Package 'glmSparseNet'

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Type Package

Title Network Centrality Metrics for Elastic-Net Regularized Models

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Description glmSparseNet is an R-package that generalizes sparse regression models when the features (e.g. genes) have a graph structure (e.g. protein-protein interactions), by including network-based regularizers. glmSparseNet uses the glmnet R-package, by including centrality measures of the network as penalty weights in the regularization. The current version implements regularization based on node degree, i.e. the strength and/or number of its associated edges, either by promoting hubs in the solution or orphan genes in the solution. All the glmnet distribution families are supported, namely ``gaussian", ``poisson", ``binomial", ``multinomial", ``cox", and ``mgaussian".

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BugReports https://www.github.com/sysbiomed/glmSparseNet/issues

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`glmSparseNet-package` *glmSparseNet: Network Centrality Metrics for Elastic-Net Regularized Models*

Description

glmSparseNet is an R-package that generalizes sparse regression models when the features (e.g. genes) have a graph structure (e.g. protein-protein interactions), by including network-based regularizers. glmSparseNet uses the glmnet R-package, by including centrality measures of the network as penalty weights in the regularization. The current version implements regularization based on node degree, i.e. the strength and/or number of its associated edges, either by promoting hubs in the solution or orphan genes in the solution. All the glmnet distribution families are supported, namely "gaussian", "poisson", "binomial", "multinomial", "cox", and "mgaussian".

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See Also

Useful links:

- https://www.github.com/sysbiomed/glmSparseNet
- Report bugs at https://www.github.com/sysbiomed/glmSparseNet/issues

---

`.baseDir` *Change base dir for '.runCache*

Description

Change base dir for '.runCache

Usage

```
.baseDir(path = NULL)
```

Arguments

`path`	to base directory where cache is saved

Value

the new path

Examples

```
glmSparseNet:::.baseDir("/tmp/cache")
```

`.biomartLoad` *Common call to biomaRt to avoid repetitive code*

Description

Common call to biomaRt to avoid repetitive code

Usage

```
.biomartLoad(attributes, filters, values, useCache, verbose)
```

Arguments

attributes	Attributes you want to retrieve. A possible list of attributes can be retrieved using the function biomaRt::listAttributes.
filters	Filters (one or more) that should be used in the query. A possible list of filters can be retrieved using the function biomaRt::listFilters.
values	Values of the filter, e.g. vector of affy IDs. If multiple filters are specified then the argument should be a list of vectors of which the position of each vector corresponds to the position of the filters in the filters argument
useCache	Boolean indicating if biomaRt cache should be used
verbose	When using biomaRt in webservice mode and setting verbose to TRUE, the XML query to the webservice will be printed.

Value

data.frame with attributes as columns and values translated to them

See Also

geneNames

ensemblGeneNames

protein2EnsemblGeneNames

biomaRt::getBM()

biomaRt::useEnsembl()

Examples

```
glmSparseNet:::.biomartLoad(
    attributes = c("external_gene_name", "ensembl_gene_id"),
    filters = "external_gene_name",
    values = c("MOB1A", "RFLNB", "SPIC", "TP53"),
    useCache = TRUE,
    verbose = FALSE
)
```

.buildFunctionDigest *Build digest of function from the actual code*

Description

Build digest of function from the actual code

Usage

```
.buildFunctionDigest(fun)
```

Arguments

fun	function call name

Value

a digest

Examples

```
glmSparseNet:::.buildFunctionDigest(sum)
glmSparseNet:::.buildFunctionDigest(c)
```

.cacheCompression *Change cache.compression for run_cache*

Description

Change cache.compression for run_cache

Usage

```
.cacheCompression(compression = NULL)
```

Arguments

compression	see compression parameter in save function

Value

the new compression

Examples

```
glmSparseNet:::.cacheCompression("bzip2")
```

.calcPenalty *Calculate penalty based on data*

Description

Internal method to calculate the network using data-dependant methods

Usage

```
.calcPenalty(xdata, penaltyType, options = networkOptions())
```

Arguments

xdata	input data
penaltyType	which method to use
options	options to be used

Value

vector with penalty weights

Examples

```
xdata <- matrix(rnorm(1000), ncol = 200)
glmSparseNet:::.calcPenalty(xdata, "none")
glmSparseNet:::.calcPenalty(
    xdata, "correlation",
    networkOptions(cutoff = .6)
)
glmSparseNet:::.calcPenalty(xdata, "correlation")
glmSparseNet:::.calcPenalty(
    xdata, "covariance",
    networkOptions(cutoff = .6)
)
glmSparseNet:::.calcPenalty(xdata, "covariance")
```

.calculateResult *Calculate/load result and save if necessary*

Description

This is where the actual work is done

Usage

```
.calculateResult(path, compression, forceRecalc, showMessage, fun, ...)
```

Arguments

path	path to save cache
compression	compression used in save
forceRecalc	force to recalculate cache
showMessage	boolean to show messages
fun	function to be called
...	arguments to said function ,

Value

result of fun(...)

