Package 'scrapper'

January 20, 2025

Version 1.0.3

Date 2024-12-17

Title Bindings to C++ Libraries for Single-Cell Analysis

Description Implements R bindings to C++ code for analyzing single-

cell (expression) data, mostly from various libscran libraries.

Each function performs an individual step in the single-

cell analysis workflow, ranging from quality control to clustering and marker detection.

It is mostly intended for other Bioconductor package developers to build more user-friendly end-to-end workflows.

License MIT + file LICENSE

Imports methods, Rcpp, beachmat (>= 2.21.6), DelayedArray, BiocNeighbors (>= 1.99.0), igraph, parallel

Suggests testthat, knitr, rmarkdown, BiocStyle, MatrixGenerics, sparseMatrixStats, Matrix, scRNAseq

LinkingTo Rcpp, assorthead, beachmat, BiocNeighbors

biocViews Normalization, RNASeq, Software, GeneExpression, Transcriptomics, SingleCell, BatchEffect, QualityControl, DifferentialExpression, FeatureExtraction, PrincipalComponent, Clustering

VignetteBuilder knitr

Encoding UTF-8

RoxygenNote 7.3.2

git_url https://git.bioconductor.org/packages/scrapper

git_branch RELEASE_3_20

git_last_commit 732a984

git_last_commit_date 2024-12-17

Repository Bioconductor 3.20

Date/Publication 2025-01-19

Author Aaron Lun [cre, aut]

Maintainer Aaron Lun <infinite.monkeys.with.keyboards@gmail.com>

2 adt_quality_control

Contents

adt_quality_control	. 2
aggregateAcrossCells	. 4
buildSnnGraph	. 5
centerSizeFactors	
chooseHighlyVariableGenes	
choosePseudoCount	
clusterGraph	
clusterKmeans	
combineFactors	
computeClrm1Factors	
correctMnn	
crispr_quality_control	
fitVarianceTrend	
modelGeneVariances	
normalizeCounts	
rna_quality_control	
runAllNeighborSteps	
runPca	
runTsne	
runUmap	
sanitizeSizeFactors	
scaleByNeighbors	
scoreGeneSet	
scoreMarkers	
subsampleByNeighbors	
summarizeEffects	
Summunizedinetts	. 50
Index	38

adt_quality_control Quality control for ADT count data

Description

Compute per-cell QC metrics from an initialized matrix of ADT counts, and use the metrics to suggest filter thresholds to retain high-quality cells.

Usage

```
computeAdtQcMetrics(x, subsets, num.threads = 1)
suggestAdtQcThresholds(
  metrics,
  block = NULL,
  min.detected.drop = 0.1,
  num.mads = 3
)
filterAdtQcMetrics(thresholds, metrics, block = NULL)
```

adt_quality_control 3

Arguments

x A matrix-like object where rows are ADTs and columns are cells. Values are

expected to be counts.

subsets List of vectors specifying tag subsets of interest, typically control tags like IgGs.

Each vector may be logical (whether to keep each row), integer (row indices) or

character (row names).

num. threads Integer scalar specifying the number of threads to use.

metrics List with the same structure as produced by computeAdtQcMetrics.

block Factor specifying the block of origin (e.g., batch, sample) for each cell in metrics.

Alternatively NULL if all cells are from the same block.

For filterAdtQcMetrics, a blocking factor should be provided if block was

used to construct thresholds.

min.detected.drop

Minimum drop in the number of detected features from the median, in order to

consider a cell to be of low quality.

num.mads Number of median from the median, to define the threshold for outliers in each

metric.

thresholds List with the same structure as produced by suggestAdtQcThresholds.

Value

For computeAdtQcMetrics, a list is returned containing:

- sum, a numeric vector containing the total ADT count for each cell.
- detected, an integer vector containing the number of detected tags per cell.
- subsets, a list of numeric vectors containing the total count of each control subset.

Each vector is of length equal to the number of cells.

For suggestAdtQcThresholds with block!=NULL, a list is returned containing:

- detected, a numeric vector containing the lower bound on the number of detected tags for each blocking level.
- subsets, a list of numeric vectors containing the upper bound on the sum of counts in each control subset for each blocking level.

Each vector is of length equal to the number of levels in block.

For suggestAdtQcThresholds with block=NULL, a list is returned containing:

- detected, a numeric scalar containing the lower bound on the number of detected tags.
- subsets, a numeric vector containing the upper bound on the sum of counts in each control subset.

For filterAdtQcMetrics, a logical vector of length ncol(x) is returned indicating which cells are of high quality.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_qc/, for the rationale of QC filtering on ADT counts.

Examples

```
# Mocking a matrix:
library(Matrix)
x <- round(abs(rsparsematrix(1000, 100, 0.1) * 100))
# Mocking up a control set.
sub <- list(IgG=rbinom(nrow(x), 1, 0.1) > 0)

qc <- computeAdtQcMetrics(x, sub)
str(qc)

filt <- suggestAdtQcThresholds(qc)
str(filt)

keep <- filterAdtQcMetrics(filt, qc)
summary(keep)</pre>
```

Description

Aggregate expression values across cells based on one or more grouping factors. This is primarily used to create pseudo-bulk profiles for each cluster/sample combination.

Usage

```
aggregateAcrossCells(x, factors, num.threads = 1)
```

Arguments

A matrix-like object where rows correspond to genes or genomic features and columns correspond to cells. Values are typically expected to be counts.

A list or data frame containing one or more grouping factors, see combineFactors.

Integer specifying the number of threads to be used for aggregation.

Value

A list containing:

- sums, a numeric matrix where each row corresponds to a gene and each column corresponds to a unique combination of grouping levels. Each entry contains the summed expression across all cells with that combination.
- detected, an integer matrix where each row corresponds to a gene and each column corresponds to a unique combination of grouping levels. Each entry contains the number of cells with detected expression in that combination.
- combinations, a data frame describing the levels for each unique combination of factors. Rows of this data frame correspond to columns of sums and detected, while columns correspond to the factors in factors.

buildSnnGraph 5

• counts, the number of cells associated with each combination. Each entry corresponds to a row of combinations.

• index, an integer vector of length equal to the number of cells in x. This specifies the combination in combinations to which each cell was assigned.

Author(s)

Aaron Lun

Examples

```
# Mocking a matrix:
library(Matrix)
x <- round(abs(rsparsematrix(1000, 100, 0.1) * 100))

# Simple aggregation:
clusters <- sample(LETTERS, 100, replace=TRUE)
agg <- aggregateAcrossCells(x, list(cluster=clusters))
str(agg)

# Multi-factor aggregation
samples <- sample(1:5, 100, replace=TRUE)
agg2 <- aggregateAcrossCells(x, list(cluster=clusters, sample=samples))
str(agg2)</pre>
```

buildSnnGraph

Build a shared nearest neighbor graph

Description

Build a shared nearest neighbor (SNN) graph where each node is a cell. Edges are formed between cells that share one or more nearest neighbors, weighted by the number or importance of those shared neighbors.

