# Package 'dnet'

March 22, 2014

Title Dynamic networks via an integrative analysis of network and expression data

Type Package

<b>Version</b> 0.99.0
<b>Date</b> 2014-3-21
Author Hai Fang and Julian Gough
Maintainer Hai Fang <hfang@cs.bris.ac.uk></hfang@cs.bris.ac.uk>
<b>Depends</b> R (>= 3.0.1), igraph, supraHex, ape
Imports
Suggests affy, limma
<b>Description</b> The 'dnet' package is initiased to fill in the need of an open-source tool for analysing biological networks and high-throughput biological data in an integrative manner. More specifically, dnet intends to analyse the biological network whose nodes/genes are associated with dynamic information such as expression levels across samples. Also, dnet aims to deliver an eye-intuitive tool for network-based sample stratifications.
<pre>URL http://dnet.r-forge.r-project.org</pre>
Collate 'eCal.r' 'eView.r' 'eWrite.r' 'visRunES.r' 'dPvalAggregate.r' 'dNetInduce.r' 'dBUMfit.r' 'dBUMscore.r' 'dNetFind.r' 'dNetPipeline.r' 'dNetConfidence.r' 'visNet.r' 'visNetMul.r' 'visNetReorder.r' 'dNetReorder.r' 'visNetArc.r' 'visNetCircle.r' 'dRWR.r' 'dContrast.r' 'dCommSignif.r' 'dSVDSignif.r' 'dFDRscore.r' 'visColormap.r' 'visColoralpha.r' 'visHeatmap.r' 'visHeatmapAdv.r' 'visTreeBootstrap.r'
License GPL-2
biocViews Bioinformatics
R topics documented:
CLL       2         dBUMfit       3         dBUMscore       5         dCommSignif       6         dContrast       7         dFDRscore       8

2 CLL

CLL		Transo from (			hron	ic L	ymp	ohoc	cytic	c L	euk	xem	ia	pai	tien	ts
Index																59
	visTreeBootstrap .		 	 	 											54
	visRunES		 	 	 											53
	visNetReorder		 	 	 											51
	visNetMul		 	 	 											49
	visNetCircle		 	 	 											47
	visNetArc															
	visNet															
	visHeatmapAdv															
	visHeatmap															38
	visColormap															37
	visColoralpha															36
	org.Mm.string															35
	org.Mm.egPS															34
	org.Mm.egMP															34
																33
	org.Mm.egHPON .															31
	org.Mm.egGOMF . org.Mm.egHPMI .															31
	org.Mm.egGOCC .															30 31
	org.Mm.egGOBP .															29
	org.Mm.egDO															28
	org.Mm.eg															28
	org.Hs.string															27
	org.At.string															26
	Hiratani_TableS1 .		 	 	 											25
	eWrite		 	 	 											24
	eView		 	 	 											23
	eCal															20
	dSVDSignif															19
	dRWR															18
	dPvalAggregate															17
	dNetReorder															15
	dNetInduce dNetPipeline															12 13
	dNetFind															10
	dNetConfidence															9
																_

# Description

This dataset involves 130 patients with chronic lymphocytic leukemia (CLL). When enrolled in the study, these CLL patients had not received prior therapy for CLL. Additional covariate about the time to treatment (i.e. prognosis) is available. The dataset has been normalised and log2-transformed, and provided as an 'ExpressionSet' object.

dBUMfit 3

### Usage

```
load(url("http://dnet.r-forge.r-project.org/data/CLL.RData"))
```

#### Value

an object of class "ExpressionSet". It has slots for "assayData", "phenoData", and "featureData":

- assayData: a matrix of 54675 features X 130 samples
- phenoData: variables describing sample phenotypes (i.e. columns in assayData), including information about samples: "Name" for sample names, "Time" for sampling time to first treatment (years) and "Treatment" for treatment event (1:yes, 0:no)
- featureData: variables describing features (i.e. rows in assayData), including information about features/genes: "EntrezID" for gene EntrezID, "Symbol" for gene symbol and "Desc" for gene description

#### References

Chuang et al. (2012). Subnetwork-based analysis of chronic lymphocytic leukemia identifies pathways that associate with disease progression. *Blood*, 120(13):2639-49.

### **Examples**

```
# not run
if(0){
load(url("http://dnet.r-forge.r-project.org/data/CLL.RData"))
CLL
# extract information about the first 5 samples
pData(CLL)[1:5,]
# extract information about the first 5 features
fData(CLL)[1:5,]
}
```

dBUMfit

Function to fit a p-value distribution under beta-uniform mixture model

# Description

dBUMfit is supposed to take as input a vector of p-values for deriving their distribution under betauniform mixture model (see Note below). The density distribution of input p-values is expressed as a mixture of two components: one for the null hypothesis (the noise component) and the other for the alternative hypothesis (the signal component). The noise component is the uniform density, while the signal component is the remainder of the mixture distribution. It returns an object of class "BUM".

```
dBUMfit(x, ntry = 1, hist.bum = T, contour.bum = T,
  verbose = T)
```

4 dBUMfit

# **Arguments**

x a vector containing input p-values

ntry an integeter specifying how many trys are used to find the optimised parameters

by maximum likelihood estimation

hist.bum logical to indicate whether the histogram graph should be drawn

contour.bum logical to indicate whether a contour plot should be drawn to show the log likeli-

hood as a function of two parameters (a and lambda) in the beta-uniform mixture

model

verbose logical to indicate whether the messages will be displayed in the screen. By

default, it sets to true for display

#### Value

an object of class "BUM", a list with following elements:

• lambda: estimated mixture parameter

• a: estimated shape parameter

• NLL: Negative log-likelihood

• pvalues: the input pvalues

• call: the call that produced this result

#### Note

The probability density function of p-values under the Beta-Uniform Mixture model is formulated as:  $f(x|\lambda, a) = \lambda + (1 - \lambda) * a * x^{a-1}$ . The model names after mixing two distributions:

- the uniform distribution with the density function as  $\frac{1}{b-a}|_{a=0}^{b=1}=1$ 

• the beta distribution with the density function as  $\frac{\Gamma(a+b)}{\Gamma(a)+\Gamma(b)}*x^{a-1}*(1-x)^{b-1}|_{b=1}=a*x^{a-1}$ 

Both are mixed via  $\lambda$ . The mixture parameter  $\lambda$  measures the contribution from the uniform distribution. Accordingly,  $1 - \lambda$  measures the contribution from the beta distribution. Notably, the probability density function of the beta distribution can be splitted into two parts (rather than the exclusitive signal):

• the constant part as noise:  $a * x^{a-1}|_{x=1} = a$ 

• the rest part as signal:  $a * (x^{a-1} - 1)$ 

In other words, there is no signal at x=1 but all being noise. It is a conservative, upper bound estimation of the noise. Therefore, the probability density function in the model can be decomposed into signal-noise components:

• the signal component:  $(1 - \lambda) * a * (x^{a-1} - 1)$ 

• the noise component:  $\lambda + (1 - \lambda) * a$ 

It is misleading to simply view  $\lambda$  as the noise component and  $(1 - \lambda) * a * x^{a-1}$  as the signal component, just as wrongly do in the literatures (e.g. http://www.ncbi.nlm.nih.gov/pubmed/18586718)

### See Also

dBUMscore

dBUMscore 5

### **Examples**

```
# 1) generate an vector consisting of random values from beta distribution
x <- rbeta(1000, shape1=0.5, shape2=1)
# 2) fit a p-value distribution under beta-uniform mixture model
fit <- dBUMfit(x)
fit$lambda
fit$a</pre>
```

dBUMscore

Function to transform p-values into scores according to the fitted betauniform mixture model and/or after controlling false discovery rate

# Description

dBUMscore is supposed to take as input a vector of p-values, which are transformed into scores according to the fitted beta-uniform mixture model. Also if the FDR threshold is given, it is used to make sure that p-values below this are considered significant and thus scored positively. Instead, those p-values above the given FDR are considered insigificant and thus scored negatively.

# Usage

```
dBUMscore(fit, method = c("pdf", "cdf"), fdr = NULL,
    scatter.bum = T)
```

# **Arguments**

fit	an object of class "BUM"
method	the method used for the transformation. It can be either "pdf" for the method based on the probability density function of the fitted model, or "cdf" for the method based on the cumulative distribution function of the fitted model
fdr	the given FDR threshold. By default, it is set to NULL, meaning there is no constraint. If given, those p-values with the FDR below this are considered significant and thus scored positively. Instead, those p-values with the FDR above this given FDR are considered insigificant and thus scored negatively
scatter.bum	logical to indicate whether the scatter graph of scores against p-values should be drawn. Also indicated is the p-value (called tau) corresponding to the given FDR threshold (if any)

#### Value

• scores: a vector of scores

#### Note

The transformation from the input p-value x to the score S(x) is based on the fitted beta-uniform mixture model with two parameters  $\lambda$  and a:  $f(x|\lambda,a)=\lambda+(1-\lambda)*a*x^{a-1}$ . Specifically, it considers the log-likehood ratio between the signal and noise component of the model. The probability density function (pdf) of the signal component and the noise component are  $(1-\lambda)*a*(x^{a-1}-1)$  and  $\lambda+(1-\lambda)*a$ , respectively. Accordingly, the cumulative distribution function (cdf) of the signal component and the noise component are  $\int_0^x (1-\lambda)*a*(x^{a-1}-1)\,\mathrm{d}x$  and

6 dCommSignif

 $\int_0^x \lambda + (1-\lambda) *a \, \mathrm{d}x$ . In order to take into account the significance of the p-value, the fdr threshold is also used for down-weighting the score. According to how to measure both components, there are two methods implemented for deriving the score S(x):

- The method "pdf":  $S(x) = log_2 \frac{(1-\lambda)*a*(x^{a-1}-1)}{\lambda+(1-\lambda)*a} log_2 \frac{(1-\lambda)*a*(\tau^{a-1}-1)}{\lambda+(1-\lambda)*a} = log_2 \left(\frac{x^{a-1}-1}{\tau^{a-1}-1}\right)$ . For the purpose of down-weighting scores, it must ensure  $log_2 \frac{(1-\lambda)*a*(\tau^{a-1}-1)}{\lambda+(1-\lambda)*a} \geq 0$ , that is, the constraint via  $\tau \leq \left(\frac{\lambda+2*a*(1-\lambda)}{a*(1-\lambda)}\right)^{\frac{1}{a-1}}$
- The method "cdf":  $S(x) = log_2 \frac{\int_0^x (1-\lambda)*a*(x^{a-1}-1) \,\mathrm{d}x}{\int_0^x \lambda + (1-\lambda)*a \,\mathrm{d}x} log_2 \frac{\int_0^\tau (1-\lambda)*a*(\tau^{a-1}-1) \,\mathrm{d}x}{\int_0^\tau \lambda + (1-\lambda)*a \,\mathrm{d}x} = log_2 \frac{(1-\lambda)*(x^{a-1}-a)}{\lambda + (1-\lambda)*a} log_2 \frac{(1-\lambda)*(\tau^{a-1}-a)}{\lambda + (1-\lambda)*a} = log_2 \frac{(1-\lambda)*(\tau^{a-1}-a)}{\tau^{a-1}-a}.$  For the purpose of down-weighting scores, it must ensure  $log_2 \frac{(1-\lambda)*(\tau^{a-1}-a)}{\lambda + (1-\lambda)*a} \geq 0, \text{ that is, the constraint via } \tau \leq \left(\frac{\lambda + 2*a*(1-\lambda)}{1-\lambda}\right)^{\frac{1}{a-1}}$
- Where  $au = \left[\frac{\lambda + (1-\lambda)*a f dr * \lambda}{f dr * (1-\lambda)}\right]^{\frac{1}{a-1}}$ , i.e. the p-value corresponding to the exact f dr threshold. It can be duduced from the definition of the false discovery rate:  $f dr \doteq \frac{\int_0^\tau \lambda + (1-\lambda)*a \, \mathrm{d}x}{\int_0^\tau \lambda + (1-\lambda)*a * x^{a-1} \, \mathrm{d}x}$ . Notably, if the calculated  $\tau$  exceeds the contraint, it will be reset to the maximum end of that constraint

#### See Also

dBUMfit

### **Examples**

```
# 1) generate an vector consisting of random values from beta distribution
x <- rbeta(1000, shape1=0.5, shape2=1)

# 2) fit a p-value distribution under beta-uniform mixture model
fit <- dBUMfit(x)

# 3) calculate the scores according to the fitted BUM and fdr=0.01
# using "pdf" method
scores <- dBUMscore(fit, method="pdf", fdr=0.01)
# using "cdf" method
scores <- dBUMscore(fit, method="cdf", fdr=0.01)</pre>
```

dCommSignif

Function to test the significance of communities within a graph

### **Description**

dCommSignif is supposed to test the significance of communities within a graph. For a community of the graph, it first calculates two types of degrees for each node: degrees based on parters only within the community itself, and the degrees based on its parters NOT in the community but in the graph. Then, it performs two-sample Wilcoxon tests on these two types of degrees to produce the significance level (p-value)

```
dCommSignif(g, comm)
```

dContrast 7

# **Arguments**

g an object of class "igraph" or "graphNEL"

comm an object of class "communities". Details on this class can be found at http:
//igraph.sourceforge.net/doc/R/communities.html

#### Value

• significance: a vector of p-values (significance)

#### Note

none

### See Also

dCommSignif

# **Examples**

```
# 1) generate an vector consisting of random values from beta distribution
x <- rbeta(1000, shape1=0.5, shape2=1)
# 2) fit a p-value distribution under beta-uniform mixture model
fit <- dBUMfit(x, ntry=1, hist.bum=FALSE, contour.bum=FALSE)</pre>
# 3) calculate the scores according to the fitted BUM and fdr=0.01
# using "pdf" method
scores <- dBUMscore(fit, method="pdf", fdr=0.05, scatter.bum=FALSE)</pre>
names(scores) <- as.character(1:length(scores))</pre>
# 4) generate a random graph according to the ER model
g <- erdos.renyi.game(1000, 1/100)</pre>
# 5) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, V(g), knn=0)</pre>
# 6) find the module with the maximum score
module <- dNetFind(subg, scores)</pre>
# 7) find the module and test its signficance
comm <- walktrap.community(module, modularity=TRUE)</pre>
significance <- dCommSignif(module, comm)</pre>
```

dContrast

Function to help build the contrast matrix

# **Description**

dContrast is used to help build the contrast matrix

```
dContrast(level_sorted,
  contrast.type = c("average", "zero", "sequential", "pairwise"))
```

8 dFDRscore

# **Arguments**

level\_sorted a vector of levels (usually sorted) which are contrated to each other

contrast.type the type of the contrast. It can be one of either 'average' for the contrast against

the average of all levels, 'zero' for the contrast against the zero, 'sequential' for the contrast in a sequential order (it requires the levels being sorted properly), or

'pairwise' for the pairwise contrast.