Examples

```
glmSparseNet:::.calculateResult(
    file.path(tempdir(), "calculate_result.Rdata"),
    "gzip",
    FALSE,
    TRUE,
    sum,
    1, 2, 3
)
```

.combinedScore *Calculate combined score for STRINGdb interactions*

Description

Please note that all the interactions have duplicates as it's a two way interaction (score(ProteinA-Protein) == score(ProteinB, PorteinA))

Usage

```
.combinedScore(allInteractions, scoreThreshold, removeText)
```

Arguments

allInteractions	table with score of all interactions
scoreThreshold	threshold to keep interactions
removeText	remove text-based interactions

Details

To better understand how the score is calculated, please see: https://string-db.org/help/faq/#how-are-the-scores-computed

Value

table with combined score

`.createDirectoryForCache` *Create directories for cache*

Description

Create directories for cache

Usage

```
.createDirectoryForCache(baseDir, parentPath)
```

Arguments

`baseDir`	tentative base dir to create.
`parentPath`	first 4 characters of digest that will become parent directory for the actual cache file (this reduces number of files per folder)

Value

a list of updated baseDir and parentDir

Examples

```
glmSparseNet:::.createDirectoryForCache(tempdir(), "abcd")

glmSparseNet:::.createDirectoryForCache(
    file.path(getwd(), "run-cache"), "abcd"
)
```

`.curlWorkaround` *Workaround for bug with curl when fetching specific ensembl mirror*

Description

Should be solved in issue #39, will test to remove it.

Usage

```
.curlWorkaround(expr)
```

Arguments

expr	expression

Value

result of expression

Examples

```
glmSparseNet:::.curlWorkaround({
    biomaRt::useEnsembl(
        biomart = "genes",
        dataset = "hsapiens_gene_ensembl"
    )
})
```

.degreeGeneric *Generic function to calculate degree based on data*

Description

The assumption to use this function is that the network represented by a matrix is symetric and without any connection the node and itself.

Usage

```
.degreeGeneric(
  fun = stats::cor,
  funPrefix = "operator",
  xdata,
  cutoff = 0,
  considerUnweighted = FALSE,
  chunks = 1000,
  forceRecalcDegree = FALSE,
  forceRecalcNetwork = FALSE,
  nCores = 1,
  ...
)
```

Arguments

fun	function that will calculate the edge weight between 2 nodes
funPrefix	used to store low-level information on network as it can become to large to be stored in memory
xdata	calculate correlation matrix on each column
cutoff	positive value that determines a cutoff value

`considerUnweighted`
: consider all edges as 1 if they are greater than 0

`chunks`
: calculate function at batches of this value (default is 1000)

`forceRecalcDegree`
: force recalculation of penalty weights (but not the network), instead of going to cache

`forceRecalcNetwork`
: force recalculation of network and penalty weights, instead of going to cache

`nCores`
: number of cores to be used

`...`
: extra parameters for fun

Value

a vector of the degrees

.digestCache *Default digest method*

Description

Sets a default caching algorithm to use with `.runCache`

Usage

```
.digestCache(val)
```

Arguments

`val`
: object to calculate hash over

Value

a hash of the sha256

Examples

```
glmSparseNet:::.digestCache(c(1, 2, 3, 4, 5))
glmSparseNet:::.digestCache("some example")
```

`.glmSparseNetPrivate` *Calculate GLM model with network-based regularization*

Description

Calculate GLM model with network-based regularization

Usage

```
.glmSparseNetPrivate(
  fun,
  xdata,
  ydata,
  network,
  experiment = NULL,
  options = networkOptions(),
  ...
)
```

Arguments

`fun`	function to be called (glmnet or cv.glmnet)
`xdata`	input data, can be a matrix or MultiAssayExperiment
`ydata`	response data compatible with glmnet
`network`	type of network, see below
`experiment`	when xdata is a MultiAssayExperiment object this parameter is required
`options`	options to calculate network
`...`	parameters that glmnet accepts

Value

an object just as glmnet network parameter accepts:

- string to calculate network based on data (correlation, covariance)
- matrix representing the network
- vector with already calculated penalty weights (can also be used directly with glmnet)

`.networkGenericParallel` *Calculate the upper triu of the matrix*

Description

Calculate the upper triu of the matrix

Usage

```
.networkGenericParallel(
  fun,
  funPrefix,
  xdata,
  buildOutput = "matrix",
  nCores = 1,
  forceRecalcNetwork = FALSE,
  showMessage = FALSE,
  ...
)
```

Arguments

`fun`	function that will calculate the edge weight between 2 nodes
`funPrefix`	used to store low-level information on network as it can become to large to be stored in memory
`xdata`	base data to calculate network
`buildOutput`	if output returns a 'matrix', 'vector' of the upper triu without the diagonal or NULL with any other argument
`nCores`	number of cores to be used
`forceRecalcNetwork`	force recalculation, instead of going to cache
`showMessage`	shows cache operation messages
`...`	extra parameters for fun

Value

depends on buildOutput parameter

.networkWorker *Worker to calculate edge weight for each pair of ixI node and following*

Description

Note that it assumes it does not calculate for index below and equal to ixI

Usage

```
.networkWorker(fun, xdata, ixI, ...)
```

Arguments

fun	function to be used, can be cor, cov or any other defined function
xdata	original data to calculate the function over
ixI	starting index, this can be used to save ony upper triu
...	extra parameters for fun