Usage

```
buildSnnGraph(
    x,
    num.neighbors = 10,
    weight.scheme = "ranked",
    num.threads = 1,
    BNPARAM = AnnoyParam()
)
```

Arguments

Х

For buildSnnGraph, a numeric matrix where rows are dimensions and columns are cells, typically containing a low-dimensional representation from, e.g., runPca. Alternatively, a named list of nearest-neighbor search results. This should contain index, an integer matrix where rows are neighbors and columns are cells.

6 buildSnnGraph

Each column contains 1-based indices for the nearest neighbors of the corresponding cell, ordered by increasing distance. The number of neighbors for each cell should be equal to num.neighbors, otherwise a warning is raised.

Alternatively, an index constructed by buildIndex.

num.neighbors Integer scalar specifying the number of neighbors to use to construct the graph.

weight.scheme String specifying the weighting scheme to use for constructing the SNN graph.

This can be "ranked" (default), "jaccard" or "number".

num. threads Integer scalar specifying the number of threads to use. Only used if x is not a

list of existing nearest-neighbor search results.

BNPARAM A BiocNeighborParam object specifying the algorithm to use. Only used if x is

not a list of existing nearest-neighbor search results.

Value

If as.pointer=FALSE, a list is returned containing:

- vertices, an integer scalar specifying the number of vertices in the graph (i.e., cells in x).
- edges, an integer vector of 1-based indices for graph edges. Pairs of values represent the endpoints of an (undirected) edge, i.e., edges[1:2] form the first edge, edges[3:4] form the second edge and so on.
- weights, a numeric vector of weights for each edge. This has length equal to half the length of edges.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_graph_cluster/, for details on the underlying implementation.

```
data <- matrix(rnorm(10000), ncol=1000)
out <- buildSnnGraph(data)
str(out)

# We can use this to make an igraph::graph.
g <- igraph::make_undirected_graph(out$edges, n = out$vertices)
igraph::E(g)$weight <- out$weight</pre>
```

centerSizeFactors 7

centerSizeFactors

Center size factors

Description

Scale the size factors so they are centered at unity, which ensures that the scale of the counts are preserved (on average) after normalization.

Usage

```
centerSizeFactors(size.factors, block = NULL, mode = c("lowest", "per-block"))
```

Arguments

block Vector or factor of length equal to size. factors, specifying the block of origin

for each cell. Alternatively NULL, in which case all cells are assumed to be in the

same block.

mode String specifying how to scale size factors across blocks. "lowest" will scale

all size factors by the the lowest per-block average. "per-block" will center

the size factors in each block separately.

Value

Numeric vector of length equal to size. factors, containing the centered size factors.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_norm/, for the rationale behind centering the size factors.

```
centerSizeFactors(runif(100))
centerSizeFactors(runif(100), block=sample(3, 100, replace=TRUE))
```

chooseHighlyVariableGenes

Choose highly variable genes

Description

Choose highly variable genes based on a variance-related statistic.

Usage

```
chooseHighlyVariableGenes(stats, top = 4000, larger = TRUE, keep.ties = TRUE)
```

Arguments

stats	Numeric vector of variances (or a related statistic) across all genes. Typically the residuals from modelGeneVariances are used here.
top	Integer specifying the number of top genes to retain.
larger	Logical scalar indicating whether larger values of stats correspond to more variable genes.
keep.ties	Logical scalar indicating whether to keep tied values of stats, even if top may

Value

Integer vector containing the indices of genes in stats that are considered to be highly variable.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_variances/, for the underlying implementation.

Examples

```
resids <- rexp(1000)
str(chooseHighlyVariableGenes(resids))</pre>
```

be exceeded.

choosePseudoCount 9

|--|

Description

Choose a suitable pseudo-count to control the bias introduced by log-transformation of normalized counts.

Usage

```
choosePseudoCount(size.factors, quantile = 0.05, max.bias = 1, min.value = 1)
```

Arguments

size.factors	Numeric vector of size factors for all cells.
quantile	Numeric scalar specifying the quantile to use for defining extreme size factors.
max.bias	Numeric scalar specifying the maximum allowed bias.
min.value	Numeric scalar specifying the minimum value for the pseudo-count.

Value

A choice of pseudo-count for normalizeCounts.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_norm/, for the motivation behind calculating a larger pseudocount.

```
sf <- runif(100)
choosePseudoCount(sf)
choosePseudoCount(sf, quantile=0.01)
choosePseudoCount(sf, max.bias=0.5)</pre>
```

10 clusterGraph

clusterGraph

Graph-based clustering of cells

Description

Identify clusters of cells using a variety of community detection methods from a graph where similar cells are connected.

Usage

```
clusterGraph(
   x,
   method = c("multilevel", "leiden", "walktrap"),
   multilevel.resolution = 1,
   leiden.resolution = 1,
   leiden.objective = c("modularity", "cpm"),
   walktrap.steps = 4,
   seed = 42
)
```

Arguments

x List containing graph information or an external pointer to a graph, as returned by buildSnnGraph. Alternatively, an igraph object with edge weights.

method String specifying the algorithm to use.

multilevel.resolution

Numeric scalar specifying the resolution when method="multilevel".

leiden.resolution

Numeric scalar specifying the resolution when method="leiden".

leiden.objective

String specifying the objective function when method="leiden".

 $walktrap.\,steps\ \ Integer\,scalar\,specifying\,the\,number\,of\,steps\,to\,use\,when\,method = "walktrap".$

seed Integer scalar specifying the random seed to use for method="multilevel" or "leiden".

Value

A list containing membership, an integer vector containing the cluster assignment for each cell; and status, an integer scalar indicating whether the algorithm completed successfully (0) or not (non-zero). Additional fields may be present depending on the method:

- For method="multilevel", the levels list contains the clustering result at each level of the algorithm. A modularity numeric vector also contains the modularity at each level, the highest of which corresponds to the reported membership.
- For method="leiden", a quality numeric scalar containg the quality of the partitioning.
- For method="walktrap", a merges matrix specifies the pair of cells or clusters that were merged at each step of the algorithm. A modularity numeric scalar also contains the modularity of the final partitioning.

clusterKmeans 11

Author(s)

Aaron Lun

See Also

```
https://igraph.org, for the underlying implementation of each clustering method.
https://libscran.github.io/scran_graph_cluster/, for wrappers around the igraph code.
```

Examples

```
data <- matrix(rnorm(10000), ncol=1000)
gout <- buildSnnGraph(data)
str(gout)

str(clusterGraph(gout))
str(clusterGraph(gout, method="leiden"))
str(clusterGraph(gout, method="walktrap"))</pre>
```

clusterKmeans

K-means clustering

Description

Perform k-means clustering using kmeans++ initialization with the Hartigan-Wong algorithm.

Usage

```
clusterKmeans(
    x,
    k,
    init.method = c("var-part", "kmeans++", "random"),
    refine.method = c("hartigan-wong", "lloyd"),
    var.part.optimize.partition = TRUE,
    var.part.size.adjustment = 1,
    lloyd.iterations = 100,
    hartigan.wong.iterations = 10,
    hartigan.wong.quick.transfer.iterations = 50,
    hartigan.wong.quit.quick.transfer.failure = FALSE,
    seed = 5489L,
    num.threads = 1
)
```

Arguments

X	Numeric matrix where rows are dimensions and columns are cells.
k	Integer scalar specifying the number of clusters.
init.method	String specifying the initialization method: variance partitioning ("var-part"), kmeans++ ("kmeans++") or random initialization ("random").
refine.method	String specifying the refinement method: Lloyd's algorithm ("lloyd") or the Hartigan-Wong algorithm ("hartigan-wong").