#### Value

a list with following components:

· each: the contrast being specified

• name: the name of the contrast

#### Note

none

#### **Examples**

```
level_sorted <- c("L1","L2","L3","L4")

# the contrast against the average of all levels
contrasts <- dContrast(level_sorted, contrast.type="average")

# the contrast against the zero
contrasts <- dContrast(level_sorted, contrast.type="zero")

# the contrast in a sequential order
contrasts <- dContrast(level_sorted, contrast.type="sequential")

# the pairwise contrast
contrasts <- dContrast(level_sorted, contrast.type="pairwise")</pre>
```

dFDRscore

Function to transform fdr into scores according to log-likelihood ratio between the true positives and the false positivies and/or after controlling false discovery rate

# **Description**

dFDRscore is supposed to take as input a vector of fdr, which are transformed into scores according to according to log-likelihood ratio between the true positives and the false positivies. Also if the FDR threshold is given, it is used to make sure that fdr below threshold are considered significant and thus scored positively. Instead, those fdr above the given threshold are considered insigificant and thus scored negatively.

```
dFDRscore(fdr, fdr.threshold = NULL, scatter = F)
```

dNetConfidence 9

### **Arguments**

fdr a vector containing a list of input fdr

fdr.threshold the given FDR threshold. By default, it is set to NULL, meaning there is no

constraint. If given, those fdr with the FDR below threshold are considered significant and thus scored positively. Instead, those fdr with the FDR above

given threshold are considered insigificant and thus scored negatively

scatter logical to indicate whether the scatter graph of scores against p-values should be

drawn. Also indicated is the score corresponding to the given FDR threshold (if

any)

#### Value

· scores: a vector of scores

#### Note

none

#### See Also

dSVDSignif, dNetPipeline

# **Examples**

```
# 1) generate data with three different distributions, each with an iid normal random matrix of 1000 x 3
data <- cbind(matrix(rnorm(1000*3,mean=0,sd=1), nrow=1000, ncol=3),
matrix(rnorm(1000*3,mean=0.5,sd=1), nrow=1000, ncol=3),
matrix(rnorm(1000*3,mean=-0.5,sd=1), nrow=1000, ncol=3))

# 2) calculate the significance according to SVD
# using "fdr" significance
fdr <- dSVDSignif(data, signif="fdr", num.permutation=100)

# 3) calculate the scores according to the fitted BUM and fdr=0.01
# no fdr threshold
scores <- dFDRscore(fdr)
# using fdr threshold of 0.01</pre>
```

scores <- dFDRscore(fdr, fdr.threshold=0.1, scatter=TRUE)</pre>

dNetConfidence

Function to append the confidence information from the source graphs into the target graph

# Description

eConsensusGraph is supposed to append the confidence information (extracted from a list of the source graphs) into the target graph. The confidence information is about how often a node (or an edge) in the target graph that can be found in the input source graphs. The target graph is an object of class "igraph" or "graphNEL", and the source graphs are a list of objects of class "igraph" or "graphNEL". It also returns an object of class "igraph" or "graphNEL"; specifically, the same as the input target graph but appended with the "nodeConfidence" attribute to the nodes and the "edgeConfidence" attribute to the edges.

10 dNetFind

# Usage

```
dNetConfidence(target, sources, plot = F)
```

# **Arguments**

the target graph, an object of class "igraph" or "graphNEL"

sources a list of the source graphs, each with an object of class "igraph" or "graphNEL".

These source graphs will be used to calculate how often a node (or an edge) in

the target graph that can be found with them.

plot logical to indicate whether the returned graph (i.e. the target graph plus the

confidence information on nodes and edges) should be plotted. If it sets true, the plot will display the returned graph with the size of nodes indicative of the node confidence (the frequency that a node appears in the source graphs), and with the width of edges indicative of the edge confidence (the frequency that an edge

appears in the source graphs)

#### Value

an object of class "igraph" or "graphNEL", which is a target graph but appended with the "node-Confidence" attribute to the nodes and the "edgeConfidence" attribute to the edges

#### Note

None

# See Also

visNet

# **Examples**

```
# 1) generate a target graph according to the ER model
g <- erdos.renyi.game(100, 1/100)
target <- dNetInduce(g, V(g), knn=0)

# 2) generate a list source graphs according to the ER model
sources <- lapply(1:100, function(x) erdos.renyi.game(100*runif(1), 1/10))

# 3) append the confidence information from the source graphs into the target graph
g <- dNetConfidence(target=target, sources=sources)

# 4) visualise the confidence target graph
visNet(g, vertex.size=V(g)$nodeConfidence/10, edge.width=E(g)$edgeConfidence)</pre>
```

dNetFind

Function to find heuristically maximum scoring module

dNetFind 11

### **Description**

dNetFind is supposed to find the maximum scoring module from an input graph and scores imposed on its nodes. The input graph and the output module are both of "igraph" or "graphNET" object. The input scores imposed on the nodes in the input graph can be divided into two parts: the positive nodes and the negative nodes. The searching for maximum scoring module is deduced to find the connected subgraph containing the positive nodes as many as possible, but the negative nodes as few as possible. To this end, a heuristic search is used (see Note below).

### Usage

```
dNetFind(g, scores)
```

### **Arguments**

g an object of class "igraph" or "graphNEL"

scores a vector of scores. For each element, it must have the name that could be mapped onto the input graph. Also, the names in input "scores" should contain all those

in the input graph "g", but the reverse is not necessary

#### Value

a module with a maximum score, an object of class "igraph" or "graphNEL"

#### Note

The search procedure is heuristic to find the module with the maximum score:

- i) transform the input graph into a new graph by collapsing connected positive nodes into a meta-node. As such, meta-nodes are isolated to each other but are linked via negative nodes (single-nodes). Clearly, meta-nodes have positive scores, and negative scores for the single-nodes.
- ii) append the weight attribute to the edges in the transformed graph. There are two types of edges: 1) the single-single edge with two single-nodes as two ends, and 2) single-meta edge with a single-node as one end and a meta-node as the other end. The weight for a single-single edge is the absolute sum of the scores in its two-end single-nodes but normalised by their degrees. The weight for a single-meta edge is simply the absolute score in its single-node end normalised by the degree. As such, weights are all non-negative.
- iii) find minimum spanning tree (MST) in the weighted transformed graph using Prim's greedy algorithm. A spanning tree of the weighted graph is a subgraph that is tree and connects all the node together. The MST is a spanning tree with the sum of its edge weights minimised among all possible spanning trees.
- iv) find all shortest paths between any pair of meta-nodes in the MST. Within the weighted transformed graph in ii), a subgraph is induced containing nodes (only occuring in these shortest paths) and all edges between them.
- v) within the induced subgraph, identify single-nodes that are direct neighbors of meta-nodes.
   For each of these single-nodes, also make sure it has the absolute scores no more than the sum of scores in its neighboring meta-nodes. These single-nodes meeting both criteria are called "linkers".
- vi) still within the induced subgraph in v), find the linker graph that contains only linkers and
  edges between them. Similarly to iii), find MST of the linker graph, called 'linker MST'.
  Notably, this linker MST serves as the scaffold, which only contains linkers but has metanodes being directly attached to.

12 dNetInduce

• vii) in linker MST plus its attached meta-nodes, find the optimal path that has the sum of scores of its nodes and attached meta-nodes maximised amongest all possible paths. Nodes along this optimal path plus their attached meta-nodes are called 'module nodes'.

• viii) finally, from the input graph extract a subgraph (called 'module') that only contains module nodes and edges betwen them. This module is the maximum scoring module containing the positive nodes as many as possible, but the negative nodes as few as possible.

### See Also

dNetFind

### **Examples**

```
# 1) generate an vector consisting of random values from beta distribution
x <- rbeta(1000, shape1=0.5, shape2=1)

# 2) fit a p-value distribution under beta-uniform mixture model
fit <- dBUMfit(x, ntry=1, hist.bum=FALSE, contour.bum=FALSE)

# 3) calculate the scores according to the fitted BUM and fdr=0.01
# using "pdf" method
scores <- dBUMscore(fit, method="pdf", fdr=0.05, scatter.bum=FALSE)
names(scores) <- as.character(1:length(scores))

# 4) generate a random graph according to the ER model
g <- erdos.renyi.game(1000, 1/100)

# 5) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, V(g), knn=0)

# 6) find the module with the maximum score
module <- dNetFind(subg, scores)</pre>
```

dNetInduce

Function to generate a subgraph induced by given vertices and their k nearest neighbors

# **Description**

dNetInduce is supposed to produce a subgraph induced by given vertices and its k nearest neighbors. The input is a graph of "igraph" or "graphNET" object, a list of the vertices of the graph, and a k value for finding k nearest neighbors for these vertices. The output is a subgraph induced by given veretices plus their k neighbours. The resultant subgraph inherits the class from the input one. The induced subgraph conatins exactly the vertices of interest, and all the edges between them.

```
dNetInduce(g, nodes_query, knn = 0, remove.loops = T,
    largest.comp = T)
```

dNetPipeline 13

### **Arguments**

g an object of class "igraph" or "graphNEL"

nodes\_query the vertices for which the calculation is performed

knn an integeter specifying how many k steps are used to find the nearest neighbours of the given vertices. By default, knn is set to zero; it means no neighbors will be considered. When knn is 1, the immediate neighbors of the given vertices will be also considered for inducing the subgraph. The same is true when knn is 2, etc

remove.loops logical to indicate whether the loop edges are to be removed. By default, it sets to true for self-loops being removed

largest.comp logical to indicate whether the largest component is only retained. By default, it sets to true for the largest component being left

#### Value

• subg: an induced subgraph, an object of class "igraph" or "graphNEL"

#### Note

The given vertices plus their k nearest neighbors will be used to induce the subgraph.

#### See Also

dNetInduce

# **Examples**

```
# 1) generate a random graph according to the ER model
g <- erdos.renyi.game(100, 1/100)

# 2) select the frist 10 vertices as the query nodes
nodes_query <- V(g)[1:10]

# 3) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, nodes_query, knn=0)

# 4) produce the induced subgraph based on the nodes in query ane their immediate neighbours
subg <- dNetInduce(g, nodes_query, knn=1)</pre>
```

dNetPipeline	Function to setup the pipeline for finding maximum-scoring module
	from an input graph and the signficance imposed on its nodes

# **Description**

dNetPipeline is supposed to finish ab inito maximum-scoring module identification for the input graph with the node information on the significance (p-values). It returns an object of class "igraph" or "graphNEL".

14 dNetPipeline

### Usage

```
dNetPipeline(g, pval, method = c("pdf", "cdf", "fdr"),
  fdr = NULL, nsize = NULL, plot = F, verbose = T)
```

# **Arguments**

g	an object of class "igraph" or "graphNEL"
pval	a vector containing input p-values. For each element, it must have the name that could be mapped onto the input graph. Also, the names in input "pval" should contain all those in the input graph "g", but the reverse is not necessary
method	the method used for the transformation. It can be either "pdf" for the method based on the probability density function of the fitted model, or "cdf" for the method based on the cumulative distribution function of the fitted model
fdr	the given FDR threshold. By default, it is set to NULL, meaning there is no constraint. If given, those p-values with the FDR below this are considered significant and thus scored positively. Instead, those p-values with the FDR above this given FDR are considered insigificant and thus scored negatively
nsize	the desired number of nodes constrained to the resulting module. It is not nulll, a wide range of FDR will be scanned to find the FDR threshold leading to the desired number of nodes in the resulting module. Notably, the given FDR threshold will be overwritten.
plot	logical to indicate whether the histogram plot, contour plot and scatter plot should be drawn. By default, it sets to false for no plotting
verbose	logical to indicate whether the messages will be displayed in the screen. By default, it sets to true for display

# Value

a module with a maximum score, an object of class "igraph" or "graphNEL"

### Note

The pipeline sequentially consists of:

- i) dBUMfit used to fit the p-value distribution under beta-uniform mixture model.
- ii) if there is the desired number of nodes constrained to the resulting module, a wide range of FDR (including rough stage with large intervals, and finetune stage with smaller intervals) will be scanned to find the FDR threshold to meet the desired number of nodes.
- iii) dBUMscore used to calculate the scores according to the fitted BUM and FDR threshold.
- iv) dNetFind used to find maximum-scoring module from the input graph and scores imposed on its nodes.

# See Also

dBUMfit, dBUMscore, dNetFind

dNetReorder 15

#### **Examples**

```
# 1) generate an vector consisting of random values from beta distribution
x <- rbeta(1000, shape1=0.5, shape2=1)
names(x) <- as.character(1:length(x))

# 2) generate a random graph according to the ER model
g <- erdos.renyi.game(1000, 1/100)

# 3) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, V(g), knn=0)

# 4) find maximum-scoring module based on fdr=0.1 threshold
module <- dNetPipeline(g=subg, pval=x, fdr=0.1)

# 5) find maximum-scoring module with the desired node number nsize=20
# module <- dNetPipeline(g=subg, pval=x, nsize=20)</pre>
```

dNetReorder

Function to reorder the multiple graph colorings within a sheet-shape rectangle grid

### **Description**

dNetReorder is reorder the multiple graph colorings within a sheet-shape rectangle grid

# Usage

```
dNetReorder(g, data, feature = c("node", "edge"),
  node.normalise = c("none", "degree"), xdim = NULL,
  ydim = NULL, amplifier = NULL,
  metric = c("none", "pearson", "spearman", "kendall", "euclidean", "manhattan", "cos", "mi"),
  init = c("linear", "uniform", "sample"),
  algorithm = c("sequential", "batch"),
  alphaType = c("invert", "linear", "power"),
  neighKernel = c("gaussian", "bubble", "cutgaussian", "ep", "gamma"))
```

### **Arguments**

g an object of class "igraph" or "graphNEL"

data an input data matrix used to color-code vertices/nodes. One column corresponds

to one graph node coloring. The input matrix must have row names, and these names should include all node names of input graph, i.e. V(g)\$name, since there is a mapping operation. After mapping, the length of the patern vector should be the same as the number of nodes of input graph. The way of how to color-code is to map values in the pattern onto the whole colormap (see the next arguments:

colormap, ncolors, zlim and colorbar)

feature the type of the features used. It can be one of either 'edge' for the edge feature

or 'node' for the node feature.

node.normalise the normalisation of the nodes. It can be one of either 'none' for no normalisa-

tion or 'degree' for a node being penalised by its degree.

