

Package ‘glmSparseNet’

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Type Package

Title Network Centrality Metrics for Elastic-Net Regularized Models

Version 1.18.0

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``.

License GPL-3

Encoding UTF-8

LazyData false

NeedsCompilation no

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Depends R (>= 4.1), Matrix, MultiAssayExperiment, glmnet

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R topics documented:

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| | |
|---------------------------|----------------------------------------|
| <code>.calcPenalty</code> | <i>Calculate penalty based on data</i> |
|---------------------------|----------------------------------------|

Description

Internal method to calculate the network using data-dependant methods

Usage

```
.calcPenalty(xdata, penalty.type, network.options = networkOptions())
```

Arguments

| | |
|------------------------------|---------------------|
| <code>xdata</code> | input data |
| <code>penalty.type</code> | which method to use |
| <code>network.options</code> | 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')
```

.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 symmetric and without any connection the node and itself.

Usage

```
.degreeGeneric(
  fun = stats::cor,
  fun.prefix = "operator",
  xdata,
  cutoff = 0,
  consider.unweighted = FALSE,
  chunks = 1000,
  force.recalc.degree = FALSE,
  force.recalc.network = FALSE,
  n.cores = 1,
  ...
)
```

Arguments

| | |
|---------------------|-------------------------------------------------------------------------------------------------|
| fun | function that will calculate the edge weight between 2 nodes |
| fun.prefix | 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 |
| consider.unweighted | consider all edges as 1 if they are greater than 0 |
| chunks | calculate function at batches of this value (default is 1000) |
| force.recalc.degree | force recalculation of penalty weights (but not the network), instead of going to cache |

force.recalc.network force recalculation of network and penalty weights, instead of going to cache
n.cores number of cores to be used
... extra parameters for fun

Value

a vector of the degrees

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

Description

Calculate GLM model with network-based regularization

Usage

```
.glmSparseNetPrivate(
  fun,
  xdata,
  ydata,
  network,
  experiment.name = NULL,
  network.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.name when xdata is a MultiAssayExperiment object this parameter is required
network.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,  
  fun.prefix,  
  xdata,  
  build.output = "matrix",  
  n.cores = 1,  
  force.recalc.network = FALSE,  
  show.message = FALSE,  
  ...  
)
```

Arguments

| | |
|-----------------------------------|---------------------------------------------------------------------------------------------------------------|
| <code>fun</code> | function that will calculate the edge weight between 2 nodes |
| <code>fun.prefix</code> | used to store low-level information on network as it can become to large to be stored in memory |
| <code>xdata</code> | base data to calculate network |
| <code>build.output</code> | if output returns a 'matrix', 'vector' of the upper triu without the diagonal or NULL with any other argument |
| <code>n.cores</code> | number of cores to be used |
| <code>force.recalc.network</code> | force recalculation, instead of going to cache |
| <code>show.message</code> | shows cache operation messages |
| <code>...</code> | extra parameters for fun |

Value

depends on `build.output` parameter

| | |
|----------------|---------------------------------------------------------------------------------|
| .networkWorker | <i>Worker to calculate edge weight for each pair of ix.i node and following</i> |
|----------------|---------------------------------------------------------------------------------|

Description

Note that it assumes it does not calculate for index below and equal to ix.i

Usage

```
.networkWorker(fun, xdata, ix.i, ...)
```

Arguments

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

Value

a vector with size 'ncol(xdata) - ix.i'

| | |
|-------------------|---------------------------------------------------|
| balanced.cv.folds | <i>Create balanced folds for cross validation</i> |
|-------------------|---------------------------------------------------|

Description

Create balanced folds for cross validation

Usage