12 clusterKmeans

var.part.optimize.partition

Logical scalar indicating whether each partition boundary should be optimized to reduce the sum of squares in the child partitions.

var.part.size.adjustment

Numeric scalar between 0 and 1, specifying the adjustment to the cluster size when prioritizing the next cluster to partition. Setting this to 0 will ignore the cluster size while setting this to 1 will generally favor larger clusters.

lloyd.iterations

Integer scalar specifying the maximmum number of iterations for the Lloyd algorithm.

hartigan.wong.iterations

Integer scalar specifying the maximmum number of iterations for the Hartigan-Wong algorithm.

hartigan.wong.quick.transfer.iterations

Integer scalar specifying the maximmum number of quick transfer iterations for the Hartigan-Wong algorithm.

hartigan.wong.quit.quick.transfer.failure

Logical scalar indicating whether to quit the Hartigan-Wong algorithm upon convergence failure during quick transfer iterations.

seed Integer scalar specifying the seed to use for random or kmeans++ initialization.

num. threads Integer scalar specifying the number of threads to use.

Value

By default, a list is returned containing:

- clusters, a factor containing the cluster assignments for each cell.
- centers, a numeric matrix with the coordinates of the cluster centroids (dimensions in rows, centers in columns).
- iterations, an integer scalar specifying the number of refinement iterations that were performed
- status, an integer scalar specifying the convergence status. Any non-zero value indicates a convergence failure though the exact meaning depends on the choice of refine.method.

Author(s)

Aaron Lun

```
x <- t(as.matrix(iris[,1:4]))
clustering <- clusterKmeans(x, k=3)
table(clustering$clusters, iris[,"Species"])</pre>
```

combineFactors 13

|--|--|--|

Description

Combine multiple categorical factors based on the unique combinations of levels from each factor.

Usage

```
combineFactors(factors, keep.unused = FALSE)
```

Arguments

factors List of vectors or factors of the same length. Corresponding elements across

all vectors/factors represent the combination of levels for a single observation. For factors, any existing levels are respected. For other vectors, the sorted and

unique values are used as levels.

Alternatively, a data frame where each column is a vector or factor and each row

corresponds to an observation.

keep. unused Logical scalar indicating whether to report unused combinations of levels.

Value

List containing levels, a data frame containing the sorted and unique combinations of levels from factors; and index, an integer vector specifying the index into levels for each observation.

Author(s)

Aaron Lun

```
combineFactors(list(
    sample(LETTERS[1:5], 100, replace=TRUE),
    sample(3, 100, replace=TRUE)
))

combineFactors(list(
    factor(sample(LETTERS[1:5], 10, replace=TRUE), LETTERS[1:5]),
    factor(sample(5, 10, replace=TRUE), 1:5)
), keep.unused=TRUE)
```

14 correctMnn

computeClrm1Factors

Compute size factors for ADT counts

Description

Compute size factors from an ADT count matrix using the CLRm1 method.

Usage

```
computeClrm1Factors(x, num.threads = 1)
```

Arguments

x A matrix-like object containing ADT count data. Rows correspond to tags and

columns correspond to cells.

num. threads Number of threads to use.

Value

Numeric vector containing the CLRm1 size factor for each cell.

Author(s)

Aaron Lun

See Also

https://github.com/libscran/clrm1, for a description of the CLRm1 method.

Examples

```
library(Matrix)
x <- abs(rsparsematrix(1000, 100, 0.1) * 10)
head(computeClrm1Factors(x))</pre>
```

correctMnn

Batch correction with mutual nearest neighbors

Description

Apply mutual nearest neighbor (MNN) correction to remove batch effects from a low-dimensional matrix.

correctMnn 15

Usage

```
correctMnn(
    x,
    block,
    num.neighbors = 15,
    num.mads = 3,
    robust.iterations = 2,
    robust.trim = 0.25,
    mass.cap = NULL,
    order = NULL,
    reference.policy = c("max-rss", "max-size", "max-variance", "input"),
    BNPARAM = AnnoyParam(),
    num.threads = 1
)
```

Arguments

Numeric matrix where rows are dimensions and columns are cells, typically

containing low-dimensional coordinates (e.g., from runPca).

block Factor specifying the block of origin (e.g., batch, sample) for each cell in x.

num.neighbors Integer scalar specifying the number of neighbors to use when identifying MNN

pairs.

num.mads Numeric scalar specifying the number of median absolute deviations to use for

removing outliers in the center-of-mass calculations.

robust.iterations

Integer scalar specifying the number of iterations for robust calculation of the

center of mass.

robust.trim Numeric scalar in [0, 1) specifying the trimming proportion for robust calcula-

tion of the center of mass.

mass.cap Integer scalar specifying the cap on the number of observations to use for center-

of-mass calculations on the reference dataset. A value of 100,000 may be appropriate for speeding up correction of very large datasets. If NULL, no cap is

used.

order Vector containing levels of batch in the desired merge order. If NULL, a suitable

merge order is automatically determined.

reference.policy

String specifying the policy to use to choose the first reference batch. This can be based on the largest batch ("max-size"), the most variable batch ("max-variance"), the batch with the largest residual sum of squares ("max-rss"), or the first specified input ("input"). Only used for automatic merges, i.e., when order=NULL.

BNPARAM A BiocNeighborParam object specifying the nearest-neighbor algorithm to use.

num. threads Integer scalar specifying the number of threads to use.

Value

List containing:

• corrected, a numeric matrix of the same dimensions as x, containing the corrected values.

- merge.order, character vector containing the unique levels of batch in the automatically determined merge order. The first level in this vector is used as the reference batch; all other batches are iteratively merged to it.
- num.pairs, integer vector of length equal to the number of batches minus 1. This contains the number of MNN pairs at each merge.

Author(s)

Aaron Lun

Examples

```
# Mocking up a dataset with multiple batches.
x <- matrix(rnorm(10000), nrow=10)
b <- sample(3, ncol(x), replace=TRUE)
x[,b==2] <- x[,b==2] + 3
x[,b==3] <- x[,b==3] + 5
lapply(split(colMeans(x), b), mean) # indeed the means differ...
corrected <- correctMnn(x, b)
str(corrected)
lapply(split(colMeans(corrected$corrected), b), mean) # now merged.</pre>
```

```
crispr_quality_control
```

Quality control for CRISPR count data

Description

Compute per-cell QC metrics from an initialized matrix of CRISPR counts, and use the metrics to suggest filter thresholds to retain high-quality cells.

