# Package 'dnet'

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Type Package

**Title** an open-source R package for integrative analysis of digitised data in terms of network, ontology and evolution

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**Depends** R (>= 3.1.0), igraph, supraHex

Imports graph, Rgraphviz, Matrix, Biobase

Suggests limma, survival

**Description** The 'dnet' package is initiated to fill in the need of an open-source tool for digitised data in an integrative manner in terms of network, ontology and evolution. More specifically, dnet intends to analyse the biological network whose nodes/genes are associated with digitised information such as expression levels across samples. To help make sense of identified networks, enrichment analysis is also supported using a wide variety of pre-compiled ontologies and phylostratific age information in major organisms including human, mouse, rat, chicken, C.elegans, fruit fly, zebrafish and arabidopsis. In summary, dnet aims to deliver an eye-intuitive tool with rich visuals but less inputs.

URL http://dnet.r-forge.r-project.org

Collate 'dGSEA.r' 'dGSEAview.r' 'dGSEAwrite.r' 'visGSEA.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' 'dRWRcontact.r' 'dRWRpipeline.r' 'dContrast.r' 'dCommSignif.r' 'dSVDsignif.r' 'dFDRscore.r' 'dDAGinduce.r' 'dDAGreverse.r' 'dDAGroot.r' 'dDAGlevel.r' 'dDAGannotate.r' 'dDAGancestor.r' 'dDAGtermSim.r' 'visDAG.r' 'dEnricher.r' 'dEnricherView.r' 'visBoxplotAdv.r' 'dRDataLoader.r'

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biocViews Bioinformatics

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Transcriptome dataset for Chronic Lymphocytic Leukemia patients from Chuang et al. (2012)
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# 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.

# Usage

CLL <- dRDataLoader(RData=CLL)</pre>

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#### 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**

```
CLL <- dRDataLoader(RData=CLL)
CLL
# extract information about the first 5 samples
pData(CLL)[1:5,]
# extract information about the first 5 features
fData(CLL)[1:5,]</pre>
```

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".

#### Usage

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

#### **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

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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

```
# 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>
```

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dBUMscore	Function to transform p-values into scores according to the fitted beta-
	uniform 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 insignificant 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-likelyhood 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  $\int_0^x \lambda+(1-\lambda)*a\,\mathrm{d}x$ . 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  $log_2 \frac{(1-\lambda)*a*(\tau^{a-1}-1)}{\lambda+(1-\lambda)*a} \geq 0, \text{ that is, the constraint via } \tau \leq \left(\frac{\lambda+2*a*(1-\lambda)}{a*(1-\lambda)}\right)^{\frac{1}{a-1}}$ 

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- $\text{ The method "cdf": } S(x) = log_2 \frac{\int_0^x (1-\lambda)*a*(x^{a-1}-1)\,\mathrm{d}x}{\int_0^x \lambda + (1-\lambda)*a\,\mathrm{d}x} log_2 \frac{\int_0^\tau (1-\lambda)*a*(\tau^{a-1}-1)\,\mathrm{d}x}{\int_0^\tau \lambda + (1-\lambda)*a\,\mathrm{d}x} = log_2 \frac{(1-\lambda)*(x^{a-1}-a)\,\mathrm{d}x}{\lambda + (1-\lambda)*a} log_2 \frac{(1-\lambda)*(\tau^{a-1}-a)}{\lambda + (1-\lambda)*a} = log_2 \frac{(1-\lambda)*(\tau^{a-1}-a)}{\lambda + (1-\lambda)*a} log_2 \frac{(1-\lambda)*(\tau^{a-1}-a)}{\lambda + (1-\lambda)*a} = log_2 \frac{(1-\lambda)*(\tau^{a-1}-a)}{\tau^{a-1}-a} \right).$  For the purpose of down-weighting scores, it must ensure  $log_2 \frac{(1-\lambda)*(\tau^{a-1}-a)}{\lambda + (1-\lambda)*a} \geq 0$ , 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 deduced 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)

### Usage

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

### 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
```

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#### Value

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

#### Note

none

#### See Also

dCommSignif

### **Examples**

```
\sharp 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>
```

 $\mathsf{dContrast}$ 

Function to help build the contrast matrix

### **Description**

dContrast is used to help build the contrast matrix

### Usage

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#### **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>
```

dDAGancestor

Function to find common ancestors of two terms/nodes from a direct acyclic graph (DAG)

### **Description**

dDAGancestor is supposed to find a list of common ancestors shared by two terms/nodes, given a direct acyclic graph (DAG; an ontology). If two terms are given as NULL, then a sparse matrix of children x ancestors is built for all terms. If one of them is null, then a sparse matrix of children x ancestors is built but only for non-null input terms.

### Usage

```
dDAGancestor(g, term1 = NULL, term2 = NULL, verbose = T)
```

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#### **Arguments**

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

term1 the first term/node as input term2 the second term/node as input

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

default, it sets to true for display

#### Value

• When two terms are given: a list of terms/nodes that are common ancestors for two input terms/nodes

- When two terms are given as NULL: a sparse matrix of children x ancestors is built for all terms, with '1' for the reachable and otherwise '0'.
- When one of terms is given as NULL: a sparse matrix of children x ancestors is built but only for non-null input terms, with '1' for the reachable and otherwise '0'.

#### Note

none

#### See Also

dDAGinduce

### **Examples**

```
# 1) load HPPA as igraph object
data(ig.HPPA)
g <- ig.HPPA
# 2) randomly give two terms
term1 <- sample(V(g)$name,1)
term2 <- sample(V(g)$name,1)
# 3) find common ancestors
dDAGancestor(g, term1, term2)</pre>
```

dDAGannotate

Function to generate a subgraph of a direct acyclic graph (DAG) induced by the input annotation data

### **Description**

dDAGannotate is supposed to produce a subgraph induced by the input annotation data, given a direct acyclic graph (DAG; an ontology). The input is a graph of "igraph" or "graphNET" object, a list of the vertices containing annotation data, and the mode defining the paths to the root of DAG. The induced subgraph contains vertices (with annotation data) and their ancestors along with the defined paths to the root of DAG. The annotations at these vertices (including their ancestors) are also updated according to the true-path rule: a gene annotated to a term should also be annotated by its all ancestor terms.

dDAGannotate

#### **Usage**

```
dDAGannotate(g, annotations, path.mode = c("all_paths",
   "shortest_paths",
   "all_shortest_paths"), verbose = TRUE)
```

#### **Arguments**

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

annotations the vertices/nodes for which annotation data are provided

path. mode the mode of paths induced by vertices/nodes with input annotation data. It can be

"all\_paths" for all possible paths to the root, "shortest\_paths" for only one path to the root (for each node in query), "all\_shortest\_paths" for all shortest paths to the root (i.e. for each node, find all shortest paths with the equal lengths)

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

default, it sets to true for display

#### Value

• subg: an induced subgraph, an object of class "igraph". In addition to the original attributes to nodes and edges, the return subgraph is also appended by new node attributes: "annotations", which contains a list of genes either as original annotations or inherited annotations; "IC", which stands for information content defined as negative 10-based log-transformed frequency of genes annotated to that term.

#### Note

For the mode "shortest\_paths", the induced subgraph is the most concise, and thus informative for visualisation when there are many nodes in query, while the mode "all\_paths" results in the complete subgraph.

#### See Also

```
dDAGinduce, dDAGlevel
```

```
# 1) load HPPA as igraph object
data(ig.HPPA)
g <- ig.HPPA

# 2) load human genes annotated by HPPA
data(org.Hs.egHPPA)
GS <- org.Hs.egHPPA # as GS object

# 3) prepare for annotation data
# randomly select vertices with annotation data
annotations <- GS$gs[sample(1:length(GS$gs),5)]

# 4) obtain the induced subgraph
# 4a) based on all possible paths (i.e. the complete subgraph induced)
dDAGannotate(g, annotations, path.mode="all_paths", verbose=TRUE)
# 4b) based on shortest paths (i.e. the most concise subgraph induced)
dag <- dDAGannotate(g, annotations, path.mode="shortest_paths", verbose=TRUE)</pre>
```

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```
# 5) color-code nodes/terms according to the number of annotations
data <- sapply(V(dag)$annotations, length)</pre>
names(data) <- V(dag)$name</pre>
visDAG(g=dag, data=data, node.info="both")
```

dDAGgeneSim

Function to calculate pair-wise semantic similarity between genes based on a direct acyclic graph (DAG) with annotated data

### **Description**

dDAGgeneSim is supposed to calculate pair-wise semantic similarity between genes based on a direct acyclic graph (DAG) with annotated data.

#### Usage

```
dDAGgeneSim(g, genes = NULL, method.gene = c("BM.average", "BM.max",
"BM.complete", "average", "max"), method.term = c("Resnik", "Lin", "Schlicker", "Jiang", "Pesquita"), force = TRUE, fast = TRUE,
verbose = TRUE)
```

### Arguments

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

the genes between which pair-wise semantic similarity is calculated. If NULL, genes

all genes annotatable in the input dag will be used for calcluation, which is very

prohibitively expensive!

method.gene

the method used for how to derive semantic similarity between genes from semantic similarity between terms. It can be "average" for average similarity between any two terms (one from gene 1, the other from gene 2), "max" for the maximum similarity between any two terms, "BM.average" for best-matching (BM) based average similarity (i.e. for each term of either gene, first calculate maximum similarity to any term in the other gene, then take average of maximum similarity; the final BM-based average similiary is the pre-calculated average between two genes in pair), "BM.max" for BM based maximum similarity (i.e. the same as "BM.average", but the final BM-based maximum similiary is the maximum of the pre-calculated average between two genes in pair), "BM.complete" for BM-based complete-linkage similarity (inspired by complete-linkage concept: the least of any maximum similarity between a term of one gene and a term of the other gene). When comparing BM-based similarity between genes, "BM.average" and "BM.max" are sensitive to the number of terms invovled; instead, "BM.complete" is much robust in this aspect. By default, it uses "BM.average".

method.term

the method used to measure semantic similarity between terms. It can be "Resnik" for information content (IC) of most informative information ancestor (MICA) (see http://arxiv.org/pdf/cmp-lg/9511007.pdf), "Lin" for 2\*IC at MICA divided by the sum of IC at pairs of terms (see http://webdocs.cs.ualberta. ca/~lindek/papers/sim.pdf), "Schlicker" for weighted version of 'Lin' by the 1-prob(MICA) (see http://www.ncbi.nlm.nih.gov/pubmed/16776819), "Jiang" for 1 - difference between the sum of IC at pairs of terms and 2\*IC

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at MICA (see http://arxiv.org/pdf/cmp-lg/9709008.pdf), "Pesquita" for graph information content similarity related to Tanimoto-Jacard index (ie. summed information content of common ancestors divided by summed information content of all ancestors of term1 and term2 (see http://www.ncbi.nlm.nih.gov/pubmed/18460186))

force logical to indicate whether the only most specific terms (for each gene) will be

used. By default, it sets to true. It is always advisable to use this since it is computationally fast but without compromising accuracy (considering the fact

that true-path-rule has been applied when running dDAGannotate)

fast logical to indicate whether a vectorised fast computation is used. By default, it

sets to true. It is always advisable to use this vectorised fast computation; since

the conventional computation is just used for understanding scripts

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

default, it sets to true for display

#### Value

It returns a sparse matrix containing pair-wise semantic similarity between input terms

#### Note

For the mode "shortest\_paths", the induced subgraph is the most concise, and thus informative for visualisation when there are many nodes in query, while the mode "all\_paths" results in the complete subgraph.

### See Also