```
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

```
glmSparseNet::balanced.cv.folds(seq(10), seq(11, 15), nfolds = 2)
# will give a warning
glmSparseNet::balanced.cv.folds(seq(10), seq(11, 13), nfolds = 10)
glmSparseNet::balanced.cv.folds(seq(100), seq(101, 133), nfolds = 10)
```

| | |
|----------|--------------------------------------|
| base.dir | <i>change base.dir for run.cache</i> |
|----------|--------------------------------------|

Description

change base.dir for run.cache

Usage

```
base.dir(path = NULL)
```

Arguments

path to base directory where cache is saved

Value

the new path

Examples

```
glmSparseNet::base.dir('/tmp/cache')
```

| | |
|--------------|--------------------------------------------------------|
| biomart.load | <i>Common call to biomaRt to avoid repetitive code</i> |
|--------------|--------------------------------------------------------|

Description

Common call to biomaRt to avoid repetitive code

Usage

```
biomart.load(attributes, filters, values, use.cache, verbose)
```


Arguments

| | |
|------------|----------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| attributes | Attributes you want to retrieve. A possible list of attributes can be retrieved using the function <code>biomaRt::listAttributes</code> . |
| filters | Filters (one or more) that should be used in the query. A possible list of filters can be retrieved using the function <code>biomaRt::listFilters</code> . |
| 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 |
| use.cache | Boolean indicating if <code>biomaRt</code> cache should be used |
| verbose | When using <code>biomaRt</code> in webservice mode and setting <code>verbose</code> to <code>TRUE</code> , 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::biomart.load(  
  attributes = c("external_gene_name", "ensembl_gene_id"),  
  filters = "external_gene_name",  
  values = c('MOB1A', 'RFLNB', 'SPIC', 'TP53'),  
  use.cache = TRUE,  
  verbose = FALSE  
)
```

`build.function.digest` *Build digest of function from the actual code*

Description

Build digest of function from the actual code

Usage

```
build.function.digest(fun)
```

Arguments

fun function call name

Value

a digest

Examples

```
glmSparseNet:::build.function.digest(sum)
glmSparseNet:::build.function.digest(c)
```

buildLambda

Auxiliary function to generate suitable lambda parameters

Description

Auxiliary function to generate suitable lambda parameters

Usage

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

Arguments

lambda.largest 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

orders.of.magnitude.smaller
 minimum value for lambda ($\text{lambda.largest} / 10^{\text{orders.of.magnitude.smaller}}$)

lambda.per.order.magnitude
 how many lambdas to create for each order of magnitude

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(string.tbl, use.names = "protein")
```

Arguments

| | |
|-------------------------|------------------------------------------------------------------------------------------------------------------------------------|
| <code>string.tbl</code> | matrix with colnames and rownames as ensembl peptide id (same order) |
| <code>use.names</code> | default is to use protein names ('protein'), other options are 'ensembl' for ensembl gene id or 'external' for external gene names |

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

```
all.interactions.700 <- stringDBhomoSapiens(score_threshold = 700)
string.network       <- buildStringNetwork(all.interactions.700,
                                           use.names = 'external')

# number of edges
sum(string.network != 0)
```

| | |
|-------------------|-----------------------------------------------|
| cache.compression | <i>change cache.compression for run.cache</i> |
|-------------------|-----------------------------------------------|

Description

change cache.compression for run.cache

Usage

```
cache.compression(compression = NULL)
```

Arguments

compression see compression parameter in save function

Value

the new compression

Examples

```
glmSparseNet:::cache.compression('bzip2')
```

```
calculate.combined.score
```

Calculate combined score for STRINGdb interactions

Description

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

Usage

```
calculate.combined.score(all.interactions, score_threshold, remove.text)
```

Arguments

all.interactions
 table with score of all interactions

score_threshold
 threshold to keep interactions

remove.text 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

| | |
|------------------|----------------------------------------------------|
| calculate.result | <i>Calculate/load result and save if necessary</i> |
|------------------|----------------------------------------------------|

Description

This is where the actual work is done

Usage

```
calculate.result(path, compression, force.recalc, show.message, fun, ...)
```

Arguments

| | |
|--------------|------------------------------|
| path | path to save cache |
| compression | compression used in save |
| force.recalc | force to recalculate cache |
| show.message | boolean to show messages |
| fun | function to be called |
| ... | arguments to said function , |

Value

result of fun(...)