xdim an integer specifying x-dimension of the grid

16 dNetReorder

ydim an integer specifying y-dimension of the grid

amplifier an integer specifying the amplifier (3 by default) of the number of component

planes. The product of the component number and the amplifier constitutes the

number of rectangles in the sheet grid

metric distance metric used to define the similarity between component planes. It can

be "none", which means directly using column-wise vectors of codebook/data matrix. Otherwise, first calculate the covariance matrix from the codebook/data matrix. The distance metric used for calculating the covariance matrix between component planes can be: "pearson" for pearson correlation, "spearman" for spearman rank correlation, "kendall" for kendall tau rank correlation, "euclidean" for euclidean distance, "manhattan" for cityblock distance, "cos" for

cosine similarity, "mi" for mutual information.

init an initialisation method. It can be one of "uniform", "sample" and "linear" ini-

tialisation methods

algorithm the training algorithm. Currently, only "sequential" algorithm has been imple-

mented

alphaType the alpha type. It can be one of "invert", "linear" and "power" alpha types

neighKernel the training neighbor kernel. It can be one of "gaussian", "bubble", "cutgaus-

sian", "ep" and "gamma" kernels

#### Value

an object of class "sReorder", a list with following components:

- nHex: the total number of rectanges in the grid
- xdim: x-dimension of the grid
- ydim: y-dimension of the grid
- uOrder: the unique order/placement for each component plane that is reordered to the "sheet"-shape grid with rectangular lattice
- coord: a matrix of nHex x 2, with each row corresponding to the coordinates of each "uOrder" rectangle in the 2D map grid
- call: the call that produced this result

# Note

According to which features are used and whether nodes should be penalised by degrees, the feature data are constructed differently from the input data and input graph. When the node features are used, the feature data is the input data (or penalised data) with the same dimension. When the edge featrues are used, each entry (i.e. given an edge and a sample) in the feature data is the absolute difference between its two-end nodes (or after being penalised). Then, the constructed feature are subject to sample correlation analysis by supraHex. That is, a map grid (with sheet shape consisting of a rectangular lattice) is used to train either column-wise vectors of the feature data matrix or the covariance matrix thereof. As a result, similar samples are placed closer to each other within this map grid. More precisely, to ensure the unique placement, each sample mapped to the "sheet"-shape grid with rectangular lattice is determined iteratively in an order from the best matched to the next compromised one. If multiple samples are hit in the same rectangular lattice, the worse one is always sacrificed by moving to the next best one till all samples are placed somewhere exclusively on their own.

dPvalAggregate 17

#### See Also

visNetReorder

#### **Examples**

```
# 1) generate a random graph according to the ER model
g <- erdos.renyi.game(100, 1/100)

# 2) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, V(g), knn=0)

# 3) reorder the module with vertices being color-coded by input data
nnodes <- vcount(subg)
nsamples <- 10
data <- matrix(runif(nnodes*nsamples), nrow=nnodes, ncol=nsamples)
rownames(data) <- V(subg)$name
sReorder <- dNetReorder(g=subg, data, feature="node", node.normalise="none")</pre>
```

dPvalAggregate

Function to aggregate p values

# Description

dPvalAggregate is supposed to aggregate a input matrix p-values into a vector of aggregated p-values. The aggregate operation is applied to each row of input matrix, each resulting in an aggregated p-value. The method implemented can be based on the order statistics of p-values or according to Fisher's method.

# Usage

```
dPvalAggregate(pmatrix,
  method = c("orderStatistic", "fishers"),
  order = ncol(pmatrix))
```

# **Arguments**

pmatrix a data frame or matrix of p-values

method the method used. It can be either "orderStatistic" for the method based on the

order statistics of p-values, or "fishers" for Fisher's method

order an integeter specifying the order used for the aggregation according to on the

order statistics of p-values

### Value

• ap: a vector with the length nrow(pmatrix), containing aggregated p-values

# Note

For each row of input matrix with the c columns, there are c p-values that are uniformly independently distributed over [0,1] under the null hypothesis (uniform distribution). According to the order statistics, they follow the Beta distribution with the paramters a = order and b = c - order + 1. According to the Fisher's method, after transformation by  $-2 * \sum^c log(pvalue)$ , they follow Chi-Squared distribution.

dRWR

#### See Also

```
dPvalAggregate
```

### **Examples**

```
# 1) generate an iid uniformly-distributed random matrix of 1000x3
pmatrix <- cbind(runif(1000), runif(1000), runif(1000))
# 2) aggregate according to the ordre statistics
ap <- dPvalAggregate(pmatrix, method="orderStatistic")
# 3) aggregate according to the Fishers method
ap <- dPvalAggregate(pmatrix, method="fishers")</pre>
```

dRWR

Function to implement Random Walk with Restart (RWR) to precompute affinity matrix for the input graph

# **Description**

dRWR is supposed to implement Random Walk with Restart (RWR) to pre-compute affinity matrix for for nodes in the input graph with respect to the starting node (loop over every node in the graph)

### Usage

```
dRWR(g,
  normalise = c("laplacian", "row", "column", "none"),
  restart = 0.75)
```

### **Arguments**

g an object of class "igraph" or "graphNEL"

normalise the way to normalise the adjacency matrix of the input graph. It can be 'lapla-

cian' for laplacian normalisation, 'row' for row-wise normalisation, 'column'

for column-wise normalisation, or 'none'

restart the restart probability used for RWR

### Value

• PTmatrix: affinity matrix with the dimension of n X n, where n is the number of nodes in the input graph. Columns stand for starting nodes walking from, and rows for ending nodes walking to. Therefore, a column for a starting node represents a steady-state affinity vector that the starting node will visit all the ending nodes in the graph

# Note

The input graph will treat as an unweighted graph if there is no 'weight' edge attribute assocaited

### See Also

dNetInduce

dSVDSignif 19

### **Examples**

```
# 1) generate a random graph according to the ER model
g <- erdos.renyi.game(100, 1/100)

# 2) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, V(g), knn=0)

# 3) calculate the affinity matrix
PTmatrix <- dRWR(subg, normalise="laplacian", restart=0.75)

# 4) visualise affinity matrix
graphics::image(PTmatrix, col=visColormap("wyr")(64), zlim=c(0,1))</pre>
```

dSVDSignif

Function to obtain SVD-based gene significance from the input genesample matrix

# **Description**

dSVDSignif is supposed to obtain gene signficance from the given gene-sample matrix according to singular value decomposition (SVD)-based method. The method includes: 1) singular value decomposition of the input matrix; 2) determination of the eigens in consideration (if not given); 3) construction of the gene-specific project vector based on the considered eigens; 4) calculation of the distance statistic from the projection vector to zero point vector; and 5) based on distance statistic to obtain the gene significance.

# Usage

```
dSVDSignif(data, num.eigen = NULL, pval.eigen = 0.01,
    signif = c("fdr", "pval"),
    orient.permutation = c("row", "column", "both"),
    num.permutation = 100,
    fdr.procedure = c("stepup", "stepdown"), verbose = T)
```

### **Arguments**

data	an input gene-sample data matrix used for singular value decomposition
num.eigen	an integer specifying the number of eigens in consideration. If NULL, this number will be automatically decided on based on the observed relative eigenexpression against randomised relative eigenexpression calculated from a list (here 100) of permutated input matrix
pval.eigen	p-value used to call those eigens as dominant. This parameter is used only when parameter 'num.eigen' is NULL. Here, p-value is calcualted to assess how likely the observed relative eigenexpression are more than the maximum relative eigenexpression calculated from permutated matrix
signif	the singificance to return. It can be either "pval" for using the p-value as the gene significance, or "fdr" for using the fdr as the gene significance
orient.permuta	tion

the orientation of matrix being permutated. It can be either "row" to permutate values within each row, or "column" to permutate values within each column, or

20 eCal

"both" to permutate values both within rows and columns. Notably, when using the p-value as the gene significance, it is always to permutate values within each row

num.permutation

an integer specifying how many permutations are used

fdr.procedure

the procedure to adjust the fdr. To ensure that the high distance statistic the more significance, the fdr should be adjusted either using "stepup" for step-up procedure (from the most significant to the least significant) or using "stepdown" for step-down procedure (from the least significant to the most significant)

verhose

logical to indicate whether the messages will be displayed in the screen. By

default, it sets to true for display

#### Value

a vector storing gene significance

### Note

none

### See Also

dFDRscore

# **Examples**

```
# 1) generate data with three different distributions, each with an iid normal random matrix of 1000 x 3
data <- cbind(matrix(rnorm(1000*3,mean=0,sd=1), nrow=1000, ncol=3),
matrix(rnorm(1000*3,mean=0.5,sd=1), nrow=1000, ncol=3),
matrix(rnorm(1000*3,mean=-0.5,sd=1), nrow=1000, ncol=3))
# 2) calculate the significance according to SVD
# using "pval" significance
pval <- dSVDSignif(data, signif="pval", num.permutation=100)
# using "fdr" significance
fdr <- dSVDSignif(data, signif="fdr", num.permutation=100)</pre>
```

eCal

Function to conduct gene set enrichment analysis given the input data and the ontology in query

### **Description**

eCal is supposed to conduct gene set enrichment analysis given the input data and the ontology in query. It returns an object of class "eTerm".

```
eCal(data, identity = c("symbol", "entrez"),
  genome = c("mm", "hs"),
  ontology = c("GOBP", "GOMF", "GOCC", "MP", "DO", "PS"),
  sizeRange = c(10, 1000), which_distance = NULL,
  weight = 1, nperm = 1000, fast = T,
  sigTail = c("two-tails", "one-tail"), verbose = T)
```

eCal 21

### **Arguments**

data a data frame or matrix of input data. It must have row names, either Entrez Gene ID or Symbol identity the type of gene identity (i.e. row names of input data), either "symbol" for gene symbols (by default) or "entrez" for Entrez Gene ID. The option "symbol" is preferred as it is relatively stable from one update to another; when gene symbols cannot be matched, synonyms will be searched against the genome identity. It can be either "mm" for mouse genome or "hs" for human genome genome the ontology supported currently. For mouse genome, it can be "GOBP" for ontology Gene Ontology Biological Process, "GOMF" for Gene Ontology Molecular Function, "GOCC" for Gene Ontology Cellular Component, or "MP" for Mammalian Phenotype. For human genome, it can be "GOBP" for Gene Ontology Biological Process, "GOMF" for Gene Ontology Molecular Function, "GOCC" for Gene Ontology Cellular Component, "HP" for Human Phenotype, or "DO" for Disease Ontology. sizeRange the minimum and maximum size of members of each gene set in consideration. By default, it sets to a minimum of 10 but no more than 1000 which\_distance which distance of terms in the ontology is used to restrict terms in consideration. By default, it sets to 'NULL' to consider all distances weight type of score weigth. It can be "0" for unweighted (an equivalent to Kolmogorov-Smirnov, only considering the rank), "1" for weighted by input gene score (by default), and "2" for over-weighted, and so on the number of random permutations. For each permutation, gene-score associanperm tions will be permutated so that permutation of gene-term associations is realised fast logical to indicate whether to fast calculate expected results from permutated data. By default, it sets to true sigTail the tail used to calculate the statistical significance. It can be either "two-tails" for the significance based on two-tails or "one-tail" for the significance based on one tail verbose logical to indicate whether the messages will be displayed in the screen. By default, it sets to false for no display

### Value

an object of class "eTerm", a list with following components:

- set\_info: a matrix of nSet X 4 containing gene set information, where nSet is the number of gene set in consideration, and the 4 columns are "setID" (i.e. "Term ID"), "name" (i.e. "Term Name"), "namespace" and "distance"
- gs: a list of gene sets, each storing gene members. Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"
- data: a matrix of nGene X nSample containing input data in consideration. It is not always the same as the input data as only those mappable are retained
- es: a matrix of nSet X nSample containing enrichment score, where nSample is the number of samples (i.e. the number of columns in input data
- nes: a matrix of nSet X nSample containing normalised enrichment score. It is the version of enrichment score but after being normalised by gene set size

22 eCal

- pvalue: a matrix of nSet X nSample containing nominal p value
- adjp: a matrix of nSet X nSample containing adjusted p value. It is the p value but after being adjusted for multiple comparisons
- gadjp: a matrix of nSet X nSample containing globally adjusted p value in terms of all samples
- fdr: a matrix of nSet X nSample containing false discovery rate (FDR). It is the estimated probability that the normalised enrichment score represents a false positive finding
- qvalue: a matrix of nSet X nSample containing q value. It is the monotunically increasing FDR
- call: the call that produced this result

#### Note

The interpretation of returned components:

- "es": enrichment score for the gene set is the degree to which this gene set is overrepresented at the top or bottom of the ranked list of genes in each column of input data;
- "nes": normalised enrichment score for the gene set is enrichment score that has already normalised by gene set size. It is comparable across analysed gene sets;
- "pvalue": nominal p value is the statistical significance of the enrichment score. It is not adjusted for multiple hypothesis testing, and thus is of limited use in comparing gene sets;
- "adjp": adjusted p value by Benjamini & Hochberg method. It is comparable across gene sets;
- "gadjp": globally adjusted p value by Benjamini & Hochberg method. Unlike "adjp", it is adjusted in terms of all samples;
- "fdr": false discovery rate is the estimated probability that the normalised enrichment score represents a false positive finding. Unlike "adjp" or "gadjp" (also aliased as "fdr") that is derived from a list of p values, this version of fdr is directly calculate from the statistic (i.e. normalised enrichment score);
- "qvalue": q value is the monotunically increasing FDR so that the higher "nes", the lower "qvalue".

### See Also

```
eView, eWrite
```

```
#load("~/Databases/ChipSeq/supraHex/Bioinformatics_AN/RTiming/GSE18019/TableS1T.RData")
#data <- RT_LR[,1:2]
#eTerm <- eCal(data=data, identity="symbol", genome="mm", ontology="MP", sizeRange=c(10,1000), which_distant</pre>
```

eView 23

eView Function to view enrichment results in a sample-specific manner
---

### **Description**

eView is supposed to view results of gene set enrichment analysis but for a specific sample.

# Usage

```
eView(eTerm, which_sample = 1, top_num = 10,
  sortBy = c("adjp", "gadjp", "ES", "nES", "pvalue", "FWER", "FDR", "qvalue"),
  decreasing = NULL, details = F)
```

### **Arguments**

eTerm an object of class "eTerm" which\_sample which sample will be viewed

top\_num the maximum number of gene sets will be viewed

sortBy which statistics will be used for sorting and viewing gene sets. It can be "adjp"

for adjusted p value, "gadjp" for globally adjusted p value, "ES" for enrichment score, "nES" for normalised enrichment score, "pvalue" for p value, "FWER" for family-wise error rate, "FDR" for false discovery rate, "qvalue" for q value

decreasing logical to indicate whether to sort in a decreasing order. If it is null, it would be

true for "ES" or "nES"; otherwise it would be false

details logical to indicate whether the detail information of gene sets is also viewed. By

default, it sets to false for no inclusion

### Value

a data frame with following components:

• setID: term ID

• ES: enrichment score

• nES: normalised enrichment score

• pvalue: nominal p value

• adjp: adjusted p value

• gadjp: globally adjusted p value

• FDR: false discovery rate

• qvalue: q value

• setSize: the number of genes in the set; optional, it is only appended when "details" is true

• name: term name; optional, it is only appended when "details" is true

• namespace: term namespace; optional, it is only appended when "details" is true

• distance: term distance; optional, it is only appended when "details" is true

# Note

none

eWrite

#### See Also

eCal

#### **Examples**

```
#eView(eTerm, which_sample=1, top_num=10, sortBy="adjp", decreasing=F, details=T)
```

eWrite

Function to write out enrichment results

# **Description**

eWrite is supposed to write out enrichment results.

