0.1,
   logFC.th = 1,
   enrichment_qval = 0.1,
   doBatchCorr = FALSE,
   batch_sels = NULL,
   control_samples_CNA = NULL,
   genes_to_plot = c("Krt8","Krt5","Tgfb1", "Foxq1", "Cdkn2b",
                      "Cdkn2a", "chr7:15000000-20000000")
   )
Arguments
    input_data_folder
                    Directory containing the input data.
                    Name given to the analysis.
    analysis_name
    output_directory
                    Directory where to create the analysis and the HTML report.
    input_data_type
                    The type of input data.
    feature_count_on
                    For raw data type, on which features to count the cells.
    feature_count_parameter
```

Additional parameter corresponding to the 'feature\_count\_on' parameter. E.g. for 'bins' must be a numeric, e.g. 50000, for 'peaks' must be a character containing path towards a BED peak file.

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rebin\_sparse\_matrix

A boolean specifying if the SparseMatrix should be rebinned on features (see

feature\_count\_on and feature\_count\_parameter).

ref\_genome The genome of reference.

run What steps to run. By default runs everything. Some steps are required in order

to run downstream steps.

min\_reads\_per\_cell

Minimum number of reads per cell.

max\_quantile\_read\_per\_cell

Upper quantile above which to consider cells doublets.

n\_top\_features Number of features to keep in the analysis.

norm\_type Normalization type.

subsample\_n Number of cells per condition to downsample to, for performance principally.

exclude\_regions

Path towards a BED file containing CNA to exclude from the analysis (optional).

n\_clust Number of clusters to force choice of clusters.

corr\_threshold Quantile of correlation above which two cells are considered as correlated.

percent\_correlation

Percentage of the total cells that a cell must be correlated with in order to be

kept in the analysis.

maxK Upper cluster number to rest for ConsensusClusterPlus.

qval.th Adjusted p-value below which to consider features differential.

logFC.th Log2-fold-change above/below which to consider a feature depleted/enriched.

enrichment\_qval

Adjusted p-value below which to consider a gene set as significantly enriched in

differential features.

doBatchCorr Logical indicating if batch correction using fastMNN should be run.

batch\_sels If doBatchCorr is TRUE, a named list containing the samples in each batch.

control\_samples\_CNA

If running CopyNumber Analysis, a character vector of the sample names that

are 'normal'.

genes\_to\_plot A character vector containing genes of interest of which to plot the coverage.

### Value

Creates a ChromSCape-readable directory and saved objects, as well as a multi-tabbed HTML report resuming the analysis.

```
## Not run:
generate_analysis("/path/to/data/", "Analysis_1")
## End(Not run)
```

generate\_count\_matrix Generate count matrix

### **Description**

Generate count matrix

### Usage

```
generate_count_matrix(cells, features, sparse, cell_names, feature_names)
```

### **Arguments**

cells Number of cells
features Number of features
sparse Is matrix sparse?
cell\_names Cell names

feature\_names Feature names

#### Value

A matrix or a sparse matrix

generate\_coverage\_tracks

Generate cell cluster pseudo-bulk coverage tracks

# **Description**

Generate cell cluster pseudo-bulk coverage tracks. First, scBED files are concatenated into cell clusters contained in the 'by' column of your SingleCellExperiment object. To do so, for each sample in the given list, the barcodes of each cluster are grepped and BED files are merged into pseudo-bulk of clusters (C1,C2...). Two cells from different can have the same barcode ID as cell affectation is done sample by sample. Then coverage of pseudo-bulk BED files is calculated by averaging & smoothing reads on small genomic window (150bp per default). The pseudo bulk BED and BigWigs coverage tracks are writtend to the output directory. This functionality is not available on Windows as it uses the 'cat' and 'gzip' utilities from Unix OS.

### Usage

```
generate_coverage_tracks(
   scExp_cf,
   input,
   odir,
   format = "scBED",
   ref_genome = c("hg38", "mm10", "ce11")[1],
   bin_width = 150,
   n_smoothBin = 5,
   read_size = 101,
   quantile_for_peak_calling = 0.85,
   by = "cell_cluster",
   progress = NULL
)
```

#### **Arguments**

	scExp_cf	A SingleCellExperiment with cluster selected.	(see choose cluster scExp).
--	----------	---	-----------------------------

It is recommended having a minimum of ~100 cells per cluster in order to obtain

smooth tracks.

input Either a named list of character vector of path towards single-cell BED files or

a sparse raw matrix of small bins («500bp). If a named list specifying scBED the names MUST correspond to the 'sample\_id' column in your SingleCellExperiment object. The single-cell BED files names MUST match the barcode names in your SingleCellExperiment (column 'barcode'). The scBED files can

be gzipped or not.

odir The output directory to write the cumulative BED and BigWig files.

format File format, either "raw\_mat", "BED" or "BAM"

ref\_genome The genome of reference, used to constrain to canonical chromosomes. Either

'hg38' or 'mm10'. 'hg38' per default.

bin\_width The width of the bin to create the coverage track. The smaller the greater the

resolution & runtime. Default to 150.

n\_smoothBin Number of bins left & right to average ('smooth') the signal on. Default to 5.

read\_size The estimated size of reads. Default to 101.

quantile\_for\_peak\_calling

The quantile to define the threshold above which signal is considered as a peak.

by A character specifying a categorical column of scExp\_cf metadata by which to

group cells and generate coverage tracks and peaks.

progress A Progress object for Shiny. Default to NULL.

#### Value

Generate coverage tracks (.bigwig) for each group in the SingleCellExperiment "by" column.

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### **Examples**

```
## Not run:
data(scExp)
input_files_coverage = list(
   "scChIP_Jurkat_K4me3" = paste0("/path/to/",scExp$barcode[1:51],".bed"),
   "scChIP_Ramos_K4me3" = paste0("/path/to/",scExp$barcode[52:106],".bed")
)
generate_coverage_tracks(scExp, input_files_coverage, "/path/to/output",
ref_genome = "hg38")
## End(Not run)
```

generate\_feature\_names

Generate feature names

# **Description**

Generate feature names

### Usage

```
generate_feature_names(featureType, ref, features)
```

# Arguments

featureType Type of feature

ref Reference genome

features Number of features to generate

### Value

A character vector of feature names

generate\_report

From a ChromSCape analysis directory, generate an HTML report.

# Description

From a ChromSCape analysis directory, generate an HTML report.

### **Usage**

```
generate_report(
   ChromSCape_directory,
   prefix = NULL,
   run = c("filter", "CNA", "cluster", "consensus", "peak_call", "coverage", "DA", "GSA",
        "report")[c(1, 3, 6, 7, 8, 9)],
   genes_to_plot = c("Krt8", "Krt5", "Tgfb1", "Foxq1", "Cdkn2b", "Cdkn2a",
        "chr7:15000000-200000000"),
   control_samples_CNA = NULL
)
```

### **Arguments**

ChromSCape\_directory

Path towards the ChromSCape directory of which you want to create the report.

The report will be created at the root of this directory.

prefix Name of the analysis with the filtering parameters (e.g. Analysis\_3000\_100000\_99\_uncorrected).

You will find the prefix in the Filtering Normalize Reduce subfolder.

run Which steps to report ("filter", "CNA", "cluster", "consensus", "peak\_call", "cov-

erage", "DA", "GSA", "report"). Only indicate steps that were done in the anal-

ysis. By default do not report CNA, consensus and peak calling.

genes\_to\_plot For the UMAP, which genes do you want to see in the report.

control\_samples\_CNA

If running the Copy Number Alteration (CNA) part, which samples are the con-

trols

#### Value

Generate an HTML report at the root of the analysis directory.

# **Examples**

```
## Not run:
generate_analysis("/path/to/data/", "Analysis_1")
## End(Not run)
```

```
gene_set_enrichment_analysis_scExp
```

Runs Gene Set Enrichment Analysis on genes associated with differential features

# **Description**

This function takes previously calculated differential features and runs hypergeometric test to look for enriched gene sets in the genes associated with differential features, for each cell cluster. This functions takes as input a SingleCellExperiment object with consclust, the type of comparison, either 'one\_vs\_rest' or 'pairwise', the adjusted p-value threshold (qval.th) and the fold-change threshold (logFC.th). It outputs a SingleCellExperiment object containing a differential list.

# Usage

# Arguments

scExp	A SingleCellExperiment object containing list of differential features.	
enrichment_qval		
	Adjusted p-value threshold for gene set enrichment. (0.1)	
ref	A reference annotation, either 'hg38', 'mm10', 'ce11'. ('hg38')	
GeneSets	A named list of gene sets. If NULL will automatically load MSigDB list of gene sets for specified reference genome. (NULL)	
GeneSetsDf	A dataframe containing gene sets & class of gene sets. If NULL will automatically load MSigDB dataframe of gene sets for specified reference genome. (NULL)	
GenePool	The pool of genes to run enrichment in. If NULL will automatically load Gencode list of genes fro specified reference genome. (NULL)	
qval.th	Adjusted p-value threshold to define differential features. (0.01)	
logFC.th	Fold change threshold to define differential features. (1)	
min.percent	Minimum fraction of cells having the feature active to consider it as significantly differential. $(0.01)$	
peak_distance	Maximum distance ToTSS of feature to gene TSS to consider associated, in bp. $(1000)$	
use_peaks	Use peak calling method (must be calculated beforehand). (FALSE)	

getExperimentNames 55

```
GeneSetClasses Which classes of MSIGdb to look for.
```

progress A shiny Progress instance to display progress bar.

#### Value

Returns a SingleCellExperiment object containing list of enriched Gene Sets for each cluster, either in depleted features, enriched features or simply differential features (both).

# **Examples**