Examples

```
glmSparseNet::calculate.result(
  file.path(tempdir(), 'calculate.result.Rdata'),
  'gzip',
  FALSE,
  TRUE,
  sum,
  1, 2, 3
)
```

```
create.directory.for.cache
```

Create directories for cache

Description

Create directories for cache

Usage

```
create.directory.for.cache(base.dir, parent.path)
```

Arguments

| | |
|-------------|------------------------------------------------------------------------------------------------------------------------------------|
| base.dir | tentative base dir to create. |
| parent.path | 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 base.dir and parent.dir

Examples

```
glmSparseNet::create.directory.for.cache(tempdir(), 'abcd')
```

```
glmSparseNet::create.directory.for.cache(  
  file.path(getwd(), 'run-cache'), 'abcd'  
)
```

```
curl.workaround
```

Workaround for bug with curl when fetching specific ensembl mirror

Description

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

Usage

```
curl.workaround(expr)
```

Arguments

| | |
|------|------------|
| expr | expression |
|------|------------|

Value

result of expression

Examples

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

cv.glmDegree

GLMNET cross-validation model penalizing nodes with small degree

Description

This function overrides the ‘trans.fun’ options in ‘network.options’ with the inverse of a degree described in Verissimo et al. (2015) that penalizes nodes with small degree.

Usage

```
cv.glmDegree(xdata, ydata, network, network.options = networkOptions(), ...)
```

Arguments

| | |
|-----------------|-----------------------------------------------------|
| xdata | input data, can be a matrix or MultiAssayExperiment |
| ydata | response data compatible with glmnet |
| network | type of network, see below |
| network.options | options to calculate network |
| ... | parameters that glmnet accepts |

Value

see cv.glmSparseNet

See Also

glmNetSparse

Examples

```
xdata <- matrix(rnorm(100), ncol = 5)
cv.glmDegree(xdata, rnorm(nrow(xdata)), 'correlation',
  family = 'gaussian',
  nfolds = 5,
  network.options = networkOptions(min.degree = .2))
```

`cv.glmHub`*GLMNET cross-validation model penalizing nodes with small degree*

Description

This function overrides the ‘trans.fun’ options in ‘network.options’ with an heuristic described in Verissimo et al. that penalizes nodes with small degree.

Usage

```
cv.glmHub(xdata, ydata, network, network.options = networkOptions(), ...)
```

Arguments

| | |
|------------------------------|-----------------------------------------------------|
| <code>xdata</code> | input data, can be a matrix or MultiAssayExperiment |
| <code>ydata</code> | response data compatible with glmnet |
| <code>network</code> | type of network, see below |
| <code>network.options</code> | options to calculate network |
| <code>...</code> | parameters that glmnet accepts |

Value

see `cv.glmSparseNet`

See Also

`glmNetSparse`

Examples

```
xdata <- matrix(rnorm(100), ncol = 5)
cv.glmHub(xdata, rnorm(nrow(xdata)), 'correlation',
          family = 'gaussian',
          nfolds = 5,
          network.options = networkOptions(min.degree = .2))
```

| | |
|--------------|------------------------------------------------------------------------|
| cv.glmOrphan | <i>GLMNET cross-validation model penalizing nodes with high degree</i> |
|--------------|------------------------------------------------------------------------|

Description

This function overrides the ‘trans.fun’ options in ‘network.options’ with an heuristic described in Verissimo et al. that penalizes nodes with high degree.