### Usage

```
eWrite(eTerm,
  which_content = c("gadjp", "adjp", "pvalue", "FWER", "FDR", "qvalue", "nES", "ES"),
  which_score = c("gadjp", "adjp", "FWER", "FDR", "qvalue"),
  cutoff = 0.1, filename = NULL, keep.significance = T)
```

### **Arguments**

eTerm an object of class "eTerm"

which\_content the content will be written out. It includes two categories: i) based on "adjp"

for adjusted p value, "gadjp" for globally adjusted p value, "pvalue" for p value, "FWER" for family-wise error rate, "FDR" for false discovery rate, "qvalue" for q value; ii) based on "ES" for enrichment score, "nES" for normalised enrichment score. For the former, the content is : first -1\*log10-transformed, and then

multiplied by -1 if nES is negative.

which\_score which statistics/score will be used for declaring the significance. It can be "adjp"

for adjusted p value, "gadjp" for globally adjusted p value, "FWER" for family-

wise error rate, "FDR" for false discovery rate, "qvalue" for q value

cutoff a cutoff to declare the signficance. It should be used together with 'which\_score'

filename a character string naming a filename

keep.significance

logical to indicate whether or not to mask those insignficant by NA. By default,

it sets to true to mask those insignfiicant by NA

# Value

a data frame with following components:

• setID: term ID

• setSize: the number of genes in the set

· name: term name

• namespace: term namespace

• distance: term distance

• sample names: sample names in the next columns

Hiratani\_TableS1 25

#### Note

If "filename" is not NULL, a tab-delimited text file will be also written out.

### See Also

eCal

#### **Examples**

#output <- eWrite(eTerm, which\_content="gadjp", which\_score="gadjp", cutoff=0.05, filename="eWrite\_output.</pre>

Hiratani\_TableS1

Mouse multilayer omics dataset from Hiratani et al. (2010)

### **Description**

This multilayer omics dataset involves the information on DNA replication timing, promoter CpG classification and gene expression. It consists of digitised replication timing, promoter CpG status and expression levels of 17,292 genes in a variety of samples.

# Usage

```
load(url("http://dnet.r-forge.r-project.org/data/Hiratani_TableS1.RData"))
```

# Value

- RT: a replication timing matrix of 17,292 genes X 22 samples. These 22 samples come from 22 cell lines during early mouse embryogenesis, and they can be categorised into: 1) pluripotent cells, including ESCs (ESC\_46C, ESC\_D3 and ESC\_TT2) and iPSCs (iPSC, iPSC\_1D4 and iPSC\_2D4); 2) partially-reprogrammed iPSCs (piPSC\_1A2, piPSC\_1B3 and piPSC\_V3); 3) early epiblast (EPL and EMB3\_D3); 4) late epiblast (EpiSC5 and EpiSC7); 5) Ectoderm (EBM6\_D3, EBM9\_D3, NPC\_46C and NPC\_TT2); 6) Mesoderm and Endoderm; and 7) late Mesoderm (Myoblast, MEF\_female and MEF\_male).
- CpG: a matrix of 17,292 genes X 1 containing gene additional information on promoter CpG classification, with '1' for HCP (high CpG density promoters), '-1' for LCP (low CpG density promoters), '0' for ICP (intermediate CpG density promoters), and 'NA' for unclassified.
- EX: an expression matrix of 17,292 genes X 8 samples, and samples include pluripotent cells (ESC\_D3); early epiblast (EMB3\_D3); late epiblast (EpiSC7); Ectoderm (EBM6\_D3 and EBM9\_D3); Mesoderm and Endoderm.

### References

Mikkelsen et al. (2007). Genome-wide maps of chromatin state in pluripotent and lineage-committed cells. *Nature*, 448:553-560.

Hiratani et al. (2010). Genome-wide dynamics of replication timing revealed by in vitro models of mouse embryogenesis. *Genome Research*, 20:155-169.

```
load(url("http://dnet.r-forge.r-project.org/data/Hiratani_TableS1.RData"))
ls() # you should see three variables: RT, CpG and EX
```

26 org.At.string

org.At.string Arabidopsis functional protein associatio (version 9.0.5).	network from STRING
--	---------------------

### **Description**

An igraph object that contains a functional protein association network in arabidopsis. The network is extracted from the STRING database (version 9.0.5). Only those associations with medium confidence (score>=0.4) are retained.

# Usage

```
load(url("http://dnet.r-forge.r-project.org/data/org.At.string.RData"))
```

#### Value

an object of class "igraph" (see http://igraph.sourceforge.net/doc/R/aaa-igraph-package. html). It has attributes for both vertices and edges. Below are attributes for the vertices:

- name: unique id for the vertices
- seqid: protein seqid for the vertices
- geneid: Entrez geneid (if any) for the vertices
- symbol: gene symbol (if any) for the vertices
- description: gene description (if any) for the vertices

Below are attributes for the edges:

- neighborhood\_score: predictive score based on neighborhood data
- fusion\_score: predictive score based on fusion data
- cooccurence\_score: predictive score based on cooccurence data
- coexpression\_score: predictive score based on coexpression
- experimental\_score: predictive score based on experimental data
- database\_score: predictive score based on database
- textmining\_score: predictive score based on text mining
- combined\_score: combined score from all above predictive scores

# References

Franceschini et al. (2013) STRING v9.1: protein-protein interaction networks, with increased coverage and integration. *Nucleic Acids Res*, 41:D808-D815.

```
# not run
if(0){
load(url("http://dnet.r-forge.r-project.org/data/org.At.string.RData"))
org.At.string
}
```

org.Hs.string 27

org.Hs.string	Human functional protein association network from STRING (version 9.0.5).

### **Description**

An igraph object that contains a functional protein association network in human. The network is extracted from the STRING database (version 9.0.5). Only those associations with medium confidence (score>=0.4) are retained.

# Usage

```
load(url("http://dnet.r-forge.r-project.org/data/org.Hs.string.RData"))
```

#### Value

an object of class "igraph" (see http://igraph.sourceforge.net/doc/R/aaa-igraph-package. html). It has attributes for both vertices and edges. Below are attributes for the vertices:

- name: unique id for the vertices
- seqid: protein seqid for the vertices
- geneid: Entrez geneid (if any) for the vertices
- symbol: gene symbol (if any) for the vertices
- description: gene description (if any) for the vertices

Below are attributes for the edges:

- neighborhood\_score: predictive score based on neighborhood data
- fusion\_score: predictive score based on fusion data
- cooccurence\_score: predictive score based on cooccurence data
- coexpression\_score: predictive score based on coexpression
- experimental\_score: predictive score based on experimental data
- database\_score: predictive score based on database
- textmining\_score: predictive score based on text mining
- combined\_score: combined score from all above predictive scores

# References

Franceschini et al. (2013) STRING v9.1: protein-protein interaction networks, with increased coverage and integration. *Nucleic Acids Res*, 41:D808-D815.

```
# not run
if(0){
load(url("http://dnet.r-forge.r-project.org/data/org.Hs.string.RData"))
org.Hs.string
}
```

28 org.Mm.egDO

org.Mm.eg

Mouse Entrez Genes (EG).

# **Description**

An R object that contains Entrez Gene information for the mouse. This data is prepared based on ftp://ftp.ncbi.nih.gov/gene/DATA/gene\_info.gz.

# Usage

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.eg.RData"))
```

#### Value

an object of class "EG", a list with following components:

• gene\_info: a matrix of nGene X 7 containing gene information, where nGene is the number of Entrez Genes, and the 7 columns are "GeneID", "Symbol", "description", "chromosome", "map\_location", "Synonyms" and "dbXrefs"

#### References

Maglott et al. (2011) Entrez Gene: gene-centered information at NCBI. *Nucleic Acids Res*, 39:D52-57.

# **Examples**

```
# not run
if(0){
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.eg.RData"))
names(org.Mm.eg)
org.Mm.eg$gene_info[1:5,]
}
```

org.Mm.egDO

Annotations of Mouse Entrez Genes (EG) by Disease Ontology (DO).

### **Description**

An R object that contains associations between Disease Ontology terms and Mouse Entrez Genes. This data is first prepared based on http://sourceforge.net/p/diseaseontology/code/HEAD/tree/trunk/HumanDO.obo and http://dga.nubic.northwestern.edu/ajax/Download.ajax.php, which results in annotations of Human Entrez Genes. Then, these annotations are transferred to Mouse Entrez Genes based on ftp://anonymous@ftp.ncbi.nih.gov/pub/HomoloGene/build67/homologene.data.

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egDO.RData"))
```

org.Mm.egGOBP 29

#### Value

an object of class "GS", a list with following components:

• set\_info: a matrix of nSet X 4 containing gene set information, where nSet is the number of gene sets (i.e. DO terms), and the 4 columns are "setID" (i.e. "Term ID"), "name" (i.e. "Term Name"), "namespace" and "distance"

• gs: a list of gene sets, each storing gene members thereof. Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"

#### References

Schriml et al. (2012) Disease Ontology: a backbone for disease semantic integration. *Nucleic Acids Res*, 40:D940-946.

Peng et al. (2012) The Disease and Gene Annotations (DGA): an annotation resource for human disease. *Nucleic Acids Res*, 41:D553-560.

Sayers et al. (2011) Database resources of the National Center for Biotechnology Information. *Nucleic Acids Res*, 39:D38-51.

# **Examples**

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egDO.RData"))
names(org.Mm.egDO)
```

org.Mm.egGOBP

Annotations of Mouse Entrez Genes (EG) by Gene Ontology Biological Process (GOBP).

# Description

An R object that contains associations between Gene Ontology Biological Process terms and Mouse Entrez Genes. This data is prepared based on http://www.geneontology.org/ontology/obo\_format\_1\_2/gene\_ontology.1\_2.obo and ftp://ftp.ncbi.nih.gov/gene/DATA/gene2go.gz.

### Usage

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egGOBP.RData"))
```

# Value

an object of class "GS", a list with following components:

- set\_info: a matrix of nSet X 4 containing gene set information, where nSet is the number of gene sets (i.e. GOBP terms), and the 4 columns are "setID" (i.e. "Term ID"), "name" (i.e. "Term Name"), "namespace" and "distance"
- gs: a list of gene sets, each storing gene members thereof. Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"

#### References

Maglott et al. (2011) Entrez Gene: gene-centered information at NCBI. *Nucleic Acids Res*, 39:D52-57

Ashburner et al. (2000) Gene ontology: tool for the unification of biology. *Nat Genet*, 25:25-29.

30 org.Mm.egGOCC

# **Examples**

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egGOBP.RData"))
names(org.Mm.egGOBP)
```

org.Mm.egGOCC

Annotations of Mouse Entrez Genes (EG) by Gene Ontology Cellular Component (GOCC).

# **Description**

An R object that contains associations between Gene Ontology Cellular Component terms and Mouse Entrez Genes. This data is prepared based on  $http://www.geneontology.org/ontology/obo_format_1_2/gene_ontology.1_2.obo and ftp://ftp.ncbi.nih.gov/gene/DATA/gene2go.gz.$ 

# Usage

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egGOCC.RData"))
```

# Value

an object of class "GS", a list with following components:

- set\_info: a matrix of nSet X 4 containing gene set information, where nSet is the number of gene sets (i.e. GOCC terms), and the 4 columns are "setID" (i.e. "Term ID"), "name" (i.e. "Term Name"), "namespace" and "distance"
- gs: a list of gene sets, each storing gene members thereof. Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"

# References

Maglott et al. (2011) Entrez Gene: gene-centered information at NCBI. *Nucleic Acids Res*, 39:D52-57.

Ashburner et al. (2000) Gene ontology: tool for the unification of biology. Nat Genet, 25:25-29.

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egGOCC.RData"))
names(org.Mm.egGOCC)
```

org.Mm.egGOMF

org.Mm.egGOMF	Annotations of Mouse Entrez Genes (EG) by Gene Ontology Molecular Function (GOMF).

# **Description**

An R object that contains associations between Gene Ontology Molecular Function terms and Mouse Entrez Genes. This data is prepared based on http://www.geneontology.org/ontology/obo\_format\_1\_2/gene\_ontology.1\_2.obo and ftp://ftp.ncbi.nih.gov/gene/DATA/gene2go.gz.

# Usage

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egGOMF.RData"))
```

#### Value

an object of class "GS", a list with following components:

- set\_info: a matrix of nSet X 4 containing gene set information, where nSet is the number of gene sets (i.e. GOMF terms), and the 4 columns are "setID" (i.e. "Term ID"), "name" (i.e. "Term Name"), "namespace" and "distance"
- gs: a list of gene sets, each storing gene members thereof. Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"

# References

Maglott et al. (2011) Entrez Gene: gene-centered information at NCBI. *Nucleic Acids Res*, 39:D52-57.

Ashburner et al. (2000) Gene ontology: tool for the unification of biology. Nat Genet, 25:25-29.

# **Examples**

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egGOMF.RData"))
names(org.Mm.egGOMF)
```

org.Mm.egHPMI	Annotations of Mouse Entrez Genes (EG) by Human Phenotype Mode of Inheritance (HPMI).
---------------	---

# **Description**

An R object that contains associations between HPMI terms and Mouse Entrez Genes. This data is first prepared based on http://compbio.charite.de/svn/hpo/trunk/src/ontology/human-phenotype-ontology obo and http://compbio.charite.de/hudson/job/hpo.annotations.monthly/lastStableBuild/artifact/annotation/ALL\_SOURCES\_ALL\_FREQUENCIES\_genes\_to\_phenotype.txt, which results in annotations of Human Entrez Genes. Then, these annotations are transferred to Mouse Entrez Genes based on ftp://anonymous@ftp.ncbi.nih.gov/pub/HomoloGene/build67/homologene.data.

32 org.Mm.egHPON

### Usage

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egHPMI.RData"))
```

#### Value

an object of class "GS", a list with following components:

- set\_info: a matrix of nSet X 4 containing gene set information, where nSet is the number of gene sets (i.e. HPMI terms), and the 4 columns are "setID" (i.e. "Term ID"), "name" (i.e. "Term Name"), "namespace" and "distance"
- gs: a list of gene sets, each storing gene members thereof. Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"

#### References

Robinson et al. (2012) The Human Phenotype Ontology: a tool for annotating and analyzing human hereditary disease. *Am J Hum Genet*, 83:610-615.