```
dDAGtermSim, dDAGinduce, dDAGtip
```

```
## Not run:
# 1) load HPPA as igraph object
data(ig.HPPA)
g <- ig.HPPA

# 2) load human genes annotated by HPPA
data(org.Hs.egHPPA)

# 3) prepare for ontology and its annotation information
dag <- dDAGannotate(g, annotations=org.Hs.egHPPA,
path.mode="all_paths", verbose=TRUE)

# 4) calculate pair-wise semantic similarity between 5 randomly chosen genes
allgenes <- unique(unlist(V(dag)$annotations))
genes <- sample(allgenes,5)
sim <- dDAGgeneSim(g=dag, genes=genes, method.gene="BM.average",
method.term="Resnik", verbose=TRUE)
sim

## End(Not run)</pre>
```

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dDAGinduce Function to generate a subgraph of a direct acyclic graph (DAG) in- duced by given vertices	dDAGinduce	
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#### **Description**

dDAGinduce is supposed to produce a subgraph induced by given vertices, given a direct acyclic graph (DAG; an ontology). The input is a graph of "igraph" or "graphNET" object, a list of the vertices of the graph, and the mode defining the paths to the root of DAG. The resultant subgraph inherits the class from the input one. The induced subgraph contains exactly the vertices of interest and their defined paths to the root of DAG.

# Usage

```
dDAGinduce(g, nodes_query, path.mode = c("all_paths", "shortest_paths",
    "all_shortest_paths"))
```

### **Arguments**

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

nodes\_query the vertices for which the calculation is performed

the mode of paths induced by nodes in query. It can be "all\_paths" for all possible paths to the root, "shortest\_paths" for only one path to the root (for each node in query), "all\_shortest\_paths" for all shortest paths to the root (i.e. for each node, find all shortest paths with the equal lengths)

### Value

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

#### Note

For the mode "shortest\_paths", the induced subgraph is the most concise, and thus informative for visualisation when there are many nodes in query, while the mode "all\_paths" results in the complete subgraph.

### See Also

dDAGroot

```
# 1) load HPPA as igraph object
data(ig.HPPA)
g <- ig.HPPA

# 2) randomly select vertices as the query nodes
# the query nodes can be igraph vertex sequences
nodes_query <- sample(V(g),5)
# more commonly, the query nodes can be term id
nodes_query <- sample(V(g),5)$name</pre>
```

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```
# 3) obtain the induced subgraph
# 3a) based on all possible paths (i.e. the complete subgraph induced)
subg <- dDAGinduce(g, nodes_query, path.mode="all_paths")
# 3b) based on shortest paths (i.e. the most concise subgraph induced)
subg <- dDAGinduce(g, nodes_query, path.mode="shortest_paths")</pre>
```

dDAGlevel

Function to define/calculate the level of nodes in a direct acyclic graph (DAG)

## Description

dDAGlevel is supposed to calculate the level of nodes, given a direct acyclic graph (DAG; an ontology). The input is a graph of "igraph" or "graphNET" object, and the definition of the node level. The return can be the level for each node or the nodes for each level.

### Usage

```
dDAGlevel(g, level.mode = c("longest_path", "shortest_path"),
return.mode = c("node2level", "level2node"))
```

### **Arguments**

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

level.mode the mode of how to define the level of nodes in DAG. It can be "longest\_path"

for defining the node level as the length of the longest path from the node to the root, and "shortest\_paths" for defining the node level as the length of the shortest

path from the node to the root

return.mode the mode of how to return the node level information. It can be "node2level"

for returning a named vector (i.e. the level for each node), and "level2node" for

returning a named list (i.e. nodes for each level)

#### Value

When "return.mode" is "node2level", it returns a named vector: for each named node (i.e. Term ID), it stores its level When "return.mode" is "level2node", it returns a named list: for each named level, it contains the names (i.e. Term ID) of nodes belonging to this level

### Note

The level for the root is 1. The level based on the longest path will ensure that nodes at the same level will never be reachable (i.e. in the same path), while the level based on the shortest path will not be necessary. The "longest path" based level can be useful in visiting nodes from the tipmost level to the root: 1) for the current node, all children have been visited; 2) nodes at the same level can be looked at independantly. The "shortest path" based level can be useful in deriving nodes according to their closeness to the root.

#### See Also

dDAGroot, dDAGreverse

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### **Examples**

```
# 1) load HPPA as igraph object
data(ig.HPPA)
g <- ig.HPPA

# 2) randomly select vertices as the query nodes
nodes_query <- sample(V(g),5)$name

# 3) obtain the complete subgraph induced
subg <- dDAGinduce(g, nodes_query)

# 4) calculate the node levels
# 4a) definition based on the longest path
dDAGlevel(subg, level.mode="longest_path")
# 4b) definition based on the shortest path
dDAGlevel(subg, level.mode="shortest_path")
# 4c) definition based on the longest path, and return nodes for each level
dDAGlevel(subg, level.mode="longest_path", return.mode="level2node")</pre>
```

dDAGreverse

Function to reverse the edge direction of a direct acyclic graph (DAG)

### **Description**

dDAGreverse is supposed to reverse the edge direction of a direct acyclic graph (DAG; an ontology). The return graph remains all attributes associated on nodes and edges.

#### Usage

dDAGreverse(g)

### **Arguments**

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

### Value

• gr: a graph being reversed, an object of class "igraph" or "graphNEL"

### Note

none

#### See Also

dDAGreverse

18 dDAGroot

### **Examples**

```
# 1) load HPPA as igraph object
data(ig.HPPA)
g <- ig.HPPA

# 2) the graph with reverse edge direction
gr <- dDAGreverse(g)
gr</pre>
```

 ${\tt dDAGroot}$ 

Function to find the root node of a direct acyclic graph (DAG)

# Description

dDAGroot is supposed to find the root node of a direct acyclic graph (DAG; an ontology). It return the name (i.e Term ID) of the root node.

### Usage

```
dDAGroot(g)
```

### **Arguments**

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

### Value

• root: the root name (i.e. Term ID)

### Note

none

### See Also

dDAGroot

```
# 1) load HPPA as igraph object
data(ig.HPPA)
g <- ig.HPPA

# 2) find the root
root <- dDAGroot(g)
root</pre>
```

dDAGtermSim

dDAGtermSim	Function to calculate pair-wise semantic similarity between input terms based on a direct acyclic graph (DAG) with annotated data

#### **Description**

dDAGtermSim is supposed to calculate pair-wise semantic similarity between input terms based on a direct acyclic graph (DAG) with annotated data.

#### Usage

```
dDAGtermSim(g, terms = NULL, method = c("Resnik", "Lin", "Schlicker",
"Jiang", "Pesquita"), fast = T, verbose = T)
```

#### **Arguments**

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

terms the terms/nodes between which pair-wise semantic similarity is calculated. If

NULL, all terms in the input DAG will be used for calcluation, which is very

prohibitively expensive!

method the method used to measure semantic similarity between input terms. It can be

"Resnik" for information content (IC) of most informative information ancestor (MICA) (see http://arxiv.org/pdf/cmp-lg/9511007.pdf), "Lin" for 2\*IC at MICA divided by the sum of IC at pairs of terms (see http://webdocs.cs.ualberta.ca/~lindek/papers/sim.pdf), "Schlicker" for weighted version of 'Lin' by the 1-prob(MICA) (see http://www.ncbi.nlm.nih.gov/pubmed/16776819), "Jiang" for 1 - difference between the sum of IC at pairs of terms and 2\*IC

at MICA (see http://arxiv.org/pdf/cmp-lg/9709008.pdf), "Pesquita" for graph information content similarity related to Tanimoto-Jacard index (ie. summed information content of common ancestors divided by summed information content of all ancestors of term1 and term2 (see http://www.ncbi.nlm.nih.gov/

pubmed/18460186)). By default, it uses "Schlicker" method

fast logical to indicate whether a vectorised fast computation is used. By default, it

sets to true. It is always advisable to use this vectorised fast computation; since

the conventional computation is just used for understanding scripts

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

default, it sets to true for display

### Value

It returns a sparse matrix containing pair-wise semantic similarity between input terms

#### Note

none

#### See Also

dDAGinduce, dDAGancestor, dDAGgeneSim

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### **Examples**

```
## Not run:
# 1) load HPPA as igraph object
data(ig.HPPA)
g <- ig.HPPA

# 2) load human genes annotated by HPPA
data(org.Hs.egHPPA)

# 3) prepare for ontology and its annotation information
dag <- dDAGannotate(g, annotations=org.Hs.egHPPA,
path.mode="all_paths", verbose=TRUE)

# 4) calculate pair-wise semantic similarity between 5 randomly chosen terms
terms <- sample(V(dag)$name, 5)
sim <- dDAGtermSim(g=dag, terms=terms, method="Schlicker")
sim

## End(Not run)</pre>
```

dDAGtip

Function to find the tip node(s) of a direct acyclic graph (DAG)

### Description

dDAGtip is supposed to find the tip node(s) of a direct acyclic graph (DAG; an ontology). It return the name (i.e Term ID) of the tip node(s).