```
data("scExp")
#Usually recommanding qval.th = 0.01 & logFC.th = 1 or 2
## Not run: scExp_cf = gene_set_enrichment_analysis_scExp(scExp,
    qval.th = 0.4, logFC.th = 0.3)
## End(Not run)
```

getExperimentNames

Get experiment names from a SingleCellExperiment

### **Description**

Get experiment names from a SingleCellExperiment

# Usage

```
getExperimentNames(scExp)
```

# **Arguments**

scExp

A SingleCellExperiment with named mainExp and altExps.

# Value

Character vector of unique experiment names

```
data(scExp)
getExperimentNames(scExp)
```

getMainExperiment

Get Main experiment of a SingleCellExperiment

# Description

Get Main experiment of a SingleCellExperiment

# Usage

```
getMainExperiment(scExp)
```

# Arguments

scExp

A SingleCellExperiment with named mainExp and altExps.

#### Value

The swapped SingleCellExperiment towards "main" experiment

# **Examples**

```
data(scExp)
getMainExperiment(scExp)
```

# **Description**

Get color dataframe from shiny::colorInput

# Usage

```
get_color_dataframe_from_input(
  input,
  levels_selected,
  color_by = c("sample_id", "total_counts"),
  input_id_prefix = "color_"
)
```

get\_cyto\_features 57

# Arguments

input Shiny input object

levels\_selected

Names of the features

color\_by Which feature color to retrieve

input\_id\_prefix

Prefix in front of the feature names

#### Value

A data frame with the feature levels and the colors of each level of this feature.

get\_cyto\_features

Map features onto cytobands

# Description

Map the features of a SingleCellExperiment onto the cytobands of a given genome. Some features might not be mapped to any cytobands (e.g. if they are not in the canconical chromosomes), and are removed from the returned object.

### Usage

```
get_cyto_features(scExp, ref_genome = c("hg38", "mm10", "ce11")[1])
```

# **Arguments**

scExp A SingleCellExperiment with genomic coordinate as features (peaks or bins)

ref\_genome Reference genome ('hg38' or 'mm10')

#### **Details**

The cytobands are an arbitrary cutting of the genome that dates back to staining metaphase chromosomes with Giemsa.

#### Value

A data frame of the SCE features with their corresponding cytoband name

```
data("scExp")
matching_cyto = get_cyto_features(scExp, ref_genome="hg38")
```

```
get_genomic_coordinates
```

Get SingleCellExperiment's genomic coordinates

# **Description**

Get SingleCellExperiment's genomic coordinates

# Usage

```
get_genomic_coordinates(scExp)
```

#### **Arguments**

scExp

A SingleCellExperiment object.

#### Value

A GRanges object of genomic coordinates.

### **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
feature_GRanges = get_genomic_coordinates(scExp)</pre>
```

```
get_most_variable_cyto
```

Retrieve the cytobands with the most variable fraction of reads

# Description

Given a SingleCellExperiment object with the slot "cytoBand" containing the fraction of reads in each cytoband, calculates the variance of each cytoband and returns a data.frame with the top variables cytobands. Most cytobands are expected to be unchanged between normal and tumor samples, therefore focusing on the top variable cytobands enable to focus on the most interseting regions.

### Usage

```
get_most_variable_cyto(scExp, top = 50)
```

### **Arguments**

scExp A SingleCellExperiment with "cytoBand" reducedDim slot filled.

top Number of cytobands to return (50).

# Value

A data.frame of the top variable cytoBands and their variance

# **Examples**

```
data("scExp")
scExp = calculate_cyto_mat(scExp, ref_genome="hg38")
get_most_variable_cyto(scExp, top=50)
```

```
get_pathway_mat_scExp Get pathway matrix
```

# **Description**

Get pathway matrix

# Usage

```
get_pathway_mat_scExp(
    scExp,
    pathways,
    max_distanceToTSS = 1000,
    ref = "hg38",
    GeneSetClasses = c("c1_positional", "c2_curated", "c3_motif", "c4_computational",
        "c5_G0", "c6_oncogenic", "c7_immunologic", "hallmark"),
    progress = NULL
)
```

### **Arguments**

scExp A SingleCellExperiment

pathways A character vector specifying the pathways to retrieve the cell count for.

max\_distanceToTSS

Numeric. Maximum distance to a gene's TSS to consider a region linked to a gene. (1000)#' @param ref

ref Reference genome, either mm10 or hg38

GeneSetClasses Which classes of MSIGdb to load

progress A shiny Progress instance to display progress bar.

#### Value

A matrix of cell to pathway

60 groupMat

# **Examples**

```
data(scExp)
mat = get_pathway_mat_scExp(scExp, pathways = "KEGG_T_CELL_RECEPTOR_SIGNALING_PATHWAY")
```

gg\_fill\_hue

gg\_fill\_hue

# Description

gg\_fill\_hue

# Usage

gg\_fill\_hue(n)

# Arguments

n

num hues

# Value

A color in HEX format

groupMat

groupMat

# Description

groupMat

# Usage

```
groupMat(mat = NA, margin = 1, groups = NA, method = "mean")
```

# Arguments

mat A matrix

margin By row or columns?

groups Groups

method Method to group

### Value

A grouped matrix

H1proportion 61

H1proportion

**H1proportion** 

# Description

H1proportion

# Usage

```
H1proportion(pv = NA, lambda = 0.5)
```

# Arguments

pv P.value vector lambda Lambda value

### Value

H1 proportion value

 $\verb|has_genomic_coordinates||$ 

Does SingleCellExperiment has genomic coordinates in features?

# Description

Does SingleCellExperiment has genomic coordinates in features?

# Usage

has\_genomic\_coordinates(scExp)

# Arguments

scExp

A SingleCellExperiment object

### Value

TRUE or FALSE

### **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
has_genomic_coordinates(scExp)
raw_genes = create_scDataset_raw(featureType="gene")
scExp_gene = create_scExp(raw_genes$mat, raw_genes$annot)
has_genomic_coordinates(scExp_gene)</pre>
```

hclustAnnotHeatmapPlot

hclust Annot Heatmap Plot

# Description

hclust Annot Heat map Plot

# Usage

```
hclustAnnotHeatmapPlot(
    x = NULL,
    hc = NULL,
    hmColors = NULL,
    anocol = NULL,
    xpos = c(0.1, 0.9, 0.114, 0.885),
    ypos = c(0.1, 0.5, 0.5, 0.6, 0.62, 0.95),
    dendro.cex = 1,
    xlab.cex = 0.8,
    hmRowNames = FALSE,
    hmRowNames.cex = 0.5
)
```

### **Arguments**

A correlation matrix Х hc An hclust object A color palette hmColors anocol A matrix of colors xpos **Xpos** Ypos ypos dendro.cex Size of denro names xlab.cex Size of x label hmRowNames Write rownames? hmRowNames.cex Size of rownames?

hg38.chromosomes 63

# Value

A heatmap

hg38.chromosomes

Data.frame of chromosome length - hg38

### **Description**

This data frame provides the length of each "canonical" chromosomes of Homo Sapiens genome build hg38.

# Usage

```
data("hg38.chromosomes")
```

### **Format**

hg38.chromosomes - a data frame with 24 rows and 3 variables:

chr Chromosome - character

start Start of the chromosome (bp) - integer

end End of the chromosome (bp) - integer

### Value

hg38.chromosomes - a data frame with 24 rows and 3 variables.

hg38.cytoBand

Data.frame of cytoBandlocation - hg38

### **Description**

This data frame provides the location of each cytoBands of Homo Sapiens genome build hg38.

#### Usage

```
data("hg38.cytoBand")
```

# **Format**

hg38.cytoBand - a data frame with 862 cytobands and 4 variables:

chr Chromosome - character

start Start of the chromosome (bp) - integer

end End of the chromosome (bp) - integer

cytoBand Name of the cytoBand - character

imageCol

# Value

hg38.cytoBand - a data frame with 862 cytobands and 4 variables.

hg38.GeneTSS

Data.frame of gene TSS - hg38

# **Description**

This dataframe was extracted from Gencode v25 and report the Transcription Start Site of each gene in the Homo Sapiens genome build hg38.

### Usage

```
data("hg38.GeneTSS")
```

#### **Format**

hg38.GeneTSS - a data frame with 32,937 genes and 5 variables:

```
chr Chromosome - character
start Start of the gene (TSS) - integer
end End of the gene - integer
Gene Gene symbol - character
Strand Strand - character
```

### Value

hg38.GeneTSS - a data frame with 32,937 genes and 5 variables

imageCol

imageCol

# **Description**

imageCol

# Usage

```
imageCol(
  matcol = NULL,
  strat = NULL,
  xlab.cex = 0.5,
  ylab.cex = 0.5,
  drawLines = c("none", "h", "v", "b")[1],
  ...
)
```

### **Arguments**

matcol A matrix of colors
strat Strat
xlab.cex X label size
ylab.cex Y label size
drawLines Draw lines?
... Additional parameters

### Value

A rectangular image

```
import_count_input_files
```

Import and count input files depending on their format

# **Description**

Import and count input files depending on their format

# Usage

```
import_count_input_files(
  files_dir_list,
  file_type,
  which,
  ref,
  verbose,
  progress,
  BPPARAM = BiocParallel::bpparam()
)
```

### **Arguments**

files\_dir\_list A named list of directories containing the input files.

file\_type Input file type.

which A GRanges object of features.

ref Reference genome.

verbose Print?

progress A progress object for Shiny.

BPPARAM BPPARAM object for multiprocessing. See bpparam for more informations.

Will take the default BPPARAM set in your R session.

import\_scExp

#### Value

A list with the feature indexes data.frame containing non-zeroes entries in the count matrix and the cell names

import\_scExp

Read single-cell matrix(ces) into scExp

# **Description**

Combine one or multiple matrices together to create a sparse matrix and cell annotation data.frame.

# Usage

