Package 'SCArray.sat'

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Description Extends the Seurat classes and functions to support Genomic Data Structure (GDS) files as a DelayedArray backend for data representation. It relies on the implementation of GDS-based DelayedMatrix in the SCArray package to represent single cell RNA-seq data. The common optimized algorithms leveraging GDS-based and single cell-specific DelayedMatrix (SC_GDSMatrix) are implemented in the SCArray package. SCArray.sat introduces a new SCArrayAssay class (derived from the Seurat Assay), which wraps raw counts, normalized expressions and scaled data matrix based on GDS-specific DelayedMatrix. It is designed to integrate seamlessly with the Seurat package to provide common data analysis in the SeuratObject-based workflow. Compared with Seurat, SCArray.sat significantly reduces the memory usage without downsampling and can be applied to very large datasets.

License GPL-3

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`SCArray.sat-package` *Large-scale single-cell RNA-seq data analysis using GDS files and Seurat*

Description

The package extends the Seurat classes and functions to support GDS files as a DelayedArray backend for data representation. It introduces a new SCArrayAssay class (derived from the Seurat Assay), which wraps raw counts, normalized expressions and scaled data matrix based on DelayedMatrix. It is designed to integrate seamlessly with the SeuratObject and Seurat packages to provide common data analysis, with the optimized algorithms for GDS data files.

Details

Package:	SCArray.sat
Type:	Package
License:	GPL version 3

Author(s)

Xiuwen Zheng

`CreateAssayObject2` *Create an Assay object*

Description

Create an Assay5 object with DelayedMatrix layers from counts or prenormalized data.

Usage

```
CreateAssayObject2(counts, data, min.cells=0, min.features=0,
    key=NULL, check.matrix=FALSE, ...)
```

Arguments

`counts`	Unnormalized raw counts (matrix, dgCMatrix or DelayedMatrix)
`data`	Prenormalized data (matrix, dgCMatrix or DelayedMatrix)
`min.cells`	if > 0, a lower cutoff for filtering cells
`min.features`	if > 0, a lower cutoff for filtering features
`check.matrix`	Check counts matrix for NA, NaN, Inf, and non-integer values
`key`	Key name for the assay
`...`	Arguments passed to `as.sparse` when `counts` or `data` is matrix or dgCMatrix

Details

Similar to `SeuratObject::CreateAssayObject()`, except allowing DelayedMatrix `counts` or `data`, and returning an `Assay5` object. `counts` and `data` should not be provided at the same time.

Value

Return an instance of `Assay5`.

Author(s)

Xiuwen Zheng

See Also

`CreateAssayObject`, `scGetFiles`

Examples

```
fn <- system.file("extdata", "example.gds", package="SCArray")

x <- scArray(fn, "counts")
colnames(x) <- paste0("c", 1:ncol(x))
rownames(x) <- paste0("g", 1:nrow(x))
x

a <- CreateAssayObject2(x)
a

scGetFiles(x)
scGetFiles(a)

remove(x, a)
```

`NormalizeData` *Normalize Count Data*

Description

Normalizes the count data in the Seurat assay.

Usage

```
# NormalizeData(object, ...)
## S3 method for class 'SC_GDSMatrix'
NormalizeData(object,
    normalization.method="LogNormalize", scale.factor=1e4, margin=1,
    verbose=TRUE, ...)
```

Arguments

`object`	input R object (e.g., a SC_GDSMatrix object)
`normalization.method`	"LogNormalize", "CLR" or "RC"; see `NormalizeData.Seurat` for more details
`scale.factor`	the scale factor for cell-level normalization
`margin`	only applicable when `normalization.method="CLR"`, normalize across features (`margin=1`) or cells (`margin=2`)
`verbose`	if `TRUE`, show information
`...`	additional arguments passed to specific methods

Details

`NormalizeData()` does not store the normalized data in a GDS file, since the calculation is "delayed" until it is needed.

Value

Returns a SC_GDSMatrix matrix.

Author(s)

Xiuwen Zheng

See Also

NormalizeData

Examples

```
fn <- system.file("extdata", "example.gds", package="SCArray")

d <- scNewSeuratGDS(fn)
d
d <- NormalizeData(d)

remove(d)
```

RunPCA *Run PCA*

Description

Performs PCA on a Seurat SCArrayAssay, Assay5 (with DelayedMatrix layers) or a DelayedMatrix object.