# Usage

```
dDAGtip(g)
```

### Arguments

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

### Value

• tip: the tip name (i.e. Term ID)

### Note

none

### See Also

```
dDAGtip
```

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#### **Examples**

```
# 1) load HPPA as igraph object
data(ig.HPPA)
g <- ig.HPPA
# 2) find tips
tips <- dDAGtip(g)
tips</pre>
```

dEnricher

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

### **Description**

dEnricher is supposed to conduct enrichment analysis given the input data and the ontology in query. It returns an object of class "eTerm". Enrichment analysis is based on either Fisher's exact test or Hypergeometric test. The test can respect the hierarchy of the ontology.

### Usage

```
dEnricher(data, identity = c("symbol", "entrez"),
check.symbol.identity = FALSE, genome = c("Hs", "Mm", "Rn", "Gg", "Ce",
"Dm", "Da", "At"), ontology = c("GOBP", "GOMF", "GOCC", "PS", "PS2",
"SF"
"DO", "HPPA", "HPMI", "HPON", "MP", "MsigdbC1", "MsigdbC2CGP",
"MsigdbC2CP"
"MsigdbC2KEGG", "MsigdbC2REACTOME", "MsigdbC2BIOCARTA", "MsigdbC3TFT",
"MsigdbC3MIR", "MsigdbC4CGN", "MsigdbC4CM", "MsigdbC5BP", "MsigdbC5MF",
"MsigdbC5CC", "MsigdbC6", "MsigdbC7", "DGIdb"), sizeRange = c(10,
min.overlap = 3, which_distance = NULL, test = c("HypergeoTest",
"FisherTest", "BinomialTest"), p.adjust.method = c("BH", "BY",
"bonferroni",
"holm", "hochberg", "hommel"), ontology.algorithm = c("none", "pc",
"elim",
"lea"), elim.pvalue = 0.01, lea.depth = 2, verbose = T,
RData.location = "http://dnet.r-forge.r-project.org/data")
```

### **Arguments**

data an input vector. It contains 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; also it is possible to search against synonyms (see the next parameter)

check.symbol.identity

logical to indicate whether synonyms will be searched against when gene symbols cannot be matched. By default, it sets to FALSE since it may take a while to do such check using all possible synoyms

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genome

the genome identity. It can be one of "Hs" for human, "Mm" for mouse, "Rn" for rat, "Gg" for chicken, "Ce" for c.elegans, "Dm" for fruitfly, "Da" for zebrafish, and "At" for arabidopsis

ontology

the ontology supported currently. It can be "GOBP" for Gene Ontology Biological Process, "GOMF" for Gene Ontology Molecular Function, "GOCC" for Gene Ontology Cellular Component, "PS" for phylostratific age information, "PS2" for the collapsed PS version (inferred ancestors being collapsed into one with the known taxonomy information), "SF" for domain superfamily assignments, "DO" for Disease Ontology, "HPPA" for Human Phenotype Phenotypic Abnormality, "HPMI" for Human Phenotype Mode of Inheritance, "HPON" for Human Phenotype ONset and clinical course, "MP" for Mammalian Phenotype, and Drug-Gene Interaction database (DGIdb) and the molecular signatures database (Msigdb) only in human (including "MsigdbC1", "MsigdbC2CGP", "MsigdbC2CP", "MsigdbC2KEGG", "MsigdbC2REACTOME", "MsigdbC2BIOCARTA", "MsigdbC3TFT", "MsigdbC3MIR", "MsigdbC4CGN", "MsigdbC4CM", "MsigdbC5BP", "MsigdbC5MF", "MsigdbC5CC", "MsigdbC6", "MsigdbC7"). Note: These four ("GOBP", "GOMF", "GOCC" and "PS") are available for all genomes/species; for "Hs" and "Mm", these five ("DO", "HPPA", "HPMI", "HPON" and "MP") are also supported; all "Msigdb" are only supported in "Hs". For details on the eligibility for pairs of input genome and ontology, please refer to the online Documentations at http://dnet.r-forge.r-project.org/docs.html

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

min.overlap

the minimum number of overlaps. Only those gene sets that overlap with input data at least min.overlap (3 by default) will be processed

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

test

the statistic test used. It can be "FisherTest" for using fisher's exact test, "HypergeoTest" for using hypergeometric test, or "BinomialTest" for using binomial test. Fisher's exact test is to test the independence between gene group (genes belonging to a group or not) and gene annotation (genes annotated by a term or not), and thus compare sampling to the left part of background (after sampling without replacement). Hypergeometric test is to sample at random (without replacement) from the background containing annotated and non-annotated genes, and thus compare sampling to background. Unlike hypergeometric test, binomial test is to sample at random (with replacement) from the background with the constant probability. In terms of the ease of finding the significance, they are in order: hypergeometric test > binomial test > fisher's exact test. In other words, in terms of the calculated p-value, hypergeometric test < binomial test < fisher's exact test

p.adjust.method

the method used to adjust p-values. It can be one of "BH", "BY", "bonferroni", "holm", "hochberg" and "hommel". The first two methods "BH" (widely used) and "BY" control the false discovery rate (FDR: the expected proportion of false discoveries amongst the rejected hypotheses); the last four methods "bonferroni", "holm", "hochberg" and "hommel" are designed to give strong control of the family-wise error rate (FWER). Notes: FDR is a less stringent condition than FWER

ontology.algorithm

the algorithm used to account for the hierarchy of the ontology. It can be one of "none", "pc", "elim" and "lea". For details, please see 'Note'

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elim.pvalue the parameter only used when "ontology.algorithm" is "elim". It is used to con-

trol how to declare a signficantly enriched term (and subsequently all genes in

this term are eliminated from all its ancestors)

lea.depth the parameter only used when "ontology.algorithm" is "lea". It is used to con-

trol how many maximum depth is uded to consider the children of a term (and subsequently all genes in these children term are eliminated from the use for the

recalculation of the signifiance at this term)

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

default, it sets to false for no display

RData.location the characters to tell the location of built-in RData files. By default, it re-

motely locates at "http://dnet.r-forge.r-project.org/data". For the user equipped with fast internet connection, this option can be just left as default. But it is always advisable to download these files locally. Especially when the user needs to run this function many times, there is no need to ask the function to remotely download every time (also it will unnecessarily increase the runtime). For examples, these files (as a whole or part of them) can be first downloaded into your current working directory, and then set this option as: RData.location = "." Surely, the location can be anywhere as long as the user provides the correct path pointing to (otherwise, the script will have to remotely download each time). Here is the UNIX command for downloading all RData files (preserving the directory structure): wget-r-l2-A" \*.RData" – np-nH--cut-dirs=0" http://dnet.r-forge.r-project.org/data"

#### 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 vector containing input data in consideration. It is not always the same as the input data as only those mappable are retained
- overlap: a list of overlapped gene sets, each storing genes overlapped between a gene set and the given input data (i.e. the genes of interest). Always, gene sets are identified by "setID" and gene members identified by "Entrez ID"
- · zscore: a vector containing z-scores
- pvalue: a vector containing p-values
- adjp: a vector containing adjusted p-values. It is the p value but after being adjusted for multiple comparisons
- call: the call that produced this result

### Note

The interpretation of the algorithms used to account for the hierarchy of the ontology is:

• "none": does not consider the ontology hierarchy at all.

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• "lea": computers the significance of a term in terms of the significance of its children at the maximum depth (e.g. 2). Precisely, once genes are already annotated to any children terms with a more significance than itself, then all these genes are eliminated from the use for the recalculation of the significance at that term. The final p-values takes the maximum of the original p-value and the recalculated p-value.

- "elim": computers the significance of a term in terms of the significance of its all children. Precisely, once genes are already annotated to a significantly enriched term under the cutoff of e.g. pvalue<1e-2, all these genes are eliminated from the ancestors of that term).
- "pc": requires the significance of a term not only using the whole genes as background but also using genes annotated to all its direct parents/ancestors as background. The final p-value takes the maximum of both p-values in these two calculations.
- "Notes": the order of the number of significant terms is: "none" > "lea" > "elim" > "pc".

#### See Also

dEnricher

#### **Examples**

```
## Not run:
# load data
data(Fang)
data <- as.character(Fang.geneinfo$Symbol[1:50])</pre>
data
# enrichment analysis
eTerm <- dEnricher(data, identity="symbol", genome="Hs", ontology="DO")
dEnricherView(eTerm, top_num=10, sortBy="adjp", decreasing=FALSE,
details=TRUE)
# visualise the top significant terms in the ontology heirarchy
ig.D0 <- dRDataLoader(RData=ig.D0)</pre>
g \leftarrow ig.D0
nodes_query <- names(sort(eTerm$adjp)[1:5])</pre>
nodes.highlight <- rep("red", length(nodes_query))</pre>
names(nodes.highlight) <- nodes_query</pre>
subg <- dDAGinduce(g, nodes_query)</pre>
# color-code terms according to the adjust p-values (taking the form of 10-based negative logarithm)
visDAG(g=subg, data=-1*log10(eTerm$adjp[V(subg)$name]),
node.info="both", zlim=c(0,2), node.attrs=list(color=nodes.highlight))
# color-code terms according to the z-scores
visDAG(g=subg, data=eTerm$zscore[V(subg)$name], node.info="both",
colormap="darkblue-white-darkorange",
node.attrs=list(color=nodes.highlight))
## End(Not run)
```

dEnricherView

Function to view enrichment results of dEnricher

### **Description**

dEnricherView is supposed to view results of enrichment analysis by dEnricher.

