Package 'MultiAssayExperiment'

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Title Software for the integration of multi-omics experiments in Bioconductor

Version 1.28.0

Description MultiAssayExperiment harmonizes data management of multiple experimental assays performed on an overlapping set of specimens. It provides a familiar Bioconductor user experience by extending concepts from SummarizedExperiment, supporting an open-ended mix of standard data classes for individual assays, and allowing subsetting by genomic ranges or rownames. Facilities are provided for reshaping data into wide and long formats for adaptability to graphing and downstream analysis.

Depends R (>= 4.2.0), SummarizedExperiment (>= 1.3.81)

Imports methods, GenomicRanges (>= 1.25.93), BiocBaseUtils, BiocGenerics, DelayedArray, S4Vectors (>= 0.23.19), IRanges, Biobase, stats, tidyr, utils

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VignetteBuilder knitr, R.rsp

URL http://waldronlab.io/MultiAssayExperiment/

Video https://youtu.be/w6HWAHaDpyk, https://youtu.be/Vh0hVVUKKFM

BugReports https://github.com/waldronlab/MultiAssayExperiment/issues

Collate 'ExperimentList-class.R' 'MultiAssayExperiment-class.R'

'MatchedAssayExperiment-class.R' 'subsetBy-methods.R'

'MultiAssayExperiment-subset.R'

'MultiAssayExperiment-methods.R'

'MultiAssayExperiment-helpers.R' 'MultiAssayExperiment-pkg.R'

'MultiAssayExperimentToMAF.R' 'assay-methods.R' 'data.R'

'hasAssay.R' 'listToMap.R' 'mapToList.R' 'prepMultiAssay.R'

'reexports.R' 'saveHDF5MultiAssayExperiment.R' 'upsetSamples.R' 'utilities.R'

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```

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MultiAssayExperiment-package

MultiAssayExperiment: Build an integrative multi-assay container

Description

MultiAssayExperiment allows the manipulation of related multiassay datasets with partially overlapping samples, associated metadata at the level of an entire study, and at the level of the "biological unit". The biological unit may be a patient, plant, yeast strain, etc.

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Details

The package hierarchy of information:

- study
- · experiments
- samples

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See Also

Useful links:

- http://waldronlab.io/MultiAssayExperiment/
- Report bugs at https://github.com/waldronlab/MultiAssayExperiment/issues

ExperimentList

Represent multiple experiments as a List-derivative ExperimentList

Description

The ExperimentList class can contain several different types of data. The only requirements for an ExperimentList class are that the objects contained have the following set of methods: dim, [, dimnames

Usage

```
ExperimentList(...)
```

Arguments

... A named list class object

Value

A ExperimentList class object of experiment data

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Examples

```
## Create an empty ExperimentList instance
ExperimentList()
## Create array matrix and AnnotatedDataFrame to create an ExpressionSet class
arraydat <- matrix(data = seq(101, length.out = 20), ncol = 4,
    dimnames = list(
        c("ENST00000294241", "ENST00000355076",
        "ENST00000383706", "ENST00000234812", "ENST00000383323"),
        c("array1", "array2", "array3", "array4")
    ))
colDat <- data.frame(slope53 = rnorm(4),</pre>
    row.names = c("array1", "array2", "array3", "array4"))
## SummarizedExperiment constructor
exprdat <- SummarizedExperiment::SummarizedExperiment(arraydat,</pre>
    colData = colDat)
## Create a sample methylation dataset
methyldat <- matrix(data = seq(1, length.out = 25), ncol = 5,</pre>
    dimnames = list(
        c("ENST00000355076", "ENST00000383706", 
"ENST00000383323", "ENST00000234812", "ENST00000294241"),
        c("methyl1", "methyl2", "methyl3",
          "methyl4", "methyl5")
   ))
## Create a sample RNASeqGene dataset
rnadat <- matrix(</pre>
    data = sample(c(46851, 5, 19, 13, 2197, 507,
        84318, 126, 17, 21, 23979, 614), size = 20, replace = TRUE),
   ncol = 4,
    dimnames = list(
        c("XIST", "RPS4Y1", "KDM5D", "ENST00000383323", "ENST00000234812"),
        c("samparray1", "samparray2", "samparray3", "samparray4")
   ))
## Create a mock RangedSummarizedExperiment from a data.frame
rangedat <- data.frame(chr="chr2", start = 11:15, end = 12:16,
    strand = c("+", "-", "+", "*", "."),
    samp0 = c(0,0,1,1,1), samp1 = c(1,0,1,0,1), samp2 = c(0,1,0,1,0),
    row.names = c(paste0("ENST", "00000", 135411:135414), "ENST00000383323"))
rangeSE <- SummarizedExperiment::makeSummarizedExperimentFromDataFrame(rangedat)</pre>
## Combine to a named list and call the ExperimentList constructor function
assayList <- list(Affy = exprdat, Methyl450k = methyldat, RNASeqGene = rnadat,
                GISTIC = rangeSE)
## Use the ExperimentList constructor
ExpList <- ExperimentList(assayList)</pre>
```

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ExperimentList-class ExperimentList - A container for multi-experiment data

Description

The ExperimentList class is a container that builds on the SimpleList with additional checks for consistency in experiment names and length. It contains a SimpleList of experiments with sample identifiers. One element present per experiment performed.