Sayers et al. (2011) Database resources of the National Center for Biotechnology Information. *Nucleic Acids Res*, 39:D38-51.

### **Examples**

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egHPMI.RData"))
names(org.Mm.egHPMI)
```

org.Mm.egHPON

Annotations of Mouse Entrez Genes (EG) by Human Phenotype ONset and clinical course (HPON).

# **Description**

An R object that contains associations between HPON terms and Mouse Entrez Genes. This data is first prepared based on http://compbio.charite.de/svn/hpo/trunk/src/ontology/human-phenotype-ontology.obo and http://compbio.charite.de/hudson/job/hpo.annotations.monthly/lastStableBuild/artifact/annotation/ALL\_SOURCES\_ALL\_FREQUENCIES\_genes\_to\_phenotype.txt, which results in annotations of Human Entrez Genes. Then, these annotations are transferred to Mouse Entrez Genes based on ftp://anonymous@ftp.ncbi.nih.gov/pub/HomoloGene/build67/homologene.data.

### Usage

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egHPON.RData"))
```

# Value

an object of class "GS", a list with following components:

- set\_info: a matrix of nSet X 4 containing gene set information, where nSet is the number of gene sets (i.e. HPON terms), and the 4 columns are "setID" (i.e. "Term ID"), "name" (i.e. "Term Name"), "namespace" and "distance"
- gs: a list of gene sets, each storing gene members thereof. Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"

org.Mm.egHPPA 33

#### References

Robinson et al. (2012) The Human Phenotype Ontology: a tool for annotating and analyzing human hereditary disease. *Am J Hum Genet*, 83:610-615.

Sayers et al. (2011) Database resources of the National Center for Biotechnology Information. *Nucleic Acids Res*, 39:D38-51.

### **Examples**

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egHPON.RData"))
names(org.Mm.egHPON)
```

org.Mm.egHPPA

Annotations of Mouse Entrez Genes (EG) by Human Phenotype Phenotypic Abnormality (HPPA).

# **Description**

An R object that contains associations between Human Phenotype Phenotypic Abnormality terms and Mouse Entrez Genes. This data is first prepared based on http://compbio.charite.de/svn/hpo/trunk/src/ontology/human-phenotype-ontology.obo and http://compbio.charite.de/hudson/job/hpo.annotations.monthly/lastStableBuild/artifact/annotation/ALL\_SOURCES\_ALL\_FREQUENCIES\_genes\_to\_phenotype.txt, which results in annotations of Human Entrez Genes. Then, these annotations are transferred to Mouse Entrez Genes based on ftp://anonymous@ftp.ncbi.nih.gov/pub/HomoloGene/build67/homologene.data.

# Usage

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egHPPA.RData"))
```

### Value

an object of class "GS", a list with following components:

- set\_info: a matrix of nSet X 4 containing gene set information, where nSet is the number of gene sets (i.e. HPPA terms), and the 4 columns are "setID" (i.e. "Term ID"), "name" (i.e. "Term Name"), "namespace" and "distance"
- gs: a list of gene sets, each storing gene members thereof. Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"

### References

Robinson et al. (2012) The Human Phenotype Ontology: a tool for annotating and analyzing human hereditary disease. *Am J Hum Genet*, 83:610-615.

Sayers et al. (2011) Database resources of the National Center for Biotechnology Information. *Nucleic Acids Res*, 39:D38-51.

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egHPPA.RData"))
names(org.Mm.egHPPA)
```

34 org.Mm.egPS

org.Mm.egMP	Annotations Mouse Entrez Genes (EG) by Mammalian Phenotype (MP).

### **Description**

An R object that contains associations between Mammalian Phenotype terms and Mouse Entrez Genes. This data is prepared based on ftp://ftp.informatics.jax.org/pub/reports/MPheno\_OBO.ontology and ftp://ftp.informatics.jax.org/pub/reports/MGI\_PhenoGenoMP.rpt.

### Usage

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egMP.RData"))
```

### Value

an object of class "GS", a list with following components:

- set\_info: a matrix of nSet X 4 containing gene set information, where nSet is the number of gene sets (i.e. MP terms), and the 4 columns are "setID" (i.e. "Term ID"), "name" (i.e. "Term Name"), "namespace" and "distance"
- gs: a list of gene sets, each storing gene members thereof. Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"

#### References

Smith et al. (2009) The Mammalian Phenotype Ontology: enabling robust annotation and comparative analysis. *Wiley Interdiscip Rev Syst Biol Med*, 1:390-399.

### **Examples**

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egMP.RData"))
names(org.Mm.egMP)
```

org.Mm.egPS

Annotations Mouse Entrez Genes (EG) by phylostratific age (PS).

# Description

An R object that contains associations between phylostratific age and Mouse Entrez Genes. This data is prepared based on 1) SUPERFAMILY database which providing domain architecture assignments to all completely sequenced genomes including eukaryotic genomes; 2) ancestral domain architecture repertoires inferred by applying Dollo parsimony to eukaryotic part of sTOL, from which the most recent common ancestor of each domain architecture is determined. The domain architecture for an Entrez gene is the protein product with the longest length of amino acids. Thus, phylostratific age for a Mouse Entrez gene is the first appearance of its domain architecture along the branch from the eukaryotic ancestor to the mouse: the most recent common ancestor, how many steps it is away starting from the eukaryotic ancestor, and how far it is in the terms of the branch length from the eukaryotic ancestor.

org.Mm.string 35

#### **Usage**

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egPS.RData"))
```

#### Value

an object of class "GS", a list with following components:

- set\_info: a matrix of nSet X 4 containing gene set information, where nSet is the number of gene sets (i.e. phylogenetic placement along the branch starting from the eukaryotic ancestor), and the 4 columns are "setID" (i.e. "phylogenetic placement ID"), "name" (i.e. name for that placement in the form of "TaxonID:Name"), "namespace" (i.e. Rank for that placement) and "distance" (i.e. the branch length from the eukaryotic ancestor). Notably, since the sTOL is bifurcating with exactly two descendants (unlike the multifurcating nature of the NCBI taxonomy), an internal node in sTOL is either mapped onto a unique taxonomic identifier or left empty (assumedly a hypothetical unknown ancestor). In the latter case, hypothetical unknown ancestor is filled with the information in its next child with known taxonomic information.
- gs: a list of gene sets, each storing gene members thereof. Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"

#### References

Morais et all. (2011) SUPERFAMILY 1.75 including a domain-centric gene ontology method. *Nucleic Acids Res*, 39(Database issue):D427-34.

Fang et al. (2013) A daily-updated tree of (sequenced) life as a reference for genome research. *Scientific reports*, 3:2015.

# Examples

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.egPS.RData"))
names(org.Mm.egPS)
```

org.Mm.string

Mouse functional protein association network from STRING (version 9.0.5).

# Description

An igraph object that contains a functional protein association network in mouse. The network is extracted from the STRING database (version 9.0.5). Only those associations with medium confidence (score>=0.4) are retained.

```
load(url("http://dnet.r-forge.r-project.org/data/org.Mm.string.RData"))
```

36 visColoralpha

#### Value

an object of class "igraph" (see http://igraph.sourceforge.net/doc/R/aaa-igraph-package.html). It has attributes for both vertices and edges. Below are attributes for the vertices:

- name: unique id for the vertices
- seqid: protein seqid for the vertices
- geneid: Entrez geneid (if any) for the vertices
- symbol: gene symbol (if any) for the vertices
- description: gene description (if any) for the vertices

Below are attributes for the edges:

- neighborhood\_score: predictive score based on neighborhood data
- fusion\_score: predictive score based on fusion data
- cooccurence\_score: predictive score based on cooccurence data
- coexpression\_score: predictive score based on coexpression
- experimental\_score: predictive score based on experimental data
- database\_score: predictive score based on database
- textmining\_score: predictive score based on text mining
- combined\_score: combined score from all above predictive scores

#### References

Franceschini et al. (2013) STRING v9.1: protein-protein interaction networks, with increased coverage and integration. *Nucleic Acids Res*, 41:D808-D815.

### **Examples**

```
# not run
if(0){
load(url("http://dnet.r-forge.r-project.org/data/org.Hs.string.RData"))
org.Mm.string
}
```

visColoralpha

Function to add transparent (alpha) into colors

### **Description**

visColoralpha is supposed to add transparent (alpha) into colors.

### Usage

```
visColoralpha(col, alpha)
```

# **Arguments**

col	input colors. It can be vector of R color specifications, such as a color name (as listed by 'colors()), a hexadecimal string of the form "#rrggbb" or "#rrggbbaa"
alpha	numeric vector of values in the range [0, 1] for alpha transparency channel (0 means transparent and 1 means opaque)

visColormap 37

## Value

a vector of colors (after transparent being added)

## Note

none

## See Also

```
visColormap
```

## **Examples**

```
# 1) define "blue-white-red" colormap
palette.name <- visColormap(colormap="bwr")
# 2) use the return function "palette.name" to generate 10 colors spanning "bwr"
col <- palette.name(10)
# 3) add transparent (alpha=0.5)
cols <- visColoralpha(col, alpha=0.5)</pre>
```

visColormap

Function to define a colormap

# Description

visColormap is supposed to define a colormap. It returns a function, which will take an integer argument specifying how many colors interpolate the given colormap.

## Usage

```
visColormap(colormap = c("bwr", "jet", "gbr", "wyr", "br", "yr", "rainbow", "wb"))
```

# **Arguments**

colormap

short name for the colormap

## Value

• palette.name: a function that takes an integer argument for generating that number of colors interpolating the given sequence

## Note

The input colormap includes:

• "jet": jet colormap

• "bwr": blue-white-red

• "gbr": green-black-red

• "wyr": white-yellow-red

38 visHeatmap

- "br": black-red
- "yr": yellow-red
- "wb": white-black
- "rainbow": rainbow colormap, that is, red-yellow-green-cyan-blue-magenta
- Alternatively, any hyphen-separated HTML color names, e.g. "blue-black-yellow", "royalblue-white-sandybrown", "darkblue-lightblue-lightyellow-darkorange", "darkgreen-white-darkviolet", "darkgreen-lightgreen-lightpink-darkred". A list of standard color names can be found in http://html-color-codes.info/color-names

#### See Also

visColoralpha

#### **Examples**

```
# 1) define "blue-white-red" colormap
palette.name <- visColormap(colormap="bwr")
# 2) use the return function "palette.name" to generate 10 colors spanning "bwr"
palette.name(10)</pre>
```

visHeatmap

Function to visualise input data matrix using heatmap

#### **Description**

visHeatmap is supposed to visualise input data matrix using heatmap. Note: this heatmap displays matrix in a bottom-to-top direction

# Usage

```
visHeatmap(data, scale = c("none", "row", "column"),
  row.metric = c("none", "pearson", "spearman", "kendall", "euclidean", "manhattan", "cos", "n
  row.method = c("ward", "single", "complete", "average", "mcquitty", "median", "centroid"),
  column.metric = c("none", "pearson", "spearman", "kendall", "euclidean", "manhattan", "cos",
  column.method = c("ward", "single", "complete", "average", "mcquitty", "median", "centroid")
  colormap = c("bwr", "jet", "gbr", "wyr", "br", "yr", "rainbow", "wb"),
  ncolors = 64, zlim = NULL, row.cutree = NULL,
  row.colormap = c("rainbow"), column.cutree = NULL,
  column.colormap = c("rainbow"), ...)
```

# **Arguments**

an input gene-sample data matrix used for heatmap

a character indicating when the input matrix should be centered and scaled. It
can be one of "none" (no scaling), "row" (being scaled in the row direction),
"column" (being scaled in the column direction)

row.metric distance metric used to calculate the distance metric between rows. It can be one
of "none" (i.e. no dendrogram between rows), "pearson", "spearman", "kendall",
"euclidean", "manhattan", "cos" and "mi". See details at http://suprahex.
r-forge.r-project.org/sDistance.html

visHeatmap 39

row.method the agglomeration method used to cluster rows. This should be one of "ward",

"single", "complete", "average", "mcquitty", "median" or "centroid". See 'Note'

below for details

column.metric distance metric used to calculate the distance metric between columns. It can

be one of "none" (i.e. no dendrogram between rows), "pearson", "spearman", "kendall", "euclidean", "manhattan", "cos" and "mi". See details at http://

suprahex.r-forge.r-project.org/sDistance.html

column.method the agglomeration method used to cluster columns. This should be one of

"ward", "single", "complete", "average", "mcquitty", "median" or "centroid".

See 'Note' below for details

colormap short name for the colormap. It can be one of "jet" (jet colormap), "bwr" (blue-

white-red colormap), "gbr" (green-black-red colormap), "wyr" (white-yellow-red colormap), "br" (black-red colormap), "yr" (yellow-red colormap), "wb" (white-black colormap), and "rainbow" (rainbow colormap, that is, red-yellow-green-cyan-blue-magenta). Alternatively, any hyphen-separated HTML color names, e.g. "blue-black-yellow", "royalblue-white-sandybrown", "darkgreen-white-darkviolet". A list of standard color names can be found in http://

html-color-codes.info/color-names

ncolors the number of colors specified over the colormap

zlim the minimum and maximum z/patttern values for which colors should be plotted,

defaulting to the range of the finite values of z. Each of the given colors will be used to color an equispaced interval of this range. The midpoints of the intervals

cover the range, so that values just outside the range will be plotted

row.cutree an integer scalar specifying the desired number of groups being cut from the

row dendrogram. Note, this optional is only enabled when the row dengrogram

is built

row.colormap short name for the colormap to color-code the row groups (i.e. sidebar colors

used to annotate the rows)

column.cutree an integer scalar specifying the desired number of groups being cut from the

column dendrogram. Note, this optional is only enabled when the column den-

grogram is built

column.colormap

short name for the colormap to color-code the column groups (i.e. sidebar colors

used to annotate the columns)

... additional graphic parameters. Type ?heatmap for the complete list.

#### Value

invisible

## Note

The clustering methods are provided:

- "ward": Ward's minimum variance method aims at finding compact, spherical clusters
- "single": The single linkage method (which is closely related to the minimal spanning tree) adopts a 'friends of friends' clustering strategy
- "complete": The complete linkage method finds similar clusters

40 visHeatmapAdv

• "average", "mcquitty", "median", "centroid": These methods can be regarded as aiming for clusters with characteristics somewhere between the single and complete link methods. Two methods "median" and "centroid" are not leading to a monotone distance measure, or equivalently the resulting dendrograms can have so called inversions (which are hard to interpret)

#### See Also

visHeatmap

## **Examples**