Usage

```
# RunPCA(object, ...)
## S3 method for class 'SCArrayAssay'
RunPCA(object, assay=NULL, features=NULL, npcs=50,
    rev.pca=FALSE, weight.by.var=TRUE, verbose=TRUE, ndims.print=1:5,
    nfeatures.print=30, reduction.key="PC_", seed.use=42, ...)
## S3 method for class 'Assay5'
RunPCA(object, assay=NULL, features=NULL,
    layer="scale.data", npcs=50, rev.pca=FALSE, weight.by.var=TRUE,
    verbose=TRUE, ndims.print=1:5, nfeatures.print=30, reduction.key="PC_",
    seed.use=42, ...)
## S3 method for class 'SC_GDSMatrix'
RunPCA(object, assay=NULL, npcs=50, rev.pca=FALSE,
    weight.by.var=TRUE, verbose=TRUE, ndims.print=1:5,
    nfeatures.print=30, reduction.key="PC_", seed.use=42, approx=TRUE,
    BPPARAM, ...)
```

Arguments

object	input R object (e.g., a SCArrayAssay or Assay5 object)
assay	NULL for using the active assay, or an assay name
features	if NULL, PCA will be run on the scaled data; otherwise, features to compute PCA on
layer	the layer in the Assay5 object used for PCA
npcs	# of top PCs to be calculated
rev.pca	By default (FALSE), perform PCA on the cell x gene matrix; otherwise, compute it on gene x cell matrix
weight.by.var	if TRUE, weight the cell embeddings (when rev.pca=FALSE) or the the gene loadings (when rev.pca=TRUE) by the variance of each PC
verbose	if TRUE, show information
ndims.print	which PCs to print genes for
nfeatures.print	# of genes to print for each PC
reduction.key	dimensional reduction key
seed.use	a random seed; or NULL for not setting a seed internally
approx	if TRUE, use IrlbaSVD; otherwise use ExactSVD
BPPARAM	NULL for non-parallel execution, or a BiocParallelParam object for parallelization; if it is missing, getAutoBPPARAM() will be used
...	additional arguments passed to specific methods

Details

RunPCA() computes the covariance matrix of genes (if # of genes <= # of cells) or the cell covariance matrix for the PCA calculation, which can reduce the times of accessing on-disk data.

RunPCA.Assay5() is used when the layer is a DelayedMatrix (e.g., SC_GDSMatrix); otherwise, it calls the method for StdAssay in the Seurat package.

Value

Return a data frame for reduction data (via CreateDimReducObject).

Author(s)

Xiuwen Zheng

See Also

RunPCA, CreateDimReducObject, BiocParallelParam, getAutoBPPARAM

Examples

```
fn <- system.file("extdata", "example.gds", package="SCArray")

d <- scNewSeuratGDS(fn)

d <- NormalizeData(d)
d <- FindVariableFeatures(d, nfeatures=250)
d <- ScaleData(d)

d <- RunPCA(d, ndims.print=1:2)
DimPlot(d, reduction="pca")

remove(d)
```

`ScaleData` *Scale and Center the Data*

Description

Scales and centers features or residuals in the dataset.

Usage

```
# ScaleData(object, ...)
## S3 method for class 'SC_GDSMatrix'
ScaleData(object, features=NULL, vars.to.regress=NULL,
    latent.data=NULL, split.by=NULL, model.use='linear', use.umi=FALSE,
    do.scale=TRUE, do.center=TRUE, scale.max=10, block.size=1000,
    min.cells.to.block=3000, verbose=TRUE, use_gds=TRUE, rm_tmpfile=TRUE, ...)
```

Arguments

`object`	input R object (e.g., a SC_GDSMatrix object)
`features`	if `NULL`, to use the variable features (found via `FindVariableFeatures()`); or features names to scale/center
`vars.to.regress`	NULL or variable names to regress out
`latent.data`	NULL or a `data.frame` to regress out the covariates
`split.by`	variable name in the metadata, or a vector or factor defining grouping of cells
`model.use`	regression model: `"linear"` (default), `"poisson"` or `"negbinom"`
`use.umi`	only applicable when the covariates are given in `vars.to.regress` for regression; default is `FALSE` for linear regression, `TRUE` for negbinom and poisson models
`do.scale`	if `TRUE`, scale the data
`do.center`	if `TRUE`, center the data

scale.max	max value in the resulting scaled data; see ScaleData.Seurat for more details
block.size	not used
min.cells.to.block	not used
verbose	if TRUE, show information
use_gds	if TRUE, use SC_GDSMatrix for the scaled data; if FALSE, to use a dense in-memory scaled matrix; or a GDS file name for storing the resulting data matrix; see details
rm_tmpfile	if TRUE, remove any temporary GDS file after the calculation; the temporary file will be created when calculating the residuals
...	additional arguments passed to specific methods