Convert from SimpleList or list to the multi-experiment data container. When using the **mergeReplicates** method, additional arguments are passed to the given simplify function argument (e.g., na.rm = TRUE)

```
## S4 method for signature 'ExperimentList'
show(object)
## S4 method for signature 'ExperimentList'
isEmpty(x)
## S4 method for signature 'ExperimentList'
dimnames(x)
## S4 method for signature 'ExperimentList'
colnames(x, do.NULL = TRUE, prefix = "col")
## S4 method for signature 'ExperimentList'
rownames(x, do.NULL = TRUE, prefix = "row")
## S4 method for signature 'ExperimentList'
mergeReplicates(x, replicates = list(), simplify = BiocGenerics::mean, ...)
## S4 method for signature 'ANY, missing'
assay(x, i, withDimnames = TRUE, ...)
## S4 method for signature 'ExperimentList'
assays(x, withDimnames = TRUE, ...)
## S4 method for signature 'ExperimentList, missing'
assay(x, i, withDimnames = TRUE, ...)
## S4 method for signature 'ExperimentList, numeric'
assay(x, i, withDimnames = TRUE, ...)
## S4 method for signature 'ExperimentList, character'
assay(x, i, withDimnames = TRUE, ...)
```

Arguments

object, x
do.NULL, prefix
See ?base::rownames for a description of these arguments.

replicates mergeReplicates: A list or LogicalList where each element represents a sample and a vector of repeated measurements for the sample

simplify A function for merging columns where duplicates are indicated by replicates

... Additional arguments. See details for more information.

i A scalar character or integer index

withDimnames logical (default TRUE) whether to return dimension names

Value

An ExperimentList class object

Methods (by generic)

- show(ExperimentList): Show method for ExperimentList class
- isEmpty(ExperimentList): check for zero length across all experiments
- dimnames (ExperimentList): Get the dimension names for an ExperimentList using CharacterList
- colnames(ExperimentList): Get the column names for an ExperimentList as a CharacterList slightly more efficiently
- rownames(ExperimentList): Get the row names for an ExperimentList as a CharacterList slightly more efficiently
- mergeReplicates(ExperimentList): Apply the mergeReplicates method on the ExperimentList elements
- assay(x = ANY, i = missing): Obtain the specified assay with a numeric or character reference
- assays(ExperimentList): Get the assay data from each element in the ExperimentList

coercion

Convert a list or S4 List to an ExperimentList using the as() function.

In the following example, x is either a list or List:

```
\code{as(x, "ExperimentList")}
```

Examples

ExperimentList()

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hasAssay

Checking assay method for any class

Description

The hasAssay function is intended for developers who would like to include new classes into a MultiAssayExperiment instance. It checks the methods tables of the assay function for the specified class of the argument.

Usage

```
hasAssay(object)
```

Arguments

object

A MultiAssayExperiment or named list object instance

Value

A logical value indicating method availability

Examples

```
lst <- structure(list(), .Names=character())
hasAssay(lst)</pre>
```

listToMap

Convert map from data.frame or DataFrame to list and vice versa

Description

The mapToList function provides a convenient way of reordering a data.frame to a list. The listToMap function does the opposite by taking a list and converting it to DataFrame.

```
listToMap(listmap, fill = TRUE)
mapToList(dfmap, assayCol = "assay")
```

Arguments

listmap	A named list object containing DataFrames with "primary" and "colname" columns
fill	$\log i cal(1)$ Whether to fill the map with an empty DataFrame when empty elements are present in the input list
dfmap	A data.frame or DataFrame object with identifiers in the first column
assayCol	A character vector of length one indicating the assay names column

Value

A DataFrame class object of names

A list object of DataFrames for each assay

Functions

• listToMap(): The inverse of the listToMap operation

Examples

```
example("MultiAssayExperiment")
## Create a sampleMap from a list using the listToMap function
sampMap <- listToMap(maplist)
## The inverse operation is also available
maplist <- mapToList(sampMap)</pre>
```

MatchedAssayExperiment-class

 ${\it Matched Assay Experiment - A matched - samples \ Multi Assay Experiment \ class}$

Description

This class supports the use of matched samples where an equal number of observations per biological unit are present in all assays.

Usage

```
MatchedAssayExperiment(...)
```

Arguments

... Either a single MultiAssayExperiment or the components to create a valid MultiAssayExperiment

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Value

A MatchedAssayExperiment object

Functions

MatchedAssayExperiment(): Construct a MatchedAssayExperiment class from MultiAssayExperiment

See Also

MultiAssayExperiment

Examples

```
data("miniACC")
acc <- as(miniACC, "MatchedAssayExperiment")
acc</pre>
```

miniACC

Adrenocortical Carcinoma (ACC) MultiAssayExperiment

Description

A MultiAssayExperiment object providing a reduced version of the TCGA ACC dataset for all 92 patients. RNA-seq, copy number, and somatic mutations are included only for genes whose proteins are included in the reverse-phase protein array. The MicroRNA-seq dataset is also included, with infrequently expressed microRNA removed. Clinical, pathological, and subtype information are provided by colData(miniACC), and some additional details are provided by metadata(miniACC).

Usage

```
data("miniACC")
```

Format

A MultiAssayExperiment with 5 experiments, providing:

RNASeq2GeneNorm RNA-seq count data: an ExpressionSet with 198 rows and 79 columns gisting Processing and Proceedings identified by GISTIC2: a Summarized Experiment with 16

gistict Reccurent copy number lesions identified by GISTIC2: a SummarizedExperiment with 198 rows and 90 columns

RPPAArray Reverse Phase Protein Array: an ExpressionSet with 33 rows and 46 columns.

Rows are indexed by genes, but protein annotations are available from featureData(miniACC[["RPPAArray"]]).

The source of these annotations is noted in abstract(miniACC[["RPPAArray"]])

Mutations Somatic mutations: a matrix with 223 rows and 90 columns. 1 for any kind of non-silent mutation, zero for silent (synonymous) or no mutation.

miRNASeqGene microRNA sequencing: an ExpressionSet with 471 rows and 80 columns. Rows not having at least 5 counts in at least 5 samples were removed.

Author(s)

Levi Waldron < lwaldron.research@gmail.com>

Source

https://github.com/waldronlab/multiassayexperiment-tcga

References

Zheng S *et al.*: Comprehensive Pan-Genomic Characterization of Adrenocortical Carcinoma. Cancer Cell 2016, 29:723-736.