```
# 1) generate data with three different distributions, each with an iid normal random matrix of 100 x 3
data <- cbind(matrix(rnorm(100*3,mean=0,sd=1), nrow=100, ncol=3),
matrix(rnorm(100*3,mean=0.5,sd=1), nrow=100, ncol=3),
matrix(rnorm(100*3,mean=-0.5,sd=1), nrow=100, ncol=3))
colnames(data) <- c("S1","S1","S2","S2","S2","S2","S3","S3","S3")

# 2) prepare colors for the column sidebar
lvs <- unique(colnames(data))
lvs_color <- visColormap(colormap="rainbow")(length(lvs))
my_ColSideColors <- sapply(colnames(data), function(x) lvs_color[x==lvs])

# 3) heatmap with row dendrogram (with 10 color-coded groups)
visHeatmap(data, row.metric="euclidean", row.method="average", colormap="gbr", zlim=c(-2,2),
ColSideColors=my_ColSideColors, row.cutree=10, row.colormap="jet", labRow=NA)</pre>
```

visHeatmapAdv

Function to visualise input data matrix using advanced heatmap

## **Description**

visHeatmapAdv is supposed to visualise input data matrix using advanced heatmap. It allows for adding multiple sidecolors in both columns and rows. Besides, the sidecolor can be automatically added via cutting histogram into groups. Note: this heatmap displays matrix in a top-to-bottom direction

```
visHeatmapAdv(data, scale = c("none", "row", "column"),
  Rowv = T, Colv = T,
  dendrogram = c("both", "row", "column", "none"),
  dist.metric = c("euclidean", "pearson", "spearman", "kendall", "manhattan", "cos", "mi"),
  linkage.method = c("complete", "ward", "single", "average", "mcquitty", "median", "centroid'
  colormap = c("bwr", "jet", "gbr", "wyr", "br", "yr", "rainbow", "wb"),
  ncolors = 64, zlim = NULL, RowSideColors = NULL,
  row.cutree = NULL, row.colormap = c("jet"),
  ColSideColors = NULL, column.cutree = NULL,
  column.colormap = c("jet"), ...)
```

visHeatmapAdv 41

## **Arguments**

data an input gene-sample data matrix used for heatmap

scale a character indicating when the input matrix should be centered and scaled. It

can be one of "none" (no scaling), "row" (being scaled in the row direction),

"column" (being scaled in the column direction)

Rowv determines if and how the row dendrogram should be reordered. By default, it

is TRUE, which implies dendrogram is computed and reordered based on row means. If NULL or FALSE, then no dendrogram is computed and no reordering is done. If a dendrogram, then it is used "as-is", ie without any reordering. If a vector of integers, then dendrogram is computed and reordered based on the

order of the vector

Colv determines if and how the column dendrogram should be reordered. Has the

options as the Rowv argument above and additionally when x is a square matrix, Colv = "Rowv" means that columns should be treated identically to the rows

dendrogram character string indicating whether to draw 'none', 'row', 'column' or 'both'

dendrograms. Defaults to 'both'. However, if Rowv (or Colv) is FALSE or NULL and dendrogram is 'both', then a warning is issued and Rowv (or Colv)

arguments are honoured

dist.metric distance metric used to calculate the distance metric between columns (or rows).

It can be one of "none" (i.e. no dendrogram between rows), "pearson", "spearman", "kendall", "euclidean", "manhattan", "cos" and "mi". See details at <a href="http://doi.org/10.1081/j.j.gov/">http://doi.org/10.1081/j.j.gov/</a>

//suprahex.r-forge.r-project.org/sDistance.html

linkage.method the agglomeration method used to cluster/linkages columns (or rows). This

should be one of "ward", "single", "complete", "average", "mcquitty", "median"

or "centroid". See 'Note' below for details

colormap short name for the colormap. It can be one of "jet" (jet colormap), "bwr" (blue-

white-red colormap), "gbr" (green-black-red colormap), "wyr" (white-yellow-red colormap), "br" (black-red colormap), "yr" (yellow-red colormap), "wb" (white-black colormap), and "rainbow" (rainbow colormap, that is, red-yellow-green-cyan-blue-magenta). Alternatively, any hyphen-separated HTML color names, e.g. "blue-black-yellow", "royalblue-white-sandybrown", "darkgreen-white-darkviolet". A list of standard color names can be found in http://

html-color-codes.info/color-names

ncolors the number of colors specified over the colormap

zlim the minimum and maximum z/patttern values for which colors should be plotted,

defaulting to the range of the finite values of z. Each of the given colors will be used to color an equispaced interval of this range. The midpoints of the intervals

cover the range, so that values just outside the range will be plotted

 $\label{lem:nowsidebars} \textbf{RowSideColors} \quad \textbf{NULL or a matrix of "numRowsidebars"} \ \textbf{X} \ \textbf{nrow}(\textbf{x}), \ \textbf{where "numRowsidebars"}$ 

stands for the number of sidebars annotating rows of x. This matrix contains the color names for vertical sidebars. By default, it sets to NULL. In this case, sidebars in rows can still be enabled by cutting the row dendrogram into several

clusters (see the next two parameters)

row.cutree an integer scalar specifying the desired number of groups being cut from the

row dendrogram. Note, this optional is only enabled when the ColSideColors is

**NULL** 

row.colormap short name for the colormap to color-code the row groups (i.e. sidebar colors

used to annotate the rows)

42 *visHeatmapAdv* 

ColSideColors

NULL or a matrix of ncol(x) X "numColsidebars", where "numColsidebars" stands for the number of sidebars annotating the columns of x. This matrix contains the color names for horizontal sidebars. By default, it sets to NULL. In this case, sidebars in columns can still be enabled by cutting the column dendrogram into several clusters (see the next two parameters)

column.cutree

an integer scalar specifying the desired number of groups being cut from the column dendrogram. Note, this optional is only enabled when the column dengrogram is built

column.colormap

short name for the colormap to color-code the column groups (i.e. sidebar colors used to annotate the columns)

. . .

additional graphic parameters. For most parameters, please refer to <a href="http://www.inside-r.org/packages/cran/gplots/docs/heatmap.2">http://www.inside-r.org/packages/cran/gplots/docs/heatmap.2</a>. For example, the parameters "srtRow" and "srtCol" to control the angle of row/column labels (in degrees from horizontal: 45 degrees for the column, 0 degrees for the row, by default), i.e. string rotation. The parameters "offsetRow" and "offsetCol" to indicate the number of character-width spaces to place between row/column labels and the edge of the plotting region. Unique to this function, there are two parameters "RowSideWidth" and RowSideLabelLocation, to respectively indicate the fraction of the row side width and the location (either bottom or top) of the row side labelling; the other two parameters "ColSideHeight" and "ColSideLabelLocation" for the column side height and the location (either left or right) of the column side labelling; and two parameters "RowSideBox" and "ColSideBox" to indicate whether there are boxes outside.

#### Value

invisible

# Note

The clustering/linkage methods are provided:

- "ward": Ward's minimum variance method aims at finding compact, spherical clusters
- "single": The single linkage method (which is closely related to the minimal spanning tree) adopts a 'friends of friends' clustering strategy
- "complete": The complete linkage method finds similar clusters
- "average", "mcquitty", "median", "centroid": These methods can be regarded as aiming for clusters with characteristics somewhere between the single and complete link methods. Two methods "median" and "centroid" are not leading to a monotone distance measure, or equivalently the resulting dendrograms can have so called inversions (which are hard to interpret)

#### See Also

visHeatmapAdv

# Examples

```
# 1) generate data with three different distributions, each with an iid normal random matrix of 100 x 3
data <- cbind(matrix(rnorm(100*3,mean=0,sd=1), nrow=100, ncol=3),
matrix(rnorm(100*3,mean=0.5,sd=1), nrow=100, ncol=3),
matrix(rnorm(100*3,mean=-0.5,sd=1), nrow=100, ncol=3))</pre>
```

visNet 43

colnames(data) <- c("S1\_R1", "S1\_R2", "S1\_R3", "S2\_R1", "S2\_R2", "S2\_R3", "S3\_R1", "S3\_R2", "S3\_R3")

```
# 2) heatmap after clustering both rows and columns
# 2a) shown with row and column dendrograms
visHeatmapAdv(data, dendrogram="both", colormap="gbr", zlim=c(-2,2), KeyValueName="log2(Ratio)",
add.expr=abline(v=(1:(ncol(data)+1))-0.5,col="white"),
lmat=rbind(c(4,3), c(2,1)), lhei=c(1,5), lwid=c(1,3))
# 2b) shown with row dendrogram only
visHeatmapAdv(data, dendrogram="row", colormap="gbr", zlim=c(-2,2))
# 2c) shown with column dendrogram only
visHeatmapAdv(data, dendrogram="column", colormap="gbr", zlim=c(-2,2))
# 3) heatmap after only clustering rows (with 2 color-coded groups)
visHeatmapAdv(data, Colv=FALSE, colormap="gbr", zlim=c(-2,2), row.cutree=2, row.colormap="jet", labRow=NA)
# 4) prepare colors for the column sidebar
# color for stages (S1-S3)
stages <- sub("_.*","",colnames(data))</pre>
lvs <- unique(stages)</pre>
lvs_color <- visColormap(colormap="rainbow")(length(lvs))</pre>
col_stages <- sapply(stages, function(x) lvs_color[x==lvs])</pre>
# color for replicates (R1-R3)
replicates <- sub(".*_","",colnames(data))</pre>
lvs <- unique(replicates)</pre>
lvs_color <- visColormap(colormap="rainbow")(length(lvs))</pre>
col_replicates <- sapply(replicates, function(x) lvs_color[x==lvs])</pre>
# combine both color vectors
ColSideColors <- cbind(col_stages,col_replicates)</pre>
colnames(ColSideColors) <- c("Stages", "Replicates")</pre>
# 5) heatmap without clustering on rows and columns but with the two sidebars in columns
visHeatmapAdv(data, Rowv=FALSE, Colv=FALSE, colormap="gbr", zlim=c(-2,2), density.info="density", tracecol-
```

visNet

Function to visualise a graph object of class "igraph" or "graphNEL"

# Description

visNet is supposed to visualise a graph object of class "igraph" or "graphNEL". It also allows the color-coding of vertices by providing the input pattern.

```
visNet(g, pattern = NULL,
  colormap = c("bwr", "jet", "gbr", "wyr", "br", "yr", "rainbow", "wb"),
  ncolors = 40, zlim = NULL, colorbar = T, newpage = T,
  glayout = layout.fruchterman.reingold,
  vertex.frame.color = NA, vertex.size = NULL,
  vertex.color = NULL, vertex.shape = NULL,
  vertex.label = NULL, vertex.label.cex = NULL,
  vertex.label.dist = NULL, vertex.label.color = "black",
  ...)
```

44 visNet

#### **Arguments**

g an object of class "igraph" or "graphNEL"

pattern a numeric vector used to color-code vertices/nodes. Notably, if the input vector

contains names, then these names should include all node names of input graph, i.e. V(g)\$name, since there is a mapping operation. After mapping, the length of the patern vector should be the same as the number of nodes of input graph; otherwise, this input pattern will be ignored. The way of how to color-code is to map values in the pattern onto the whole colormap (see the next arguments:

colormap, ncolors, zlim and colorbar)

colormap short name for the colormap. It can be one of "jet" (jet colormap), "bwr" (blue-

white-red colormap), "gbr" (green-black-red colormap), "wyr" (white-yellow-red colormap), "br" (black-red colormap), "yr" (yellow-red colormap), "wb" (white-black colormap), and "rainbow" (rainbow colormap, that is, red-yellow-green-cyan-blue-magenta). Alternatively, any hyphen-separated HTML color names, e.g. "blue-black-yellow", "royalblue-white-sandybrown", "darkgreen-white-darkviolet". A list of standard color names can be found in http://

html-color-codes.info/color-names

ncolors the number of colors specified over the colormap

zlim the minimum and maximum z/patttern values for which colors should be plotted,

defaulting to the range of the finite values of z. Each of the given colors will be used to color an equispaced interval of this range. The midpoints of the intervals

cover the range, so that values just outside the range will be plotted

colorbar logical to indicate whether to append a colorbar. If pattern is null, it always sets

to false

newpage logical to indicate whether to open a new page. By default, it sets to true for

opening a new page

glayout either a function or a numeric matrix configuring how the vertices will be placed

on the plot. If layout is a function, this function will be called with the graph as the single parameter to determine the actual coordinates. This function can be one of "layout.auto", "layout.random", "layout.circle", "layout.sphere", "layout

out.fruchterman.reingold", "layout.kamada.kawai", "layout.spring", "layout.reingold.tilford",

"layout.fruchterman.reingold.grid", "layout.lgl", "layout.graphopt", "layout.svd" and "layout.norm". A full explanation of these layouts can be found in  ${\tt http}$ :

//igraph.sourceforge.net/doc/R/layout.html

vertex.frame.color

the color of the frame of the vertices. If it is NA, then there is no frame

vertex. size the size of each vertex. If it is a vector, each vertex may differ in size

vertex.color the fill color of the vertices. If it is NA, then there is no fill color. If the pattern

is given, this setup will be ignored

vertex.shape the shape of each vertex. It can be one of "circle", "square", "csquare", "rect-

angle", "crectangle", "vrectangle", "pie" (http://igraph.sourceforge.net/doc/R/vertex.shape.pie.html), "sphere", and "none". If it sets to NULL,

these vertices with negative will be "csquare" and the rest "circle".

vertex.label the label of the vertices. If it is NA, then there is no label. The default vertex

labels are the name attribute of the nodes

vertex.label.cex

the font size of vertex labels.

visNetArc 45

```
vertex.label.dist
```

the distance of the label from the center of the vertex. If it is 0 then the label is centered on the vertex. If it is 1 then the label is displayed beside the vertex.

vertex.label.color

the color of vertex labels.

additional graphic parameters. See <a href="http://igraph.sourceforge.net/doc/R/plot.graph.html">http://igraph.sourceforge.net/doc/R/plot.graph.html</a> for the complete list.

#### Value

invisible

#### Note

none

#### See Also

dNetFind

## **Examples**

```
# 1) generate a random graph according to the ER model
g <- erdos.renyi.game(100, 1/100)

# 2) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, V(g), knn=0)

# 3) visualise the subg with vertices being color-coded by the pattern
pattern <- runif(vcount(subg))
names(pattern) <- V(subg)$name
visNet(g=subg, pattern=pattern, colormap="bwr", vertex.shape="sphere")</pre>
```

visNetArc

Function to visualise an igraph object via arc diagram

# **Description**

visNetArc is supposed to visualise a graph object of class "igraph" via arc diagram in one-dimensional layout. More precisely, it displays vertices (nodes) along an axis, with edges linked by arcs. With proper ordering of vertices (e.g. according to communities and degrees), arc diagram is able to identify clusters and bridges (as effective as two-dimensional layout). One advantage of using arc diagram is to allow for easy annotations along vertices.