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#### Usage

```
dEnricherView(eTerm, top_num = 10, sortBy = c("adjp", "pvalue",
"zscore",
"nSet", "nOverlap", "none"), decreasing = NULL, details = F)
```

#### **Arguments**

eTerm an object of class "eTerm"

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

sortBy which statistics will be used for sorting and viewing gene sets (terms). It can

be "adjp" for adjusted p value, "pvalue" for p value, "zscore" for enrichment z-score, "nSet" for the number of sets (terms), "nOverlap" for the number in

overlaps, and "none" for ordering according to ID of gene sets (terms)

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

true for "zscore", "nSet" or "nOverlap"; otherwise it would be false

details logical to indicate whether the detail information of gene sets (terms) is also

viewed. By default, it sets to false for no inclusion

#### Value

a data frame with following components:

• setID: term ID

• nSet: number of sets (terms)

• nOverlap: number in overlaps

• zscore: enrichment z-score

• pvalue: nominal p value

• adjp: adjusted p value

• 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

#### See Also

dEnricher

```
## Not run:
dEnricherView(eTerm, top_num=10, sortBy="adjp", decreasing=FALSE,
details=TRUE)
dEnricherView(eTerm, top_num=NULL, sortBy="none")
## End(Not run)
```

26 dFDRscore

dFDRscore	Function to transform fdr into scores according to log-likelihood ratio
	between the true positives and the false positivies and/or after control- ling 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.

### Usage

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

#### **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
```

```
# 1) generate data with an iid matrix of 1000 x 9
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=10)
# 3) calculate the scores according to the fitted BUM and fdr=0.01</pre>
```

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```
# no fdr threshold
scores <- dFDRscore(fdr)
# using fdr threshold of 0.01
scores <- dFDRscore(fdr, fdr.threshold=0.1, scatter=TRUE)</pre>
```

dGSEA

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

### **Description**

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

#### Usage

```
dGSEA(data, identity = c("symbol", "entrez"), check.symbol.identity =
FALSE,
genome = c("Hs", "Mm", "Rn", "Gg", "Ce", "Dm", "Da", "At"),
ontology = c("GOBP", "GOMF", "GOCC", "PS", "PS2", "SF", "DO", "HPPA",
"HPMI", "HPON", "MP", "MsigdbC1", "MsigdbC2CGP", "MsigdbC2CP",
"MsigdbC2KEGG",
"MsigdbC2REACTOME", "MsigdbC2BIOCARTA", "MsigdbC3TFT", "MsigdbC3MIR",
"MsigdbC4CGN", "MsigdbC4CM", "MsigdbC5BP", "MsigdbC5MF", "MsigdbC5CC",
"MsigdbC6", "MsigdbC7", "DGIdb", "Customised"), customised.genesets =
NULL,
sizeRange = c(10, 20000), which_distance = NULL, weight = 1,
nperm = 1000, fast = T, sigTail = c("two-tails", "one-tail"),
p.adjust.method = c("BH", "BY", "bonferroni", "holm", "hochberg",
"hommel"),
verbose = T, RData.location = "http://dnet.r-forge.r-project.org/data")
```

### **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; also it is possible

to search against synonyms (see the next parameter)

check.symbol.identity

logical to indicate whether synonyms will be searched against when gene symbols cannot be matched. By default, it sets to FALSE since it may take a while

to do such check using all possible synoyms

genome the genome identity. It can be one of "Hs" for human, "Mm" for mouse, "Rn" for

rat, "Gg" for chicken, "Ce" for c.elegans, "Dm" for fruitfly, "Da" for zebrafish,

and "At" for arabidopsis

ontology the ontology supported currently. It can be "GOBP" for Gene Ontology Bio-

logical Process, "GOMF" for Gene Ontology Molecular Function, "GOCC" for Gene Ontology Cellular Component, "PS" for phylostratific age information,

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"PS2" for the collapsed PS version (inferred ancestors being collapsed into one with the known taxonomy information), "SF" for domain superfamily assignments, "DO" for Disease Ontology, "HPPA" for Human Phenotype Phenotypic Abnormality, "HPMI" for Human Phenotype Mode of Inheritance, "HPON" for Human Phenotype ONset and clinical course, "MP" for Mammalian Phenotype, and Drug-Gene Interaction database (DGIdb) and the molecular signatures database (Msigdb) in human (including "MsigdbC1", "MsigdbC2CGP", "MsigdbC2CP", "MsigdbC2KEGG", "MsigdbC2REACTOME", "MsigdbC2BIOCARTA", "MsigdbC3TFT", "MsigdbC3MIR", "MsigdbC4CGN", "MsigdbC4CM", "MsigdbC5BP", "MsigdbC5MF", "MsigdbC5CC", "MsigdbC6", "MsigdbC7"). Note: These four ("GOBP", "GOMF", "GOCC" and "PS") are availble for all genomes/species; for "Hs" and "Mm", these five ("DO", "HPPA", "HPMI", "HPON" and "MP") are also supported; all "Msigdb" are only supported in "Hs". For details on the eligibility for pairs of input genome and ontology, please refer to the online Documentations at http://dnet.r-forge.r-project.org/docs.html. Also supported are the user-customised gene sets; in doing so, the option "Customised" should be used together with the input of the next parameter "customised.genesets"

customised.genesets

an input vector/matrix/list which only works when the user chooses "Customised" in the previous parameter "ontology". It contains either Entrez Gene ID or Symbol

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 v

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 weight. 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

nperm

the number of random permutations. For each permutation, gene-score associations 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

p.adjust.method

the method used to adjust p-values. It can be one of "BH", "BY", "bonferroni", "holm", "hochberg" and "hommel". The first two methods "BH" (widely used) and "BY" control the false discovery rate (FDR: the expected proportion of false discoveries amongst the rejected hypotheses); the last four methods "bonferroni", "holm", "hochberg" and "hommel" are designed to give strong control of the family-wise error rate (FWER). Notes: FDR is a less stringent condition than FWER

verbose

logical to indicate whether the messages will be displayed in the screen. By default, it sets to false for no display

RData.location

the characters to tell the location of built-in RData files. By default, it remotely locates at "http://dnet.r-forge.r-project.org/data". For the user equipped with fast internet connection, this option can be just left as default. But it is always advisable to download these files locally. Especially when

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the user needs to run this function many times, there is no need to ask the function to remotely download every time (also it will unnecessarily increase the runtime). For examples, these files (as a whole or part of them) can be first downloaded into your current working directory, and then set this option as: RData.location = ".". Surely, the location can be anywhere as long as the user provides the correct path pointing to (otherwise, the script will have to remotely download each time). Here is the UNIX command for downloading all RData files (preserving the directory structure): wget-r-l2-A"\*.RData" – np-nH--cut-dirs=0" http://dnet.r-forge.r-project.org/data"

#### 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
- 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;

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• "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

```
dGSEAview, dGSEAwrite, visGSEA
```

#### **Examples**

```
## Not run:
load(url("http://dnet.r-forge.r-project.org/data/Datasets/Hiratani_TableS1.RData"))
data <- RT[1:1000,1:2]
eTerm <- dGSEA(data, identity="symbol", genome="Mm", ontology="MP",
which_distance=c(1,2))
res <- dGSEAview(eTerm, which_sample=1, top_num=5, sortBy="adjp",
decreasing=FALSE, details=TRUE)
visGSEA(eTerm, which_sample=1, which_term=rownames(res)[1])
output <- dGSEAwrite(eTerm, which_content="gadjp", which_score="gadjp",
filename="eTerm.txt")

## based on customised gene sets
eTerm <- dGSEA(data, identity="symbol", genome="Mm",
ontology="Customised", customised.genesets=rownames(data)[1:100])

## End(Not run)</pre>
```

dGSEAview

Function to view enrichment results in a sample-specific manner

### **Description**

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

### Usage

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

### Arguments

sortBy

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

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

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

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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

#### See Also

dGSEA

### **Examples**

```
## Not run:
dGSEAview(eTerm, which_sample=1, top_num=10, sortBy="adjp",
decreasing=FALSE, details=TRUE)
## End(Not run)
```

dGSEAwrite

Function to write out enrichment results

#### **Description**

dGSEAwrite is supposed to write out enrichment results.

### Usage

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

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### **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 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 insignfiicant 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

### Note

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

### See Also

dGSEA

```
## Not run:
output <- dGSEAwrite(eTerm, which_content="gadjp", which_score="gadjp",
filename="eTerm.txt")
## End(Not run)</pre>
```

dNetConfidence 33

dNetConfidence Function to append the confidence information from the source graph into the target graph	hs
--	----

#### **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"; specifically, the same as the input target graph but appended with the "nodeConfidence" attribute to the nodes and the "edgeConfidence" attribute to the edges.

### 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

```
# 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),</pre>
```

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```
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 subgraph

### **Description**

dNetFind is supposed to find the maximum scoring subgraph from an input graph and scores imposed on its nodes. The input graph and the output subgraph 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 subgraph 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

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

### Value

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

#### Note

The search procedure is heuristic to find the subgraph 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 singlenodes.
- 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.

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

#### See Also

dNetFind

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

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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 vertices plus their k neighbours. The resultant subgraph inherits the class from the input one. The induced subgraph contains exactly the vertices of interest, and all the edges between them.

