

Package ‘glmSparseNet’

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Type Package

Title Network Centrality Metrics for Elastic-Net Regularized Models

Version 1.10.0

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 true

NeedsCompilation no

biocViews Software, StatisticalMethod, DimensionReduction, Regression, Classification, Survival, Network, GraphAndNetwork

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

Imports SummarizedExperiment, `biomaRt`, `futile.logger`, `sparsebn`, `sparsebnUtils`, `forcats`, `dplyr`, `glue`, `readr`, `httr`, `ggplot2`, `survminer`, `reshape2`, `stringr`, `parallel`, `methods`, `loose.rock` (>= 1.0.12)

Suggests `testthat`, `knitr`, `rmarkdown`, `survival`, `survcomp`, `pROC`, `VennDiagram`, `BiocStyle`, `curatedTCGAData`, `TCGAutils`

VignetteBuilder `knitr`

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

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.calcPenalty	<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

xdata	input data
penalty.type	which method to use
network.options	options to be used

Value

vector with penalty weights

Examples

```
xdata <- matrix(rnorm(100), ncol = 20)
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')
glmSparseNet::::calcPenalty(xdata, 'sparsebn')
```

.degreeGeneric	<i>Generic function to calculate degree based on data</i>
----------------	---

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 only upper triu
...	extra parameters for fun

Value

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

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 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
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

`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
)
```

`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

<code>lambda.largest</code>	numeric value for largest number of lambda to consider (usually with a target of 1 selected variable)
<code>xdata</code>	X parameter for <code>glmnet</code> function
<code>ydata</code>	Y parameter for <code>glmnet</code> function

```

family      family parameter to glmnet function
orders.of.magnitude.smaller
            minimum value for lambda (lambda.largest / 10^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	<i>Build gene network from peptide ids</i>
--------------------	--

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

string.tbl	matrix with colnames and rownames as ensembl peptide id (same order)
use.names	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)

```

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 (score(ProteinA-Protein) == score(ProteinB, ProteinA))

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

curl.workaround

Workaround for bug with curl when fetching specific ensembl mirror

Description

<https://github.com/grimbough/biomaRt/issues/39>

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 Veríssimo 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 Veríssimo 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

GLMNET cross-validation model penalizing nodes with high degree

Description

This function overrides the ‘trans.fun’ options in ‘network.options’ with an heuristic described in Veríssimo 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
xdata <- MultiAssayExperiment::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)
```

degreeSparsebn	<i>Calculate degree of correlation matrix</i>
----------------	---

Description

Calculate degree of correlation matrix

Usage

```
degreeSparsebn(
  xdata,
  type = "continuous",
  levels = NULL,
  ivn = NULL,
  n = NULL,
  object = NULL,
  cutoff = 0,
```

```

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

```

Arguments

xdata	calculate correlation matrix on each column
type	either "discrete" or "continuous", see <code>sparsebnUtils::sparsebnData</code>
levels	(optional) list of levels for each node. see <code>sparsebnUtils::sparsebnData</code>
ivn	(optional) list of interventions for each observation, see <code>sparsebnUtils::sparsebnData</code>
n	(optional) number of rows from data matrix to print, see <code>sparsebnUtils::sparsebnData</code>
object	(optional) an object of type <code>sparsebnData</code> , see <code>sparsebnUtils::sparsebnData</code>
cutoff	positive value that determines a cutoff value
consider.unweighted	consider all edges as 1 if they are greater than 0
n.cores	number of cores to be used
show.message	shows cache operation messages
force.recalc.degree	force recalculation, instead of going to cache
force.recalc.network	force recalculation of network and penalty weights, instead of going to cache
...	parameters for <code>sparsebn::estimate.dag</code>

Value

a vector of the degrees

Examples

```

# generate a random matrix of observations
xdata <- matrix(rnorm(1000), nrow = 20)
degreeSparsebn(xdata)

```

downloadFileLocal	<i>Download files to local temporary path</i>
-------------------	---

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 Veríssimo et al. (2015) that penalizes nodes with small degree.

Usage

```
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 `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 Veríssimo 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 Veríssimo 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
xdata <- MultiAssayExperiment::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.name = 'RNASeq2GeneNorm')

```

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 chosen 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 npm

Examples

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

heuristicScale *Heuristic function to use in high dimensions*

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: ‘ $10^0 - \text{sub.exp10} = 1 - \text{sub.exp10}$ ’
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, ‘ $\exp(0) - \text{sub.exp} = 1 - \text{sub.exp}$ ’

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))
```

networkCorParallel *Calculates the correlation network*

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)
```

<code>networkCovParallel</code>	<i>Calculates the covariance network</i>
---------------------------------	--

Description

Calculates the covariance network

Usage

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

Arguments

<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
...	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	cuttoff 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
trans.fun	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	<i>Heuristic function to penalize nodes with high degree</i>
-----------------	--

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	<i>Retrieve ensembl gene ids from proteins</i>
--------------------------	--

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'
))
```

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,
  ...
)
```

Arguments

chosen.btas	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)
no.plot	Only calculate p-value and do not generate survival curve plot
plot.title	Name of file if
xlim	Optional argument to limit the x-axis view
ylim	Optional argument to limit the y-axis view
expand.yzero	expand to y = 0

```

legend.outside If TRUE legend will be outside plot, otherwise inside
stop.when.overlap
    when probs vector allows for overlapping of samples in both groups, then stop.
    Otherwise it will calculate with duplicate samples, i.e. simply adding them to
    xdata and ydata (in a different group)
...
    additional parameters to survminer::ggsurvplot

```

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

```

data('cancer', package = 'survival')
xdata <- ovarian[,c('age', 'resid.ds')]
ydata <- data.frame(time = ovarian$futime, status = ovarian$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

`string.network.700.cache`

Format

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

References

<https://string-db.org/>

Examples

```
head(string.network.700.cache)
```

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)
```

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