Examples

```
data("miniACC")
metadata(miniACC)
colnames(colData(miniACC))
table(miniACC$vital_status)
longFormat(
    miniACC["MAPK3", , ],
    colDataCols = c("vital_status", "days_to_death")
)
wideFormat(
    miniACC["MAPK3", , ],
    colDataCols = c("vital_status", "days_to_death")
)
```

MultiAssayExperiment Construct an integrative representation of multi-omic data with MultiAssayExperiment

Description

The constructor function for the MultiAssayExperiment-class combines multiple data elements from the different hierarchies of data (study, experiments, and samples). It can create instances where neither a sampleMap or a colData set is provided. Please see the MultiAssayExperiment API documentation for more information.

```
MultiAssayExperiment(
   experiments = ExperimentList(),
   colData = S4Vectors::DataFrame(),
   sampleMap = S4Vectors::DataFrame(assay = factor(), primary = character(), colname =
      character()),
   metadata = list(),
   drops = list()
)
```

Arguments

experiments A list or ExperimentList of all combined experiments

colData A DataFrame or data. frame of characteristics for all biological units

sampleMap A DataFrame or data.frame of assay names, sample identifiers, and colname

samples

metadata An optional argument of "ANY" class (usually list) for content describing the

experiments

drops A list of unmatched information (included after subsetting)

Value

A MultiAssayExperiment object that can store experiment and phenotype data

colData

The 'colData' input can be either 'DataFrame' or 'data.frame' with subsequent coercion to DataFrame. The rownames in the 'colData' must match the colnames in the experiments if no sampleMap is provided.

experiments

The 'experiments' input can be of class SimpleList or 'list'. This input becomes the ExperimentList. Each element of the input 'list' or 'List' must be named, rectangular with two dimensions, and have 'dimnames'.

sampleMap

The sampleMap can either be input as 'DataFrame' or 'data.frame' with eventual coercion to 'DataFrame'. The 'sampleMap' relates biological units and biological measurements within each assay. Each row in the 'sampleMap' is a single such link. The standard column names of the 'sampleMap' are "assay", "primary", and "colname". Note that the "assay" column is a factor corresponding to the names of each experiment in the 'ExperimentList'. In the case where these names do not match between the 'sampleMap' and the experiments, the documented experiments in the 'sampleMap' take precedence and experiments are dropped by the harmonization procedure. The constructor function will generate a 'sampleMap' in the case where it is not provided and this method may be a 'safer' alternative for creating the 'MultiAssayExperiment' (so long as the rownames are identical in the 'colData', if provided). An empty 'sampleMap' may produce empty experiments if the levels of the "assay" factor in the 'sampleMap' do not match the names in the 'ExperimentList'.

See Also

MultiAssayExperiment-class

Examples

```
## Run the example ExperimentList
example("ExperimentList")
```

```
## Create sample maps for each experiment
exprmap <- data.frame(</pre>
    primary = c("Jack", "Jill", "Barbara", "Bob"),
    colname = c("array1", "array2", "array3", "array4"),
    stringsAsFactors = FALSE)
methylmap <- data.frame(</pre>
    primary = c("Jack", "Jack", "Jill", "Barbara", "Bob"),
    colname = c("methyl1", "methyl2", "methyl3", "methyl4", "methyl5"),
    stringsAsFactors = FALSE)
rnamap <- data.frame(</pre>
    primary = c("Jack", "Jill", "Bob", "Barbara"),
    colname = c("samparray1", "samparray2", "samparray3", "samparray4"),
    stringsAsFactors = FALSE)
gistmap <- data.frame(</pre>
    primary = c("Jack", "Bob", "Jill"),
    colname = c("samp0", "samp1", "samp2"),
    stringsAsFactors = FALSE)
## Combine as a named list and convert to a DataFrame
maplist <- list(Affy = exprmap, Methyl450k = methylmap,</pre>
    RNASeqGene = rnamap, GISTIC = gistmap)
## Create a sampleMap
sampMap <- listToMap(maplist)</pre>
## Create an example phenotype data
colDat <- data.frame(sex = c("M", "F", "M", "F"), age = 38:41,
    row.names = c("Jack", "Jill", "Bob", "Barbara"))
## Create a MultiAssayExperiment instance
mae <- MultiAssayExperiment(experiments = ExpList, colData = colDat,</pre>
    sampleMap = sampMap)
```

MultiAssayExperiment-class

MultiAssayExperiment - An integrative multi-assay class for experiment data

Description

The MultiAssayExperiment class can be used to manage results of diverse assays on a collection of specimen. Currently, the class can handle assays that are organized instances of SummarizedExperiment, ExpressionSet, matrix, RaggedExperiment (inherits from GRangesList), and RangedVcfStack. Create new MultiAssayExperiment instances with the homonymous constructor, minimally with the argument ExperimentList, potentially also with the arguments colData (see section below) and sampleMap.

Usage

```
## S4 method for signature 'MultiAssayExperiment'
show(object)
## S4 method for signature 'MultiAssayExperiment'
length(x)
## S4 method for signature 'MultiAssayExperiment'
names(x)
## S4 method for signature 'MultiAssayExperiment'
updateObject(object, ..., verbose = FALSE)
## S4 method for signature 'MultiAssayExperiment'
dimnames(x)
## S4 method for signature 'MultiAssayExperiment'
c(x, ..., sampleMap = NULL, mapFrom = NULL)
## S4 method for signature 'MultiAssayExperiment'
exportClass(
 object,
 dir = tempdir(),
  fmt,
  ext,
 match = FALSE,
 verbose = TRUE,
)
## S4 method for signature 'MultiAssayExperiment'
assays(x, withDimnames = TRUE, ...)
## S4 method for signature 'MultiAssayExperiment,missing'
assay(x, i, withDimnames = TRUE, ...)
## S4 method for signature 'MultiAssayExperiment, numeric'
assay(x, i, withDimnames = TRUE, ...)
## S4 method for signature 'MultiAssayExperiment,character'
assay(x, i, withDimnames = TRUE, ...)
```

Arguments

```
object, x A MultiAssayExperiment object
... Additional arguments for supporting functions. See details.

verbose logical(1) Whether to print additional information (default TRUE)
```

sampleMap	c method: a sampleMap list or DataFrame to guide merge
mapFrom	Either a logical, character, or integer vector indicating the experiment(s) that have an identical colname order as the experiment input(s). If using a character input, the name must match exactly.
dir	character(1) A directory for saving exported data (default: 'tempdir()')
fmt	character(1) or function() Either a format character atomic as supported by 'write.table' either ('csv', or 'tsv') or a function whose first two arguments are 'object to save' and 'file location'
ext	character(1) A file extension supported by the format argument
match	logical(1) Whether to coerce the current object to a 'MatchedAssayExperiment' object (default: FALSE)
withDimnames	logical (default TRUE) whether to return dimension names included in the object

Details

i

The dots (...) argument allows the user to specify additional arguments in several instances.