Value

a vector with size `ncol(xdata) - ixI`

.runCache *Run function and save cache*

Description

This method saves the function that's being called

Usage

```
.runCache(
  fun,
  ...,
  seed = NULL,
  baseDir = NULL,
  cachePrefix = "generic_cache",
  cacheDigest = list(),
  showMessage = NULL,
  forceRecalc = FALSE,
  addToHash = NULL
)

## S4 method for signature 'function'
```

```
.runCache(
  fun,
  ...,
  seed = NULL,
  baseDir = NULL,
  cachePrefix = "generic_cache",
  cacheDigest = list(),
  showMessage = NULL,
  forceRecalc = FALSE,
  addToHash = NULL
)
```

Arguments

fun	function call name
...	parameters for function call
seed	when function call is random, this allows to set seed beforehand
baseDir	directory where data is stored
cachePrefix	prefix for file name to be generated from parameters (...)
cacheDigest	cache of the digest for one or more of the parameters
showMessage	show message that data is being retrieved from cache
forceRecalc	force the recalculation of the values
addToHash	something to add to the filename generation

Value

the result of fun(...)

Functions

- `.runCache(`function`)`: accepts function as first argument and save cache

Examples

```
# [optional] save cache in a temporary directory
#
glmSparseNet:::.baseDir(tempdir())
glmSparseNet:::.runCache(c, 1, 2, 3, 4)
#
# next three should use the same cache
#  note, the middle call should be a little faster as digest is not
#  calculated
#   for the first argument
glmSparseNet:::.runCache(c, 1, 2, 3, 4)
glmSparseNet:::.runCache(c, a = 1, 2, c = 3, 4)

# Using a local folder
# glmSparseNet:::.runCache(c, 1, 2, 3, 4, baseDir = "runcache")
```

.saveRunCache *Saving the cache*

Description

Saving the cache

Usage

```
.saveRunCache(result, path, compression, showMessage)
```

Arguments

result	main result to save
path	path to the file to save
compression	compression method to be used
showMessage	TRUE to show messages, FALSE otherwise

Value

result of save operation

Examples

```
glmSparseNet:::.saveRunCache(
    35, file.path(tempdir(), "save_run_cache.Rdata"), FALSE, TRUE
)
```

.showMessage *Show messages option in .runCache*

Description

Show messages option in .runCache

Usage

```
.showMessage(showMessage = NULL)
```

Arguments

showMessage	boolean indicating to show messages or not

Value

the show.message option

Examples

```
glmSparseNet:::.showMessage(FALSE)
```

`.tempdirCache` *Temporary directory for runCache*

Description

Temporary directory for runCache

Usage

```
.tempdirCache()
```

Value

a path to a temporary directory used by runCache

`.writeReadme` *Write a file in run-cache directory to explain the origin*

Description

Write a file in run-cache directory to explain the origin

Usage

```
.writeReadme(baseDir)
```

Arguments

`baseDir`	directory where to build this file

Value

the path to the file it has written

Examples

```
glmSparseNet:::.writeReadme(tempdir())
```

balancedCvFolds *Create balanced folds for cross validation using stratified sampling*

Description

Create balanced folds for cross validation using stratified sampling

Usage

```
balancedCvFolds(..., nfolds = 10)

# deprecated, please use balancedCvFolds()
balanced.cv.folds(..., nfolds = 10)
```

Arguments

...	vectors representing data
nfolds	number of folds to be created

Value

list with given input, nfolds and result. The result is a list matching the input with foldid attributed to each position.

Examples

```
balancedCvFolds(seq(10), seq(11, 15), nfolds = 2)

# will give a warning
balancedCvFolds(seq(10), seq(11, 13), nfolds = 10)

balancedCvFolds(seq(100), seq(101, 133), nfolds = 10)
```

buildLambda *Auxiliary function to generate suitable lambda parameters*

Description

Auxiliary function to generate suitable lambda parameters

Usage

```
buildLambda(
  lambdaLargest = NULL,
  xdata = NULL,
  ydata = NULL,
  family = NULL,
  ordersOfMagnitudeSmaller = 3,
  lambdaPerOrderMagnitude = 150,
  lambda.largest = deprecated(),
  orders.of.magnitude.smaller = deprecated(),
  lambda.per.order.magnitude = deprecated()
)
```

Arguments

`lambdaLargest`	numeric value for largest number of lambda to consider (usually with a target of 1 selected variable)
`xdata`	X parameter for glmnet function
`ydata`	Y parameter for glmnet function
`family`	family parameter to glmnet function
`ordersOfMagnitudeSmaller`	minimum value for lambda (lambda.largest / 10^orders.of.magnitude.smaller)
`lambdaPerOrderMagnitude`	how many lambdas to create for each order of magnitude
`lambda.largest`	**[Deprecated]**
`orders.of.magnitude.smaller`	**[Deprecated]**
`lambda.per.order.magnitude`	**[Deprecated]**

Value

a numeric vector with suitable lambdas

Examples

```
buildLambda(5.4)
```

`buildStringNetwork` *Build gene network from peptide ids*

Description

This can reduce the dimension of the original network, as there may not be a mapping between peptide and gene id

Usage

```
buildStringNetwork(
  stringTbl,
  useNames = c("protein", "ensembl", "external"),
  string.tbl = deprecated(),
  use.names = deprecated()
)
```

Arguments

`stringTbl`	`data.frame` or `tibble` with colnames and rownames as ensembl peptide id *(same order)*.
`useNames`	`character(1)` that defaults to use protein names _('protein'), other options are 'ensembl' for ensembl gene id or 'external' for external gene names.
`string.tbl`	**[Deprecated]**
`use.names`	**[Deprecated]**

Value

a new matrix with gene ids instead of peptide ids. The size of matrix can be different as there may not be a mapping or a peptide mapping can have multiple genes.