# Usage

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

### **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 false
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

```
# 1) generate a random graph according to the ER model
g <- erdos.renyi.game(100, 1/100)
# 2) select the first 10 vertices as the query nodes
nodes_query <- V(g)[1:10]
# 3) produce the induced subgraph only based on the nodes in query</pre>
```

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```
subg <- dNetInduce(g, nodes_query, knn=0)</pre>
```

# 4) produce the induced subgraph based on the nodes in query ane their immediate neighbours subg <- dNetInduce(g, nodes\_query, knn=1)

dNetPipeline

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

## **Description**

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

### Usage

```
dNetPipeline(g, pval, method = c("pdf", "cdf", "customised"),
significance.threshold = NULL, nsize = NULL, plot = F, verbose = T)
```

## **Arguments**

g	an object	of class	"igraph"	or "graphNEL"
6	un object	or crass	151upii	or grapin tel

pval a vector containing input p-values (or fdr). 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

significance.threshold

the given significance threshold. By default, it is set to NULL, meaning there is no constraint. If given, those p-values below this are considered significant and thus scored positively. Instead, those p-values above this given significance

threshold are considered insigificant and thus scored negatively

nsize the desired number of nodes constrained to the resulting subgraph. It is not

nulll, a wide range of significance thresholds will be scanned to find the optimal significance threshold leading to the desired number of nodes in the resulting subgraph. Notably, the given significance threshold will be overwritten by this

option.

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

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#### Note

The pipeline sequentially consists of:

• ia) if the method is either "pdf" or "cdf", dBUMfit used to fit the p-value distribution under beta-uniform mixture model, and dBUMscore used to calculate the scores according to the fitted BUM and the significance threshold.

- ib) if the method is either "customised", then the user input list of fdr (or p-values) and the significance threshold will be directly used for score transformation by dFDRscore.
- ii) if there is the desired number of nodes constrained to the resulting subgraph, a wide range of significance thresholds (including rough stage with large intervals, and finetune stage with smaller intervals) will be scanned to find the significance threshold to meet the desired number of nodes
- iii) dNetFind used to find maximum-scoring subgraph from the input graph and scores imposed on its nodes.

#### See Also

```
dBUMfit, dBUMscore, dFDRscore, dNetFind
```

#### **Examples**

```
# 1) generate an vector consisting of random values from beta distribution
x <- rbeta(1000, shape1=0.5, shape2=1)
names(x) \leftarrow as.character(1:length(x))
# 2) generate a random graph according to the ER model
g <- erdos.renyi.game(1000, 1/100)</pre>
# 3) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, V(g), knn=0)</pre>
# 4) find maximum-scoring subgraph based on the given significance threshold
# 4a) assume the input is a list of p-values (controlling fdr=0.1)
subgraph <- dNetPipeline(g=subg, pval=x, significance.threshold=0.1)</pre>
# 4b) assume the input is a list of customised significance (eg FDR directly)
subgraph <- dNetPipeline(g=subg, pval=x, method="customised",</pre>
significance.threshold=0.1)
## Not run:
# 5) find maximum-scoring subgraph with the desired node number nsize=20
subgraph <- dNetPipeline(g=subg, pval=x, nsize=20)</pre>
## End(Not run)
```

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

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#### 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 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

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- 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.

#### 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.

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#### **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.

## 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>
```

dRDataLoader

Function to load dnet built-in RData

# **Description**

dRDataLoader is supposed to load dnet built-in RData.

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#### Usage

```
dRDataLoader(RData = c("CLL", "Hiratani_TableS1", "TCGA_mutations",
"org.Gg.string", "org.Gg.egPS", "org.Gg.egGOBP", "org.Gg.egGOCC",
"org.Gg.egSF", "org.Gg.eg", "org.Gg.egGOMF", "org.Rn.egGOCC",
"org.Rn.egGOMF"
"org.Rn.egSF", "org.Rn.string", "org.Rn.egPS", "org.Rn.eg",
"org.Rn.egGOBP",
"org.Mm.egHPMI", "org.Mm.eg", "org.Mm.egGOCC", "org.Mm.string",
"org.Mm.egGOBP", "org.Mm.egGOMF", "org.Mm.egHPPA", "org.Mm.egSF",
"org.Mm.egMP", "org.Mm.egHPON", "org.Mm.egPS", "org.Mm.egDO", "ig.MP",
"ig.GOBP", "ig.DO", "ig.HPON", "ig.HPPA", "ig.GOCC", "ig.GOMF",
"ig.HPMI", "org.Ce.egPS", "org.Ce.egGOMF", "org.Ce.egGOCC",
"org.Ce.egGOBP",
"org.Ce.eg", "org.Ce.string", "org.Ce.egSF", "org.Hs.egMsigdbC5MF", "org.Hs.egMsigdbC1", "org.Hs.egMsigdbC3TFT", "org.Hs.egMsigdbC3MIR",
"org.Hs.egMsigdbC2REACTOME", "org.Hs.egMsigdbC7",
"org.Hs.egMsigdbC5BP",
"org.Hs.egMsigdbC6", "org.Hs.egMsigdbC2KEGG", "org.Hs.egMsigdbC2CP",
"org.Hs.egMsigdbC5CC", "org.Hs.egMsigdbC4CGN", "org.Hs.egMsigdbC4CM",
"org.Hs.egMsigdbC2BIOCARTA", "org.Hs.egMsigdbC2CGP", "org.At.egSF",
"org.At.eg", "org.At.egPS", "org.At.egGOBP", "org.At.egGOMF",
"org.At.string",
"org.At.egGOCC", "org.Da.egGOBP", "org.Da.egSF", "org.Da.eg",
"org.Da.egPS",
"org.Da.string", "org.Da.egGOMF", "org.Da.egGOCC", "org.Dm.egPS",
"org.Dm.egGOBP", "org.Dm.egSF", "org.Dm.eg", "org.Dm.string",
"org.Dm.egGOCC",
"org.Dm.egGOMF", "org.Hs.eg", "org.Hs.egMP", "org.Hs.egDGIdb",
"org.Hs.egHPPA", "org.Hs.egSF", "org.Hs.egGOCC", "org.Hs.string", "org.Hs.egGOBP", "org.Hs.egGOMF", "org.Hs.egHPMI", "org.Hs.egDO",
"org.Hs.egHPON", "org.Hs.egPS"),
RData.location = "http://dnet.r-forge.r-project.org/data")
```

## Arguments

RData

which built-in RData to load. It can be one of 'CLL', 'Hiratani\_TableS1', 'TCGA\_mutations', 'org.Gg.string', 'org.Gg.egPS', 'org.Gg.egGOBP', 'org.Gg.egGOCC', 'org.Gg.egSF', 'org.Gg.eg', 'org.Gg.egGOMF', 'org.Rn.egGOCC', 'org.Rn.egGOMF', 'org.Rn.egSF', 'org.Rn.string', 'org.Rn.egPS', 'org.Rn.eg', 'org.Rn.egGOBP', 'org.Mm.egHPMI', 'org.Mm.eg', 'org.Mm.egGOCC', 'org.Mm.string', 'org.Mm.egGOBP', 'org.Mm.egGOMF', 'org.Mm.egHPPA', 'org.Mm.egSF', 'org.Mm.egMP', 'org.Mm.egHPON', 'org.Mm.egPS', 'org.Mm.egDO', 'ig.MP', 'ig.GOBP', 'ig.DO', 'ig.HPON', 'ig.HPPA', 'ig.GOCC', 'ig.GOMF', 'ig.HPMI', 'org.Ce.egPS', 'org.Ce.egGOMF', 'org.Ce.egGOCC', 'org.Ce.egGOBP', 'org.Ce.eg', 'org.Ce.string', 'org.Ce.egSF', 'org.Hs.egMsigdbC5MF', 'org.Hs.egMsigdbC1', 'org.Hs.egMsigdbC3TFT', 'org.Hs.egMsigdbC3MIR', 'org.Hs.egMsigdbC2R 'org.Hs.egMsigdbC7', 'org.Hs.egMsigdbC5BP', 'org.Hs.egMsigdbC6', 'org.Hs.egMsigdbC2KEGG 'org.Hs.egMsigdbC2CP', 'org.Hs.egMsigdbC5CC', 'org.Hs.egMsigdbC4CGN', 'org.Hs.egMsigdbC4CM', 'org.Hs.egMsigdbC2BIOCARTA', 'org.Hs.egMsigdbC2CGP', 'org.At.egSF', 'org.At.eg', 'org.At.egPS', 'org.At.egGOBP', 'org.At.egGOMF', 'org.At.string', 'org.At.egGOCC', 'org.Da.egGOBP', 'org.Da.egSF', 'org.Da.eg', 'org.Da.egPS', 'org.Da.string', 'org.Da.egGOMF', 'org.Da.egGOCC', 'org.Dm.egPS', 'org.Dm.egGOBP', 'org.Dm.egSF', 'org.Dm.eg', 'org.Dm.string', 'org.Dm.egGOCC',

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> 'org.Dm.egGOMF', 'org.Hs.eg', 'org.Hs.egMP', 'org.Hs.egDGIdb', 'org.Hs.egHPPA', 'org.Hs.egSF', 'org.Hs.egGOCC', 'org.Hs.string', 'org.Hs.egGOBP', 'org.Hs.egGOMF', 'org.Hs.egHPMI', 'org.Hs.egDO', 'org.Hs.egHPON', 'org.Hs.egPS'. On the meanings, please refer to the Documentations

RData.location the characters to tell the location of built-in RData files. By default, it remotely locates at http://dnet.r-forge.r-project.org/data or http://supfam. org/dnet/data. For the user equipped with fast internet connection, this option can be just left as default. But it is always advisable to download these files locally. Especially when the user needs to run this function many times, there is no need to ask the function to remotely download every time (also it will unnecessarily increase the runtime). For examples, these files (as a whole or part of them) can be first downloaded into your current working directory, and then set this option as: RData.location = ".". Surely, the location can be anywhere as long as the user provides the correct path pointing to (otherwise, the script will have to remotely download each time). Here is the UNIX command for downloading all RData files (preserving the directory structure): wget - r l2-A" \* .RData" -np-nH--cut-dirs=0" http://dnet.r-forge.rproject.org/data"

#### Value

any use-specified variable that is given on the right side of the assignment sign '<-', which contains the loaded RData.

### Note

If there are no use-specified variable that is given on the right side of the assignment sign '<-', then no RData will be loaded onto the working environment.

#### See Also

dRDataLoader

### **Examples**

```
org.Hs.egSF <- dRDataLoader(RData=org.Hs.egSF)</pre>
org.Hs.eg <- dRDataLoader(RData=org.Hs.eg)</pre>
org.Hs.egDGIdb <- dRDataLoader(RData=org.Hs.egDGIdb)</pre>
org.Hs.egMsigdbC2KEGG <- dRDataLoader(RData=org.Hs.egMsigdbC2KEGG)</pre>
ig.MP <- dRDataLoader(RData=ig.MP)</pre>
```

dRWR

Function to implement Random Walk with Restart (RWR) on the input graph

## **Description**

dRWR is supposed to implement Random Walk with Restart (RWR) on the input graph. If the seeds (i.e. a set of starting nodes) are given, it intends to calculate the affinity score of all nodes in the graph to the seeds. If the seeds are not give, it will pre-compute affinity matrix for nodes in the input graph with respect to each starting node (as a seed) by looping over every node in the graph.