```
import_scExp(file_paths, remove_pattern = "", temp_path = NULL)
```

# **Arguments**

file\_paths A character vector of file names towards single cell epigenomic matrices (fea-

tures x cells) (must be .txt / .tsv)

remove\_pattern A string pattern to remove from the sample names. Can be a regexp.

temp\_path In case matrices are stored in temporary folder, a character vector of path to-

wards temporary files. (NULL)

### Value

A list containing:

- datamatrix: a sparseMatrix of features x cells
- annot raw: an annotation of cells as data.frame

```
mat1 = mat2 = create_scDataset_raw()$mat
tmp1 = tempfile(fileext = ".tsv")
tmp2 = tempfile(fileext = ".tsv")
write.table(as.matrix(mat1),file=tmp1,sep = "\t",
row.names = TRUE,col.names = TRUE,quote = FALSE)
write.table(as.matrix(mat2),file=tmp2, sep = "\t",
row.names = TRUE,col.names = TRUE,quote = FALSE)
file_paths = c(tmp1,tmp2)
out = import_scExp(file_paths)
```

```
index_peaks_barcodes_to_matrix_indexes
```

Read index-peaks-barcodes trio files on interval to create count indexes

# **Description**

Read index-peaks-barcodes trio files on interval to create count indexes

# Usage

```
index_peaks_barcodes_to_matrix_indexes(
  feature_file,
  matrix_file,
  barcode_file,
  binarize = FALSE
)
```

# **Arguments**

feature\_file A file containing the features genomic locations

matrix\_file A file containing the indexes of non-zeroes values and their value (respectively

i,j,x,see sparseMatrix)

barcode\_file A file containing the barcode ids

binarize Binarize matrix?

#### Value

A list containing a "feature index" data.frame, name\_cells, and a region GenomicRange object used to form the sparse matrix

```
inter_correlation_scExp
```

Calculate inter correlation between cluster or samples

# **Description**

Calculate inter correlation between cluster or samples

### Usage

```
inter_correlation_scExp(
   scExp_cf,
   by = c("sample_id", "cell_cluster")[1],
   reference_group = unique(scExp_cf[[by]])[1],
   other_groups = unique(scExp_cf[[by]]),
   fullCor = TRUE
)
```

# **Arguments**

scExp\_cf A SingleCellExperiment

by On which feature to calculate correlation ("sample\_id" or "cell\_cluster")

reference\_group

Reference group to calculate correlation with. Must be in accordance with "by".

other\_groups

Groups on which to calculate correlation (can contain multiple groups, and also reference\_group). Must be in accordance with "by".

fullCor

A logical specifying if the correlation matrix was calculated on the entire set of

cells (TRUE).

### Value

A data frame of average inter-correlation of cells in other groups with cells in reference group

### **Examples**

```
data(scExp)
inter_correlation_scExp(scExp)
```

```
intra_correlation_scExp
```

Calculate intra correlation between cluster or samples

# **Description**

Calculate intra correlation between cluster or samples

# Usage

```
intra_correlation_scExp(
   scExp_cf,
   by = c("sample_id", "cell_cluster")[1],
   fullCor = TRUE
)
```

launchApp 69

# **Arguments**

scExp\_cf A SingleCellExperiment

by On which feature to calculate correlation ("sample\_id" or "cell\_cluster")

fullCor Logical specifying if the correlation matrix was run on the entire number of cells

or on a subset.

### Value

A data.frame of cell average intra-correlation

# **Examples**

```
data(scExp)
intra_correlation_scExp(scExp, by = "sample_id")
intra_correlation_scExp(scExp, by = "cell_cluster")
```

launchApp

Launch ChromSCape

# Description

Main function to launch ChromSCape in your favorite browser. You can pass additional parameters that you would pass to shiny::runApp (runApp)

# Usage

```
launchApp(launch.browser = TRUE, ...)
```

# **Arguments**

```
launch.browser Wether to launch browser or not
... Additional parameters passed to runApp
```

### Value

Launches the shiny application

```
## Not run:
launchApp()
## End(Not run)
```

load\_MSIGdb

Load and format MSIGdb pathways using msigdbr package

# **Description**

Load and format MSIGdb pathways using msigdbr package

# Usage

```
load_MSIGdb(
  ref,
  GeneSetClasses = c("c1_positional", "c2_curated", "c3_motif", "c4_computational",
        "c5_G0", "c6_oncogenic", "c7_immunologic", "hallmark")
)
```

### **Arguments**

```
ref Reference genome, either mm10 or hg38
GeneSetClasses Which classes of MSIGdb to load
```

#### Value

A list containing the GeneSet (list), GeneSetDf (data.frame) and GenePool character vector of all possible genes

merge\_MACS2\_peaks

Merge peak files from MACS2 peak caller

### **Description**

Merge peak files from MACS2 peak caller

### Usage

```
merge_MACS2_peaks(peak_file, peak_distance_to_merge, min_peak_size = 200, ref)
```

# **Arguments**

### Value

Peaks as GRanges

mm10.chromosomes 71

mm10.chromosomes

Data.frame of chromosome length - mm10

### **Description**

This data frame provides the length of each "canonical" chromosomes of Mus Musculus (Mouse) genome build mm10.

### Usage

```
data("mm10.chromosomes")
```

#### **Format**

mm10.chromosomes - a data frame with 24 chromosomes and 3 variables:

```
chr Chromosome - character
```

start Start of the chromosome (bp) - integer

end End of the chromosome (bp) - integer

#### Value

mm10.chromosomes - a data frame with 24 chromosomes and 3 variables

mm10.cytoBand

Data.frame of cytoBandlocation - mm10

# Description

This data frame provides the location of each cytoBands of Homo Sapiens genome build mm10.

#### **Usage**

```
data("mm10.cytoBand")
```

#### **Format**

mm10.cytoBand - a data frame with 403 cytobands and 4 variables:

```
chr Chromosome - character
```

start Start of the chromosome (bp) - integer

end End of the chromosome (bp) - integer

cytoBand Name of the cytoBand - character

# Value

mm10.cytoBand - a data frame with 403 cytobands and 4 variables.

72 normalize\_scExp

mm10.GeneTSS

Data.frame of gene TSS - mm10

### **Description**

This dataframe was extracted from Gencode v25 and report the Transcription Start Site of each gene in the Mus Musculus genome build mm10 (Mouse).

# Usage

```
data("mm10.GeneTSS")
```

#### **Format**

```
mm10.GeneTSS - a data frame with 27,916 genes and 5 variables:

chr Chromosome name - character

start Start of the gene (TSS) - integer

end End of the gene - integer

Gene Gene symbol - character

Strand Strand - character
```

#### Value

mm10.GeneTSS - a data frame with 27,916 genes and 5 variables

normalize\_scExp

Normalize counts

# **Description**

Normalize counts

# Usage

```
normalize_scExp(
   scExp,
   type = c("CPM", "TFIDF", "RPKM", "TPM", "feature_size_only")
)
```

### **Arguments**

scExp A S

A SingleCellExperiment object.

type

Which normalization to apply. Either 'CPM', 'TFIDF', 'RPKM', 'TPM' or 'feature\_size\_only'. Note that for all normalization by size (RPKM, TPM, feature\_size\_only), the features must have defined genomic coordinates.

## Value

A SingleCellExperiment object containing normalized counts. (See ?normcounts())

#### **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp = normalize_scExp(scExp)
head(SingleCellExperiment::normcounts(scExp))</pre>
```

# Description

Number of cells before & after correlation filtering

## Usage

```
num_cell_after_cor_filt_scExp(scExp, scExp_cf)
```

## **Arguments**

scExp SingleCellExperiment object before correlation filtering. scExp\_cf SingleCellExperiment object after correlation filtering.

#### Value

A colored kable with the number of cells per sample before and after filtering for display

```
data("scExp")
scExp_cf = correlation_and_hierarchical_clust_scExp(scExp)
scExp_cf = filter_correlated_cell_scExp(scExp_cf,
corr_threshold = 99, percent_correlation = 1)
## Not run: num_cell_after_cor_filt_scExp(scExp,scExp_cf)
```

Table of cells before / after QC

# Usage

```
num_cell_after_QC_filt_scExp(scExp, annot, datamatrix)
```

## **Arguments**

scExp A SingleCellExperiment object.

annot A raw annotation data.frame of cells before filtering.

datamatrix A matrix of cells per regions before filtering.

#### Value

A formatted kable in HTML.

#### **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp_filtered = filter_scExp(scExp)
## Not run: num_cell_after_QC_filt_scExp(
scExp_filtered,as.data.frame(SingleCellExperiment::colData(scExp)), counts(scExp))
## End(Not run)</pre>
```

```
num_cell_before_cor_filt_scExp
```

Table of number of cells before correlation filtering

# Description

Table of number of cells before correlation filtering

## Usage

```
num_cell_before_cor_filt_scExp(scExp)
```

#### **Arguments**

scExp

A SingleCellExperiment Object

#### Value

A colored kable with the number of cells per sample for display

## **Examples**

```
data("scExp")
## Not run: num_cell_before_cor_filt_scExp(scExp)
```

```
num_cell_in_cluster_scExp
```

Number of cells in each cluster

## **Description**

Number of cells in each cluster

## Usage

```
num_cell_in_cluster_scExp(scExp)
```

## **Arguments**

scExp

A SingleCellExperiment object containing chromatin groups.

## Value

A formatted kable of cell assignation to each cluster.