See Also

`stringDBhomoSapiens()`

Examples

```
interactions <- stringDBhomoSapiens(scoreThreshold = 100)
string_network <- buildStringNetwork(interactions)

# number of edges
sum(string_network != 0)
```

cv.glmDegree *Calculate cross validating GLM model with network-based regularization*

Description

network parameter accepts:

Usage

```
cv.glmDegree(
  xdata,
  ydata,
  network,
  options = networkOptions(),
  experiment = NULL,
  network.options = deprecated(),
  experiment.name = deprecated(),
  ...
)

cv.glmHub(
  xdata,
  ydata,
  network,
  options = networkOptions(),
  experiment = NULL,
  network.options = deprecated(),
  experiment.name = deprecated(),
  ...
)

cv.glmOrphan(
  xdata,
  ydata,
  network,
  options = networkOptions(),
  experiment = NULL,
  network.options = deprecated(),
  experiment.name = deprecated(),
  ...
)

cv.glmSparseNet(
  xdata,
  ydata,
  network,
```

```
  options = networkOptions(),
  experiment = NULL,
  network.options = deprecated(),
  experiment.name = deprecated(),
  ...
)
```

Arguments

xdata	input data, can be a matrix or MultiAssayExperiment.
ydata	response data compatible with glmnet.
network	type of network, see below.
options	options to calculate network.
experiment	name of experiment to use as input in MultiAssayExperiment object (only if xdata is an object of this class).
network.options	**[Deprecated]**
experiment.name	**[Deprecated]**
...	parameters that `glmnet::cv.glmnet()` accepts.

Details

- string to calculate network based on data (correlation, covariance)
- matrix representing the network
- vector with already calculated penalty weights (can also be used directly glmnet)

Value

an object just as `cv.glmnet`

Functions

- `cv.glmDegree()`: penalizes nodes with small degree *(inversion penalization* `h(x) = 1 / x`*)*.
- `cv.glmHub()`: penalizes nodes with small degree *(normalized heuristic that promotes nodes with many edges)*.
- `cv.glmOrphan()`: penalizes nodes with high degree *(normalized heuristic that promotes nodes with few edges)*.

See Also

Model with the same penalizations `glmSparseNet()`.

Examples

```
# Degree penalization

xdata <- matrix(rnorm(100), ncol = 5)
cv.glmDegree(
    xdata,
    rnorm(nrow(xdata)),
    "correlation",
    family = "gaussian",
    nfolds = 5,
    options = networkOptions(minDegree = .2)
)
# Hub penalization

xdata <- matrix(rnorm(100), ncol = 5)
cv.glmHub(
    xdata,
    rnorm(nrow(xdata)),
    "correlation",
    family = "gaussian",
    nfolds = 5,
    options = networkOptions(minDegree = .2)
)
# Orphan penalization

xdata <- matrix(rnorm(100), ncol = 5)
cv.glmOrphan(
    xdata,
    rnorm(nrow(xdata)),
    "correlation",
    family = "gaussian",
    nfolds = 5,
    options = networkOptions(minDegree = .2)
)

# Gaussian model
xdata <- matrix(rnorm(500), ncol = 5)
cv.glmSparseNet(
    xdata, rnorm(nrow(xdata)), "correlation",
    family = "gaussian"
)
cv.glmSparseNet(
    xdata, rnorm(nrow(xdata)), "covariance",
    family = "gaussian"
)


#
#
# Using MultiAssayExperiment with survival model
library(MultiAssayExperiment)
data("miniACC", package = "MultiAssayExperiment")
```

```
xdata <- miniACC

#
# build valid data with days of last follow up or to event
event.ix <- which(!is.na(xdata$days_to_death))
cens.ix <- which(!is.na(xdata$days_to_last_followup))
xdata$surv_event_time <- array(NA, nrow(colData(xdata)))
xdata$surv_event_time[event.ix] <- xdata$days_to_death[event.ix]
xdata$surv_event_time[cens.ix] <- xdata$days_to_last_followup[cens.ix]

#
# Keep only valid individuals
valid.ix <- as.vector(!is.na(xdata$surv_event_time) &
    !is.na(xdata$vital_status) &
    xdata$surv_event_time > 0)
xdata.valid <- xdata[, rownames(colData(xdata))[valid.ix]]
ydata.valid <- colData(xdata.valid)[, c("surv_event_time", "vital_status")]
colnames(ydata.valid) <- c("time", "status")

#
cv.glmSparseNet(
    xdata.valid,
    ydata.valid,
    nfolds     = 5,
    family     = "cox",
    network    = "correlation",
    experiment = "RNASeq2GeneNorm"
)
```