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#### **Usage**

```
dRWR(g, normalise = c("laplacian", "row", "column", "none"),
setSeeds = NULL, restart = 0.75, normalise.affinity.matrix = c("none",
"quantile"), verbose = T)
```

#### **Arguments**

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'

an input matrix used to define sets of starting seeds. One column corresponds setSeeds

to one set of seeds that a walker starts with. The input matrix must have row names, coming from node names of input graph, i.e. V(g)\$name, since there is a mapping operation. The non-zero entries mean that the corresonding rows (i.e. the gene/row names) are used as the seeds, and non-zero values can be viewed as how to weight the relative importance of seeds. By default, this option sets to "NULL", suggesting each node in the graph will be used as a set of the seed to pre-compute affinity matrix for the input graph. This default does not scale for large input graphs since it will loop over every node in the graph; however, the pre-computed affinity matrix can be extensively reused for obtaining affinity scores between any combinations of nodes/seeds, allows for some flexibility in the downstream use, in particular when sampling a large number of random node

combinations for statistical testing

the restart probability used for RWR. The restart probability takes the value from 0 to 1, controlling the range from the starting nodes/seeds that the walker will

explore. The higher the value, the more likely the walker is to visit the nodes centered on the starting nodes. At the extreme when the restart probability is zero, the walker moves freely to the neighbors at each step without restarting

from seeds, i.e., following a random walk (RW)

normalise.affinity.matrix

the way to normalise the output affinity matrix. It can be 'none' for no normalisation, 'quantile' for quantile normalisation to ensure that columns (if multiple)

of the output affinity matrix have the same quantiles

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

default, it sets to true for display

## Value

It returns a sparse matrix, called 'PTmatrix':

- When the seeds are NOT given: a pre-computated 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
- When the seeds are given: an affinity matrix with the dimension of n X nset, where n is the number of nodes in the input graph, and nset for the number of the sets of seeds (i.e. the number of columns in setSeeds). Each column stands for the steady probability vector, storing the affinity score of all nodes in the graph to the starting nodes/seeds. This steady probability vector can be viewed as the "influential impact" over the graph imposed by the starting nodes/seeds.

restart

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#### Note

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

#### See Also

```
dRWRcontact, dRWRpipeline
```

### **Examples**

```
# 1) generate a random graph according to the ER model
g <- erdos.renyi.game(100, 1/100)</pre>
# 2) produce the induced subgraph only based on the nodes in query
subg <- dNetInduce(g, V(g), knn=0)</pre>
V(subg)$name <- 1:vcount(subg)</pre>
# 3) obtain the pre-computated affinity matrix
PTmatrix <- dRWR(g=subg, normalise="laplacian", restart=0.75)
# visualise affinity matrix
visHeatmapAdv(PTmatrix, Rowv=FALSE, Colv=FALSE, colormap="wyr",
KeyValueName="Affinity")
# 4) obtain affinity matrix given sets of seeds
# define sets of seeds
# each seed with equal weight (i.e. all non-zero entries are 1)
aSeeds <- c(1,0,1,0,1)
bSeeds <-c(0,0,1,0,1)
setSeeds <- data.frame(aSeeds,bSeeds)</pre>
rownames(setSeeds) <- 1:5</pre>
# calcualte affinity matrix
PTmatrix <- dRWR(g=subg, normalise="laplacian", setSeeds=setSeeds,
restart=0.75)
PTmatrix
```

dRWRcontact

Function to estimate RWR-based contact strength between samples from an input gene-sample data matrix, an input graph and its precomputed affinity matrix

# Description

dRWRcontact is supposed to estimate sample relationships (ie. contact strength between samples) from an input gene-sample matrix, an input graph and its affinity matrix pre-computed according to random walk restart (RWR) of the input graph. It includes: 1) RWR-smoothed columns of input gene-sample matrix based on the pre-computed affinity matrix; 2) calculation of contact strength (inner products of RWR-smooth columns of input gene-sample matrix); 3) estimation of the contact significance by a randomalisation procedure.

### Usage

```
dRWRcontact(data, g, Amatrix, permutation = c("degree", "none"),
num.permutation = 10, p.adjust.method = c("BH", "BY", "bonferroni",
"holm", "hochberg", "hommel"), adjp.cutoff = 0.05, verbose = T)
```

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### **Arguments**

data an input gene-sample data matrix used for seeds

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

Amatrix an affinity matrix pre-computed from the input graph. Notes: columns for start-

ing nodes walking from, and rows for ending nodes walking to

permutation how to do permutation. It can be 'degree' for degree-preserving permutation,

'none' for permutation purely in random

num.permutation

the number of permutations used to for generating the distribution of contact

strength under randomalisation

p.adjust.method

the method used to adjust p-values. It can be one of "BH", "BY", "bonferroni", "holm", "hochberg" and "hommel". The first two methods "BH" (widely used) and "BY" control the false discovery rate (FDR: the expected proportion of false discoveries amongst the rejected hypotheses); the last four methods "bonferroni", "holm", "hochberg" and "hommel" are designed to give strong control of the family-wise error rate (FWER). Notes: FDR is a less stringent condition

than FWER

adjp.cutoff the cutoff of adjusted pvalue to construct the contact graph

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 "dContact", a list with following components:

- ratio: a symmetric matrix storing ratio (the observed against the expected) between pairwise samples
- zscore: a symmetric matrix storing zscore between pairwise samples
- pval: a symmetric matrix storing pvalue between pairwise samples
- adjpval: a symmetric matrix storing adjusted pvalue between pairwise samples
- cgraph: the constructed contact graph (as a 'igraph' object) under the cutoff of adjusted value
- call: the call that produced this result

### Note

none

### See Also

dRWR

```
# 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)
V(subg)$name <- 1:vcount(subg)</pre>
```

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```
# 3) pre-compute affinity matrix from the input graph
Amatrix <- dRWR(g=subg)

# 4) estimate RWR-based sample relationships
# define sets of seeds as data
# each seed with equal weight (i.e. all non-zero entries are 1)
aSeeds <- c(1,0,1,0,1)
bSeeds <- c(0,0,1,0,1)
data <- data.frame(aSeeds,bSeeds)
rownames(data) <- 1:5
# calcualte their two contacts
dContact <- dRWRcontact(data=data, g=subg, Amatrix=Amatrix)
dContact</pre>
```

dRWRpipeline

Function to setup a pipeine to estimate RWR-based contact strength between samples from an input gene-sample data matrix and an input graph

### Description

dRWRpipeline is supposed to estimate sample relationships (ie. contact strength between samples) from an input gene-sample matrix and an input graph. The pipeline includes: 1) random walk restart (RWR) of the input graph using the input matrix as seeds; 2) calculation of contact strength (inner products of RWR-smoothed columns of input matrix); 3) estimation of the contact signficance by a randomalisation procedure. It supports two methods how to use RWR: 'direct' for directly applying RWR in the given seeds; 'indirectly' for first pre-computing affinity matrix of the input graph, and then deriving the affinity score.

# Usage

```
dRWRpipeline(data, g, method = c("direct", "indirect"),
normalise = c("laplacian", "row", "column", "none"), restart = 0.5,
normalise.affinity.matrix = c("none", "quantile"),
permutation = c("degree", "none"), num.permutation = 10,
p.adjust.method = c("BH", "BY", "bonferroni", "holm", "hochberg",
"hommel"),
adjp.cutoff = 0.05, verbose = T)
```

### **Arguments**

an input gene-sample data matrix used for seeds

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

method the method used to calculate RWR. It can be 'direct' for directly applying RWR, 'indirect' for indirectly applying RWR (first pre-compute affinity matrix and then derive the affinity score)

normalise the way to normalise the adjacency matrix of the input graph. It can be 'laplacian' for laplacian normalisation, 'row' for row-wise normalisation, 'column' for column-wise normalisation, or 'none'

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restart

the restart probability used for RWR. The restart probability takes the value from 0 to 1, controlling the range from the starting nodes/seeds that the walker will explore. The higher the value, the more likely the walker is to visit the nodes centered on the starting nodes. At the extreme when the restart probability is zero, the walker moves freely to the neighbors at each step without restarting from seeds, i.e., following a random walk (RW)

normalise.affinity.matrix

the way to normalise the output affinity matrix. It can be 'none' for no normalisation, 'quantile' for quantile normalisation to ensure that columns (if multiple) of the output affinity matrix have the same quantiles

permutation

how to do permutation. It can be 'degree' for degree-preserving permutation, 'none' for permutation in random

num.permutation

the number of permutations used to for generating the distribution of contact strength under randomalisation

p.adjust.method

the method used to adjust p-values. It can be one of "BH", "BY", "bonferroni", "holm", "hochberg" and "hommel". The first two methods "BH" (widely used) and "BY" control the false discovery rate (FDR: the expected proportion of false discoveries amongst the rejected hypotheses); the last four methods "bonferroni", "holm", "hochberg" and "hommel" are designed to give strong control of the family-wise error rate (FWER). Notes: FDR is a less stringent condition than FWER

adjp.cutoff

the cutoff of adjusted pvalue to construct the contact graph

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 "dContact", a list with following components:

- ratio: a symmetric matrix storing ratio (the observed against the expected) between pairwise samples
- zscore: a symmetric matrix storing zscore between pairwise samples
- pval: a symmetric matrix storing pvalue between pairwise samples
- adjpval: a symmetric matrix storing adjusted pvalue between pairwise samples
- cgraph: the constructed contact graph (as a 'igraph' object) under the cutoff of adjusted value
- call: the call that produced this result

### Note

The choice of which method to use RWR depends on the number of seed sets and the number of permutations for statistical test. If the total product of both numbers are huge, it is better to use 'indrect' method (for a single run). However, if the user wants to re-use pre-computed affinity matrix (ie. re-use the input graph a lot), then it is highly recommended to sequentially use dRWR and dRWRcontact instead.

### See Also

dRWR, dRWRcontact

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#### **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)
V(subg)$name <- 1:vcount(subg)

# 3) estimate RWR dating based sample relationships
# define sets of seeds as data
# each seed with equal weight (i.e. all non-zero entries are 1)
aSeeds <- c(1,0,1,0,1)
bSeeds <- c(0,0,1,0,1)
data <- data.frame(aSeeds,bSeeds)
rownames(data) <- 1:5
# calcualte their two contact graph
dContact <- dRWRpipeline(data=data, g=subg)
dContact</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**

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

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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.permutation

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 "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)

verbose

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 an iid matrix of 1000 x 9
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=10)
## Not run:
# using "pval" significance
pval <- dSVDsignif(data, signif="pval", num.permutation=10)
## End(Not run)</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.

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## Usage

load(url("http://dnet.r-forge.r-project.org/data/Datasets/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.

# **Examples**

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

ig.DO

Disease Ontology (DO).

### **Description**

An R object that contains information on Disease Ontology terms. These terms are organised as a direct acyclic graph (DAG), which is further stored as an object of the class 'igraph' (see http://igraph.sourceforge.net/doc/R/aaa-igraph-package.html). This data is prepared based on http://sourceforge.net/p/diseaseontology/code/HEAD/tree/trunk/HumanDO.obo.

### Usage