Usage

```
computeCrisprQcMetrics(x, num.threads = 1)
suggestCrisprQcThresholds(
  metrics,
  block = NULL,
  min.detected.drop = 0.1,
  num.mads = 3
)
filterCrisprQcMetrics(thresholds, metrics, block = NULL)
```

Arguments

x A matrix-like object where rows are CRISPRs and columns are cells. Values are expected to be counts.

num.threads Integer scalar specifying the number of threads to use.

metrics List with the same structure as produced by computeCrisprQcMetrics.

crispr_quality_control 17

block Factor specifying the block of origin (e.g., batch, sample) for each cell in metrics.

Alternatively NULL if all cells are from the same block.

For filterCrisprQcMetrics, a blocking factor should be provided if block

was used to construct thresholds.

min.detected.drop

Minimum drop in the number of detected features from the median, in order to

consider a cell to be of low quality.

num.mads Number of median from the median, to define the threshold for outliers in each

metric.

thresholds List with the same structure as produced by suggestCrisprQcThresholds.

Value

For computeCrisprQcMetrics, a list is returned containing:

- sum, a numeric vector containing the total CRISPR count for each cell.
- detected, an integer vector containing the number of detected guides per cell.
- max.value, a numeric vector containing the count for the most abundant guide in cell.
- max.index, an integer vector containing the row index of the most abundant guide in cell.

Each vector is of length equal to the number of cells.

For suggestCrisprQcThresholds with block!=NULL, a list is returned containing:

• max.value, a numeric vector containing the lower bound on the maximum counts for each blocking level.

Each vector is of length equal to the number of levels in block.

For suggestCrisprQcThresholds with block=NULL, a list is returned containing:

• max.value, a numeric scalar containing the lower bound on the maximum counts for each blocking level.

For filterCrisprQcMetrics, a logical vector of length ncol(x) is returned indicating which cells are of high quality.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_qc/, for the rationale of QC filtering on CRISPR counts.

```
# Mocking a matrix:
library(Matrix)
x <- round(abs(rsparsematrix(100, 100, 0.1) * 100))
qc <- computeCrisprQcMetrics(x)
str(qc)
filt <- suggestCrisprQcThresholds(qc)
str(filt)</pre>
```

18 fitVarianceTrend

```
keep <- filterCrisprQcMetrics(filt, qc)
summary(keep)</pre>
```

 ${\tt fitVarianceTrend}$

Fit a mean-variance trend

Description

Fit a trend to the per-cell variances with respect to the mean.

Usage

```
fitVarianceTrend(
  means,
  variances,
  mean.filter = TRUE,
  min.mean = 0.1,
  transform = TRUE,
  span = 0.3,
  use.min.width = FALSE,
  min.width = 1,
  min.window.count = 200,
  num.threads = 1
)
```

Arguments

means	Numeric vector containing the mean (log-)expression for each gene.
variances	Numeric vector containing the variance in the (log-)expression for each gene.
mean.filter	Logical scalar indicating whether to filter on the means before trend fitting.
min.mean	Numeric scalar specifying the minimum mean of genes to use in trend fitting. Only used if mean.filter=TRUE.
transform	Logical scalar indicating whether a quarter-root transformation should be applied before trend fitting.
span	$Numeric\ scalar\ specifying\ the\ span\ of\ the\ LOWESS\ smoother.\ Ignored\ if\ use. \verb min.width=TRUE .$
use.min.width	Logical scalar indicating whether a minimum width constraint should be applied to the LOWESS smoother. Useful to avoid overfitting in high-density intervals.
min.width min.window.cou	Minimum width of the window to use when use.min.width=TRUE. nt
	Minimum number of observations in each window. Only used if use.min.width=TRUE.

Value

num.threads

List containing fitted, the fitted values of the trend for each gene; and residuals, the residuals from the trend.

Number of threads to use.

modelGeneVariances 19

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_variances/, for the underlying implementation.

Examples

```
x <- runif(1000)
y <- 2^rnorm(1000)
out <- fitVarianceTrend(x, y)

plot(x, y)
o <- order(x)
lines(x[o], out$fitted[o], col="red")</pre>
```

modelGeneVariances

Model per-gene variances in expression

Description

Compute the variance in (log-)expression values for each gene, and model the trend in the variances with respect to the mean.

Usage

```
modelGeneVariances(
    x,
    block = NULL,
    block.weight.policy = c("variable", "equal", "none"),
    variable.block.weight = c(0, 1000),
    mean.filter = TRUE,
    min.mean = 0.1,
    transform = TRUE,
    span = 0.3,
    use.min.width = FALSE,
    min.width = 1,
    min.window.count = 200,
    num.threads = 1
)
```

Arguments

Х

A matrix-like object where rows correspond to genes or genomic features and columns correspond to cells. It is typically expected to contain log-expression values, e.g., from normalizeCounts.

block

Factor specifying the block of origin (e.g., batch, sample) for each cell in x. Alternatively NULL if all cells are from the same block.

20 modelGeneVariances

block.weight.policy

String specifying the policy to use for weighting different blocks when computing the average for each statistic Only used if block is not NULL.

variable.block.weight

Numeric vector of length 2, specifying the parameters for variable block weighting. The first and second values are used as the lower and upper bounds, respectively, for the variable weight calculation. Only used if block is not NULL and

block.weight.policy = "variable".

mean.filter Logical scalar indicating whether to filter on the means before trend fitting.

min.mean Numeric scalar specifying the minimum mean of genes to use in trend fitting.

Only used if mean.filter=TRUE.

transform Logical scalar indicating whether a quarter-root transformation should be ap-

plied before trend fitting.

span Numeric scalar specifying the span of the LOWESS smoother. Ignored if use.min.width=TRUE.

use.min.width Logical scalar indicating whether a minimum width constraint should be applied

to the LOWESS smoother. Useful to avoid overfitting in high-density intervals.

min.width Minimum width of the window to use when use.min.width=TRUE.

min.window.count

Minimum number of observations in each window. Only used if use.min.width=TRUE.

num. threads Integer scalar specifying the number of threads to use.

Value

A list containing statistics. This is a data frame with the columns means, variances, fitted and residuals, each of which is a numeric vector containing the statistic of the same name across all genes.

If block is supplied, each of the column vectors described above contains the average across all blocks. The list will also contain per.block, a list of data frames containing the equivalent statistics for each block.

Author(s)

Aaron Lun

See Also

```
https://libscran.github.io/scran_variances/, for the variance modelling. https://libscran.github.io/scran_blocks/, for details on the blocking. fitVarianceTrend, which fits the mean-variance trend.
```

```
library(Matrix)
x <- abs(rsparsematrix(1000, 100, 0.1) * 10)
out <- modelGeneVariances(x)
str(out)

# Throwing in some blocking.
block <- sample(letters[1:4], ncol(x), replace=TRUE)
out <- modelGeneVariances(x, block=block)
str(out)</pre>
```

normalizeCounts 21

_					
normal	١i	700	'nι	ınt	٠.

Normalize the count matrix

Description

Apply scaling normalization to a count matrix to obtain log-transformed normalized expression values.