---

degreeCor *Calculate the degree of the correlation network based on xdata*

Description

Calculate the degree of the correlation network based on xdata

Usage

```
degreeCor(
  xdata,
  cutoff = 0,
  considerUnweighted = FALSE,
  forceRecalcDegree = FALSE,
  forceRecalcNetwork = FALSE,
  nCores = 1,
  ...,
  consider.unweighted = deprecated(),
```

```
  force.recalc.degree = deprecated(),
  force.recalc.network = deprecated(),
  n.cores = deprecated()
)
```

Arguments

xdata	calculate correlation matrix on each column.
cutoff	positive value that determines a cutoff value.
considerUnweighted	consider all edges as 1 if they are greater than 0.
forceRecalcDegree	force recalculation of penalty weights (but not the network), instead of going to cache.
forceRecalcNetwork	force recalculation of network and penalty weights, instead of going to cache.
nCores	number of cores to be used.
...	extra parameters for cor function.
consider.unweighted	**[Deprecated]**
force.recalc.degree	**[Deprecated]**
force.recalc.network	**[Deprecated]**
n.cores	**[Deprecated]**

Value

a vector of the degrees.

Examples

```
n.col <- 6
xdata <- matrix(rnorm(n.col * 4), ncol = n.col)
degreeCor(xdata)
degreeCor(xdata, cutoff = .5)
degreeCor(xdata, cutoff = .5, considerUnweighted = TRUE)
```

---

degreeCov *Calculate the degree of the covariance network based on xdata*

Description

Calculate the degree of the covariance network based on xdata

Usage

```
degreeCov(
  xdata,
  cutoff = 0,
  considerUnweighted = FALSE,
  forceRecalcDegree = FALSE,
  forceRecalcNetwork = FALSE,
  nCores = 1,
  ...,
  consider.unweighted = deprecated(),
  force.recalc.degree = deprecated(),
  force.recalc.network = deprecated(),
  n.cores = deprecated()
)
```

Arguments

xdata	calculate correlation matrix on each column.
cutoff	positive value that determines a cutoff value.
considerUnweighted	consider all edges as 1 if they are greater than 0.
forceRecalcDegree	force recalculation of penalty weights (but not the network), instead of going to cache.
forceRecalcNetwork	force recalculation of network and penalty weights, instead of going to cache.
nCores	number of cores to be used.
...	extra parameters for cov function.
consider.unweighted	**[Deprecated]**
force.recalc.degree	**[Deprecated]**
force.recalc.network	**[Deprecated]**
n.cores	**[Deprecated]**

Value

a vector of the degrees

Examples

```
n.col <- 6
xdata <- matrix(rnorm(n.col * 4), ncol = n.col)
degreeCov(xdata)
degreeCov(xdata, cutoff = .5)
degreeCov(xdata, cutoff = .5, considerUnweighted = TRUE)
```

downloadFileLocal *Download files to local temporary path*

Description

In case of new call it uses the temporary cache instead of downloading again.

Usage

```
downloadFileLocal(urlStr, oD = tempdir())
```

Arguments

urlStr	url of file to download
oD	temporary directory to store file

Details

Inspired by STRINGdb Bioconductor package, but using curl as file may be too big to handle.

Value

path to file

Examples

```
glmSparseNet:::downloadFileLocal(
    "https://string-db.org/api/tsv-no-header/version"
)
```

ensemblGeneNames *Retrieve ensembl gene names from biomaRt*

Description

Retrieve ensembl gene names from biomaRt

Usage

```
ensemblGeneNames(
  geneId,
  useCache = TRUE,
  verbose = FALSE,
  gene.id = deprecated(),
  use.cache = deprecated()
)
```

Arguments

geneId	character vector with gene names
useCache	Boolean indicating if biomaRt cache should be used
verbose	When using biomaRt in webservice mode and setting verbose to TRUE, the XML query to the webservice will be printed.
gene.id	**[Deprecated]**
use.cache	**[Deprecated]**

Value

a dataframe with external gene names, ensembl_id

Examples

```
ensemblGeneNames(c("MOB1A", "RFLNB", "SPIC", "TP53"))
```

---

geneNames *Retrieve gene names from biomaRt*

Description

Retrieve gene names from biomaRt

Usage

```
geneNames(
  ensemblGenes,
  useCache = TRUE,
  verbose = FALSE,
  ensembl.genes = deprecated(),
  use.cache = deprecated()
)
```

Arguments

ensemblGenes	character vector with gene names in ensembl_id format
useCache	Boolean indicating if biomaRt cache should be used
verbose	When using biomaRt in webservice mode and setting verbose to TRUE, the XML query to the webservice will be printed.
ensembl.genes	**[Deprecated]**
use.cache	**[Deprecated]**

Value

a dataframe with external gene names, ensembl_id

Examples

```
geneNames(c("ENSG00000114978", "ENSG00000166211", "ENSG00000183688"))
```

`glmSparseNet` *Calculate GLM model with network-based regularization*

Description

network parameter accepts:

- string to calculate network based on data (correlation, covariance)
- matrix representing the network
- vector with already calculated penalty weights (can also be used directly with glmnet)

Usage

```
glmSparseNet(
  xdata,
  ydata,
  network,
  options = networkOptions(),
  experiment = NULL,
  network.options = deprecated(),
  experiment.name = deprecated(),
  ...
)

glmDegree(
  xdata,
  ydata,
  network,
  options = networkOptions(),
  experiment = NULL,
  network.options = deprecated(),
  experiment.name = deprecated(),
  ...
)

glmHub(
  xdata,
  ydata,
  network,
  options = networkOptions(),
  experiment = NULL,
  network.options = deprecated(),
  experiment.name = deprecated(),
```

```
  ...
)

glmOrphan(
  xdata,
  ydata,
  network,
  options = networkOptions(),
  experiment = NULL,
  network.options = deprecated(),
  experiment.name = deprecated(),
  ...
)
```

Arguments

`xdata`	input data, can be a matrix or MultiAssayExperiment.
`ydata`	response data compatible with glmnet.
`network`	type of network, see below.
`options`	options to calculate network.
`experiment`	name of experiment to use as input in MultiAssayExperiment object (only if xdata is an object of this class).
`network.options`	**[Deprecated]**
`experiment.name`	**[Deprecated]**
`...`	parameters that `glmnet::glmnet()` accepts.

Value

an object just as glmnet

Functions

- `glmDegree()`: penalizes nodes with small degree *(inversion penalization* `h(x) = 1 / x`*)*.
- `glmHub()`: Penalizes nodes with small degree *(normalized heuristic that promotes nodes with many edges)*.
- `glmOrphan()`: Penalizes nodes with high degree *(normalized heuristic that promotes nodes with few edges)*.

See Also

Cross-validation functions `cv.glmSparseNet()`.

Examples

```
xdata <- matrix(rnorm(100), ncol = 20)
glmSparseNet(xdata, rnorm(nrow(xdata)), "correlation", family = "gaussian")
glmSparseNet(xdata, rnorm(nrow(xdata)), "covariance", family = "gaussian")


#
#
# Using MultiAssayExperiment
# load data
library(MultiAssayExperiment)
data("miniACC", package = "MultiAssayExperiment")

xdata <- miniACC
# TODO aking out x individuals missing values
# build valid data with days of last follow up or to event
event.ix <- which(!is.na(xdata$days_to_death))
cens.ix <- which(!is.na(xdata$days_to_last_followup))

xdata$surv_event_time <- array(NA, nrow(colData(xdata)))
xdata$surv_event_time[event.ix] <- xdata$days_to_death[event.ix]
xdata$surv_event_time[cens.ix] <- xdata$days_to_last_followup[cens.ix]

# Keep only valid individuals
valid.ix <- as.vector(!is.na(xdata$surv_event_time) &
    !is.na(xdata$vital_status) &
    xdata$surv_event_time > 0)
xdata.valid <- xdata[, rownames(colData(xdata))[valid.ix]]
ydata.valid <- colData(xdata.valid)[, c("surv_event_time", "vital_status")]
colnames(ydata.valid) <- c("time", "status")

glmSparseNet(
    xdata.valid,
    ydata.valid,
    family = "cox",
    network = "correlation",
    experiment = "RNASeq2GeneNorm"
)

# Degree penalization

xdata <- matrix(rnorm(100), ncol = 5)
glmDegree(
    xdata,
    rnorm(nrow(xdata)),
    "correlation",
    family = "gaussian",
    options = networkOptions(minDegree = .2)
)
xdata <- matrix(rnorm(100), ncol = 5)
glmHub(
    xdata,
```

```
    rnorm(nrow(xdata)),
    "correlation",
    family = "gaussian",
    options = networkOptions(minDegree = .2)
)
# Orphan penalization

xdata <- matrix(rnorm(100), ncol = 5)
glmOrphan(
    xdata,
    rnorm(nrow(xdata)),
    "correlation",
    family = "gaussian",
    options = networkOptions(minDegree = .2)
)
```

`hallmarks` *Retrieve hallmarks of cancer count for genes*

Description

[Defunct] The API has been removed and this function is no longer available.

Usage

```
hallmarks(
  genes,
  metric = "count",
  hierarchy = "full",
  generate.plot = TRUE,
  show.message = FALSE
)
```

Arguments

`genes`	gene names
`metric`	see below
`hierarchy`	see below
`generate.plot`	flag to indicate if return object has a ggplot2 object
`show.message`	flag to indicate if run_cache method shows messages

Value

data.frame with choosen metric and hierarchy It also returns a vector with genes that do not have any hallmarks.