```
ig.D0 <- dRDataLoader(RData=ig.D0)</pre>
```

#### Value

an object of class "igraph". As a direct graph, it has attributes to vertices/nodes and edges:

- vertex attributes: "name" (i.e. "Term ID"), "term\_id" (i.e. "Term ID"), "term\_name" (i.e "Term Name") and "term\_distance" (i.e. Term Distance: the distance to the root; always 0 for the root itself)
- edge attributes: "relation" (either 'is\_a' or 'part\_of')

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#### References

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

## **Examples**

```
ig.D0 <- dRDataLoader(RData=ig.D0)
ig.D0</pre>
```

ig.GOBP

Gene Ontology Biological Process (GOBP).

# **Description**

An R object that contains information on Gene Ontology Biological Process terms. These terms are organised as a direct acyclic graph (DAG), which is further stored as an object of the class 'igraph' (see http://igraph.sourceforge.net/doc/R/aaa-igraph-package.html). This data is prepared based on http://www.geneontology.org/ontology/obo\_format\_1\_2/gene\_ontology.1\_2.obo.

# Usage

```
ig.GOBP <- dRDataLoader(RData=ig.GOBP)</pre>
```

### Value

an object of class "igraph". As a direct graph, it has attributes to vertices/nodes and edges:

- vertex attributes: "name" (i.e. "Term ID"), "term\_id" (i.e. "Term ID"), "term\_name" (i.e "Term Name") and "term\_distance" (i.e. Term Distance: the distance to the root; always 0 for the root itself)
- edge attributes: "relation" (either 'is\_a' or 'part\_of')

### References

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

```
ig.GOBP <- dRDataLoader(RData=ig.GOBP)
ig.GOBP</pre>
```

ig.GOCC 53

ig.GOCC

Gene Ontology Cellular Component (GOCC).

### **Description**

An R object that contains information on Gene Ontology Cellular Component terms. These terms are organised as a direct acyclic graph (DAG), which is further stored as an object of the class 'igraph' (see http://igraph.sourceforge.net/doc/R/aaa-igraph-package.html). This data is prepared based on http://www.geneontology.org/ontology/obo\_format\_1\_2/gene\_ontology.1\_2.obo.

## Usage

```
ig.GOCC <- dRDataLoader(RData=ig.GOCC)</pre>
```

## Value

an object of class "igraph". As a direct graph, it has attributes to vertices/nodes and edges:

- vertex attributes: "name" (i.e. "Term ID"), "term\_id" (i.e. "Term ID"), "term\_name" (i.e "Term Name") and "term\_distance" (i.e. Term Distance: the distance to the root; always 0 for the root itself)
- edge attributes: "relation" (either 'is\_a' or 'part\_of')

### References

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

## **Examples**

```
ig.GOCC <- dRDataLoader(RData=ig.GOCC)
ig.GOCC</pre>
```

ig.GOMF

Gene Ontology Molecular Function (GOMF).

### **Description**

An R object that contains information on Gene Ontology Molecular Function terms. These terms are organised as a direct acyclic graph (DAG), which is further stored as an object of the class 'igraph' (see http://igraph.sourceforge.net/doc/R/aaa-igraph-package.html). This data is prepared based on http://www.geneontology.org/ontology/obo\_format\_1\_2/gene\_ontology.1\_2.obo.

## Usage

```
ig.GOMF <- dRDataLoader(RData=ig.GOMF)</pre>
```

54 ig.HPMI

#### Value

an object of class "igraph". As a direct graph, it has attributes to vertices/nodes and edges:

- vertex attributes: "name" (i.e. "Term ID"), "term\_id" (i.e. "Term ID"), "term\_name" (i.e "Term Name") and "term\_distance" (i.e. Term Distance: the distance to the root; always 0 for the root itself)
- edge attributes: "relation" (either 'is\_a' or 'part\_of')

#### References

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

### **Examples**

```
ig.GOMF <- dRDataLoader(RData=ig.GOMF)
ig.GOMF</pre>
```

ig.HPMI

Human Phenotype Mode of Inheritance (HPMI).

# Description

An R object that contains information on Human Phenotype Mode of Inheritance terms. These terms are organised as a direct acyclic graph (DAG), which is further stored as an object of the class 'igraph' (see http://igraph.sourceforge.net/doc/R/aaa-igraph-package.html). This data is prepared based on http://compbio.charite.de/svn/hpo/trunk/src/ontology/human-phenotype-ontology.obo.

#### Usage

```
ig.HPMI <- dRDataLoader(RData=ig.HPMI)</pre>
```

# Value

an object of class "igraph". As a direct graph, it has attributes to vertices/nodes and edges:

- vertex attributes: "name" (i.e. "Term ID"), "term\_id" (i.e. "Term ID"), "term\_name" (i.e "Term Name") and "term\_distance" (i.e. Term Distance: the distance to the root; always 0 for the root itself)
- edge attributes: "relation" (either 'is\_a' or 'part\_of')

#### 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.

```
ig.HPMI <- dRDataLoader(RData=ig.HPMI)
ig.HPMI</pre>
```

ig.HPON 55

ig.HPON

Human Phenotype ONset and clinical course (HPON).

### **Description**

An R object that contains information on Human Phenotype ONset and clinical course terms. These terms are organised as a direct acyclic graph (DAG), which is further stored as an object of the class 'igraph' (see http://igraph.sourceforge.net/doc/R/aaa-igraph-package.html). This data is prepared based on http://compbio.charite.de/svn/hpo/trunk/src/ontology/human-phenotype-ontology.obo.

## Usage

```
ig.HPON <- dRDataLoader(RData=ig.HPON)</pre>
```

#### Value

an object of class "igraph". As a direct graph, it has attributes to vertices/nodes and edges:

- vertex attributes: "name" (i.e. "Term ID"), "term\_id" (i.e. "Term ID"), "term\_name" (i.e "Term Name") and "term\_distance" (i.e. Term Distance: the distance to the root; always 0 for the root itself)
- edge attributes: "relation" (either 'is\_a' or 'part\_of')

## 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.

# **Examples**

```
ig.HPON <- dRDataLoader(RData=ig.HPON)
ig.HPON</pre>
```

ig.HPPA

Human Phenotype Phenotypic Abnormality (HPPA).

# Description

An R object that contains information on Human Phenotype Phenotypic Abnormality terms. These terms are organised as a direct acyclic graph (DAG), which is further stored as an object of the class 'igraph' (see http://igraph.sourceforge.net/doc/R/aaa-igraph-package.html). This data is prepared based on http://compbio.charite.de/svn/hpo/trunk/src/ontology/human-phenotype-ontology.obo.

# Usage

```
data(ig.HPPA)
```

56 ig.MP

#### Value

an object of class "igraph". As a direct graph, it has attributes to vertices/nodes and edges:

- vertex attributes: "name" (i.e. "Term ID"), "term\_id" (i.e. "Term ID"), "term\_name" (i.e "Term Name") and "term\_distance" (i.e. Term Distance: the distance to the root; always 0 for the root itself)
- edge attributes: "relation" (either 'is\_a' or 'part\_of')

#### 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.

## **Examples**

```
#ig.HPPA <- dRDataLoader(RData=ig.HPPA)
data(ig.HPPA)
ig.HPPA</pre>
```

ig.MP

Mammalian Phenotype (MP).

## **Description**

An R object that contains information on Mammalian Phenotype terms. These terms are organised as a direct acyclic graph (DAG), which is further stored as an object of the class 'igraph' (see <a href="http://igraph.sourceforge.net/doc/R/aaa-igraph-package.html">http://sourceforge.net/doc/R/aaa-igraph-package.html</a>). This data is prepared based on <a href="http://sourceforge.net/p/diseaseontology/code/HEAD/tree/trunk/HumanMP.obo">http://sourceforge.net/p/diseaseontology/code/HEAD/tree/trunk/HumanMP.obo</a>.

# Usage

```
ig.MP <- dRDataLoader(RData=ig.MP)</pre>
```

### Value

an object of class "igraph". As a direct graph, it has attributes to vertices/nodes and edges:

- vertex attributes: "name" (i.e. "Term ID"), "term\_id" (i.e. "Term ID"), "term\_name" (i.e "Term Name") and "term\_distance" (i.e. Term Distance: the distance to the root; always 0 for the root itself)
- edge attributes: "relation" (either 'is\_a' or 'part\_of')

#### References

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

```
ig.MP <- dRDataLoader(RData=ig.MP)
ig.MP</pre>
```

org.At.eg 57

org.At.eg

Arabidopsis Entrez Genes (EG).

## **Description**

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

## Usage

```
org.At.eg <- dRDataLoader(RData=org.At.eg)</pre>
```

### 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**

```
org.At.eg <- dRDataLoader(RData=org.At.eg)
names(org.At.eg)
org.At.eg$gene_info[1:5,]</pre>
```

org.At.egGOBP

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

# Description

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

### Usage

```
org.At.egGOBP <- dRDataLoader(RData=org.At.egGOBP)</pre>
```

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#### 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.

## **Examples**

```
org.At.egGOBP <- dRDataLoader(RData=org.At.egGOBP)
names(org.At.egGOBP)</pre>
```

org.At.egGOCC

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

## **Description**

An R object that contains associations between Gene Ontology Cellular Component terms and Arabidopsis 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

```
\verb|org.At.egGOCC| <- dRDataLoader(RData=org.At.egGOCC)| \\
```

## 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.

org.At.egGOMF

#### **Examples**

```
org.At.egGOCC <- dRDataLoader(RData=org.At.egGOCC)
names(org.At.egGOCC)</pre>
```

org.At.egGOMF

Annotations of Arabidopsis Entrez Genes (EG) by Gene Ontology Molecular Function (GOMF).

## **Description**

An R object that contains associations between Gene Ontology Molecular Function terms and Arabidopsis 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

```
org.At.egGOMF <- dRDataLoader(RData=org.At.egGOMF)</pre>
```

# 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.