Details

ScaleData() stores the scaled data in a GDS file when use_gds=TRUE or an output GDS file name is given via use_gds. When vars.to.regress and split.by are both NULL, an output GDS file is not needed, since the resulting DelayedMatrix can be represented as common operations on the count matrix. If use_gds=TRUE, an output file name "_scale_data.gds" will be used if it does not exists, or "_scale_data2.gds" (if not exists), "_scale_data3.gds" and so on. If use_gds is an output file name, the resulting data matrix will be saved to a GDS file. When vars.to.regress are given, a temporary GDS file (e.g., "_temp_scale_data.gds", use_gds with a prefix "_temp") will be created to store the residuals before scaling. This temporary file will be deleted after the calculation when rm_tmpfile=TRUE.

Value

Returns a SC_GDSMatrix matrix if use_gds=TRUE or use_gds is an output file name, otherwise returns an in-memory matrix.

Author(s)

Xiuwen Zheng

See Also

ScaleData

Examples

```
fn <- system.file("extdata", "example.gds", package="SCArray")

d <- scNewSeuratGDS(fn)

d <- NormalizeData(d)
d <- FindVariableFeatures(d, nfeatures=50)
d <- ScaleData(d)

GetAssayData(d, layer="scale.data")  # DelayedMatrix
```

```
# scale with split.by
ss <- rep(c(TRUE, FALSE), length.out=ncol(d))
d <- ScaleData(d, split.by=ss)

fn <- scGetFiles(d)
fn[2L]    # the file name storing scaled data

remove(d)
unlink(fn[grepl("^_scale", fn)], force=TRUE)
```

SCArrayAssay-class *GDS-specific Assay Class*

Description

The SCArrayAssay class extends the Assay class of Seurat with the new slots `counts2`, `data2` and `scale.data2` replacing `counts`, `data` and `scale.data`.

Slots

`counts2` Unnormalized raw counts (dgCMatrix or SC_GDSMatrix), replacing `Assay@counts`

`data2` Normalized expression data (dgCMatrix or SC_GDSMatrix), replacing `Assay@data`

`scale.data2` Scaled expression data (NULL, matrix or SC_GDSMatrix), replacing `Assay@scale.data`

Author(s)

Xiuwen Zheng

See Also

`Assay`, `Seurat_g-class`, `GetAssayData`, `SetAssayData`

SCArrayAssay-methods *SCArrayAssay S3 methods*

Description

Gets and sets data in the Seurat Assay object.

Usage

```
## S3 method for class 'SCArrayAssay'
GetAssayData(object,
    slot=c("data", "scale.data", "counts"), ...)
## S3 method for class 'SCArrayAssay'
SetAssayData(object, layer, new.data,
    slot=c('data', 'scale.data', 'counts'), ...)

## S3 method for class 'SCArrayAssay'
subset(x, cells=NULL, features=NULL, ...)
```

Arguments

object, x	a SCArrayAssay object (inherited from SeuratObject::Assay)
layer	layer
new.data	a new data matrix (dgCMatrix or SC_GDSMatrix)
slot	data matrix in the Assay object, "data" is used by default
cells	names or indices for selected cells
features	names or indices for selected features
...	further arguments to be passed to or from other methods

Value

Return a data matrix or an instance of SCArrayAssay.

Author(s)

Xiuwen Zheng

See Also

Assay

---

scGetFiles *File names for on-disk backend*

Description

Get a list of file names for DelayedArray with an on-disk backend.

Usage

```
scGetFiles(object, ...)
## S4 method for signature 'Assay'
scGetFiles(object, ...)
## S4 method for signature 'Assay5'
scGetFiles(object, ...)
## S4 method for signature 'SCArrayAssay'
scGetFiles(object, ...)
## S4 method for signature 'Seurat'
scGetFiles(object, ...)
```

Arguments

object	input R object (e.g., a Seurat object)
...	additional arguments passed to specific methods

Value

Return a character vector storing file names.