```
data("scExp")
scExp_cf = correlation_and_hierarchical_clust_scExp(scExp)
scExp_cf = choose_cluster_scExp(scExp_cf,nclust=3,consensus=FALSE)
## Not run: num_cell_in_cluster_scExp(scExp_cf)
```

num\_cell\_scExp

Table of cells

#### **Description**

Table of cells

## Usage

```
num_cell_scExp(annot, datamatrix)
```

# **Arguments**

annot An annotation of cells. Can be obtain through 'colData(scExp)'.

datamatrix A matrix of cells per regions before filtering.

#### Value

A formatted kable in HTML.

#### **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
## Not run: num_cell_scExp(SingleCellExperiment::colData(scExp))</pre>
```

```
pca_irlba_for_sparseMatrix
```

Run sparse PCA using irlba SVD

## **Description**

This function allows to run a PCA using IRLBA Singular Value Decomposition in a fast & memory efficient way. The increamental Lanczos bidiagonalisation algorithm allows to keep the matrix sparse as the "loci" centering is implicit. The function then multiplies by the approximate singular values (svd\$d) in order to get more importance to the first PCs proportionnally to their singular values. This step is crucial for downstream approaches, e.g. UMAP or T-SNE.

#### Usage

```
pca_irlba_for_sparseMatrix(x, n_comp, work = 3 * n_comp)
```

#### **Arguments**

x A sparse normalized matrix (features x cells)n\_compThe number of principal components to keep

work Working subspace dimension, larger values can speed convergence at the cost

of more memory use.

# Value

The rotated data, e.g. the cells x PC column in case of sc data.

```
{\tt plot\_cluster\_consensus\_scExp} \\ {\tt Plot\ cluster\ consensus}
```

# Description

Plot cluster consensus score for each k as a bargraph.

# Usage

```
plot_cluster_consensus_scExp(scExp)
```

## **Arguments**

scExp A SingleCellExperiment

# Value

The consensus score for each cluster for each k as a barplot

```
data("scExp")
plot_cluster_consensus_scExp(scExp)
```

```
plot_correlation_PCA_scExp

Plotting correlation of PCs with a variable of interest
```

Plotting correlation of PCs with a variable of interest

## Usage

```
plot_correlation_PCA_scExp(
    scExp,
    correlation_var = "total_counts",
    color_by = NULL,
    topPC = 10
)
```

## **Arguments**

```
scExp A SingleCellExperiment Object

correlation_var

A string specifying with which numeric variable from colData of scExp to calculate and plot the correlation of each PC with. ('total_counts')

color_by A string specifying with which categorical variable to color the plot. ('NULL')

topPC An integer specifying the number of PCs to plot correlation with 10
```

## Value

A ggplot histogram representing the distribution of count per cell

```
data("scExp")
plot_correlation_PCA_scExp(scExp, topPC = 25)
plot_correlation_PCA_scExp(scExp, color_by = "cell_cluster")
plot_correlation_PCA_scExp(scExp, color_by = "sample_id")
```

```
plot_coverage_BigWig Coverage plot
```

Coverage plot

## Usage

```
plot_coverage_BigWig(
  coverages,
  label_color_list,
  peaks = NULL,
  chrom,
  start,
  end,
  ref = "hg38"
)
```

# Arguments

## Value

A coverage plot annotated with genes

```
data(scExp)
```

```
{\it plot\_differential\_summary\_scExp} \\ {\it Differential\ summary\ barplot}
```

Differential summary barplot

## Usage

```
plot_differential_summary_scExp(
    scExp_cf,
    qval.th = 0.01,
    logFC.th = 1,
    min.percent = 0.01
)
```

## **Arguments**

scExp\_cf A SingleCellExperiment object
qval.th Adjusted p-value threshold. (0.01)
logFC.th Fold change threshold. (1)
min.percent Minimum fraction of cells having the feature active to consider it as significantly differential. (0.01)

#### Value

A barplot summary of differential analysis

# Examples

```
data("scExp")
plot_differential_summary_scExp(scExp)
```

## **Description**

Volcano plot of differential features

# Usage

```
plot_differential_volcano_scExp(
   scExp_cf,
   group = "C1",
   logFC.th = 1,
   qval.th = 0.01,
   min.percent = 0.01
)
```

## **Arguments**

scExp\_cf A SingleCellExperiment object
group A character indicating the group for which to plot the differential volcano plot.
("C1")
logFC.th Fold change threshold. (1)
qval.th Adjusted p-value threshold. (0.01)
min.percent Minimum fraction of cells having the feature active to consider it as significantly differential. (0.01)

## Value

A volcano plot of differential analysis of a specific cluster

## **Examples**

```
data("scExp")
plot_differential_volcano_scExp(scExp,"C1")
```

```
plot_distribution_scExp
```

Plotting distribution of signal

# **Description**

Plotting distribution of signal

## Usage

```
plot_distribution_scExp(
    scExp,
    raw = TRUE,
    log10 = FALSE,
    pseudo_counts = 1,
    bins = 150
)
```

#### **Arguments**

scExp A SingleCellExperiment Object

raw Use raw counts?

log10 Transform using log10?

pseudo\_counts Pseudo-count to add if using log10 bins Number of bins in the histogram

#### Value

A ggplot histogram representing the distribution of count per cell

## **Examples**

```
data("scExp")
plot_distribution_scExp(scExp)
```

```
plot_gain_or_loss_barplots
```

Plot Gain or Loss of cytobands of the most variables cytobands

## **Description**

Plot Gain or Loss of cytobands of the most variables cytobands Plot Gain or Loss of cytobands of the most variables cytobands

#### Usage

```
plot_gain_or_loss_barplots(scExp, cells = NULL, top = 20)
plot_gain_or_loss_barplots(scExp, cells = NULL, top = 20)
```

## **Arguments**

scExp A SingleCellExperiment with "logRatio\_cytoBand" reducedDim slot filled. See

calculate\_logRatio\_CNA

cells Cell IDs of the tumor samples to

top Number of most variables cytobands to plot

#### Value

Plot the gains/lost in the selected cells of interest as multiple barplots Plot the gains/lost in the selected cells of interest as multiple barplots plot\_heatmap\_scExp 83

#### **Examples**

```
data("scExp")
scExp = calculate_CNA(scExp, control_samples = unique(scExp$sample_id)[1],
ref_genome="hg38", quantiles_to_define_gol = c(0.05,0.95))
plot_gain_or_loss_barplots(scExp, cells = scExp$cell_id[which(
scExp$sample_id %in% unique(scExp$sample_id)[2])])

data("scExp")
scExp = calculate_CNA(scExp, control_samples = unique(scExp$sample_id)[1],
ref_genome="hg38", quantiles_to_define_gol = c(0.05,0.95))
plot_gain_or_loss_barplots(scExp, cells = scExp$cell_id[which(
scExp$sample_id %in% unique(scExp$sample_id)[2])])
```

plot\_heatmap\_scExp

Plot cell correlation heatmap with annotations

## **Description**

Plot cell correlation heatmap with annotations

#### Usage

```
plot_heatmap_scExp(
    scExp,
    name_hc = "hc_cor",
    corColors = (grDevices::colorRampPalette(c("royalblue", "white", "indianred1")))(256),
    color_by = NULL,
    downsample = 1000,
    hc_linkage = "ward.D"
)
```

## **Arguments**

scExp	A SingleCellExperiment Object
name_hc	Name of the hclust contained in the SingleCellExperiment object
corColors	A palette of colors for the heatmap
color_by	Which features to add as additional bands on top of plot
downsample	Number of cells to downsample
hc_linkage	A linkage method for hierarchical clustering. See cor. ('ward.D')

#### Value

A heatmap of cell to cell correlation, grouping cells by hierarchical clustering.

#### **Examples**

```
data("scExp")
plot_heatmap_scExp(scExp)
```

```
plot_inter_correlation_scExp
```

Violin plot of inter-correlation distribution between one or multiple groups and one reference group

# **Description**

Violin plot of inter-correlation distribution between one or multiple groups and one reference group

#### Usage

```
plot_inter_correlation_scExp(
    scExp_cf,
    by = c("sample_id", "cell_cluster")[1],
    jitter_by = NULL,
    reference_group = unique(scExp_cf[[by]])[1],
    other_groups = unique(scExp_cf[[by]]),
    downsample = 5000
)
```

#### **Arguments**

scExp\_cf A SingleCellExperiment

by Color by sample\_id or cell\_cluster

jitter\_by Add jitter points of another layer (cell\_cluster or sample\_id)

reference\_group

Character containing the reference group name to calculate correlation from.

other\_groups Character vector of the other groups for which to calculate correlation with the

reference group.

downsample Downsample for plotting

#### Value

A violin plot of inter-correlation

```
data(scExp)
plot_intra_correlation_scExp(scExp)
```

```
{\it Plot\_intra\_correlation\_scExp} \\ {\it Violin~plot~of~intra-correlation~distribution}
```

Violin plot of intra-correlation distribution

## Usage

```
plot_intra_correlation_scExp(
    scExp_cf,
    by = c("sample_id", "cell_cluster")[1],
    jitter_by = NULL,
    downsample = 5000
)
```

## Arguments

scExp\_cf A SingleCellExperiment

by Color by sample\_id or cell\_cluster

jitter\_by Add jitter points of another layer (cell\_cluster or sample\_id)

downsample Downsample for plotting

#### Value

A violin plot of intra-correlation

# **Examples**

```
data(scExp)
plot_intra_correlation_scExp(scExp)
```

```
plot_most_contributing_features
```

Plot Top/Bottom most contributing features to PCA

## **Description**

Plot Top/Bottom most contributing features to PCA

#### Usage

```
plot_most_contributing_features(
   scExp,
   component = "Component_1",
   n_top_bot = 10
)
```

## **Arguments**

scExp A SingleCellExperiment containing "PCA" in reducedDims and gene annotation

in rowRanges

component The name of the component of interest

n\_top\_bot An integer number of top and bot regions to plot

#### **Details**

If a gene TSS is within 10,000bp of the region, the name of the gene(s) will be displayed instead of the region