An integer or character scalar indicating the assay to return

- subsetting [: additional arguments sent to findOverlaps.
- mergeReplicates: used to specify arguments for the simplify functional argument
- assay: may contain withDimnames, which is forwarded to assays
- combining **c**: compatible MultiAssayExperiment classes passed on to the ExperimentList constructor, can be a list, List, or a series of named arguments. See the examples below.

Value

A MultiAssayExperiment object

Methods (by generic)

- show(MultiAssayExperiment): Show method for a MultiAssayExperiment
- length(MultiAssayExperiment): Get the length of ExperimentList
- names(MultiAssayExperiment): Get the names of the ExperimentList
- updateObject(MultiAssayExperiment): Update old serialized MultiAssayExperiment objects to new API
- dimnames(MultiAssayExperiment): Get the dimension names for a MultiAssayExperiment object
- c(MultiAssayExperiment): Add a supported data class to the ExperimentList
- exportClass(MultiAssayExperiment): Export data from class to a series of text files
- assays(MultiAssayExperiment): Obtain a SimpleList of assay data for all available experiments in the object
- assay(x = MultiAssayExperiment, i = missing): Convenience function for extracting the assay of the first element (default) in the ExperimentList. A numeric or character index can also be provided

Slots

ExperimentList A ExperimentList class object for each assay dataset colData A DataFrame of all clinical/specimen data available across experiments sampleMap A DataFrame of translatable identifiers of samples and participants metadata Additional data describing the MultiAssayExperiment object drops A metadata list of dropped information

colData

The colData slot is a collection of primary specimen data valid across all experiments. This slot is strictly of class DataFrame but arguments for the constructor function allow arguments to be of class data.frame and subsequently coerced.

ExperimentList

The ExperimentList slot is designed to contain results from each experiment/assay. It contains a SimpleList.

sampleMap

The sampleMap contains a DataFrame of translatable identifiers of samples and participants or biological units. The standard column names of the sampleMap are "assay", "primary", and "colname". Note that the "assay" column is a factor corresponding to the names of each experiment in the ExperimentList. In the case where these names do not match between the sampleMap and the experiments, the documented experiments in the sampleMap take precedence and experiments are dropped by the harmonization procedure. The constructor function will generate a sampleMap in the case where it is not provided and this method may be a 'safer' alternative for creating the MultiAssayExperiment (so long as the rownames are identical in the colData, if provided). An empty sampleMap may produce empty experiments if the levels of the "assay" factor in the sampleMap do not match the names in the ExperimentList.

coercion

Convert a list or S4 List to a MultiAssayExperiment object using the as function.

In the following example, x is either a list or List:

as(x, "MultiAssayExperiment")

Convert a MultiAssayExperiment to MAF class object using the as function.

In the following example, x is a MultiAssayExperiment:

MultiAssayExperimentToMAF(x)

See Also

MultiAssayExperiment-methods for slot modifying methods, MultiAssayExperiment API

Examples

```
example("MultiAssayExperiment")
## Subsetting
# Rows (i) Rows/Features in each experiment
mae[1, , ]
mae[c(TRUE, FALSE), , ]
# Columns (j) Rows in colData
mae[, rownames(colData(mae))[3:2], ]
# Assays (k)
mae[, , "Affy"]
## Complete cases (returns logical vector)
completes <- complete.cases(mae)</pre>
compMAE <- mae[, completes, ]</pre>
compMAE
colData(compMAE)
example("MultiAssayExperiment")
## Add an experiment
test1 <- mae[[1L]]
colnames(test1) <- rownames(colData(mae))</pre>
## Combine current MultiAssayExperiment with additional experiment
## (no sampleMap)
c(mae, newExperiment = test1)
test2 <- mae[[3L]]
c(mae, newExp = test2, mapFrom = 3L)
## Add experiment using experiment name in mapFrom
c(mae, RNASeqGeneV2 = test2, mapFrom = "RNASeqGene")
```

MultiAssayExperiment-helpers

A group of helper functions for manipulating and cleaning a MultiAssayExperiment

Description

A set of helper functions were created to help clean and manipulate a MultiAssayExperiment object. intersectRows also works for ExperimentList objects.

• complete.cases: Returns a logical vector corresponding to 'colData' rows that have data across all experiments