See http://chat.lionproject.net/api for more details on the metric and hallmarks parameters

To standardize the colors in the gradient you can use scale_fill_gradientn(limits=c(0,1), colours=topo.colors(3)) to limit between 0 and 1 for cprob and -1 and 1 for npmi

`heuristicScale` *Heuristic function to use in high dimensions*

Description

Heuristic function to use in high dimensions

Usage

```
heuristicScale(
  x,
  subExp10 = -1,
  expMult = -1,
  subExp = -1,
  sub.exp10 = deprecated(),
  exp.mult = deprecated(),
  sub.exp = deprecated()
)
```

Arguments

`x`	vector of values to scale
`subExp10`	value to subtract to base 10 exponential, for example: `10^0 - subExp10 = 1 - subExp10`
`expMult`	parameter to multiply exponential, i.e. to have a negative exponential or positive
`subExp`	value to subtract for exponentional, for example if x = 0, `exp(0) - sub.exp = 1 - sub.exp`
`sub.exp10`	**[Deprecated]**
`exp.mult`	**[Deprecated]**
`sub.exp`	**[Deprecated]**

Value

a vector of scaled values

Examples

```
heuristicScale(rnorm(1:10))
```

hubHeuristic *Heuristic function to penalize nodes with low degree*

Description

Heuristic function to penalize nodes with low degree

Usage

```
hubHeuristic(x)
```

Arguments

x	single value of vector

Value

transformed

Examples

```
hubHeuristic(rnorm(1:10))
```

myColors *Custom pallete of colors*

Description

Custom pallete of colors

Usage

```
myColors(ix = NULL)

# deprecated, please use myColors()
my.colors(ix = NULL)
```

Arguments

ix	index for a color

Value

a color

Examples

```
myColors()
myColors(5)
```

mySymbols *Custom pallete of symbols in plots*

Description

Custom pallete of symbols in plots

Usage

```
mySymbols(ix = NULL)

# deprecated, please use mySymbols()
my.symbols(ix = NULL)
```

Arguments

ix	index for symbol

Value

a symbol

Examples

```
mySymbols()
mySymbols(2)
```

networkCorParallel *Calculates the correlation network*

Description

Calculates the correlation network

Usage

```
networkCorParallel(
  xdata,
  buildOutput = "matrix",
  nCores = 1,
  forceRecalcNetwork = FALSE,
  showMessage = FALSE,
  ...,
  build.output = deprecated(),
  n.cores = deprecated(),
  force.recalc.network = deprecated(),
  show.message = deprecated()
)
```

Arguments

xdata	base data to calculate network
buildOutput	if output returns a 'matrix', 'vector' of the upper triu without the diagonal or NULL with any other argument
nCores	number of cores to be used
forceRecalcNetwork	force recalculation, instead of going to cache
showMessage	shows cache operation messages
...	extra parameters for fun
build.output	lifecycle::badge("deprecated") without the diagonal or NULL with any other argument
n.cores	lifecycle::badge("deprecated")
force.recalc.network	lifecycle::badge("deprecated")
show.message	lifecycle::badge("deprecated")

Value

depends on build.output parameter

Examples

```
n_col <- 6
xdata <- matrix(rnorm(n_col * 4), ncol = n_col)
networkCorParallel(xdata)
```

---

networkCovParallel *Calculates the covariance network*

Description

Calculates the covariance network

Usage

```
networkCovParallel(
  xdata,
  buildOutput = "matrix",
  nCores = 1,
  forceRecalcNetwork = FALSE,
  showMessage = FALSE,
  ...,
  build.output = deprecated(),
  n.cores = deprecated(),
  force.recalc.network = deprecated(),
  show.message = deprecated()
)
```

Arguments

xdata	base data to calculate network
buildOutput	if output returns a 'matrix', 'vector' of the upper triu without the diagonal or NULL with any other argument
nCores	number of cores to be used
forceRecalcNetwork	force recalculation, instead of going to cache
showMessage	shows cache operation messages
...	extra parameters for fun
build.output	lifecycle::badge("deprecated") without the diagonal or NULL with any other argument
n.cores	lifecycle::badge("deprecated")
force.recalc.network	lifecycle::badge("deprecated")
show.message	lifecycle::badge("deprecated")

Value

depends on build.output parameter

Examples

```
n.col <- 6
xdata <- matrix(rnorm(n.col * 4), ncol = n.col)
networkCovParallel(xdata)
```

---

networkOptions *Setup network options*

Description

Setup network options, such as using weighted or unweighted degree, which centrality measure to use

Usage

```
networkOptions(
  method = "pearson",
  unweighted = TRUE,
  cutoff = 0,
  centrality = "degree",
  minDegree = 0,
  nCores = 1,
  transFun = function(x) x,
  min.degree = deprecated(),
```

```
  n.cores = deprecated(),
  trans.fun = deprecated()
)
```

Arguments

method	in case of correlation and covariance, which method to use.
unweighted	calculate degree using unweighted network.
cutoff	cuttoff value in network edges to trim the network.
centrality	centrality measure to use, currently only supports degree.
minDegree	minimum value that individual penalty weight can take.
nCores	number of cores to use, default to 1.
transFun	See details below.
min.degree	**[Deprecated]**
n.cores	**[Deprecated]**
trans.fun	**[Deprecated]**
	The transFun argument takes a function definition that will apply a transformation to the penalty vector calculated from the degree. This transformation allows to change how the penalty is applied.