#### Value

A barplot of top and bottom features with the largest absolute value in the component of interest

#### **Examples**

# Description

Barplot of the % of active cells for a given features

## Usage

```
plot_percent_active_feature_scExp(
    scExp,
    gene,
    by = c("cell_cluster", "sample_id")[1],
    highlight = NULL,
    downsample = 5000,
    max_distanceToTSS = 1000
)
```

#### **Arguments**

scExp A SingleCellExperiment

gene A character specifying the gene to plot

by Color violin by cell\_cluster or sample\_id ("cell\_cluster")

highlight A specific group to highlight in a one vs all fashion

downsample Downsample for plotting (5000)

max\_distanceToTSS

Numeric. Maximum distance to a gene's TSS to consider a region linked to a

gene. (1000)

#### Value

A violin plot of intra-correlation

## **Examples**

```
data(scExp)
plot_percent_active_feature_scExp(scExp, "UBXN10")
```

```
plot_pie_most_contributing_chr
```

Pie chart of top contribution of chromosomes in the 100 most contributing features to PCA #'

# Description

Pie chart of top contribution of chromosomes in the 100 most contributing features to PCA #'

## Usage

```
plot_pie_most_contributing_chr(
   scExp,
   component = "Component_1",
   n_top_bot = 100
)
```

#### **Arguments**

scExp A SingleCellExperiment containing "PCA" in reducedDims and gene annotation

in rowRanges

component The name of the component of interest

n\_top\_bot An integer number of top and bot regions to plot (100)

## Value

A pie chart showing the distribution of chromosomes in the top features with the largest absolute value in the component of interest

# **Examples**

## **Description**

Plot reduced dimensions (PCA, TSNE, UMAP)

## Usage

```
plot_reduced_dim_scExp(
    scExp,
    color_by = "sample_id",
    reduced_dim = c("PCA", "TSNE", "UMAP"),
    select_x = NULL,
    select_y = NULL,
    downsample = 5000,
    transparency = 0.6,
    size = 1,
    max_distanceToTSS = 1000,
    annotate_clusters = "cell_cluster" %in% colnames(colData(scExp)),
    min_quantile = 0.01,
    max_quantile = 0.99
)
```

#### **Arguments**

scExp	A SingleCellExperiment Object
color_by	Character of eature used for coloration. Can be cell metadata ('total_counts', 'sample_id',) or a gene name.
reduced_dim	Reduced Dimension used for plotting
select_x	Which variable to select for x axis
select_y	Which variable to select for y axis
downsample	Number of cells to downsample
transparency	Alpha parameter, between 0 and 1
size	Size of the points.

max\_distanceToTSS

The maximum distance to TSS to consider a gene linked to a region. Used only if "color\_by" is a gene name.

annotate\_clusters

A logical indicating if clusters should be labelled. The 'cell\_cluster' column

should be present in metadata.

min\_quantile The lower threshold to remove outlier cells, as quantile of cell embeddings (be-

tween 0 and 0.5).

max\_quantile The upper threshold to remove outlier cells, as quantile of cell embeddings (be-

tween 0.5 and 1).

#### Value

A ggplot geom\_point plot of reduced dimension 2D reprensentation

#### **Examples**

```
data("scExp")
plot_reduced_dim_scExp(scExp, color_by = "sample_id")
plot_reduced_dim_scExp(scExp, color_by = "total_counts")
plot_reduced_dim_scExp(scExp, reduced_dim = "UMAP")
plot_reduced_dim_scExp(scExp, color_by = "CD52", reduced_dim = "UMAP")
```

```
plot_reduced_dim_scExp_CNA
```

Plot UMAP colored by Gain or Loss of cytobands

## **Description**

Plot UMAP colored by Gain or Loss of cytobands

#### Usage

```
plot_reduced_dim_scExp_CNA(scExp, cytoBand)
```

#### **Arguments**

scExp A SingleCellExperiment with "gainOrLoss\_cytoBand" reducedDim slot filled.

See calculate\_gain\_or\_loss

cytoBand Which cytoBand to color cells by

#### Value

Plot the gains/lost of the cytoband overlayed on the epigenetic UMAP.

90 plot\_top\_TF\_scExp

## **Examples**

```
data("scExp")
scExp = calculate_CNA(scExp, control_samples = unique(scExp$sample_id)[1],
ref_genome="hg38", quantiles_to_define_gol = c(0.05,0.95))
plot_reduced_dim_scExp_CNA(scExp, get_most_variable_cyto(scExp)$cytoBand[1])
```

plot\_top\_TF\_scExp

Barplot of top TFs from ChEA3 TF enrichment analysis

# **Description**

Barplot of top TFs from ChEA3 TF enrichment analysis

## Usage

```
plot_top_TF_scExp(
    scExp,
    group = unique(scExp$cell_cluster)[1],
    set = c("Differential", "Enriched", "Depleted")[1],
    type = c("Score", "nTargets", "nTargets_over_TF", "nTargets_over_genes")[1],
    n_top = 25
)
```

## **Arguments**

scExp	A SingleCellExperiment
group	A character string specifying the differential group to display the top TFs
set	A character string specifying the set of genes in which the TF were enriched, either 'Differential', 'Enriched' or 'Depleted'.
type	A character string specifying the Y axis of the plot, either the number of differential targets or the ChEA3 integrated mean score. E.g. either "Score", "nTargets", "nTargets_over_TF" for the number of target genes over the total number of genes targeted by the TF or "nTargets_over_genes" for the number of target genes over the number of genes in the gene set.
n_top	An integer specifying the number of top TF to display

## Value

A bar plot of top TFs from ChEA3 TF enrichment analysis

#### **Examples**

```
data("scExp")

plot_top_TF_scExp(
    scExp,
    group = "C1",
    set = "Differential",
        type = "Score",
        n_top = 10)

plot_top_TF_scExp(
    scExp,
    group = "C1",
    set = "Enriched",
        type = "nTargets_over_genes",
        n_top = 20)
```

## **Description**

Violin plot of features

#### Usage

```
plot_violin_feature_scExp(
   scExp,
   gene,
   by = c("cell_cluster", "sample_id")[1],
   downsample = 5000,
   max_distanceToTSS = 1000
)
```

# **Arguments**

scExp A SingleCellExperiment
gene A character specifying the gene to plot
by Color violin by cell\_cluster or sample\_id ("cell\_cluster")
downsample Downsample for plotting (5000)
max\_distanceToTSS

Numeric. Maximum distance to a gene's TSS to consider a region linked to a gene. (1000)

#### Value

A violin plot of intra-correlation

## **Examples**

```
data(scExp)
plot_violin_feature_scExp(scExp, "UBXN10")
```

```
preprocessing_filtering_and_reduction
```

Preprocess and filter matrix annotation data project folder to SCE

## **Description**

Preprocess and filter matrix annotation data project folder to SCE

#### Usage

```
preprocessing_filtering_and_reduction(
  datamatrix,
  annot_raw,
  min_reads_per_cell = 1600,
  max_quantile_read_per_cell = 95,
  n_top_features = 40000,
  norm_type = "CPM",
  n_dims = 10,
  remove_PC = NULL,
  subsample_n = NULL,
  ref_genome = "hg38",
  exclude_regions = NULL,
  doBatchCorr = FALSE,
  batch_sels = NULL
)
```

## **Arguments**

preprocess\_CPM 93

n\_dims An integer specifying the number of dimensions to keep for PCA

remove\_PC A vector of string indicating which principal components to remove before

downstream analysis as probably correlated to library size. Should be under the form: 'Component\_1', 'Component\_2', ... Recommended when using 'TFIDF'

normalization method. (NULL)

subsample\_n Number of cells to subsample.

ref\_genome Reference genome ("hg38" or "mm10").

exclude\_regions

GenomicRanges with regions to remove from the object.

doBatchCorr Run batch correction? TRUE or FALSE

batch\_sels If doBatchCorr is TRUE, List of characters. Names are batch names, characters

are sample names.

#### Value

A SingleCellExperiment object containing feature spaces.

#### **Examples**

```
raw <- create_scDataset_raw()
scExp = preprocessing_filtering_and_reduction(raw$mat, raw$annot)</pre>
```

preprocess\_CPM

Preprocess scExp - Counts Per Million (CPM)

#### **Description**

Preprocess scExp - Counts Per Million (CPM)

#### Usage

```
preprocess_CPM(scExp)
```

#### **Arguments**

scExp A SingleCellExperiment Object

#### Value

A SingleCellExperiment object.