- isEmpty: Returns a logical TRUE value for zero length MultiAssayExperiment objects
- intersectRows: Takes all common rows across experiments, excludes experiments with empty rownames
- intersectColumns: A wrapper for complete.cases to return a MultiAssayExperiment with only those biological units that have measurements across all experiments
- replicated: Identifies, via logical vectors, colnames that originate from a single biological unit within each assay
- replicates: Provides the replicate colnames found with the replicated function by their name, empty list if none
- anyReplicated: Whether the assay has replicate measurements
- showReplicated: Displays the actual columns that are replicated per assay and biological unit, i.e., primary value (colData rowname) in the sampleMap
- mergeReplicates: A function that combines replicated / repeated measurements across all experiments and is guided by the replicated return value
- longFormat: A MultiAssayExperiment method that returns a small and skinny DataFrame. The colDataCols arguments allows the user to append colData columns to the data.
- wideFormat: A function to reshape the data in a MultiAssayExperiment to a "wide" format DataFrame. Each row in the DataFrame represents an observation (corresponding to an entry in the colData). If replicates are present, their data will be appended at the end of the corresponding row and will generate additional NA data. It is recommended to remove or consolidate technical replicates with mergeReplicates. Optional colDataCols can be added when the original object is a MultiAssayExperiment.
- hasRowRanges: A function that identifies ExperimentList elements that have a rowRanges method
- getWithColData: A convenience function for extracting an assay and associated colData
- renamePrimary: A convenience function to rename the primary biological units as represented in the rownames(colData)
- renameColname: A convenience function to rename the colnames of a particular assay

```
## S4 method for signature 'MultiAssayExperiment'
complete.cases(...)

## S4 method for signature 'MultiAssayExperiment'
isEmpty(x)

intersectRows(x)

intersectColumns(x)

replicated(x)

## S4 method for signature 'MultiAssayExperiment'
replicated(x)
```

```
anyReplicated(x)
## S4 method for signature 'MultiAssayExperiment'
anyReplicated(x)
showReplicated(x)
## S4 method for signature 'MultiAssayExperiment'
showReplicated(x)
replicates(x, ...)
## S4 method for signature 'MultiAssayExperiment'
replicates(x, ...)
mergeReplicates(x, replicates = list(), simplify = BiocGenerics::mean, ...)
## S4 method for signature 'MultiAssayExperiment'
mergeReplicates(
 х,
  replicates = replicated(x),
  simplify = BiocGenerics::mean,
)
## S4 method for signature 'ANY'
mergeReplicates(x, replicates = list(), simplify = BiocGenerics::mean, ...)
longFormat(object, colDataCols = NULL, i = 1L)
wideFormat(
  object,
  colDataCols = NULL,
  check.names = TRUE,
  collapse = "_",
  i = 1L
)
hasRowRanges(x)
## S4 method for signature 'MultiAssayExperiment'
hasRowRanges(x)
## S4 method for signature 'ExperimentList'
hasRowRanges(x)
getWithColData(x, i, mode = c("append", "replace"), verbose = FALSE)
```

```
renamePrimary(x, value)
renameColname(x, i, value)
splitAssays(x, hitList)
## S4 method for signature 'MultiAssayExperiment'
splitAssays(x, hitList)
makeHitList(x, patternList)
```

Arguments

... Additional arguments. See details for more information.

x A MultiAssayExperiment or ExperimentList

replicates A list of LogicalLists indicating multiple / duplicate entries for each biological

unit per assay, see replicated (default replicated(x)).

simplify A function for merging repeat measurements in experiments as indicated by the

replicated method for MultiAssayExperiment

object Any supported class object

colDataCols A character, logical, or numeric index for colData columns to be included

i longFormat: The i-th assay in SummarizedExperiment-like objects. A vector in-

put is supported in the case that the SummarizedExperiment object(s) has more than one assay (default 1L), renameColname: Either a numeric or character

index indicating the assay whose colnames are to be renamed

check.names (logical default TRUE) Column names of the output DataFrame will be checked

for syntactic validity and made unique, if necessary

collapse (character default "_") A single string delimiter for output column names. In

wideFormat, experiments and rownames (and when replicate samples are present,

colnames) are seperated by this delimiter

mode String indicating how MultiAssayExperiment column-level metadata should be

 $added \ to \ the \ {\color{blue} Summarized Experiment \ colData}.$

verbose logical(1) Whether to 'suppressMessages' on subsetting operations in 'getWith-

ColData' (default FALSE)

value renamePrimary: A character vector of the same length as the existing rownames(colData)

to use for replacement, renameColname: A CharacterList or list with match-

ing lengths to replace colnames(x)

hitList a named list or List of logical vectors that indicate groupings in the assays

patternList a named list or List of atomic character vectors that are the input to grepl

for identifying groupings in the assays

Details

The replicated function finds replicate measurements in each assay and returns a list of LogicalLists. Each element in a single LogicalList corresponds to a biological or *primary* unit as in the sampleMap. Below is a small graphic for one particular biological unit in one assay, where the logical vector corresponds to the number of measurements/samples in the assay:

```
> replicated(MultiAssayExperiment)
(list str) '-- $ AssayName
(LogicalList str) '-- [[ "Biological Unit" ]]
Replicated if sum(...) > 1 '-- TRUE TRUE FALSE FALSE
```

anyReplicated determines if any of the assays have at least one replicate. *Note*. These methods are not available for the ExperimentList class due to a missing sampleMap structure (by design). showReplicated returns a list of CharacterLists where each element corresponds to the the biological or *primary* units that are replicated in that assay element. The values in the inner list are the colnames in the assay that are technical replicates.

The replicates function (noun) returns the colnames from the sampleMap that were identified as replicates. It returns a list of CharacterLists for each assay present in the MultiAssayExperiment and an inner entry for each biological unit that has replicate observations in that assay.

The mergeReplicates function is a house-keeping method for a MultiAssayExperiment where only complete.cases are returned. This by-assay operation averages replicate measurements (by default) and columns are aligned by the row order in colData. Users can provide their own function for merging replicates with the simplify functional argument. Additional inputs . . . are sent to the 'simplify' function.

The mergeReplicates "ANY" method consolidates duplicate measurements for rectangular data structures, returns object of the same class (endomorphic). The ellipsis or . . . argument allows the user to provide additional arguments to the simplify functional argument.

The longFormat "ANY" class method, works with classes such as ExpressionSet and Summarized-Experiment as well as matrix to provide a consistent long and skinny DataFrame.

The hasRowRanges method identifies assays that support a rowRanges method *and* return a GRanges object.