Usage

```
cv.glmOrphan(xdata, ydata, network, network.options = networkOptions(), ...)
```

Arguments

| | |
|-----------------|-----------------------------------------------------|
| xdata | input data, can be a matrix or MultiAssayExperiment |
| ydata | response data compatible with glmnet |
| network | type of network, see below |
| network.options | options to calculate network |
| ... | parameters that glmnet accepts |

Value

see cv.glmSparseNet

See Also

glmNetSparse

Examples

```
xdata <- matrix(rnorm(100), ncol = 5)
cv.glmOrphan(xdata, rnorm(nrow(xdata)), 'correlation',
             family = 'gaussian',
             nfolds = 5,
             network.options = networkOptions(min.degree = .2))
```

| | |
|-----------------|-------------------------------------------------------------------------------|
| cv.glmSparseNet | <i>Calculate cross validating GLM model with network-based regularization</i> |
|-----------------|-------------------------------------------------------------------------------|

Description

network parameter accepts:

Usage

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

Arguments

| | |
|-----------------|-----------------------------------------------------|
| xdata | input data, can be a matrix or MultiAssayExperiment |
| ydata | response data compatible with glmnet |
| network | type of network, see below |
| network.options | options to calculate network |
| experiment.name | Name of experiment in MultiAssayExperiment |
| ... | parameters that 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

Examples

```
# 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

#
# load data
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.name = '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,

```

```

    consider.unweighted = FALSE,
    force.recalc.degree = FALSE,
    force.recalc.network = FALSE,
    n.cores = 1,
    ...
)

```

Arguments

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

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, consider.unweighted = 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,
  consider.unweighted = FALSE,
  force.recalc.degree = FALSE,
  force.recalc.network = FALSE,
  n.cores = 1,
  ...
)

```

Arguments

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

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, consider.unweighted = TRUE)

```

| | |
|--------------|------------------------------|
| digest.cache | <i>Default digest method</i> |
|--------------|------------------------------|

Description

Sets a default caching algorithm to use with run.cache

Usage

```
digest.cache(val)
```

Arguments

val object to calculate hash over

Value

a hash of the sha256

Examples

```
glmSparseNet:::digest.cache(c(1,2,3,4,5))  
glmSparseNet:::digest.cache('some example')
```

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 | <i>Retrieve ensembl gene names from biomaRt</i> |
|------------------|-------------------------------------------------|

Description

Retrieve ensembl gene names from biomaRt

Usage

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

Arguments

| | |
|-----------|---------------------------------------------------------------------------------------------------------------------|
| gene.id | character vector with gene names |
| use.cache | 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

a dataframe with external gene names, ensembl_id

Examples

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

| | |
|-----------|-----------------------------------------|
| geneNames | <i>Retrieve gene names from biomaRt</i> |
|-----------|-----------------------------------------|

Description

Retrieve gene names from biomaRt

Usage

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

Arguments

| | |
|---------------|---------------------------------------------------------------------------------------------------------------------|
| ensembl.genes | character vector with gene names in ensembl_id format |
| use.cache | 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

a dataframe with external gene names, `ensembl_id`

Examples

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

glmDegree

GLMNET model penalizing nodes with small degree

Description

This function overrides the ‘`trans.fun`’ options in ‘`network.options`’ with the inverse of a degree described in Verissimo et al. (2015) that penalizes nodes with small degree.

Usage

```
glmDegree(xdata, ydata, network, network.options = networkOptions(), ...)
```

Arguments

| | |
|------------------------------|------------------------------------------------------------------|
| <code>xdata</code> | input data, can be a matrix or <code>MultiAssayExperiment</code> |
| <code>ydata</code> | response data compatible with <code>glmnet</code> |
| <code>network</code> | type of network, see below |
| <code>network.options</code> | options to calculate network |
| <code>...</code> | parameters that <code>glmnet</code> accepts |

Value

see `glmNetSparse`

See Also

`glmNetSparse`

Examples

```
xdata <- matrix(rnorm(100), ncol = 5)
glmDegree(xdata, rnorm(nrow(xdata)), 'correlation',
          family = 'gaussian',
          network.options = networkOptions(min.degree = .2))
```

glmHub *GLMNET model penalizing nodes with small degree*

Description

This function overrides the ‘trans.fun’ options in ‘network.options’ with an heuristic described in Verissimo et al. that penalizes nodes with small degree.