Usage

```
normalizeCounts(
   x,
   size.factors,
   log = TRUE,
   pseudo.count = 1,
   log.base = 2,
   preserve.sparsity = FALSE
)
```

Arguments

A matrix-like object where rows correspond to genes or genomic features and columns correspond to cells. Values are typically expected to be counts. Alternatively, an external pointer created by initializeCpp.

size.factors

A numeric vector of length equal to the number of cells in x, containing positive size factors for all cells.

log

Logical scalar indicating whether log-transformation should be performed.

Numeric scalar specifying the positive pseudo-count to add before any log-transformation. Ignored if log=FALSE.

log.base

Numeric scalar specifying the base of the log-transformation. Ignored if log=FALSE.

preserve.sparsity

Logical scalar indicating whether to preserve sparsity for pseudo.count != 1. If TRUE, users should manually add log(pseudo.count, log.base) to the returned matrix to obtain the desired log-transformed expression values.

Value

If x is a matrix-like object, a DelayedArray is returned containing the (log-transformed) normalized expression matrix.

If x is an external pointer produced by initializeCpp, a new external pointer is returned containing the normalized expression matrix.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_norm/, for the rationale behind normalization.

22 rna_quality_control

Examples

```
# Mocking a matrix:
library(Matrix)
x <- round(abs(rsparsematrix(1000, 100, 0.1) * 100))
sf <- centerSizeFactors(colSums(x))
normed <- normalizeCounts(x, size.factors=sf)
normed

# Passing a pointer.
ptr <- beachmat::initializeCpp(x)
optr <- normalizeCounts(ptr, sf)
optr</pre>
```

rna_quality_control

Quality control for RNA count data

Description

Compute per-cell QC metrics from an initialized matrix of RNA counts, and use the metrics to suggest filter thresholds to retain high-quality cells.

Usage

```
computeRnaQcMetrics(x, subsets, num.threads = 1)
suggestRnaQcThresholds(metrics, block = NULL, num.mads = 3)
filterRnaQcMetrics(thresholds, metrics, block = NULL)
```

Arguments

х	A matrix-like object where rows are genes and columns are cells. Values are expected to be counts.
subsets	List of vectors specifying gene subsets of interest, typically for control-like features like mitochondrial genes or spike-in transcripts. Each vector may be logical (whether to keep each row), integer (row indices) or character (row names).
num.threads	Integer scalar specifying the number of threads to use.
metrics	List with the same structure as produced by computeRnaQcMetrics.
block	Factor specifying the block of origin (e.g., batch, sample) for each cell in metrics. Alternatively NULL if all cells are from the same block.
	For filterRnaQcMetrics, a blocking factor should be provided if block was used to construct thresholds.
num.mads	Number of median from the median, to define the threshold for outliers in each metric.
thresholds	List with the same structure as produced by suggestRnaQcThresholds.

rna_quality_control 23

Value

For computeRnaQcMetrics, a list is returned containing:

- sum, a numeric vector containing the total RNA count for each cell.
- detected, an integer vector containing the number of detected genes per cell.
- subsets, a list of numeric vectors containing the proportion of counts in each feature subset.

Each vector is of length equal to the number of cells.

For suggestRnaQcThresholds with block!=NULL, a list is returned containing:

- sum, a numeric vector containing the lower bound on the sum for each blocking level.
- detected, a numeric vector containing the lower bound on the number of detected genes for each blocking level.
- subsets, a list of numeric vectors containing the upper bound on the sum of counts in each feature subset for each blocking level.

Each vector is of length equal to the number of levels in block.

For suggestRnaQcThresholds with block=NULL, a list is returned containing:

- sum, a numeric scalar containing the lower bound on the sum.
- detected, a numeric scalar containing the lower bound on the number of detected genes.
- subsets, a numeric vector containing the upper bound on the sum of counts in each feature subset.

For filterRnaQcMetrics, a logical vector of length ncol(x) is returned indicating which cells are of high quality.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_qc/, for the rationale of QC filtering on RNA counts.

```
# Mocking a matrix:
library(Matrix)
x <- round(abs(rsparsematrix(1000, 100, 0.1) * 100))
# Mocking up a control set.
sub <- list(mito=rbinom(nrow(x), 1, 0.1) > 0)

qc <- computeRnaQcMetrics(x, sub)
str(qc)

filt <- suggestRnaQcThresholds(qc)
str(filt)

keep <- filterRnaQcMetrics(filt, qc)
summary(keep)</pre>
```

24 runAllNeighborSteps

runAllNeighborSteps

Run all neighbor-related steps

Description

Run all steps that require a nearest-neighbor search. This includs runUmap, runTsne and buildSnnGraph with clusterGraph. The idea is to build the index once, perform the neighbor search, and run each task in parallel to save time.

Usage

```
runAllNeighborSteps(
    x,
    runUmap.args = list(),
    runTsne.args = list(),
    buildSnnGraph.args = list(),
    clusterGraph.args = list(),
    BNPARAM = AnnoyParam(),
    collapse.search = FALSE,
    num.threads = 3
)
```

Arguments

x Numeric matrix where rows are dimensions and columns are cells, typically

containing a low-dimensional representation from, e.g., runPca.

Alternatively, an index constructed by buildIndex.

runUmap.args Named list of further arguments to pass to runUmap. This can be set to NULL to

omit the UMAP.

runTsne.args Named list of further arguments to pass to runTsne. This can be set to NULL to

omit the t-SNE.

buildSnnGraph.args

Named list of further arguments to pass to buildSnnGraph.

clusterGraph.args

Named list of further arguments to pass to clusterGraph. This can be set to

NULL to omit the clustering.

BNPARAM A BiocNeighborParam instance specifying the nearest-neighbor search algo-

rithm to use.

collapse.search

Logical scalar indicating whether to collapse the nearest-neighbor search for each step into a single search. Steps that need fewer neighbors will take a subset of the neighbors from the collapsed search. This is faster but may not give the same results as separate searches for some algorithms (e.g., approximate

searches).

num. threads Integer scalar specifying the number of threads to use. At least one thread should

be available for each step.

Value

A named list containing the results of each step. See each individual function for the format of the results.

runPca 25

Author(s)

Aaron Lun

Examples

```
x <- t(as.matrix(iris[,1:4]))
# (Turning down the number of threads so that R CMD check is happy.)
res <- runAllNeighborSteps(x, num.threads=2)
str(res)</pre>
```

runPca

Principal components analysis

Description

Run a PCA on the gene-by-cell log-expression matrix to obtain a low-dimensional representation for downstream analyses.

Usage

```
runPca(
    x,
    number = 25,
    scale = FALSE,
    block = NULL,
    block.weight.policy = c("variable", "equal", "none"),
    variable.block.weight = c(0, 1000),
    components.from.residuals = FALSE,
    extra.work = 7,
    iterations = 1000,
    seed = 5489,
    realized = TRUE,
    num.threads = 1
)
```

Arguments

x A matrix-like object where rows correspond to genes or genomic features and columns correspond to cells. Typically, the matrix is expected to contain log-expression values, and the rows should be filtered to relevant (e.g., highly vari-

able) genes.

number Integer scalar specifying the number of PCs to retain.

scale Logical scalar indicating whether to scale all genes to have the same variance.

block Factor specifying the block of origin (e.g., batch, sample) for each cell in x.