Value

See the itemized list in the description section for details.

mergeReplicates

The mergeReplicates function makes use of the output from replicated which will point out the duplicate measurements by biological unit in the MultiAssayExperiment. This function will return a MultiAssayExperiment with merged replicates. Additional arguments can be provided to the simplify argument via the ellipsis (...). For example, when replicates "TCGA-B" and "TCGA-A" are found in the assay, the name of the first appearing replicate is taken (i.e., "B"). Note that a typical use case of merging replicates occurs when there are multiple measurements on the **same** sample (within the same assay) and can therefore be averaged.

longFormat

The 'longFormat' method takes data from the ExperimentList in a MultiAssayExperiment and returns a uniform DataFrame. The resulting DataFrame has columns indicating primary, rowname, colname and value. This method can optionally include columns of the MultiAssayExperiment colData named by colDataCols character vector argument. (MultiAssayExperiment method only). The i argument allows the user to specify the assay value for the SummarizedExperiment assay function's i argument.

wideFormat

The wideFormat function returns standardized wide DataFrame where each row represents a biological unit as in the colData. Depending on the data and setup, biological units can be patients, tumors, specimens, etc. Metadata columns are generated based on the names produced in the wide format DataFrame. These can be accessed via the mcols function. See the wideFormat section for description of the colDataCols and i arguments.

hasRowRanges

The hasRowRanges method identifies assays with associated ranged row data by directly testing the method on the object. The result from the test must be a GRanges class object to satisfy the test.

getWithColData

The getWithColData function allows the user to conveniently extract a particular assay as indicated by the i index argument. It will also attempt to provide the colData along with the extracted object using the colData<- replacement method when possible. Typically, this method is available for SummarizedExperiment and RaggedExperiment classes.

The setting of mode determines how the colData is added. If mode="append", the MultiAssayExperiment metadata is appended onto that of the SummarizedExperiment. If any fields are duplicated by name, the values in the SummarizedExperiment are retained, with a warning emitted if the values are different. For mode="replace", the MultiAssayExperiment metadata replaces that of the SummarizedExperiment, while for mode="none", no replacement or appending is performed.

rename*

The renamePrimary function allows the user to conveniently change the actual names of the primary biological units as seen in rownames(colData). renameColname allows the user to change the names of a particular assay based on index i. i can either be a single numeric or character value. See colnames<- method for renaming multiple colnames in a MultiAssayExperiment.

splitAssays

The splitAssays method separates columns in each of the assays based on the hitList input. The hitList can be generated using the makeHitList helper function. To use the makeHitList helper, the user should input a list of patterns that will match on the column names of each assay. These matches should be mutually exclusive as to avoid repetition of columns across assays. See the examples section.

Examples

```
example(MultiAssayExperiment)
complete.cases(mae)
isEmpty(MultiAssayExperiment())
## renaming biological units (primary)
mae2 <- renamePrimary(mae, paste0("pt", 1:4))</pre>
colData(mae2)
sampleMap(mae2)
## renaming observational units (colname)
mae2 <- renameColname(mae, i = "Affy", paste0("ARRAY", 1:4))</pre>
colnames(mae2)
sampleMap(mae2)
patts <- list(</pre>
    normals = TCGA-[A-Z0-9]{2}-[A-Z0-9]{4}-11,
    tumors = "TCGA-[A-Z0-9]{2}-[A-Z0-9]{4}-01"
)
data("miniACC")
hits <- makeHitList(miniACC, patts)</pre>
## only turmors present
splitAssays(miniACC, hits)
```

MultiAssayExperiment-methods

Accessing and modifying information in MultiAssayExperiment

Description

A set of accessor and setter generic functions to extract either the sampleMap, the ExperimentList, colData, or metadata slots of a MultiAssayExperiment object

```
## S4 method for signature 'MultiAssayExperiment'
sampleMap(x)
```

```
## S4 method for signature 'MultiAssayExperiment'
experiments(x)
## S4 method for signature 'MultiAssayExperiment'
colData(x, ...)
## S4 method for signature 'MultiAssayExperiment'
drops(x)
## S4 replacement method for signature 'MultiAssayExperiment,DataFrame'
sampleMap(object) <- value</pre>
## S4 replacement method for signature 'MultiAssayExperiment,ANY'
sampleMap(object) <- value</pre>
drops(x, ...) \leftarrow value
## S4 replacement method for signature 'MultiAssayExperiment,ExperimentList'
experiments(object) <- value</pre>
## S4 replacement method for signature 'MultiAssayExperiment,List'
experiments(object) <- value</pre>
## S4 replacement method for signature 'MultiAssayExperiment,DataFrame'
colData(x) \leftarrow value
## S4 replacement method for signature 'MultiAssayExperiment,ANY'
colData(x) <- value</pre>
## S4 replacement method for signature 'MultiAssayExperiment'
drops(x, ...) \leftarrow value
## S4 replacement method for signature 'MultiAssayExperiment'
x$name <- value
## S4 replacement method for signature 'MultiAssayExperiment'
names(x) \leftarrow value
## S4 replacement method for signature 'MultiAssayExperiment,List'
colnames(x) \leftarrow value
## S4 replacement method for signature 'MultiAssayExperiment,list'
colnames(x) \leftarrow value
## S4 method for signature 'MultiAssayExperiment'
x$name
```

```
## S4 method for signature 'MultiAssayExperiment'
metadata(x, ...)
## S4 replacement method for signature 'MultiAssayExperiment'
metadata(x, ...) <- value</pre>
```

Arguments

... Argument not in use

object, x $\hspace{1cm} A \hspace{1cm} ext{MultiAssayExperiment object}$

value See details.

name A column in colData

Value

Accessors: Either a sampleMap, ExperimentList, or DataFrame object

Setters: A MultiAssayExperiment object

Accessors

Eponymous names for accessing MultiAssayExperiment slots with the exception of the ExperimentList accessor named experiments.

- colData: Access the colData slot
- sampleMap: Access the sampleMap slot
- experiments: Access the ExperimentList slot
- '[[': Access the ExperimentList slot
- '\$': Access a column in colData
- 'drops': Get a vector of dropped ExperimentList names

Setters

Setter method values (i.e., 'function(x) <- value'):

- experiments<-: An ExperimentList object containing experiment data of supported classes
- sampleMap<-: A DataFrame object relating samples to biological units and assays
- colData<-: A DataFrame object describing the biological units
- metadata<-: A list object of metadata
- '[[<-': Equivalent to the experiments<- setter method for convenience
- '\$<-': A vector to replace the indicated column in colData
- 'drops<-': Trace ExperimentList names that have been removed