Author(s)

Xiuwen Zheng

See Also

scGetFiles

Examples

```
fn <- system.file("extdata", "example.gds", package="SCArray")

a <- scNewAssayGDS(fn)
d <- Seurat::CreateSeuratObject(a)

scGetFiles(a)
scGetFiles(d)

remove(a, d)
```

---

scMemory *Load Data to Memory*

Description

Loads the internal data to memory for any on-disk object.

Usage

```
scMemory(x, ...)
## S4 method for signature 'SCArrayAssay'
scMemory(x, slot=NULL, ...)
## S4 method for signature 'Seurat'
scMemory(x, assay=NULL, slot=NULL, ...)
```

Arguments

x	input R object (e.g., a Seurat object)
assay	NULL for using the active assay, or a list of assay names
slot	NULL for all "counts", "data" and "scale.data"; or a character vector including "counts", "data" or "scale.data"; see details
...	additional arguments passed to specific methods

Details

If slot=NULL, return a Assay object instead of SCArrayAssay object, so it can downgrade a SCArrayAssay object to a Assay object.

Value

Return an object (it maybe a different type from class(x)).

Author(s)

Xiuwen Zheng

See Also

scMemory

Examples

```
fn <- system.file("extdata", "example.gds", package="SCArray")

d1 <- scNewSeuratGDS(fn)
is(GetAssay(d1))

d2 <- scMemory(d1)
is(GetAssay(d2))

remove(d1, d2)
```

scNewAssayGDS *Create Assay Object*

Description

Creates a new Seurat Assay object (SCArrayAssay) from a GDS file.

Usage

```
scNewAssayGDS(gdsfile, name="counts", key="rna_", row_data=TRUE, check=TRUE,
    verbose=TRUE)
```

Arguments

gdsfile	a file name for the GDS file, or a SCArrayFileClass object
name	characters for the name of data matrix in the GDS file; if NA_character_, to use the first assay
key	characters for the Assay key
row_data	if TRUE, add rowData() to the feature-level meta data of the Seurat Assay
check	if TRUE, check the feature names
verbose	if TRUE, show information

Value

Return an instance of SCArrayAssay.

Author(s)

Xiuwen Zheng

See Also

SCArrayAssay, SCArrayFileClass, scExperiment, scNewSeuratGDS

Examples

```
# raw count data in a GDS file
fn <- system.file("extdata", "example.gds", package="SCArray")

a <- scNewAssayGDS(fn)
a
class(a)

d <- Seurat::CreateSeuratObject(a)
d

rm(a, d)
```

scNewSeuratGDS *Create Seurat Object*

Description

Creates a new Seurat object from a GDS file.

Usage

```
scNewSeuratGDS(gdsfile, assay.name=NULL, key=c(counts="rna_"), row_data=TRUE,
    col_data=TRUE, check=TRUE, verbose=TRUE)
```

Arguments

gdsfile	a file name for the GDS file, or a SCArrayFileClass object
assay.name	characters for the name of data matrix in the GDS file; if NULL, to use all of the assays
key	a character vector for an assay key map, where its names are the GDS node names
row_data	if TRUE, add rowData() to the feature-level meta data of the first Seurat Assay
col_data	if TRUE, add colData() to the cell-level meta data of the Seurat object
check	if TRUE, check the feature names
verbose	if TRUE, show information

Details

"counts" must be in the input GDS file and it is used as the raw count data in the active Seurat assay. If "logcounts" exists, it is used as normalized data associated with "counts". If there are other data matrices in the GDS file, they will be added to the assay list.

Value

Return an instance of Seurat.

Author(s)

Xiuwen Zheng

See Also

SCArrayAssay, SCArrayFileClass, scExperiment, scNewAssayGDS

Examples

```
# raw count data in a GDS file
fn <- system.file("extdata", "example.gds", package="SCArray")

d <- scNewSeuratGDS(fn)
d
class(d)

rm(d)
```

`Seurat_g-class` *GDS-based Seurat Class*

Description

The Seurat_g class inherits directly from "Seurat".

Slots

```
setClass("Seurat_g", contains="Seurat")
```

Author(s)

Xiuwen Zheng

See Also

SCArrayAssay-class

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