Alternatively NULL if all cells are from the same block.

block.weight.policy

String specifying the policy to use for weighting different blocks when computing the average for each statistic Only used if block is not NULL.

26 runPca

variable.block.weight

Numeric vector of length 2, specifying the parameters for variable block weighting. The first and second values are used as the lower and upper bounds, respectively, for the variable weight calculation. Only used if block is not NULL and block.weight.policy = "variable".

components.from.residuals

Logical scalar indicating whether to compute the PC scores from the residuals in the presence of a blocking factor. By default, the residuals are only used to compute the rotation matrix, and the original expression values of the cells are projected onto this new space. Only used if block is not NULL.

extra.work Integer scalar specifying the extra dimensions for the IRLBA workspace.

iterations Integer scalar specifying the maximum number of restart iterations for IRLBA.

seed Integer scalar specifying the seed for the initial random vector in IRLBA.

realized Logical scalar indicating whether to realize x into an optimal memory layout for

IRLBA. This improves computation time at the cost of increased memory usage.

num. threads Number of threads to use.

Value

List containing:

- components, a matrix of PC scores. Rows are dimensions (i.e., PCs) and columns are cells.
- rotation, the rotation matrix. Rows are genes and columns are dimensions.
- variance.explained, the vector of variances explained by each PC.
- total.variance, the total variance in the dataset.
- center, a numeric vector containing the mean for each gene. If block is provided, this is instead a matrix containing the mean for each gene (column) in each block (row).
- scale, a numeric vector containing the scaling for each gene. Only reported if scale=TRUE.

Author(s)

Aaron Lun

See Also

```
https://libscran.github.io/scran_pca/, for more details on the PCA.
https://libscran.github.io/scran_blocks/, for more details on the block weighting.
```

```
library(Matrix)
x <- abs(rsparsematrix(1000, 100, 0.1) * 10)
y <- normalizeCounts(x, size.factors=centerSizeFactors(colSums(x)))
# A simple PCA:
out <- runPca(y)
str(out)
# Blocking on uninteresting factors:
block <- sample(LETTERS[1:3], ncol(y), replace=TRUE)
bout <- runPca(y, block=block)
str(bout)</pre>
```

runTsne 27

runTsne

t-stochastic neighbor embedding

Description

Compute t-SNE coordinates to visualize similarities between cells.

Usage

```
runTsne(
  х,
 perplexity = 30,
 num.neighbors = tsnePerplexityToNeighbors(perplexity),
 max.depth = 20,
  leaf.approximation = FALSE,
 max.iterations = 500,
  seed = 42,
  num.threads = 1,
 BNPARAM = AnnoyParam()
)
tsnePerplexityToNeighbors(perplexity)
```

Arguments

Numeric matrix where rows are dimensions and columns are cells, typically containing a low-dimensional representation from, e.g., runPca.

Alternatively, a named list of nearest-neighbor search results. This should contain index, an integer matrix where rows are neighbors and columns are cells. Each column contains 1-based indices for the nearest neighbors of the corresponding cell, ordered by increasing distance. The number of neighbors should be the same as num.neighbors, otherwise a warning is raised.

Alternatively, an index constructed by buildIndex.

perplexity

Numeric scalar specifying the perplexity to use in the t-SNE algorithm.

num.neighbors

Integer scalar specifying the number of neighbors, typically derived from perplexity.

max.depth

Integer scalar specifying the maximum depth of the Barnes-Hut quadtree. Smaller

values (7-10) improve speed at the cost of accuracy.

leaf.approximation

Logical scalar indicating whether to use the "leaf approximation" approach, which sacrifices some accuracy for greater speed. Only effective when max.depth is small enough for multiple cells to be assigned to the same leaf node of the

quadtree.

max.iterations Integer scalar specifying the maximum number of iterations to perform.

seed

Integer scalar specifying the seed to use for generating the initial coordinates.

num.threads

Integer scalar specifying the number of threads to use.

BNPARAM

A BiocNeighborParam object specifying the algorithm to use. Only used if x is

not a list of existing nearest-neighbor search results.

28 runUmap

Value

For runTsne, a numeric matrix where rows are cells and columns are the two dimensions of the embedding.

For tsnePerplexityToNeighbors, an integer scalar specifying the number of neighbors to use for a given perplexity.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/qdtsne/, for an explanation of the approximations.

Examples

```
x <- t(as.matrix(iris[,1:4]))
embedding <- runTsne(x)
plot(embedding[,1], embedding[,2], col=iris[,5])</pre>
```

runUmap

Uniform manifold approxation and projection

Description

Compute UMAP coordinates to visualize similarities between cells.

Usage

```
runUmap(
    x,
    num.dim = 2,
    num.neighbors = 15,
    num.epochs = -1,
    min.dist = 0.1,
    seed = 1234567890,
    num.threads = 1,
    parallel.optimization = FALSE,
    BNPARAM = AnnoyParam()
)
```

Arguments

Х

Numeric matrix where rows are dimensions and columns are cells, typically containing a low-dimensional representation from, e.g., runPca.

Alternatively, a named list of nearest-neighbor search results. This should contain index, an integer matrix where rows are neighbors and columns are cells. Each column contains 1-based indices for the nearest neighbors of the corresponding cell, ordered by increasing distance. The number of neighbors should be the same as num.neighbors, otherwise a warning is raised.

Alternatively, an index constructed by buildIndex.

sanitizeSizeFactors 29

num.dim Integer scalar specifying the number of dimensions of the output embedding. num.neighbors Integer scalar specifying the number of neighbors to use in the UMAP algorithm. Integer scalar specifying the number of epochs to perform. If set to -1, an apnum.epochs propriate number of epochs is chosen based on ncol(x). min.dist Numeric scalar specifying the minimum distance between points. Integer scalar specifying the seed to use. seed num.threads Integer scalar specifying the number of threads to use. parallel.optimization Logical scalar specifying whether to parallelize the optimization step. **BNPARAM** A BiocNeighborParam object specifying the algorithm to use. Only used if x is not a list of existing nearest-neighbor search results.

Value

A numeric matrix where rows are cells and columns are the two dimensions of the embedding.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/umappp/, for details on the underlying implementation.

Examples

```
x <- t(as.matrix(iris[,1:4]))
embedding <- runUmap(x)
plot(embedding[,1], embedding[,2], col=iris[,5])</pre>
```

sanitizeSizeFactors Sanitize size factors

Description

Replace invalid size factors, i.e., zero, negative, infinite or NaN values.

Usage

```
sanitizeSizeFactors(
    size.factors,
    replace.zero = TRUE,
    replace.negative = TRUE,
    replace.infinite = TRUE,
    replace.nan = TRUE
)
```

30 scaleByNeighbors

Arguments

size.factors Numeric vector of size factors across cells.

replace.zero Logical scalar indicating whether to replace size factors of zero with the lowest positive factor. If FALSE, zeros are retained.

replace.negative Logical scalar indicating whether to replace negative size factors with the lowest positive factor. If FALSE, negative values are retained.

replace.infinite Logical scalar indicating whether to replace infinite size factors with the largest positive factor. If FALSE, infinite values are retained.

positive factor. If FALSE, infinite values are retained.

replace.nan Logical scalar indicating whether to replace NaN size factors with unity. If FALSE, NaN values are retained.