Examples

```
## Load example MultiAssayExperiment
example(MultiAssayExperiment)
## Access the sampleMap
sampleMap(mae)
## Replacement method for a MultiAssayExperiment sampleMap
sampleMap(mae) <- S4Vectors::DataFrame()</pre>
## Access the ExperimentList
experiments(mae)
## Replace with an empty ExperimentList
experiments(mae) <- ExperimentList()</pre>
## Access the metadata
metadata(mae)
## Replace metadata with a list
metadata(mae) <- list(runDate =</pre>
    format(Sys.time(), "%B %d, %Y"))
## Access the colData
colData(mae)
## Access a column in colData
mae$age
## Replace a column in colData
mae$age <- mae$age + 1
```

MultiAssayExperimentToMAF

Convert MultiAssayExperiment to MAF class

Description

Take a MultiAssayExperiment object with specific mutation assays and convert these into a maftools representation. The names provided via synAssay and nonSynAssay must match exactly those assays in the MultiAssayExperiment.

```
MultiAssayExperimentToMAF(x, synAssay = "maf_syn", nonSynAssay = "maf_nonSyn")
```

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Arguments

Χ	A MultiAssayExperiment object
synAssay	$character (1) \ The \ name \ of the \ Experiment List \ element \ in \ the \ MultiAssay Experiment that \ identifies \ synonymous \ variant \ classifications.$
nonSynAssay	$character (1) \ The \ name \ of the \ Experiment List \ element \ in \ the \ MultiAssay Experiment that \ identifies \ non-synonymous \ variant \ classifications.$

prepMultiAssay Prepare a MultiAssayExperiment instance

Description

The purpose of this helper function is to faciltate the creation of a MultiAssayExperiment object by detecting any inconsistencies with all types of names in either the ExperimentList, the colData, or sampleMap.

Usage

```
prepMultiAssay(ExperimentList, colData, sampleMap, ...)
```

Arguments

ExperimentList A list of all combined experiments

colData A DataFrame of the phenotype data for all participants

sampleMap A DataFrame of sample identifiers, assay samples, and assay names

... Optional arguments for the MultiAssayExperiment constructor function such

as metadata and drops.

Value

A list containing all the essential components of a MultiAssayExperiment as well as a "drops" metadata element that indicates non-matched names. The names of the resulting list correspond to the arguments of the MultiAssayExperiment constructor function.

Checks

The prepMultiAssay function checks that all columns in the sampleMap are character.

It checks that all names and lengths match in both the ExperimentList and in the unique assay names of the sampleMap.

If ExperimentList names and assay names only differ by case and are not duplicated, the function will standardize all names to lowercase.

If names cannot be matched between the colname column of the sampleMap and the colnames of the ExperimentList, those unmatched will be dropped and found in the "drops" element of the resulting list.

Names in the "primary" column of the sampleMap, will be matched to those in the colData. Unmatched "primary" column rows will be dropped from the sampleMap. Suggestions for name fixes in either the ExperimentList or colnames will be made when necessary.

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Examples

```
## Run example
example("MultiAssayExperiment")

## Check if there are any inconsistencies within the different names
preparedMAE <- prepMultiAssay(ExpList, colDat, sampMap)

## Results in a list of components for the MultiAssayExperiment constructor
## function
MultiAssayExperiment(preparedMAE$experiments, preparedMAE$colData,
preparedMAE$sampleMap)

## Alternatively, use the do.call function
do.call(MultiAssayExperiment, preparedMAE)</pre>
```

reexports

Objects exported from other packages

Description

These objects are imported from other packages. Click on the function name to see their documentation.

• S4Vectors: DataFrame

Examples

DataFrame()

saveHDF5MultiAssayExperiment

Save a MultiAssayExperiment class object to HDF5 and Rds files

Description

This function takes a MultiAssayExperiment object and uses the assays functionality to obtain data matrices out of the experiments. These are then saved into the .h5 file format. This function relies heavily on the HDF5Array package whose installation is required before use. saveHDF5MultiAssayExpeirment preserves the classes contained in the ExperimentList with the exception of matrix which is converted to HDF5Matrix. Internal SummarizedExperiment assays are converted to HDF5-backed assays as in HDF5Array::saveHDF5SummarizedExperiment. SummarizedExperiment objects with multiple i-th assays will have the first assay take precedence and others assays will be dropped with a warning. If the first assay in a SummarizedExperiment contains an array, the array is preserved in the process of saving and loading the HDF5-backed MultiAssayExperiment.

Usage

```
saveHDF5MultiAssayExperiment(
    x,
    dir = "h5_mae",
    prefix = NULL,
    replace = FALSE,
    chunkdim = NULL,
    level = NULL,
    as.sparse = NA,
    verbose = NA
)

loadHDF5MultiAssayExperiment(dir = "h5_mae", prefix = NULL)
```

Arguments

x A MultiAssayExperiment object or derivative

dir The path (as a single string) to the directory where to save the HDF5-based

MultiAssayExperiment object or to load it from.

When saving, the directory will be created if it doesn't already exist. If the directory already exists and no prefix is specified and replace is set to TRUE,

then it's replaced with an empty directory.

prefix An optional prefix to add to the names of the files created inside dir. This allows

saving more than one object in the same directory. When the prefix is NULL, the name of the x input MultiAssayExperiment is used. To avoid the default setting use an empty character string i.e., "". An underscore (_) is appended to the prefix when provided; therefore, typical inputs should be words, e.g., "test".

replace When no prefix is specified, should a pre-existing directory be replaced with a

new empty one? The content of the pre-existing directory will be lost!

chunkdim, level

The dimensions of the chunks and the compression level to use for writing the assay data to disk.

Passed to the internal calls to writeHDF5Array. See ?writeHDF5Array for more information.

as.sparse Whether the assay data should be flagged as sparse or not. If set to NA (the default), then the specific as.sparse value to use for each assay is determined

by calling is_sparse() on them.

Passed to the internal calls to writeHDF5Array. See ?writeHDF5Array for more information and an IMPORTANT NOTE.

verbose Set to TRUE to make the function display progress.

In the case of saveHDF5MultiAssayExperiment(), verbose is set to NA by de-

 $fault, in which case \ verbosity \ is \ controlled \ by \ Delayed Array. \ verbose. block. processing$

option. Setting verbose to TRUE or FALSE overrides the option.