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp = preprocess_CPM(scExp)
head(SingleCellExperiment::normcounts(scExp))</pre>
```

94 preprocess\_RPKM

```
\label{eq:constraint} Preprocess \ scExp \ - \ size \ only
```

## **Description**

Preprocess scExp - size only

## Usage

```
preprocess_feature_size_only(scExp)
```

## **Arguments**

scExp

A SingleCellExperiment Object

#### Value

A SingleCellExperiment object.

## **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp = preprocess_feature_size_only(scExp)
head(SingleCellExperiment::normcounts(scExp))</pre>
```

preprocess\_RPKM

Preprocess scExp - Read per Kilobase Per Million (RPKM)

# Description

Preprocess scExp - Read per Kilobase Per Million (RPKM)

## Usage

```
preprocess_RPKM(scExp)
```

## **Arguments**

scExp

A SingleCellExperiment Object

## Value

A SingleCellExperiment object.

preprocess\_TFIDF 95

## **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp = preprocess_RPKM(scExp)
head(SingleCellExperiment::normcounts(scExp))</pre>
```

preprocess\_TFIDF

Preprocess scExp - TF-IDF

# Description

```
Preprocess scExp - TF-IDF
```

## Usage

```
preprocess_TFIDF(scExp, scale = 10000, log = TRUE)
```

# Arguments

scExp	A SingleCellExperiment Object
scale	A numeric to multiply the matrix in order to have human readeable numbers. Has no impact on the downstream analysis
log	Wether to use neperian log on the TF-IDF normalized data or not.

## Value

A SingleCellExperiment object.

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp = preprocess_TFIDF(scExp)
head(SingleCellExperiment::normcounts(scExp))</pre>
```

96 rawfile\_ToBigWig

preprocess\_TPM

Preprocess scExp - Transcripts per Million (TPM)

## **Description**

Preprocess scExp - Transcripts per Million (TPM)

## Usage

```
preprocess_TPM(scExp)
```

## **Arguments**

scExp

A SingleCellExperiment Object

#### Value

A SingleCellExperiment object.

## **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp = preprocess_TPM(scExp)
head(SingleCellExperiment::normcounts(scExp))</pre>
```

rawfile\_ToBigWig

rawfile\_ToBigWig: reads in BAM file and write out BigWig coverage file, normalized and smoothed

## **Description**

rawfile\_ToBigWig: reads in BAM file and write out BigWig coverage file, normalized and smoothed

#### Usage

```
rawfile_ToBigWig(
  input,
  BigWig_filename,
  format = "BAM",
  bin_width = 150,
  norm_factor,
  n_smoothBin = 5,
  ref = "hg38",
  read_size = 101,
```

```
original_bins = NULL,
quantile_for_peak_calling = 0.85
)
```

## **Arguments**

input Either a named list of character vector of path towards single-cell BED files or

a sparse raw matrix of small bins («500bp). If a named list specifying scBEDn the names MUST correspond to the 'sample\_id' column in your SingleCellExperiment object. The single-cell BED files names MUST match the barcode names in your SingleCellExperiment (column 'barcode'). The scBED files can

be gzipped or not.

BigWig\_filename

Path to write the output BigWig file

format File format, either "BAM" or "BED"

bin\_width Bin size for coverage

norm\_factor Then number of cells or total number of reads in the given sample, for normal-

ization

n\_smoothBin Number of bins for smoothing values

ref Reference genome.
read\_size Length of the reads.

original\_bins Original bins GenomicRanges in case the format is raw matrix.

quantile\_for\_peak\_calling

The quantile to define the threshold above which signal is considered as a peak.

#### Value

Writes in the output directory a bigwig file displaying the cumulative coverage of cells and a basic set of peaks called by taking all peaks above a given threshold

Writes a BigWig file as output

```
raw_counts_to_sparse_matrix
```

Create a sparse count matrix from various format of input data.

## **Description**

This function takes three different type of single-cell input: - Single cell BAM files (sorted) - Single cell BED files (gzipped) - A combination of an index file, a peak file and cell barcode file (The index file is composed of three column: index i, index j and value x for the non zeroes entries in the sparse matrix.)

#### Usage

```
raw_counts_to_sparse_matrix(
   files_dir_list,
   file_type = c("scBED", "scBAM", "FragmentFile"),
   use_Signac = TRUE,
   peak_file = NULL,
   n_bins = NULL,
   bin_width = NULL,
   genebody = NULL,
   extendPromoter = 2500,
   verbose = TRUE,
   ref = c("hg38", "mm10", "ce11")[1],
   progress = NULL,
   BPPARAM = BiocParallel::bpparam()
)
```

# Arguments

files_dir_list	A named character vector of directories containing the files. The names correspond to sample names.
file_type	Input file(s) type(s) ('scBED', 'scBAM', 'FragmentFile')
use_Signac	Use Signac wrapper function 'FeatureMatrix' if the Signac package is installed (TRUE).
peak_file	A file containing genomic location of peaks (NULL)
n_bins	The number of bins to tile the genome (NULL)
bin_width	The size of bins to tile the genome (NULL)
genebody	Count on genes (body + promoter) ? (NULL)
extendPromoter	If counting on genes, number of base pairs to extend up or downstream of TSS $(2500)$ .
verbose	Verbose (TRUE)
ref	reference genome to use (hg38)
progress	Progress object for Shiny
BPPARAM	BPPARAM object for multiprocessing. See bpparam for more informations. Will take the default BPPARAM set in your R session.

# **Details**

This functions re-counts signal on either fixed genomic bins, a set of user-defined peaks or around the TSS of genes.

#### Value

A sparse matrix of features x cells

#### References

Stuart el al., Multimodal single-cell chromatin analysis with Signac bioRxiv https://doi.org/10.1101/2020.11.09.373613

```
read_count_mat_with_separated_chr_start_end

Read a count matrix with three first columns (chr,start,end)
```

# Description

Read a count matrix with three first columns (chr,start,end)

## Usage

```
read_count_mat_with_separated_chr_start_end(
  path_to_matrix,
  format_test,
  separator
)
```

#### **Arguments**

```
path_to_matrix Path to the count matrix format_test Sample of the read.table separator Separator character
```

#### Value

A sparseMatrix with rownames in the form "chr1:1222-55555"

```
read_sparse_matrix Read in one or i
```

Read in one or multiple sparse matrices (10X format)

#### **Description**

Given one or multiple directories, look in each directory for a combination of the following files :

- A 'features' file containing unique feature genomic locations -in tab separated format (\*\_features.bed / .txt / .tsv / .gz), e.g. chr, start and end
- A 'barcodes' file containing unique barcode names (\_barcode.txt / .tsv / .gz)
- A 'matrix' A file containing indexes of non zero entries (\_matrix.mtx / .gz)

100 rebin\_helper

#### Usage

```
read_sparse_matrix(
  files_dir_list,
  ref = c("hg38", "mm10", "ce11")[1],
  verbose = TRUE
)
```

# **Arguments**

files\_dir\_list A named character vector containing the full path towards folders. Each folder

should contain only the Feature file, the Barcode file and the Matrix file (see

description).

ref Reference genome (used to filter non-canonical chromosomes).

verbose Print?

#### Value

Returns a list containing a datamatrix and cell annotation

# **Examples**

```
## Not run:
sample_dirs = c("/path/to/folder1/", "/path/to/folder2/")
names(sample_dirs) = c("sample_1", "sample_2")
out <- read_sparse_matrix(sample_dirs, ref = "hg38")
head(out$datamatrix)
head(out$annot_raw)
## End(Not run)</pre>
```

rebin\_helper

Rebin Helper for rebin\_matrix function

## **Description**

Rebin Helper for rebin\_matrix function

## Usage

```
rebin_helper(mat_df)
```

#### **Arguments**

mat\_df

A data.frame corresponding to sparse matrix indexes & values.

#### Value

a data.frame grouped mean-summarised by col and new\_row

rebin\_matrix 101

#### **Description**

This functions is best used to re-count large number of small bins or peaks (e.g. <= 5000bp) into equal or larger sized bins. The genome is either cut in fixed bins (e.g. 50,000bp) or into an user defined number of bins. Bins are calculated based on the canconical chromosomes. Note that if peaks are larger than bins, or if peaks are overlapping multiple bins, the signal is added to each bin. Users can increase the minimum overlap to consider peaks overlapping bins (by default 150bp, size of a nucleosome) to disminish the number of peaks overlapping multiple region. Any peak smaller than the minimum overlapp threshold will be dismissed. Therefore, library size might be slightly different from peaks to bins if signal was duplicated into multiple bins or ommitted due to peaks smaller than minimum overlap.

#### Usage

```
rebin_matrix(
  mat,
  bin_width = 50000,
  custom_annotation = NULL,
  minoverlap = 500,
  verbose = TRUE,
  ref = "hg38",
  nthreads = 1,
  rebin_function = rebin_helper
)
```

#### **Arguments**

mat A matrix of peaks x cells

bin\_width Width of bins to produce in base pairs (minimum 500) (50000)

custom\_annotation

A GenomicRanges object specifying the new features to count the matrix on instead of recounting on genomic bins. If not NULL, takes predecency over

bin\_width.

minoverlap Minimum overlap between the original bins and the new features to consider the

peak as overlapping the bin. We recommand to put this number at exactly half of the original bin size (e.g. 500bp for original bin size of 1000bp) so that no

original bins are counted twice. (500)

verbose Verbose

ref Reference genome to use (hg38)

nthreads Number of threads to use for paralell processing

rebin\_function A function to use to rebin the matrix.

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#### Value

A sparse matrix of larger bins or peaks.

#### **Examples**

```
mat = create_scDataset_raw()$mat
binned_mat = rebin_matrix(mat,bin_width = 10e6)
dim(binned_mat)
```

reduce\_dims\_scExp

Reduce dimensions (PCA, TSNE, UMAP)

# Description

Reduce dimensions (PCA, TSNE, UMAP)

## Usage

```
reduce_dims_scExp(
    scExp,
    dimension_reductions = c("PCA", "UMAP"),
    n = 10,
    batch_correction = FALSE,
    batch_list = NULL,
    remove_PC = NULL,
    verbose = TRUE
)
```

# **Arguments**

scExp A SingleCellExperiment object.

dimension\_reductions

A character vector of methods to apply. (c('PCA','TSNE','UMAP'))

n Numbers of dimensions to keep for PCA. (50)

batch\_correction

Do batch correction? (FALSE)

batch\_list List of characters. Names are batch names, characters are sample names.

remove\_PC A vector of string indicating which principal components to remove before

downstream analysis as probably correlated to library size. Should be under the form: 'Component\_1', 'Component\_2', ... Recommended when using 'TFIDF'

normalization method. (NULL)

verbose Print messages ?(TRUE)

#### Value

A SingleCellExperiment object containing feature spaces. See ?reduceDims().

#### **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp = normalize_scExp(scExp, "CPM")
scExp = reduce_dims_scExp(scExp,dimension_reductions=c("PCA","UMAP"))</pre>
```

reduce\_dim\_batch\_correction

Reduce dimension with batch corrections

# Description

Reduce dimension with batch corrections

#### Usage

```
reduce_dim_batch_correction(scExp, mat, batch_list, n)
```

## **Arguments**

scExp SingleCellExperiment

mat The normalized count matrix

batch\_list List of batches

n Number of PCs to keep

#### Value

A list containing the SingleCellExperiment with batch info and the corrected pca

remove\_chr\_M\_fun

Remove chromosome M from scExprownames

# Description

Remove chromosome M from scExprownames

#### Usage

```
remove_chr_M_fun(scExp, verbose)
```

# Arguments

scExp A SingleCellExperiment

verbose Print?