Value

a list of options

See Also

glmOrphan() and glmDegree()

Examples

```
networkOptions(unweighted = FALSE)
```

---

orphanHeuristic *Heuristic function to penalize nodes with high degree*

Description

Heuristic function to penalize nodes with high degree

Usage

```
orphanHeuristic(x)
```

Arguments

x	single value of vector

Value

transformed

Examples

```
orphanHeuristic(rnorm(1:10))
```

`protein2EnsemblGeneNames` *Retrieve ensembl gene ids from proteins*

Description

Retrieve ensembl gene ids from proteins

Usage

```
protein2EnsemblGeneNames(
  ensemblProteins,
  useCache = TRUE,
  verbose = FALSE,
  ensembl.proteins = deprecated(),
  use.cache = deprecated()
)
```

Arguments

`ensemblProteins`	character vector with gene names in ensembl_peptide_id format
`useCache`	Boolean indicating if biomaRt cache should be used
`verbose`	When using biomaRt in webservice mode and setting verbose to TRUE, the XML query to the webservice will be printed.
`ensembl.proteins`	**[Deprecated]**
`use.cache`	**[Deprecated]**

Value

a dataframe with external gene names, ensembl_peptide_id

Examples

```
protein2EnsemblGeneNames(c(
    "ENSP00000235382",
    "ENSP00000233944",
    "ENSP00000216911"
))
```

`separate2GroupsCox` *Separate data in High and Low risk groups (based on Cox model)*

Description

Draws multiple kaplan meyer survival curves (or just 1) and calculates logrank test

Usage

```
separate2GroupsCox(
  chosenBetas,
  xdata,
  ydata,
  probs = c(0.5, 0.5),
  noPlot = FALSE,
  plotTitle = "SurvivalCurves",
  xlim = NULL,
  ylim = NULL,
  expandYZero = FALSE,
  legendOutside = FALSE,
  stopWhenOverlap = TRUE,
  ...,
  chosen.btas = deprecated(),
  no.plot = deprecated(),
  plot.title = deprecated(),
  expand.yzero = deprecated(),
  legend.outside = deprecated(),
  stop.when.overlap = deprecated()
)
```

Arguments

`chosenBetas`	list of testing coefficients to calculate prognostic indexes, for example `list(Age = some_vector)`.
`xdata`	n x m matrix with n observations and m variables.
`ydata`	Survival object.
`probs`	How to separate high and low risk patients `50%-50%` is the default, but for top and bottom `40%` -> `c(.4,.6)`.
`noPlot`	Only calculate p-value and do not generate survival curve plot.
`plotTitle`	Name of file if.
`xlim`	Optional argument to limit the x-axis view.
`ylim`	Optional argument to limit the y-axis view.
`expandYZero`	expand to y = 0.
`legendOutside`	If TRUE legend will be outside plot, otherwise inside.

`stopWhenOverlap`
: when probs vector allows for overlapping of samples in both groups, then stop.

`...`
: additional parameters to survminer::ggsurvplot

`chosen.btas`
: **[Deprecated]**

`no.plot`
: **[Deprecated]**

`plot.title`
: **[Deprecated]**

`expand.yzero`
: **[Deprecated]**

`legend.outside`
: **[Deprecated]**

`stop.when.overlap`
: **[Deprecated]**
Otherwise it will calculate with duplicate samples, i.e. simply adding them to xdata and ydata (in a different group).

Value

object with logrank test and kaplan-meier survival plot

A list with plot, p-value and kaplan-meier object. The plot was drawn from survminer::ggsurvplot with only the palette, data and fit arguments being defined and keeping all other defaults that can be customized as additional parameters to this function.

See Also

`survminer::ggsurvplot()`

Examples

```
xdata <- survival::ovarian[, c("age", "resid.ds")]
ydata <- data.frame(
    time = survival::ovarian$futime,
    status = survival::ovarian$fustat
)
separate2GroupsCox(c(age = 1, 0), xdata, ydata)
separate2GroupsCox(c(age = 1, 0.5), xdata, ydata)
separate2GroupsCox(
    c(age = 1), c(1, 0, 1, 0, 1, 0),
    data.frame(time = runif(6), status = rbinom(6, 1, .5))
)
separate2GroupsCox(list(
    aa = c(age = 1, 0.5),
    bb = c(age = 0, 1.5)
), xdata, ydata)
```

string.network.700.cache

Cache of protein-protein network, as it takes some time to retrieve and process this will facilitate the vignette building

Description

It was filtered with combined_scores and individual scores below 700 without text-based scores

Usage

```
data('string.network.700.cache', package = 'glmSparseNet')
```

Format

An object of class `dgCMatrix` with 11033 rows and 11033 columns.

References

https://string-db.org/

stringDBhomoSapiens *Download protein-protein interactions from STRING DB*

Description

Download protein-protein interactions from STRING DB

Usage

```
stringDBhomoSapiens(
  version = "11.0",
  scoreThreshold = 0,
  removeText = TRUE,
  score_threshold = deprecated(),
  remove.text = deprecated()
)
```

Arguments

`version`	version of the database to use
`scoreThreshold`	remove scores below threshold
`removeText`	remove text mining-based scores
`score_threshold`	**[Deprecated]**
`remove.text`	**[Deprecated]**

Value

a data.frame with rows representing an interaction between two proteins, and columns the count of scores above the given score_threshold

Examples

```
stringDBhomoSapiens(scoreThreshold = 800)
```

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