.

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Examples

```
data("miniACC")

testDir <- file.path(tempdir(), "test_mae")

saveHDF5MultiAssayExperiment(
    miniACC, dir = testDir, verbose = TRUE, replace = TRUE
)

## inspect the files in the dir
list.files(testDir)

loadHDF5MultiAssayExperiment(
    dir = testDir
)

## remove example files
unlink(testDir, recursive = TRUE)</pre>
```

subsetBy

Subsetting a MultiAssayExperiment object

Description

A set of functions for extracting and dividing a MultiAssayExperiment

```
subsetByRow(x, y, ...)
subsetByColData(x, y)
subsetByColumn(x, y)
subsetByAssay(x, y)
## S4 method for signature 'ExperimentList, ANY'
subsetByRow(x, y, ...)
## S4 method for signature 'ExperimentList, list'
subsetByRow(x, y)
## S4 method for signature 'ExperimentList, List'
subsetByRow(x, y)
## S4 method for signature 'ExperimentList, List'
subsetByRow(x, y)
```

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```
## S4 method for signature 'ExperimentList,list'
subsetByColumn(x, y)
## S4 method for signature 'ExperimentList,List'
subsetByColumn(x, y)
## S4 method for signature 'ExperimentList,logical'
subsetByColumn(x, y)
## S4 method for signature 'ExperimentList'
subsetByAssay(x, y)
## S4 method for signature 'MultiAssayExperiment, ANY'
subsetByColData(x, y)
## S4 method for signature 'MultiAssayExperiment,character'
subsetByColData(x, y)
## S4 method for signature 'MultiAssayExperiment,ANY'
subsetByRow(x, y, ...)
## S4 method for signature 'MultiAssayExperiment,ANY'
subsetByColumn(x, y)
## S4 method for signature 'MultiAssayExperiment'
subsetByAssay(x, y)
## S4 method for signature 'MultiAssayExperiment, ANY, ANY, ANY'
x[i, j, k, ..., drop = FALSE]
## S4 method for signature 'MultiAssayExperiment,ANY,ANY'
x[[i, j, ...]]
## S4 replacement method for signature 'MultiAssayExperiment,ANY,ANY'
x[[i, j, \ldots]] \leftarrow value
## S4 replacement method for signature 'MultiAssayExperiment,ANY,ANY,ANY'
x[i, j, ...] \leftarrow value
```

Arguments

X	A MultiAssayExperiment or ExperimentList
У	Any argument used for subsetting, can be a character, logical, integer, list or List vector
	Additional arguments passed on to lower level functions.
i	Either a character, integer, logical or GRanges object for subsetting by rows

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j	Either a character, logical, or numeric vector for subsetting by colData rows. See details for more information.
k	Either a character, logical, or numeric vector for subsetting by assays
drop	logical (default FALSE) whether to drop all empty assay elements in the Experiment List
value	An assay compatible with the MultiAssayExperiment API

Details

Subsetting a MultiAssayExperiment by the \mathbf{j} index can yield a call to either subsetByColData or subsetByColumn. For vector inputs, the subset will be applied to the colData rows. For List-type inputs, the List will be applied to each of the elements in the ExperimentList. The order of the subsetting elements in the List must match that of the ExperimentList in the MultiAssayExperiment.

- subsetBycolData: Select biological units by vector input types
- subsetByColumn: Select observations by assay or for each assay
- subsetByRow: Select rows by assay or for each assay
- subsetByAssay: Select experiments

Value

 $subset By * operations \ are \ endomorphic \ and \ return \ either \ MultiAssay Experiment \ or \ Experiment List \ depending \ on \ the \ input.$

Examples

```
## Load the example MultiAssayExperiment
example("MultiAssayExperiment")
## Using experiment names
subsetByAssay(mae, "Affy")
## Using numeric indices
subsetByAssay(mae, 1:2)
## Using a logical vector
subsetByAssay(mae, c(TRUE, FALSE, TRUE))
## Subset by character vector (Jack)
subsetByColData(mae, "Jack")
## Subset by numeric index of colData rows (Jack and Bob)
subsetByColData(mae, c(1, 3))
## Subset by logical indicator of colData rows (Jack and Jill)
subsetByColData(mae, c(TRUE, TRUE, FALSE, FALSE))
subsetByColumn(mae, list(Affy = 1:2,
   Methyl450k = c(3,5,2), RNASeqGene = 2:4, GISTIC = 1))
subsetWith <- S4Vectors::mendoapply(`[`, colnames(mae),</pre>
```

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upsetSamples

Create a generalized Venn Diagram analog for sample membership in multiple assays, using the upset algorithm in UpSetR

Description

Create a generalized Venn Diagram analog for sample membership in multiple assays, using the upset algorithm in UpSetR

Usage

```
upsetSamples(
  MultiAssayExperiment,
  nsets = NULL,
  sets = names(MultiAssayExperiment),
  nintersects = NA_integer_,
  order.by = "freq",
  check.names = FALSE,
  ...
)
```

Arguments

 ${\it MultiAssayExperiment}$

A MultiAssayExperiment object

nsets numeric(1) The number of sets to analyze. If specified, sets will be ignored.

sets character() A character vector of names in MultiAssayExperiment to use. If

specified, nsets will be ignored.

nintersects numeric() The number of intersections to plot. By default, all intersections will

be plotted.

order.by How the intersections in the matrix should be ordered by. Options include fre-

quency (entered as "freq"), degree, or both in any order.

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```
check.names logical(1) Whether to munge names as in the data.frame() constructor (default FALSE).parameters passed to upset
```

Value

Produces a visualization of set intersections using the UpSet matrix design

Note

This function is intended to provide convenient visualization of assay availability configurations in MultiAssayExperiment instances. The UpSetR::upset function requires data.frame input and has many parameters to tune appearance of the result. Assay name handling is important for interpretability.

Author(s)

Vincent J Carey

Examples

```
data(miniACC)
upsetSamples(miniACC)
upsetSamples(miniACC, nsets = 3, nintersects = 3)
```

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