```
visNetArc(g, orientation = c("vertical", "horizontal"),
  newpage = T, ordering = NULL, labels = V(g)$name,
  vertex.label.color = "black", vertex.label.cex = 1,
  vertex.color = "transparent",
  vertex.frame.color = "black",
  vertex.size = log(degree(g)) + 0.1, vertex.pch = 21,
  vertex.lwd = 1, edge.color = "grey", edge.width = 1,
  edge.lty = 1, ...)
```

46 visNetArc

#### **Arguments**

an object of class "igraph" the orientation of the plots. It can be either "vertical" (default) or "horizontal" orientation newpage logical to indicate whether to open a new page. By default, it sets to true for opening a new page a numeric vector about the ordering of vertices. It is optional. It is highly recordering ommend to order vertices according to communities and degrees labels the label of the vertices. The default vertex labels are the name attribute of the vertex.label.color the color of vertex labels vertex.label.cex the font size of vertex labels vertex.color the fill color of the vertices. The default vertex colors are transparent vertex.frame.color the color of the frame of the vertices. The default vertex frame colors are black the size of each vertex. By default, it is decided according to node degrees vertex.size the shape of each vertex. Either an integer specifying a symbol or a single charvertex.pch acter to be used as the default in plotting points. See <a href="http://www.statmethods">http://www.statmethods</a>. net/advgraphs/parameters.html vertex.lwd line width for the vertices (default 1) edge.color the color of the edges (default "grey") edge.width line width for the edges (default 1) line type for the edges (default 1) edge.lty additional graphic parameters associated with 'mtext'

#### Value

invisible

# Note

none

# See Also

visNet

# Examples

```
# 1) generate a random graph according to the ER model
g <- erdos.renyi.game(100, 1/80)

# 2) produce the induced subgraph only based on the nodes in query
g <- dNetInduce(g, V(g), knn=0)

# 3) color nodes according to communities identified via a spin-glass model and simulated annealing
com <- spinglass.community(g, spins=4)
vgroups <- com$membership
palette.name <- visColormap(colormap="rainbow")</pre>
```

visNetCircle 47

```
vcolors <- palette.name(length(com))[vgroups]

# 4) size nodes according to degrees
vdegrees <- igraph::degree(g)

# 5) sort nodes: first by communities and then degrees
tmp <- data.frame(ind=1:vcount(g), vgroups, vdegrees)
ordering <- tmp[order(vgroups,vdegrees),]$ind

# 6) visualise graph using 1-dimensional arc diagram
visNetArc(g, ordering=ordering, labels=V(g)$name, vertex.label.color=vcolors, vertex.color=vcolors, vertex

# 7) as comparison, also visualise graph on 2-dimensional layout
visNet(g, colormap="bwr", layout=layout.kamada.kawai(g), vertex.label=V(g)$name, vertex.color=vcolors, vertex.color=vcolor=vcolor=vcolor=vcolor=vcolor=vcolor=vcolor=vcolor=vcolor=vcolor=vcolor=vcolor=vcolor=vcolor=vcolor=vco
```

visNetCircle

Function to visualise an igraph object via circle diagram

## **Description**

visNetCircle is supposed to visualise a graph object of class "igraph" via circle diagram. For better visualisation, ordering of vertices is determined according to communities and degrees.

# Usage

```
visNetCircle(g, com, circles = c("single", "multiple"),
  newpage = T, ordering = NULL,
  colormap = c("rainbow", "bwr", "jet", "gbr", "wyr", "br", "yr", "wb"),
  vertex.label = V(g)$name,
  vertex.size = log(igraph::degree(g)) + 2,
  vertex.label.color = "black", vertex.label.cex = 0.6,
  vertex.label.dist = 0.75, vertex.shape = "sphere",
  edge.width = 1, edge.lty = 1,
  edge.color.within = "grey",
  edge.color.crossing = "black", mark.shape = 1,
  mark.expand = 10, ...)
```

## **Arguments**

g	an object of class "igraph"
COM	an object of class "communities" (see http://igraph.sourceforge.net/doc/R/communities.html)
circles	how circles are drawn in the plot. It can be either "single" for all communities being drawn in a single circle (by default) or "multiple" for communities being drawn in the different circles (i.e. one circle per community)
newpage	logical to indicate whether to open a new page. By default, it sets to true for opening a new page
ordering	a numeric vector about the ordering of vertices. It is optional. It is highly recommend to order vertices according to communities and degrees

48 visNetCircle

colormap short name for the colormap. It can be one of "jet" (jet colormap), "bwr" (bluewhite-red colormap), "gbr" (green-black-red colormap), "wyr" (white-yellowred colormap), "br" (black-red colormap), "yr" (yellow-red colormap), "wb" (white-black colormap), and "rainbow" (rainbow colormap, that is, red-yellowgreen-cyan-blue-magenta). Alternatively, any hyphen-separated HTML color names, e.g. "blue-black-yellow", "royalblue-white-sandybrown", "darkgreenwhite-darkviolet". A list of standard color names can be found in http:// html-color-codes.info/color-names the label of the vertices. The default vertex labels are the name attribute of the vertex.label the size of each vertex. By default, it is decided according to node degrees vertex.size vertex.label.color the color of vertex labels vertex.label.cex the font size of vertex labels vertex.label.dist the distance of the label from the center of the vertex. If it is 0 then the label is centered on the vertex. If it is 1 then the label is displayed beside the vertex. the shape of each vertex. It can be one of "circle", "square", "csquare", "rectvertex.shape angle", "crectangle", "pie" (http://igraph.sourceforge.net/ doc/R/vertex.shape.pie.html), "sphere", and "none". If it sets to NULL, these vertices with negative will be "csquare" and the rest "circle". edge.width line width for the edges (default 1) edge.lty line type for the edges (default 1) edge.color.within the color for edges within a community (default "grey") edge.color.crossing the color for edges between communities (default "black") mark.shape a numeric scalar or vector controlling the smoothness of the vertex group marking polygons. Its possible values are between -1 (fully polygons) and 1 (fully smoothness) a numeric scalar or vector, the size of the border around the marked vertex mark.expand groups additional graphic parameters. See http://igraph.sourceforge.net/doc/

R/plot.graph.html for the complete list.

## Value

invisible

## Note

none

## See Also

visNet

visNetMul 49

#### **Examples**

```
# 1) generate a random graph according to the ER model
g <- erdos.renyi.game(100, 1/80)</pre>
# 2) produce the induced subgraph only based on the nodes in query
g <- dNetInduce(g, V(g), knn=0)</pre>
# 3) color nodes according to communities identified via a spin-glass model and simulated annealing
com <- spinglass.community(g, spins=4)</pre>
vgroups <- com$membership</pre>
palette.name <- visColormap(colormap="rainbow")</pre>
mcolors <- palette.name(length(com))</pre>
vcolors <- mcolors[vgroups]</pre>
# 4) size nodes according to degrees
vdegrees <- igraph::degree(g)</pre>
# 5) sort nodes: first by communities and then degrees
tmp<-data.frame(ind=1:vcount(g), vgroups, vdegrees)</pre>
ordering <- tmp[order(vgroups, vdegrees),]$ind</pre>
# 6) visualise graph using circle diagram
# 6a) drawn into a single circle
#visNetCircle(g=g, colormap="bwr", com=com, ordering=ordering, vertex.label=V(g)$name)
# 6b) drawn into multlpe circles (one circle per community)
#visNetCircle(g=g, colormap="bwr", com=com, circles="multiple", ordering=ordering, vertex.label=V(g)$name)
# 7) as comparison, also visualise graph on 2-dimensional layout
mark.groups <- communities(com)</pre>
#mark.col <- visColoralpha(mcolors, alpha=0.2)</pre>
#mark.border <- visColoralpha(mcolors, alpha=0.2)</pre>
edge.color <- c("grey", "black")[crossing(com,g)+1]</pre>
#visNet(g, colormap="bwr", glayout=layout.fruchterman.reingold, vertex.color=vcolors, vertex.frame.color=vc
```

visNetMul

Function to visualise the same graph but with multiple graph node colorings according to input data matrix

## **Description**

visNetMul is supposed to visualise the same graph but with multiple colorings according to input data matrix

```
visNetMul(g, data, height = 7, margin = rep(0.1, 4),
border.color = "#EEEEEE",
colormap = c("bwr", "jet", "gbr", "wyr", "br", "yr", "rainbow", "wb"),
ncolors = 40, zlim = NULL, colorbar = T,
colorbar.fraction = 0.25, newpage = T,
glayout = layout.fruchterman.reingold, mtext.side = 3,
```

50 visNetMul

```
mtext.adj = 0, mtext.cex = 1, mtext.font = 2,
mtext.col = "black", ...)
```

#### **Arguments**

g an object of class "igraph" or "graphNEL"

data an input data matrix used to color-code vertices/nodes. One column corresponds

to one graph node coloring. The input matrix must have row names, and these names should include all node names of input graph, i.e. V(g)\$name, since there is a mapping operation. After mapping, the length of the patern vector should be the same as the number of nodes of input graph. The way of how to color-code is to map values in the pattern onto the whole colormap (see the next arguments:

colormap, ncolors, zlim and colorbar)

height a numeric value specifying the height of device

margin margins as units of length 4 or 1 border.color the border color of each figure

colormap short name for the colormap. It can be one of "jet" (jet colormap), "bwr" (blue-

white-red colormap), "gbr" (green-black-red colormap), "wyr" (white-yellow-red colormap), "br" (black-red colormap), "yr" (yellow-red colormap), "wb" (white-black colormap), and "rainbow" (rainbow colormap, that is, red-yellow-green-cyan-blue-magenta). Alternatively, any hyphen-separated HTML color names, e.g. "blue-black-yellow", "royalblue-white-sandybrown", "darkgreen-white-darkviolet". A list of standard color names can be found in http://

html-color-codes.info/color-names

ncolors the number of colors specified over the colormap

zlim the minimum and maximum z/patttern values for which colors should be plotted,

defaulting to the range of the finite values of z. Each of the given colors will be used to color an equispaced interval of this range. The midpoints of the intervals

cover the range, so that values just outside the range will be plotted

colorbar logical to indicate whether to append a colorbar. If pattern is null, it always sets

to false

 ${\tt colorbar.fraction}$ 

the relative fraction of colorbar block against the figure block

newpage logical to indicate whether to open a new page. By default, it sets to true for

opening a new page

glayout either a function or a numeric matrix configuring how the vertices will be placed

on the plot. If layout is a function, this function will be called with the graph as the single parameter to determine the actual coordinates. This function can be one of "layout.auto", "layout.random", "layout.circle", "layout.sphere", "layout

out.fruchterman.reingold", "layout.kamada.kawai", "layout.spring", "layout.reingold.tilford",

"layout.fruchterman.reingold.grid", "layout.lgl", "layout.graphopt", "layout.svd" and "layout.norm". A full explanation of these layouts can be found in <a href="http://dx.doi.org/10.1001/j.j.graphopt">http://dx.doi.org/10.1001/j.j.graphopt</a>", "layout.svd" and "layout.norm".

//igraph.sourceforge.net/doc/R/layout.html

mtext.side on which side of the mtext plot (1=bottom, 2=left, 3=top, 4=right)

mtext.adj the adjustment for mtext alignment (0 for left or bottom alignment, 1 for right

or top alignment)

mtext.cex the font size of mtext labels
mtext.font the font weight of mtext labels

visNetReorder 51

```
mtext.col the color of mtext labels
... additional graphic parameters. See http://igraph.sourceforge.net/doc/
R/plot.graph.html for the complete list.
```

#### Value

invisible

# Note

none

## See Also

visNet

#### **Examples**

```
# 1) generate a random graph according to the ER model
g <- erdos.renyi.game(100, 1/80)

# 2) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, V(g), knn=0)

# 3) visualise the module with vertices being color-coded by scores
nnodes <- vcount(subg)
nsamples <- 10
data <- matrix(runif(nnodes*nsamples), nrow=nnodes, ncol=nsamples)
rownames(data) <- V(subg)$name
visNetMul(g=subg, colormap="bwr", data=data, glayout=layout.fruchterman.reingold)</pre>
```

visNetReorder

Function to visualise the multiple graph colorings reorded within a sheet-shape rectangle grid

# Description

visNetReorder is supposed to visualise the multiple graph colorings reorded within a sheet-shape rectangle grid

```
visNetReorder(g, data, sReorder, height = 7,
  margin = rep(0.1, 4), border.color = "#EEEEEE",
  colormap = c("bwr", "jet", "gbr", "wyr", "br", "yr", "rainbow", "wb"),
  ncolors = 40, zlim = NULL, colorbar = T,
  colorbar.fraction = 0.5, newpage = T,
  glayout = layout.fruchterman.reingold, mtext.side = 3,
  mtext.adj = 0, mtext.cex = 1, mtext.font = 2,
  mtext.col = "black", ...)
```

52 visNetReorder

## **Arguments**

g an object of class "igraph" or "graphNEL"

data an input data matrix used to color-code vertices/nodes. One column corresponds

to one graph node coloring. The input matrix must have row names, and these names should include all node names of input graph, i.e. V(g)\$name, since there is a mapping operation. After mapping, the length of the patern vector should be the same as the number of nodes of input graph. The way of how to color-code is to map values in the pattern onto the whole colormap (see the next arguments:

colormap, ncolors, zlim and colorbar)

height a numeric value specifying the height of device

sReorder an object of class "sReorder"
margin margins as units of length 4 or 1
border.color the border color of each figure

colormap short name for the colormap. It can be one of "jet" (jet colormap), "bwr" (blue-

white-red colormap), "gbr" (green-black-red colormap), "wyr" (white-yellow-red colormap), "br" (black-red colormap), "yr" (yellow-red colormap), "wb" (white-black colormap), and "rainbow" (rainbow colormap, that is, red-yellow-green-cyan-blue-magenta). Alternatively, any hyphen-separated HTML color names, e.g. "blue-black-yellow", "royalblue-white-sandybrown", "darkgreen-white-darkviolet". A list of standard color names can be found in http://

html-color-codes.info/color-names

ncolors the number of colors specified over the colormap

zlim the minimum and maximum z/patttern values for which colors should be plotted,

defaulting to the range of the finite values of z. Each of the given colors will be used to color an equispaced interval of this range. The midpoints of the intervals

cover the range, so that values just outside the range will be plotted

colorbar logical to indicate whether to append a colorbar. If pattern is null, it always sets

to false

colorbar.fraction

the relative fraction of colorbar block against the figure block

newpage logical to indicate whether to open a new page. By default, it sets to true for

opening a new page

glayout either a function or a numeric matrix configuring how the vertices will be placed

on the plot. If layout is a function, this function will be called with the graph as the single parameter to determine the actual coordinates. This function can be one of "layout.auto", "layout.random", "layout.circle", "layout.sphere", "layout.auto", "layout.a

out.fruchterman.reingold", "layout.kamada.kawai", "layout.spring", "layout.reingold.tilford", "layout.spring", "layout.reingold.tilford", "layout.spring", "l

"layout.fruchterman.reingold.grid", "layout.lgl", "layout.graphopt", "layout.svd" and "layout.norm". A full explanation of these layouts can be found in <a href="http://dx.doi.org/10.1001/j.j.graphopt">http://dx.doi.org/10.1001/j.j.graphopt</a>", "layout.graphopt", "layout.svd" and "layout.norm".

//igraph.sourceforge.net/doc/R/layout.html

mtext.side on which side of the mtext plot (1=bottom, 2=left, 3=top, 4=right)

mtext.adj the adjustment for mtext alignment (0 for left or bottom alignment, 1 for right

or top alignment)

... additional graphic parameters. See http://igraph.sourceforge.net/doc/

R/plot.graph.html for the complete list.

visRunES 53

#### Value

invisible

#### Note

none

#### See Also