Usage

```
glmHub(xdata, ydata, network, network.options = networkOptions(), ...)
```

Arguments

| | |
|-----------------|-----------------------------------------------------|
| xdata | input data, can be a matrix or MultiAssayExperiment |
| ydata | response data compatible with glmnet |
| network | type of network, see below |
| network.options | options to calculate network |
| ... | parameters that glmnet accepts |

Value

see glmNetSparse

See Also

glmNetSparse

Examples

```
xdata <- matrix(rnorm(100), ncol = 5)
glmHub(xdata, rnorm(nrow(xdata)), 'correlation', family = 'gaussian',
       network.options = networkOptions(min.degree = .2))
```

glmOrphan *GLMNET model penalizing nodes with high degree*

Description

This function overrides the ‘trans.fun’ options in ‘network.options’ with an heuristic described in Verissimo et al. that penalizes nodes with high degree.

Usage

```
glmOrphan(xdata, ydata, network, network.options = networkOptions(), ...)
```

Arguments

| | |
|-----------------|-----------------------------------------------------|
| xdata | input data, can be a matrix or MultiAssayExperiment |
| ydata | response data compatible with glmnet |
| network | type of network, see below |
| network.options | options to calculate network |
| ... | parameters that glmnet accepts |

Value

see glmNetSparse

See Also

glmNetSparse

Examples

```
xdata <- matrix(rnorm(100), ncol = 5)
glmOrphan(xdata, rnorm(nrow(xdata)), 'correlation', family = 'gaussian',
          network.options = networkOptions(min.degree = .2))
```

glmSparseNet

Calculate GLM model with network-based regularization

Description

network parameter accepts:

Usage

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

Arguments

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

Value

an object just as glmnet

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
data('miniACC', package="MultiAssayExperiment")
xdata <- miniACC
# TODO asking 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.name = 'RNASeq2GeneNorm')
```

```
glmSparseNet.options
```

Constants for 'glmSparseNet' package

Description

Log level constants and the logger options.

Usage

```
glmSparseNet.options(..., simplify = FALSE, update = list())
```

Arguments

| | |
|----------|--------------------------------|
| ... | TODO |
| simplify | TODO |
| update | pair list of update to options |

Details

The logging configuration is managed by 'glmSparseNet.options', a function generated by Option-Manager within 'futile.options'.

Value

futile.options::OptionsManager object

See Also

futile.options

```
hallmarks
```

Retrieve hallmarks of cancer count for genes

Description

Retrieve hallmarks of cancer count for genes

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

Examples

```
hallmarks(c('MOB1A', 'RFLNB', 'SPIC'))

hallmarks(c('MOB1A', 'RFLNB', 'SPIC'), metric = 'cprob')
```

| | |
|----------------|-----------------------------------------------------|
| heuristicScale | <i>Heuristic function to use in high dimensions</i> |
|----------------|-----------------------------------------------------|

Description

Heuristic function to use in high dimensions

Usage

```
heuristicScale(x, sub.exp10 = -1, exp.mult = -1, sub.exp = -1)
```

Arguments

| | |
|-----------|------------------------------------------------------------------------------------------------------|
| x | vector of values to scale |
| sub.exp10 | value to subtract to base 10 exponential, for example: '10 ⁰ - sub.exp10 = 1 - sub.exp10' |
| exp.mult | parameter to multiply exponential, i.e. to have a negative exponential or positive |
| sub.exp | value to subtract for exponential, for example if x = 0, 'exp(0) - sub.exp = 1 - sub.exp' |

Value

a vector of scaled values

Examples

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

| | |
|--------------|-------------------------------------------------------------|
| hubHeuristic | <i>Heuristic function to penalize nodes with low degree</i> |
|--------------|-------------------------------------------------------------|

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))
```

| | |
|-----------|---------------------------------|
| my.colors | <i>Custom pallete of colors</i> |
|-----------|---------------------------------|