Value

Numeric vector of length equal to size. factors, containing the sanitized size factors.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_norm/, for more details on the sanitization.

Examples

```
sf <- 2^rnorm(100)
sf[1] <- 0
sf[2] <- -1
sf[3] <- Inf
sf[4] <- NaN
sanitizeSizeFactors(sf)</pre>
```

scaleByNeighbors

Scale and combine multiple embeddings

Description

Scale multiple embeddings (usually derived from different modalities across the same set of cells) so that their within-population variances are comparable, and then combine them into a single embedding matrix for combined downstream analysis.

Usage

```
scaleByNeighbors(
    x,
    num.neighbors = 20,
    num.threads = 1,
    weights = NULL,
    BNPARAM = AnnoyParam()
)
```

scoreGeneSet 31

Arguments

x List of numeric matrices of principal components or other embeddings, one for

each modality. For each entry, rows are dimensions and columns are cells. All entries should have the same number of columns but may have different numbers

of rows.

num.neighbors Integer scalar specifying the number of neighbors to use to define the scaling

factor.

num. threads Integer scalar specifying the number of threads to use.

weights Numeric vector of length equal to that of x, specifying the weights to apply to

each modality. Each value represents a multiplier of the within-population variance of its modality, i.e., larger values increase the contribution of that modality in the combined output matrix. NULL is equivalent to an all-1 vector, i.e., all

modalities are scaled to have the same within-population variance.

BNPARAM A BiocNeighborParam object specifying how to perform the neighbor search.

Value

List containing scaling, a vector of scaling factors to be aplied to each embedding; and combined, a numeric matrix formed by scaling each entry of x and then rbinding them together.

Author(s)

Aaron Lun

Examples

```
pcs <- list(
    gene = matrix(rnorm(10000), ncol=200),
    protein = matrix(rnorm(1000, sd=3), ncol=200),
    guide = matrix(rnorm(2000, sd=5), ncol=200)
)

out <- scaleByNeighbors(pcs)
out$scaling
dim(out$combined)</pre>
```

scoreGeneSet

Score gene set activity for each cell

Description

Compute per-cell scores for a gene set, defined as the column sums of a rank-1 approximation to the submatrix for the feature set. This uses the same approach implemented in the **GSDecon** package from Jason Hackney.

32 scoreGeneSet

Usage

```
scoreGeneSet(
    x,
    set,
    rank = 1,
    scale = FALSE,
    block = NULL,
    block.weight.policy = c("variable", "equal", "none"),
    variable.block.weight = c(0, 1000),
    extra.work = 7,
    iterations = 1000,
    seed = 5489,
    realized = TRUE,
    num.threads = 1
)
```

Arguments

x A matrix-like object where rows correspond to genes or genomic features and

columns correspond to cells. Typically, the matrix is expected to contain log-expression values, and the rows should be filtered to relevant (e.g., highly vari-

able) genes.

set Integer, logical or character vector specifying the rows that belong to the gene

set.

rank Integer scalar specifying the rank of the approximation.

scale Logical scalar indicating whether to scale all genes to have the same variance.

block Factor specifying the block of origin (e.g., batch, sample) for each cell in x.

Alternatively NULL if all cells are from the same block.

block.weight.policy

String specifying the policy to use for weighting different blocks when comput-

ing the average for each statistic Only used if block is not NULL.

variable.block.weight

Numeric vector of length 2, specifying the parameters for variable block weighting. The first and second values are used as the lower and upper bounds, respectively, for the variable weight calculation. Only used if block is not NULL and

block.weight.policy = "variable".

extra.work Integer scalar specifying the extra dimensions for the IRLBA workspace.

iterations Integer scalar specifying the maximum nunber of restart iterations for IRLBA.

seed Integer scalar specifying the seed for the initial random vector in IRLBA.

realized Logical scalar indicating whether to realize x into an optimal memory layout for

IRLBA. This improves computation time at the cost of increased memory usage.

num. threads Number of threads to use.

Value

List containing scores, a numeric vector of per-cell scores for each column in x; and weights, a numeric vector of per-feature weights for each feature in set.

scoreMarkers 33

Author(s)

Aaron Lun

See Also

https://libscran.github.io/gsdecon/, for more details on the underlying algorithm.

Examples

```
library(Matrix)
x <- round(abs(rsparsematrix(1000, 100, 0.1) * 100))
normed <- normalizeCounts(x, size.factors=centerSizeFactors(colSums(x)))
scoreGeneSet(normed, set=c(1,3,5,10,20,100))</pre>
```

scoreMarkers

Score marker genes

Description

Score marker genes for each group using a variety of effect sizes from pairwise comparisons between groups. This includes Cohen's d, the area under the curve (AUC), the difference in the means (delta-mean) and the difference in the proportion of detected cells (delta-detected).

Usage

```
scoreMarkers(
    x,
    groups,
    block = NULL,
    block.weight.policy = c("variable", "equal", "none"),
    variable.block.weight = c(0, 1000),
    compute.auc = TRUE,
    threshold = 0,
    all.pairwise = FALSE,
    num.threads = 1
)
```

Arguments

A matrix-like object where rows correspond to genes or genomic features and columns correspond to cells. It is typically expected to contain log-expression

values, e.g., from normalizeCounts.

groups A vector specifying the group assignment for each cell in x.

block Factor specifying the block of origin (e.g., batch, sample) for each cell in x.

Alternatively NULL if all cells are from the same block.

block.weight.policy

String specifying the policy to use for weighting different blocks when computing the average for each statistic Only used if block is not NULL.

34 scoreMarkers

variable.block.weight

Numeric vector of length 2, specifying the parameters for variable block weighting. The first and second values are used as the lower and upper bounds, respectively, for the variable weight calculation. Only used if block is not NULL and block.weight.policy = "variable".

compute.auc Logical scalar indicating whether to compute the AUC. Setting this to FALSE

can improve speed and memory efficiency.

threshold Non-negative numeric scalar specifying the minimum threshold on the differ-

ences in means (i.e., the log-fold change, if x contains log-expression values).

This is incorporated into the effect sizes for Cohen's d and the AUC.

all.pairwise Logical scalar indicating whether to report the full effects for every pairwise

comparison between groups.

num. threads Integer scalar specifying the number of threads to use.

Value

If all.pairwise=FALSE, a list is returned containing:

- mean, a numeric matrix containing the mean expression for each group. Each row is a gene and each column is a group.
- detected, a numeric matrix containing the proportion of detected cells in each group. Each row is a gene and each column is a group.
- cohens.d, a list of data frames where each data frame corresponds to a group. Each row of each data frame represents a gene, while each column contains a summary of Cohen's d from pairwise comparisons to all other groups. This includes the min, mean, median, max and min.rank.
- auc, a list like cohens.d but containing the summaries of the AUCs from each pairwise comparison. Omitted if compute.auc=FALSE.
- delta.mean, a list like cohens.d but containing the summaries of the delta-mean from each pairwise comparison.
- delta.detected, a list like cohens.d but containing the summaries of the delta-detected from each pairwise comparison.