104 results\_enrichmentTest

## Value

A SingleCellExperiment without chromosome M (mitochondrial chr)

```
remove_non_canonical_fun
```

Remove non canonical chromosomes from scExp

# Description

Remove non canonical chromosomes from scExp

## Usage

```
remove_non_canonical_fun(scExp, verbose)
```

#### **Arguments**

scExp A SingleCellExperiment

verbose Print?

## Value

A SingleCellExperiment without non canonical chromosomes (random,unknown, contigs etc...)

```
results_enrichmentTest
```

Resutls of hypergeometric gene set enrichment test

## **Description**

Run hypergeometric enrichment test and combine significant pathways into a data.frame

## Usage

```
results_enrichmentTest(
  differentialGenes,
  enrichment_qval,
  GeneSets,
  GeneSetsDf,
  GenePool
)
```

## Arguments

```
differentialGenes
```

Genes significantly over / under expressed

enrichment\_qval

Adusted p-value threshold above which a pathway is considered significative

GeneSets List of pathways

GeneSetsDf Data.frame of pathways

GenePool Pool of possible genes for testing

#### Value

A data.frame with pathways passing q.value threshold

```
retrieve_top_bot_features_pca
```

Retrieve Top and Bot most contributing features of PCA

# Description

Retrieve Top and Bot most contributing features of PCA

#### Usage

```
retrieve_top_bot_features_pca(
  pca,
  counts,
  component,
  n_top_bot,
  absolute = FALSE
)
```

## **Arguments**

pca A matrix/data.frame of rotated data counts the normalized counts used for PCA

component the componenent of interest

n\_top\_bot the number of top & bot features to take

absolute If TRUE, return the top features in absolute values instead.

#### Value

a data.frame of top bot contributing features in PCA

run\_tsne\_scExp

run\_pairwise\_tests

Run pairwise tests

# Description

Run pairwise tests

#### Usage

```
run_pairwise_tests(
   affectation,
   by,
   counts,
   feature,
   method,
   progress = NULL,
   BPPARAM = BiocParallel::bpparam()
)
```

## Arguments

affectation An annotation data.frame with cell\_cluster and cell\_id columns

by = A character specifying the column of the object containing the groups of cells

to compare.

counts Count matrix

feature Feature data.frame

method DA method, Wilcoxon or edgeR

progress A shiny Progress instance to display progress bar.

BPPARAM object for multiprocessing. See bpparam for more informations.

Will take the default BPPARAM set in your R session.

## Value

A list containing objects for DA function

run\_tsne\_scExp

Run tsne on single cell experiment

## **Description**

Run tsne on single cell experiment

scExp 107

#### Usage

```
run_tsne_scExp(scExp, verbose = FALSE)
```

#### **Arguments**

scExp A SingleCellExperiment Object

verbose Print?

#### Value

A colored kable with the number of cells per sample for display

scExp

A SingleCellExperiment outputed by ChromSCape

## **Description**

Data from a single-cell ChIP-seq experiment against H3K4me3 active mark from two cell lines, Jurkat B cells and Ramos T cells from Grosselin et al., 2019. The count matrices, on 5kbp bins, were given to ChromSCape and the filtering parameter was set to 3% of cells active in regions and subsampled down to 150 cells per sample. After correlation filtering, the experiment is composed of respectively 51 and 55 cells from Jurkat & Ramos and 5499 5kbp-genomic bins where signal is located.

#### Usage

```
data("scExp")
```

#### **Format**

scExp - a SingleCellExperiment with 106 cells and 5499 features (genomic bins) in hg38:

chr A SingleCellExperiment

#### **Details**

The scExp is composed of:

- counts and normcounts assays, PCA, UMAP, and Correlation matrix in reducedDims(scExp)
- Assignation of genes to genomic bins in rowRanges(scExp)
- Cluster information in colData(scExp) correlation
- Hierarchical clustering dengogram in metadata\$hc\_cor
- Consensus clustering raw data in metadata\$consclust
- Consensus clustering cluster-consensus and item consensus dataframes in metadata\$icl
- Differential analysis in metadata\$diff
- Gene Set Analysis in metadata\$enr

## Value

A scExp - a SingleCellExperiment with 106 cells and 5499 features

# **Examples**

```
data("scExp")
plot_reduced_dim_scExp(scExp)
plot_reduced_dim_scExp(scExp,color_by = "cell_cluster")
plot_heatmap_scExp(scExp)
plot_differential_volcano_scExp(scExp, "C1")
plot_differential_summary_scExp(scExp)
```

```
separate_BAM_into_clusters
```

Separate BAM files into cell cluster BAM files

# **Description**

Separate BAM files into cell cluster BAM files

#### Usage

```
separate_BAM_into_clusters(affectation, odir, merged_bam)
```

## **Arguments**

affectation An annotation data.frame containing cell\_id and cell\_cluster columns

odir A valid output directory path

merged\_bam A list of merged bam file paths

@importFrom Rsamtools filterBam ScanBamParam

# Value

Create one BAM per cluster from one BAM per condition

separator\_count\_mat 109

separator\_count\_mat

Determine Count matrix separator ("tab" or ",")

# **Description**

Determine Count matrix separator ("tab" or ",")

#### Usage

```
separator_count_mat(path_to_matrix)
```

# **Arguments**

path\_to\_matrix A path towards the count matrix to check

# Value

A character separator

smoothBin

Smooth a vector of values with nb\_bins left and righ values

# Description

Smooth a vector of values with nb\_bins left and righ values

# Usage

```
smoothBin(bin_score, nb_bins = 10)
```

# **Arguments**

bin\_score A numeric vector of values to be smoothed

nb\_bins Number of values to take left and right

@importFrom BiocParallel bpvec

# Value

A smooth vector of the same size

subsample\_scExp

Subsample scExp

# **Description**

Randomly sample x cells from each sample in a SingleCellExperiment to return a subsampled SingleCellExperiment with all samples having maximum n cells. If n is higher than the number of cell in a sample, this sample will not be subsampled.

# Usage

```
subsample_scExp(scExp, n_cell_per_sample = 500, n_cell_total = NULL)
```

#### **Arguments**

```
\begin{tabular}{lll} sc Exp & A Single Cell Experiment \\ n\_cell\_per\_sample & An integer number of cells to subsample for each sample. Exclusive with n\_cells\_total. \\ (500) & \\ n\_cell\_total & An integer number of cells to subsample in total. Exclusive with n\_cell\_per\_sample \\ (NULL). & \\ \end{tabular}
```

#### Value

A subsampled SingleCellExperiment

#### **Examples**

```
raw <- create_scDataset_raw()
scExp = create_scExp(raw$mat, raw$annot)
scExp_sub = subsample_scExp(scExp,50)
## Not run: num_cell_scExp(scExp_sub)</pre>
```

subset\_bam\_call\_peaks Peak calling on cell clusters

# Description

This functions does peak calling on each cell population in order to refine gene annotation for large bins. For instance, a 50000bp bins might contain the TSS of several genes, while in reality only one or two of these genes are overlapping the signal (peak). To do so, first in-silico cell sorting is applied based on previously defined clusters contained in the SingleCellExperiment. Taking BAM files of each sample as input, samtools pools then splits reads from each cell barcode into 1 BAM file per cell cluster (pseudo-bulk). Then MACS2 calls peaks on each cluster. The peaks are aggregated and merged if closer to a certain distance defined by user (10000bp). Then,

bar-

This function takes as input a SingleCellExperiment, that must contain a 'cell\_cluster' column in it's colData, an output directory where to store temporary files, the list of BAM files corresponding to each sample and containing the cell barcode information as a tag (for instance tag CB:Z:xxx, XB:Z:xxx or else...) or single-cell BED files containing the raw reads and corresponding to the 'barcode' column metadata, the p.value used by MACS2 to distinguish significant peaks, the reference genome (either hg38 or mm10), the maximal merging distance in bp and a data.frame containing gene TSS genomic cooridnates of corresponding genome (if set to NULL, will automatically load geneTSS). The output is a SingleCellExperiment with GRanges object containing ranges of each merged peaks that falls within genomic bins of the SingleCellExperiment, saving the bin range as additional column (window\_chr, window\_start, window\_end), as well as the closests genes and their distance relative to the peak. The peaks may be present in several rows if multiple genes are close / overlap to the peaks.

Note that the user must have MACS2 installed and available in the PATH. Users can open command terminal and type 'which macs2' to verify the availability of these programs. Will only work on unix operating system. Check operating system with 'print(.Platform)'.