Description

Custom pallete of colors

Usage

```
my.colors(ix = NULL)
```

Arguments

ix index for a color

Value

a color

Examples

```
my.colors()  
my.colors(5)
```

| | |
|------------|-------------------------------------------|
| my.symbols | <i>Custom palette of symbols in plots</i> |
|------------|-------------------------------------------|

Description

Custom palette of symbols in plots

Usage

```
my.symbols(ix = NULL)
```

Arguments

ix index for symbol

Value

a symbol

Examples

```
my.symbols()  
my.symbols(2)
```

| | |
|--------------------|-------------------------------------------|
| networkCorParallel | <i>Calculates the correlation network</i> |
|--------------------|-------------------------------------------|

Description

Calculates the correlation network

Usage

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

Arguments

| | |
|----------------------|---------------------------------------------------------------------------------------------------------------|
| xdata | base data to calculate network |
| build.output | if output returns a 'matrix', 'vector' of the upper triu without the diagonal or NULL with any other argument |
| n.cores | number of cores to be used |
| force.recalc.network | force recalculation, instead of going to cache |
| show.message | shows cache operation messages |
| ... | extra parameters for fun |

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,
  build.output = "matrix",
  n.cores = 1,
  force.recalc.network = FALSE,
  show.message = FALSE,
  ...
)
```

Arguments

| | |
|----------------------|---------------------------------------------------------------------------------------------------------------|
| xdata | base data to calculate network |
| build.output | if output returns a 'matrix', 'vector' of the upper triu without the diagonal or NULL with any other argument |
| n.cores | number of cores to be used |
| force.recalc.network | force recalculation, instead of going to cache |
| show.message | shows cache operation messages |
| ... | extra parameters for fun |

Value

depends on build.output parameter

Examples

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

| | |
|----------------|------------------------------|
| networkOptions | <i>Setup network options</i> |
|----------------|------------------------------|

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",
  min.degree = 0,
  n.cores = 1,
  trans.fun = function(x) { x }
)
```

Arguments

| | |
|------------|--------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------------|
| method | in case of correlation and covariance, which method to use |
| unweighted | calculate degree using unweighted network |
| cutoff | cutoff value in network edges to trim the network |
| centrality | centrality measure to use, currently only supports degree |
| min.degree | minimum value that individual penalty weight can take |
| n.cores | number of cores to use, default to 1 |
| | The trans.fun 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. |
| trans.fun | see below |

Value

a list of options

See Also

glmOrphan 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(ensembl.proteins, use.cache = TRUE, verbose = FALSE)
```

Arguments

| | |
|------------------|---------------------------------------------------------------------------------------------------------------------|
| ensembl.proteins | character vector with gene names in ensembl_peptide_id format |
| use.cache | 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

a dataframe with external gene names, ensembl_peptide_id

Examples

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

run.cache

Run function and save cache

Description

This method saves the function that's being called

Usage

```
run.cache(
  fun,
  ...,
  seed = NULL,
  base.dir = NULL,
  cache.prefix = "generic_cache",
  cache.digest = list(),
  show.message = NULL,
  force.recalc = FALSE,
  add.to.hash = NULL
)
```

Arguments

| | |
|----------|------------------------------------------------------------------|
| fun | function call name |
| ... | parameters for function call |
| seed | when function call is random, this allows to set seed beforehand |
| base.dir | directory where data is stored |

| | |
|--------------|------------------------------------------------------------|
| cache.prefix | prefix for file name to be generated from parameters (...) |
| cache.digest | cache of the digest for one or more of the parameters |
| show.message | show message that data is being retrieved from cache |
| force.recalc | force the recalculation of the values |
| add.to.hash | something to add to the filename generation |

Value

the result of fun(...)

Examples