If all.pairwise=TRUE, a list is returned containing:

- mean, a numeric matrix containing the mean expression for each group. Each row is a gene and each column is a group.
- detected, a numeric matrix containing the proportion of detected cells in each group. Each row is a gene and each column is a group.
- cohens.d, a 3-dimensional numeric array containing the Cohen's from each pairwise comparison between groups. The extents of the first two dimensions are equal to the number of groups, while the extent of the final dimension is equal to the number of genes. The entry [i, j, k] represents Cohen's d from the comparison of group j over group i for gene k.
- auc, an array like cohens.d but containing the AUCs from each pairwise comparison. Omitted if compute.auc=FALSE.
- delta.mean, an array like cohens.d but containing the delta-mean from each pairwise comparison.
- delta.detected, an array like cohens.d but containing the delta-detected from each pairwise comparison.

See Also

https://libscran.github.io/scran_markers/, in particular the score_markers_summary function (for all.pairwise=FALSE), the score_markers_pairwise function (for all.pairwise=TRUE), and their blocked equivalents score_markers_summary_blocked and score_markers_pairwise_blocked (when block is not NULL).

summarizeEffects, to summarize the pairwise effects returned when all.pairwise=TRUE.

Examples

```
# Mocking a matrix:
library(Matrix)
x <- round(abs(rsparsematrix(1000, 100, 0.1) * 100))
normed <- normalizeCounts(x, size.factors=centerSizeFactors(colSums(x)))
g <- sample(letters[1:4], ncol(x), replace=TRUE)
scores <- scoreMarkers(normed, g)
names(scores)
head(scores$mean)
head(scores$cohens.d[["a"]])</pre>
```

subsampleByNeighbors Subsample cells based on their neighbors

Description

Subsample a dataset by selecting cells to represent all of their nearest neighbors.

Usage

```
subsampleByNeighbors(
    x,
    num.neighbors = 20,
    min.remaining = 10,
    num.threads = 1,
    BNPARAM = AnnoyParam()
)
```

Arguments

Х

A numeric matrix where rows are dimensions and columns are cells, typically containing a low-dimensional representation from, e.g., runPca.

Alternatively, an index constructed by buildIndex.

Alternatively, a list containing existing nearest-neighbor search results. This should contain:

- index, an integer matrix where rows are neighbors and columns are cells. Each column contains 1-based indices for the nearest neighbors of the corresponding cell, ordered by increasing distance.
- distance, a numeric matrix of the same dimensions as index, containing the distances to each of the nearest neighbors.

36 summarizeEffects

The number of neighbors should be equal to num. neighbors, otherwise a warning is raised.

num.neighbors

Integer scalar specifying the number of neighbors to use. Larger values result in greater downsampling. Only used if x does not contain existing nearest-neighbor results.

min.remaining

Integer scalar specifying the minimum number of remaining (i.e., unselected) neighbors that a cell must have in order to be considered for selection. This should be less than or equal to num.neighbors.

num.threads

Integer scalar specifying the number of threads to use for the nearest-neighbor search. Only used if x does not contain existing nearest-neighbor results.

BNPARAM

A BiocNeighborParam object specifying the algorithm to use. Only used if x

Value

Integer vector with the indices of the selected cells in the subsample.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/nenesub/, for more details on the underlying algorithm.

does not contain existing nearest-neighbor results.

Examples

```
x <- matrix(rnorm(10000), nrow=2)
keep <- subsampleByNeighbors(x, 10)
plot(x[1,], x[2,])
points(x[1,keep], x[2,keep], col="red")
legend('topright', col=c('black', 'red'), legend=c('all', 'subsample'), pch=1)</pre>
```

summarizeEffects

Summarize pairwise effect sizes for each group

Description

For each group, summarize the effect sizes for all pairwise comparisons to other groups. This yields a set of summary statistics that can be used to rank marker genes for each group.

Usage

```
summarizeEffects(effects, num.threads = 1)
```

summarizeEffects 37

Arguments

effects A 3-dimensional numeric containing the effect sizes from each pairwise com-

parison between groups. The extents of the first two dimensions are equal to the number of groups, while the extent of the final dimension is equal to the number of genes. The entry [i, j, k] represents Cohen's d from the comparison of group j over group i for gene k. See also the output of scoreMarkers with

all.pairwise=TRUE.

num. threads Integer scalar specifying the number of threads to use.

Value

List of data frames containing summary statistics for the effect sizes. Each data frame corresponds to a group, each row corresponds to a gene, and each column contains a single summary.

Author(s)

Aaron Lun

See Also

https://libscran.github.io/scran_markers/, for more details on the statistics. scoreMarkers, to compute the pairwise effects in the first place.

```
# Mocking a matrix:
library(Matrix)
x <- round(abs(rsparsematrix(1000, 100, 0.1) * 100))
normed <- normalizeCounts(x, size.factors=centerSizeFactors(colSums(x)))
g <- sample(letters[1:4], ncol(x), replace=TRUE)
effects <- scoreMarkers(normed, g, all.pairwise=TRUE)
summarized <- summarizeEffects(effects$cohens.d)
str(summarized)</pre>
```

Index

runUmap, 24, 28

```
adt_quality_control, 2
                                                 sanitizeSizeFactors, 29
aggregateAcrossCells, 4
                                                 scaleByNeighbors, 30
                                                 scoreGeneSet, 31
BiocNeighborParam, 6, 15, 24, 27, 29, 31, 36
                                                 scoreMarkers, 33, 37
buildIndex, 6, 24, 27, 28, 35
                                                 subsampleByNeighbors, 35
buildSnnGraph, 5, 10, 24
                                                 suggestAdtQcThresholds
                                                          (adt_quality_control), 2
centerSizeFactors, 7
                                                 suggestCrisprQcThresholds
chooseHighlyVariableGenes, 8
                                                          (crispr_quality_control), 16
choosePseudoCount, 9
                                                 {\it suggestRnaQcThresholds}
clusterGraph, 10, 24
                                                          (rna_quality_control), 22
clusterKmeans, 11
                                                 summarizeEffects, 35, 36
combineFactors, 4, 13
computeAdtQcMetrics
                                                 tsnePerplexityToNeighbors (runTsne), 27
        (adt_quality_control), 2
computeClrm1Factors, 14
computeCrisprQcMetrics
        (crispr_quality_control), 16
{\tt computeRnaQcMetrics}
        (rna_quality_control), 22
correctMnn, 14
crispr_quality_control, 16
DelayedArray, 21
filterAdtQcMetrics
        (adt_quality_control), 2
filterCrisprQcMetrics
        (crispr_quality_control), 16
filterRnaQcMetrics
        (rna_quality_control), 22
fitVarianceTrend, 18, 20
igraph, 10
initializeCpp, 21
modelGeneVariances, 8, 19
normalizeCounts, 9, 19, 21, 33
rna_quality_control, 22
runAllNeighborSteps, 24
runPca, 5, 15, 24, 25, 27, 28, 35
runTsne, 24, 27
```