#### Usage

```
subset_bam_call_peaks(
    scExp,
    odir,
    input,
    format = "BAM",
    p.value = 0.05,
    ref = "hg38",
    peak_distance_to_merge = 10000,
    geneTSS_annotation = NULL,
    run_coverage = FALSE,
    progress = NULL
)
```

# **Arguments**

scExp

odir	Output directory where to write temporary files and each cluster's BAM file
input	A character vector of file paths to each sample's BAM file, containing cell b

code information as tags. BAM files can be paired-end or single-end.

format Format of the input data, either "BAM" or "scBED".

A SingleCellExperiment object

p. value a p-value to use for MACS2 to determine significant peaks. (0.05)

ref A reference genome, either hg38, mm10 or ce11. ('hg38')

peak\_distance\_to\_merge

Maximal distance to merge peaks together after peak calling, in bp. (10000)

geneTSS\_annotation

A data.frame annotation of genes TSS. If NULL will automatically load Gen-

code list of genes fro specified reference genome.

run\_coverage Create coverage tracks (.bw) for each cluster?

progress A shiny Progress instance to display progress bar.

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#### **Details**

The BED files of the peaks called for each clusters, as well as the merged peaks are written in the output directory.

#### Value

A SingleCellExperiment with refinded annotation

# **Examples**

```
## Not run:
data("scExp")
subset_bam_call_peaks(scExp, "path/to/out/", list("sample1" =
    "path/to/BAM/sample1.bam", "sample2" = "path/to/BAM/sample2.bam"),
    p.value = 0.05, ref = "hg38", peak_distance_to_merge = 10000,
    geneTSS_annotation = NULL)
## End(Not run)
```

summary\_DA

Summary of the differential analysis

# **Description**

Summary of the differential analysis

# Usage

```
summary_DA(scExp, qval.th = 0.01, logFC.th = 1, min.percent = 0.01)
```

#### **Arguments**

scExp A SingleCellExperiment object containing consclust with selected number of

cluster.

qval.th Adjusted p-value threshold. (0.01)

logFC.th Fold change threshold. (1)

min.percent Minimum fraction of cells having the feature active to consider it as significantly

differential. (0.01)

### Value

A table summary of the differential analysis

# **Examples**

```
data('scExp')
summary_DA(scExp)
```

```
swapAltExp_sameColData
```

Swap main & alternative Experiments, with fixed colData

# **Description**

Swap main & alternative Experiments, with fixed colData

#### Usage

```
swapAltExp_sameColData(scExp, alt)
```

# **Arguments**

scExp A SingleCellExperiment
alt Name of the alternative experiment

#### Value

A swapped SingleCellExperiment with the exact same colData.

# **Examples**

```
data(scExp)
swapAltExp_sameColData(scExp, "peaks")
```

```
table_enriched_genes_scExp
```

Creates table of enriched genes sets

# Description

Creates table of enriched genes sets

# Usage

```
table_enriched_genes_scExp(
    scExp,
    set = "Both",
    group = "C1",
    enr_class_sel = c("c1_positional", "c2_curated", "c3_motif", "c4_computational",
        "c5_G0", "c6_oncogenic", "c7_immunologic", "hallmark")
)
```

114 warning\_DA

# **Arguments**

scExp	A SingleCellExperiment object containing list of enriched gene sets.
set	A character vector, either 'Both', 'Overexpressed' or 'Underexpressed'. ('Both')
group	The "group" name from differential analysis. Can be the cluster name or the custom name in case of a custom differential analysis.
enr_class_sel	Which classes of gene sets to show. (c('c1_positional', 'c2_curated',))

# Value

A DT::data.table of enriched gene sets.

# **Examples**

```
data("scExp")
## Not run: table_enriched_genes_scExp(scExp)
```

warning_DA	Warning for differential_analysis_scExp	
------------	---	--

# Description

Warning for differential\_analysis\_scExp

# Usage

```
warning_DA(scExp, by, de_type, method, block, group, ref)
```

# Arguments

scExp	A SingleCellExperiment object containing consclust with selected number of cluster.
by	= A character specifying the column of the object containing the groups of cells to compare. Exclusive with de_type == custom
de_type	Type of comparisons. Either 'one_vs_rest', to compare each cluster against all others, or 'pairwise' to make 1 to 1 comparisons. ('one_vs_rest')
method	Wilcoxon or edgerGLM
block	Use batches as blocking factors?
group	If de_type is custom, the group to compare (data.frame), must be a one-column data.frame with cell_clusters or sample_id as character in rows
ref	If de_type is custom, the reference to compare (data.frame), must be a one-column data.frame with cell_clusters or sample_id as character in rows

# Value

Warnings or Errors if the input are not correct

# **Description**

```
warning_filter_correlated_cell_scExp
```

# Usage

```
warning_filter_correlated_cell_scExp(
    scExp,
    random_iter,
    corr_threshold,
    percent_correlation,
    run_tsne,
    downsample,
    verbose
)
```

#### **Arguments**

SCEXP A SingleCellExperiment object containing 'Cor', a correlation matrix, in re-

ducedDims.

random\_iter Number of random matrices to create to calculate random correlation scores.

(50)

corr\_threshold Quantile of random correlation score above which a cell is considered to be

'correlated' with another cell. (99)

percent\_correlation

Percentage of the cells that any cell must be 'correlated' to in order to not be

filtered. (1)

run\_tsne Re-run tsne ? (FALSE)

downsample Number of cells to calculate correlation filtering threshold? (2500)

verbose (TRUE)

# Value

Warnings or Errors if the input are not correct

#### **Description**

A warning helper for plot\_reduced\_dim\_scExp

# Usage

```
warning_plot_reduced_dim_scExp(
    scExp,
    color_by,
    reduced_dim,
    downsample,
    transparency,
    size,
    max_distanceToTSS,
    annotate_clusters,
    min_quantile,
    max_quantile
)
```

# **Arguments**

scExp A SingleCellExperiment Object
color\_by Feature used for coloration
reduced\_dim Reduced Dimension used for plotting
downsample Number of cells to downsample
transparency Alpha parameter, between 0 and 1
size Size of the points.
max\_distanceToTSS
Numeric. Maximum distance to a ge

Numeric. Maximum distance to a gene's TSS to consider a region linked to a gene.

annotate\_clusters

A logical indicating if clusters should be labelled. The 'cell\_cluster' column

should be present in metadata.

min\_quantile The lower threshold to remove outlier cells, as quantile of cell embeddings (be-

tween 0 and 0.5).

max\_quantile The upper threshold to remove outlier cells, as quantile of cell embeddings (be-

tween 0.5 and 1).

#### Value

Warning or errors if the inputs are not correct

# Description

Warning for raw\_counts\_to\_sparse\_matrix

# Usage

```
warning_raw_counts_to_sparse_matrix(
   files_dir_list,
   file_type = c("scBAM", "scBED", "SparseMatrix"),
   peak_file = NULL,
   n_bins = NULL,
   bin_width = NULL,
   genebody = NULL,
   extendPromoter = 2500,
   verbose = TRUE,
   ref = "hg38"
)
```

# **Arguments**

files_dir_list	A named character vector of directory containing the raw files
file_type	Input file(s) type(s) ('scBED', 'scBAM', 'SparseMatrix')
peak_file	A file containing genomic location of peaks (NULL)
n_bins	The number of bins to tile the genome (NULL)
bin_width	The size of bins to tile the genome (NULL)
genebody	Count on genes (body + promoter) ? (NULL)
extendPromoter	If counting on genes, number of base pairs to extend up or downstream of TSS $(2500)$ .
verbose	Verbose (TRUE)
ref	reference genome to use (hg38)

# Value

Error or warnings if the input are not correct

```
wrapper_Signac_FeatureMatrix
```

Wrapper around 'FeatureMatrix' function from Signac Package

# **Description**

Wrapper around 'FeatureMatrix' function from Signac Package

# Usage

```
wrapper_Signac_FeatureMatrix(
    files_dir_list,
    which,
    ref = "hg38",
    process_n = 2000,
    set_future_plan = TRUE,
    verbose = TRUE,
    progress = NULL
)
```

#### **Arguments**

files\_dir\_list A named character vector of directories containing the files. The names corre-

spond to sample names.

which A GenomicRanges containing the features to count on.

ref Reference genome to use (hg38). Chromosomes that are not present in the canon-

ical chromosomes of the given reference genome will be excluded from the ma-

trix.

process\_n Number of regions to load into memory at a time, per thread. Processing more

regions at once can be faster but uses more memory. (2000)

set\_future\_plan

Set 'multisession' plan within the function (TRUE). If TRUE, the previous plan

(e.g. future::plan()) will be set back on exit.

verbose Verbose (TRUE).

progress Progress object for Shiny.

#### **Details**

Signac & future are not required packages for ChromSCape as they are required only for the fragment matrix calculations. To use this function, install Signac package first (future will be installed as a dependency). For the simplicity of the application & optimization, the function by defaults sets future::plan("multisession") with workers = future::availableCores(omit = 1) in order to allow parallel processing with Signac. On exit the plan is re-set to the previously set future plan. Note that future multisession may have trouble running when VPN is on. To run in parallel, first deactivate your VPN if you encounter long runtimes.

# Value

A sparse matrix of features x cells

#### References

Stuart el al., Multimodal single-cell chromatin analysis with Signac bioRxiv https://doi.org/10.1101/2020.11.09.373613

# **Examples**

```
## Not run:
gr_bins = define_feature("hg38", bin_width = 50000)
wrapper_Signac_FeatureMatrix("/path/to/dir_containing_fragment_files",
    gr_bins, ref = "hg38")
## End(Not run)
```

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