```
# [optional] save cache in a temporary directory
#
glmSparseNet::base.dir(tempdir())
glmSparseNet::run.cache(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::run.cache(c, 1, 2, 3, 4)
glmSparseNet::run.cache(c, a=1, 2, c= 3, 4)

# Using a local folder
# glmSparseNet::run.cache(c, 1, 2, 3, 4, base.dir = "runcache")
```

run.cache,function-method

Run function and save cache

Description

Run function and save cache

Usage

```
## S4 method for signature ``function``
run.cache(
  fun,
  ...,
  seed = NULL,
  base.dir = NULL,
  cache.prefix = "generic_cache",
  cache.digest = list(),
  show.message = NULL,
```

```

    force.recalc = FALSE,
    add.to.hash = NULL
  )

```

Arguments

| | |
|--------------|------------------------------------------------------------------|
| fun | function call name |
| ... | parameters for function call |
| seed | when function call is random, this allows to set seed beforehand |
| base.dir | directory where data is stored |
| cache.prefix | prefix for file name to be generated from parameters (...) |
| cache.digest | cache of the digest for one or more of the parameters |
| show.message | show message that data is being retrieved from cache |
| force.recalc | force the recalculation of the values |
| add.to.hash | something to add to the filename generation |

Value

the result of fun(...)

Examples

```

# [optional] save cache in a temporary directory
#
glmSparseNet::base.dir(tempdir())
glmSparseNet::run.cache(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::run.cache(c, 1, 2, 3, 4)
glmSparseNet::run.cache(c, a=1, 2, c= 3, 4)

# Using a local folder
# glmSparseNet::run.cache(c, 1, 2, 3, 4, base.dir = "runcache")

```

save.run.cache

Saving the cache

Description

Saving the cache

Usage

```
save.run.cache(result, path, compression, show.message)
```

Arguments

| | |
|--------------|----------------------------------------|
| result | main result to save |
| path | path to the file to save |
| compression | compression method to be used |
| show.message | TRUE to show messages, FALSE otherwise |

Value

result of save operation

Examples

```
glmSparseNet::save.run.cache(
  35, file.path(tempdir(), 'save.run.cache.Rdata'), FALSE, TRUE
)
```

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(
  chosen.btas,
  xdata,
  ydata,
  probs = c(0.5, 0.5),
  no.plot = FALSE,
  plot.title = "SurvivalCurves",
  xlim = NULL,
  ylim = NULL,
  expand.yzero = FALSE,
  legend.outside = FALSE,
  stop.when.overlap = TRUE,
  ...
)
```


show.message

Show messages option in run.cache

Description

Show messages option in run.cache

Usage

```
show.message(show.message = NULL)
```

Arguments

show.message boolean indicating to show messages or not

Value

the show.message option

Examples

```
glmSparseNet:::show.message(FALSE)
```

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 dgCMatix 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", score_threshold = 0, remove.text = TRUE)
```

Arguments

| | |
|-----------------|---------------------------------|
| version | version of the database to use |
| score_threshold | remove scores below threshold |
| remove.text | remove text mining-based scores |

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(score_threshold = 800)
```

tempdir.cache *Temporary directory for runCache*

Description

Temporary directory for runCache

Usage

```
tempdir.cache()
```

Value

a path to a temporary directory used by runCache

| | |
|--------------|------------------------------------------------------------------|
| write.readme | <i>Write a file in run-cache directory to explain the origin</i> |
|--------------|------------------------------------------------------------------|

Description

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

Usage

```
write.readme(base.dir)
```

Arguments

base.dir directory where to build this file

Value

the path to the file it has written

Examples

```
glmSparseNet::write.readme(tempdir())
```

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