```
visNet, dNetReorder
```

# **Examples**

```
# 1) generate a random graph according to the ER model
g <- erdos.renyi.game(100, 1/100)

# 2) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, V(g), knn=0)

# 3) reorder the module with vertices being color-coded by input data
nnodes <- vcount(subg)
nsamples <- 10
data <- matrix(runif(nnodes*nsamples), nrow=nnodes, ncol=nsamples)
rownames(data) <- V(subg)$name
sReorder <- dNetReorder(g=subg, data, feature="node", node.normalise="none")

# 4) visualise the module with vertices being color-coded by input data
visNetReorder(g=subg, colormap="bwr", data=data, sReorder)</pre>
```

visRunES

Function to visualise running enrichment score for a given sample and a gene set

# **Description**

eView is supposed to visualise running enrichment score for a given sample and a gene set. To help understand the underlying running enrichment score, the input gene scores are also displayed. Positions for members in the given gene set are color-coded in both displays (red line for the positive gene scores, and green line for the negative).

# Usage

```
visRunES(eTerm, which_sample = 1,
  which_term = "GO:0006281", weight = 1,
  orientation = c("vertical", "horizontal"), newpage = T)
```

# **Arguments**

```
eTerm an object of class "eTerm"

which_sample which sample will be used. It can be index or sample names

which_term which term will be used. It can be index or term ID or term names
```

weight type of score weigth. It can be "0" for unweighted (an equivalent to Kolmogorov-

Smirnov, only considering the rank), "1" for weighted by input gene score (by

default), and "2" for over-weighted, and so on

orientation the orientation of the plots. It can be either "vertical" (default) or "horizontal"

newpage logical to indicate whether to open a new page. By default, it sets to true for

opening a new page

#### Value

invisible

## Note

none

#### See Also

eCal, eView

## **Examples**

```
#visRunES(eTerm, which_sample=1, which_term=1)
```

visTreeBootstrap

Function to build and visualise the bootstrapped tree

## **Description**

visTreeBootstrap is supposed to build the tree, perform bootstrap analysis and visualise the bootstrapped tree. It returns an object of class "phylo". For easy downstream analysis, the bootstrapped tree is rerooted either at the internal node with the miminum bootstrap/confidence value or at any customised internal node.

# Usage

```
visTreeBootstrap(data,
  algorithm = c("nj", "fastme.ols", "fastme.bal"),
  metric = c("euclidean", "pearson", "spearman", "cos", "manhattan", "kendall", "mi"),
  num.bootstrap = 100, consensus = FALSE,
  consensus.majority = 0.5, reroot = "min.bootstrap",
  plot.phylo.arg = NULL, nodelabels.arg = NULL,
  visTree = T, verbose = T, ...)
```

# **Arguments**

data an input data matrix used to build the tree. The built tree describes the relation-

ships between rows of input matrix

algorithm the tree-building algorithm. It can be one of "nj" for the neighbor-joining tree es-

timation, "fastme.ols" for the minimum evolution algorithm with ordinary least-squares (OLS) fitting of a metric to a tree structure, and "fastme.bal" for the minimum evolution algorithm under a balanced (BAL) weighting scheme

metric distance metric used to calculate a distance matrix between rows of input matrix. It can be: "pearson" for pearson correlation, "spearman" for spearman rank

trix. It can be: "pearson" for pearson correlation, "spearman" for spearman rank correlation, "kendall" for kendall tau rank correlation, "euclidean" for euclidean distance, "manhattan" for cityblock distance, "cos" for cosine similarity, "mi"

for mutual information

num.bootstrap an integer specifying the number of bootstrap replicates

consensus logical to indicate whether to return the consensus tree. By default, it sets to false

for not doing so. Note: if true, there will be no visualisation of the bootstrapped

tree

consensus.majority

a numeric value between 0.5 and 1 (or between 50 and 100) giving the propor-

tion for a clade to be represented in the consensus tree

reroot determines if and how the bootstrapped tree should be rerooted. By default, it is

"min.bootstrap", which implies that the bootstrapped tree will be rerooted at the internal node with the miminum bootstrap/confidence value. If it is an integer between 1 and the number of internal nodes, the tree will be rerooted at the

internal node with this index value

 $\verb|plot.phylo.arg| a list of main parameters used in the function "ape::plot.phylo" \verb|http://www.|$ 

inside-r.org/packages/cran/ape/docs/plot.phylo. See 'Note' below for

details on the parameters

nodelabels.arg a list of main parameters used in the function "ape::nodelabels" http://www.

inside-r.org/packages/cran/ape/docs/nodelabels. See 'Note' below for

details on the parameters

visTree logical to indicate whether the bootstrap tree will be visualised. By default, it

sets to true for display. Note, the consensus tree can not be enabled for visuali-

sation

verbose logical to indicate whether the messages will be displayed in the screen. By

default, it sets to true for display

... additional "ape::plot.phylo" parameters

## Value

an object of class "phylo". It can return a bootstrapped tree or a consensus tree (if enabled): When a bootstrapped tree is returned (also visualised by default), the "phylo" object has a list with following components:

- Nnode: the number of internal nodes
- node.label: the labels for internal nodes. Here, each internal node is associated with the bootstrap value
- tip.label: the lables for tip nodes. Tip labels come from the row names of the input matrix, but are not necessarily the same order as they appear in the input matrix
- edge: a two-column matrix describing the links between tree nodes (including internal and tip nodes)
- edge.length: a vector indicating the edge length in the 'edge'
- Note: the tree structure is indexed with 1:Ntip for tip nodes, and (Ntip+1):(Ntip+Nnode) for internal nodes, where Ntip is the number of tip nodes and Nnode for the number of internal nodes. Moreover, nrow(data) = Ntip = Nnode 2.

When a consensus tree is returned (no visualisation), the "phylo" object has a list with following components:

- Nnode: the number of internal nodes
- tip.label: the lables for tip nodes. Tip labels come from the row names of the input matrix, but are not necessarily the same order as they appear in the input matrix
- edge: a two-column matrix describing the links between tree nodes (including internal and tip nodes)

#### Note

A list of main parameters used in the function "ape::plot.phylo":

- "type": a character string specifying the type of phylogeny to be drawn; it must be one of "phylogram" (the default), "cladogram", "fan", "unrooted", "radial" or any unambiguous abbreviation of these
- "direction": a character string specifying the direction of the tree. Four values are possible: "rightwards" (the default), "leftwards", "upwards", and "downwards"
- "lab4ut": (= labels for unrooted trees) a character string specifying the display of tip labels for unrooted trees: either "horizontal" where all labels are horizontal (the default), or "axial" where the labels are displayed in the axis of the corresponding terminal branches. This option has an effect only if type = "unrooted"
- "edge.color": a vector of mode character giving the colours used to draw the branches of the plotted phylogeny. These are taken to be in the same order than the component edge of phy. If fewer colours are given than the length of edge, then the colours are recycled
- "edge.width": a numeric vector giving the width of the branches of the plotted phylogeny. These are taken to be in the same order than the component edge of phy. If fewer widths are given than the length of edge, then these are recycled
- "edge.lty": same than the previous argument but for line types; 1: plain, 2: dashed, 3: dotted, 4: dotdash, 5: longdash, 6: twodash
- "font": an integer specifying the type of font for the labels: 1 (plain text), 2 (bold), 3 (italic, the default), or 4 (bold italic)
- "cex": a numeric value giving the factor scaling of the tip and node labels (Character EXpansion). The default is to take the current value from the graphical parameters
- "adj": a numeric specifying the justification of the text strings of the labels: 0 (left-justification), 0.5 (centering), or 1 (right-justification). This option has no effect if type="unrooted". If NULL (the default) the value is set with respect of direction (see details)
- "srt": a numeric giving how much the labels are rotated in degrees (negative values are allowed resulting in clock-like rotation); the value has an effect respectively to the value of direction (see Examples). This option has no effect if type="unrooted"
- "no.margin": a logical. If TRUE, the margins are set to zero and the plot uses all the space of the device
- "label.offset": a numeric giving the space between the nodes and the tips of the phylogeny and their corresponding labels. This option has no effect if type="unrooted"
- "rotate.tree": for "fan", "unrooted", or "radial" trees: the rotation of the whole tree in degrees (negative values are accepted

A list of main parameters used in the function "ape::nodelabels":

• "text": a vector of mode character giving the text to be printed. By default, the labels for internal nodes (see "node.label"), that is, the bootstrap values associated with internal nodes

• "node": a vector of mode numeric giving the numbers of the nodes where the text or the symbols are to be printed. By default, indexes for internal nodes, that is, (Ntip+1):(Ntip+Nnode), where Ntip is the number of tip nodes and Nnode for the number of internal nodes

- "adj": one or two numeric values specifying the horizontal and vertical, respectively, justification of the text or symbols. By default, the text is centered horizontally and vertically. If a single value is given, this alters only the horizontal position of the text
- "frame": a character string specifying the kind of frame to be printed around the text. This must be one of "rect" (the default), "circle", "none", or any unambiguous abbreviation of these
- "cex": a numeric value giving the factor scaling of the tip and node labels (Character EXpansion). The default is to take the current value from the graphical parameters
- "font": an integer specifying the type of font for the labels: 1 (plain text), 2 (bold), 3 (italic, the default), or 4 (bold italic)
- "col": a character string giving the color to be used for the text or the plotting symbols; this is eventually recycled
- "bg": a character string giving the color to be used for the background of the text frames or of the plotting symbols if it applies; this is eventually recycled. It can be one of "jet" (jet colormap), "bwr" (blue-white-red colormap), "gbr" (green-black-red colormap), "wyr" (white-yellow-red colormap), "br" (black-red colormap), "yr" (yellow-red colormap), "wb" (white-black colormap), and "rainbow" (rainbow colormap, that is, red-yellow-green-cyan-blue-magenta). Alternatively, any hyphen-separated HTML color names, e.g. "blue-black-yellow", "royalblue-white-sandybrown", "darkgreen-white-darkviolet". A list of standard color names can be found in http://html-color-codes.info/color-names

## See Also

visTreeBootstrap

# **Examples**

```
# 1) generate an iid normal random matrix of 100x10
data <- matrix( rnorm(100*10, mean=0, sd=1), nrow=100, ncol=10)</pre>
colnames(data) <- paste(rep(S,10), seq(1:10), sep="")</pre>
data <- t(data)</pre>
# 2) build neighbor-joining tree with bootstrap values and visualise it by default
visTreeBootstrap(data, metric="mi")
# 3) only display those internal nodes with bootstrap values > 30
# 3a) generate the bootstrapped tree (without visualisation)
tree_bs <- visTreeBootstrap(data, visTree=FALSE)</pre>
# 3b) look at the bootstrap values and ordered row names of input matrix
# the bootstrap values
tree_bs$node.label
# ordered row names of input matrix
tree_bs$tip.label
# 3c) determine internal nodes that should be displayed
Ntip <- length(tree_bs$tip.label) # number of tip nodes</pre>
Nnode <- length(tree_bs$node.label) # number of internal nodes</pre>
flag <- as.numeric(tree_bs$node.label) > 30
text <- tree_bs$node.label[flag]</pre>
node <- Ntip + (1:Nnode)[flag]</pre>
visTreeBootstrap(data, nodelabels.arg=list(text=text,node=node))
```

# 4) obtain the consensus tree
tree\_cons <- visTreeBootstrap(data, consensus=TRUE, consensus.majority=0.5)</pre>

# Index

*Topic datasets	org.Mm.egGOBP, 29	
CLL, 2	org.Mm.egGOCC, 30	
Hiratani_TableS1, 25	org.Mm.egGOMF, 31	
org.At.string,26	org.Mm.egHPMI,31	
org.Hs.string,27	org.Mm.egHPON, 32	
org.Mm.eg, 28	org.Mm.egHPPA, 33	
org.Mm.egDO, 28	org.Mm.egMP, 34	
org.Mm.egGOBP, 29	org.Mm.egPS, 34	
org.Mm.egGOCC, 30	org.Mm.string,35	
org.Mm.egGOMF, 31		
org.Mm.egHPMI, 31	RT (Hiratani_TableS1), 25	
org.Mm.egHPON, 32		
org.Mm.egHPPA, 33	visColoralpha, 36, 38	
org.Mm.egMP, 34	visColormap, 37, 37	
org.Mm.egPS, 34	visHeatmap, 38, 40	
org.Mm.string, 35	visHeatmapAdv, $40,42$	
5	visNet, 10, 43, 46, 48, 51, 53	
CLL, 2	visNetArc, 45	
CpG (Hiratani_TableS1), 25	visNetCircle,47	
	visNetMul, 49	
dBUMfit, 3, 6, 14	visNetReorder, 17,51	
dBUMscore, 4, 5, 14	visRunES, 53	
dCommSignif, 6, 7	visTreeBootstrap, 54, 57	
dContrast, 7		
dFDRscore, $8,20$		
dNetConfidence, 9		
dNetFind, 10, 12, 14, 45		
dNetInduce, 12, <i>13</i> , <i>18</i>		
dNetPipeline, 9, 13		
dNetReorder, 15, 53		
dPvalAggregate, 17, 18		
dRWR, 18		
dSVDSignif, 9, 19		
eCal, 20, 24, 25, 54		
eView, 22, 23, 54		
eWrite, 22, 24		
EX (Hiratani_TableS1), 25		
Hiratani_TableS1, 25		
org.At.string, 26		
org.Hs.string, 27		
org.Mm.eg, 28		
-		
org.Mm.egDO, 28		