```
org.At.egGOMF <- dRDataLoader(RData=org.At.egGOMF)
names(org.At.egGOMF)</pre>
```

60 org.At.egPS

org.At.egPS Annotations of Arabidopsis Entrez Genes (EG) by phylostratific age (PS).

#### **Description**

An R object that contains phylostratific age information for Arabidopsis Entrez Genes. This data is prepared based on 1) SUPERFAMILY database which provides 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 species tree of life (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 Arabidopsis Entrez gene is the first appearance of its domain architecture along the branch from the eukaryotic ancestor to the arabidopsis, and thus can be measured by: i) the most recent common ancestor, ii) 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.

### Usage

```
org.At.egPS <- dRDataLoader(RData=org.At.egPS)</pre>
```

#### 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). 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 nearest descendant 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 al. (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.

```
org.At.egPS <- dRDataLoader(RData=org.At.egPS)
names(org.At.egPS)</pre>
```

org.At.egSF

org.At.egSF	Annotations of Arabidopsis Entrez Genes (EG) by domain superfamilies (SF).

### **Description**

An R object that contains domain superfamily information for Arabidopsis Entrez Genes. This data is prepared based on SUPERFAMILY database, which provides SCOP domain architecture assignments to all completely sequenced genomes including eukaryotic genomes. The domain architecture for an Entrez gene is the protein product with the longest length of amino acids. Thus, domain superfamily information for Arabidopsis Entrez gene is a list of domain superfamilies (excluding unknown gap) appearing in its domain architecture.

#### **Usage**

```
org.At.egSF <- dRDataLoader(RData=org.At.egSF)</pre>
```

#### 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. SCOP domain superfamilies). The 4 columns are "setID" (i.e. "SCOP domain identifier"), "name" (i.e. "SCOP domain description"), "namespace" (i.e. "SCOP domain level") and "distance" (i.e. "SCOP domain classification").
- 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 al. (2011) SUPERFAMILY 1.75 including a domain-centric gene ontology method. *Nucleic Acids Res*, 39(Database issue):D427-34.

Andreeva et al. (2008) Data growth and its impact on the SCOP database: new developments. *Nucleic Acids Res*, 36(Database issue):D419-425

#### **Examples**

```
org.At.egSF <- dRDataLoader(RData=org.At.egSF)
names(org.At.egSF)</pre>
```

org.At.string

Arabidopsis functional protein association network from STRING (version 9.1).

### **Description**

An igraph object that contains a functional protein association network in arabidopsis. The network is extracted from the STRING database (version 9.1). Only those associations with medium confidence (score>=0.4) are retained.

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#### Usage

```
org.At.string <- dRDataLoader(RData=org.At.string)</pre>
```

#### 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**

```
org.At.string <- dRDataLoader(RData=org.At.string)
org.At.string</pre>
```

org.Ce.eg

C.elegans Entrez Genes (EG).

#### **Description**

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

## Usage

```
org.Ce.eg <- dRDataLoader(RData=org.Ce.eg)</pre>
```

org.Ce.egGOBP 63

#### 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**

```
org.Ce.eg <- dRDataLoader(RData=org.Ce.eg)
names(org.Ce.eg)
org.Ce.eg$gene_info[1:5,]</pre>
```

org.Ce.egGOBP

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

## **Description**

An R object that contains associations between Gene Ontology Biological Process terms and C.elegans 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

```
org.Ce.egGOBP <- dRDataLoader(RData=org.Ce.egGOBP)</pre>
```

### 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.

```
org.Ce.egGOBP <- dRDataLoader(RData=org.Ce.egGOBP)
names(org.Ce.egGOBP)</pre>
```

64 org.Ce.egGOMF

org.Ce.egGOCC	Annotations of C.elegans Entrez Genes (EG) by Gene Ontology Cellular Component (GOCC).

### **Description**

An R object that contains associations between Gene Ontology Cellular Component terms and C.elegans 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

```
org.Ce.egGOCC <- dRDataLoader(RData=org.Ce.egGOCC)</pre>
```

### 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.

## **Examples**

```
org.Ce.egGOCC <- dRDataLoader(RData=org.Ce.egGOCC)
names(org.Ce.egGOCC)</pre>
```

org.Ce.egGOMF Annotations of C.elegans Entrez Genes (EG) by Gene Ontology Molecular Function (GOMF).

### **Description**

An R object that contains associations between Gene Ontology Molecular Function terms and C.elegans 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

```
org.Ce.egGOMF <- dRDataLoader(RData=org.Ce.egGOMF)</pre>
```

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#### 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**

```
org.Ce.egGOMF <- dRDataLoader(RData=org.Ce.egGOMF)
names(org.Ce.egGOMF)</pre>
```

org.Ce.egPS

Annotations of C.elegans Entrez Genes (EG) by phylostratific age (PS).

## **Description**

An R object that contains phylostratific age information for C.elegans Entrez Genes. This data is prepared based on 1) SUPERFAMILY database which provides 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 species tree of life (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 C.elegans Entrez gene is the first appearance of its domain architecture along the branch from the eukaryotic ancestor to the c.elegans, and thus can be measured by: i) the most recent common ancestor, ii) 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.

## Usage

```
org.Ce.egPS <- dRDataLoader(RData=org.Ce.egPS)</pre>
```

#### 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). 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

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left empty (assumedly a hypothetical unknown ancestor). In the latter case, hypothetical unknown ancestor is filled with the information in its nearest descendant 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 al. (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**

```
org.Ce.egPS <- dRDataLoader(RData=org.Ce.egPS)
names(org.Ce.egPS)</pre>
```

org.Ce.egSF

Annotations of C.elegans Entrez Genes (EG) by domain superfamilies (SF).

## Description

An R object that contains domain superfamily information for C.elegans Entrez Genes. This data is prepared based on SUPERFAMILY database, which provides SCOP domain architecture assignments to all completely sequenced genomes including eukaryotic genomes. The domain architecture for an Entrez gene is the protein product with the longest length of amino acids. Thus, domain superfamily information for C.elegans Entrez gene is a list of domain superfamilies (excluding unknown gap) appearing in its domain architecture.

### Usage

```
org.Ce.egSF <- dRDataLoader(RData=org.Ce.egSF)</pre>
```

# 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. SCOP domain superfamilies). The 4 columns are "setID" (i.e. "SCOP domain identifier"), "name" (i.e. "SCOP domain description"), "namespace" (i.e. "SCOP domain level") and "distance" (i.e. "SCOP domain classification").
- 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 al. (2011) SUPERFAMILY 1.75 including a domain-centric gene ontology method. *Nucleic Acids Res*, 39(Database issue):D427-34.

Andreeva et al. (2008) Data growth and its impact on the SCOP database: new developments. *Nucleic Acids Res*, 36(Database issue):D419-425

org.Ce.string 67

## **Examples**

```
org.Ce.egSF <- dRDataLoader(RData=org.Ce.egSF)
names(org.Ce.egSF)</pre>
```

org.Ce.string

C.elegans functional protein association network from STRING (version 9.1).

## **Description**

An igraph object that contains a functional protein association network in c.elegans. The network is extracted from the STRING database (version 9.1). Only those associations with medium confidence (score>=0.4) are retained.

## Usage

```
org.Ce.string <- dRDataLoader(RData=org.Ce.string)</pre>
```

### 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.

```
org.Ce.string <- dRDataLoader(RData=org.Ce.string)
org.Ce.string</pre>
```

68 org.Da.egGOBP

org.Da.eg

Zebrafish Entrez Genes (EG).

## **Description**

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

## Usage

```
org.Da.eg <- dRDataLoader(RData=org.Da.eg)</pre>
```

### 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**

```
org.Da.eg <- dRDataLoader(RData=org.Da.eg)
names(org.Da.eg)
org.Da.eg$gene_info[1:5,]</pre>
```

org.Da.egGOBP

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

# **Description**

An R object that contains associations between Gene Ontology Biological Process terms and Zebrafish 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

```
org.Da.egGOBP <- dRDataLoader(RData=org.Da.egGOBP)</pre>
```

org.Da.egGOCC 69

#### 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.

### **Examples**

```
org.Da.egGOBP <- dRDataLoader(RData=org.Da.egGOBP)
names(org.Da.egGOBP)</pre>
```

org.Da.egGOCC

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

## **Description**

An R object that contains associations between Gene Ontology Cellular Component terms and Zebrafish 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

```
\verb|org.Da.egGOCC| <- dRDataLoader(RData=org.Da.egGOCC)| \\
```

## 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.

70 org.Da.egGOMF

#### **Examples**

```
org.Da.egGOCC <- dRDataLoader(RData=org.Da.egGOCC)
names(org.Da.egGOCC)</pre>
```

org.Da.egGOMF

Annotations of Zebrafish Entrez Genes (EG) by Gene Ontology Molecular Function (GOMF).

## **Description**

An R object that contains associations between Gene Ontology Molecular Function terms and Zebrafish 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

```
org.Da.egGOMF <- dRDataLoader(RData=org.Da.egGOMF)</pre>
```

# 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.

```
org.Da.egGOMF <- dRDataLoader(RData=org.Da.egGOMF)
names(org.Da.egGOMF)</pre>
```

org.Da.egPS 71

org.Da.egPS

Annotations of Zebrafish Entrez Genes (EG) by phylostratific age (PS).

#### **Description**

An R object that contains phylostratific age information for Zebrafish Entrez Genes. This data is prepared based on 1) SUPERFAMILY database which provides 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 species tree of life (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 Zebrafish Entrez gene is the first appearance of its domain architecture along the branch from the eukaryotic ancestor to the zebrafish, and thus can be measured by: i) the most recent common ancestor, ii) 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.

### Usage

```
org.Da.egPS <- dRDataLoader(RData=org.Da.egPS)</pre>
```

#### 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). 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 nearest descendant 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 al. (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.

```
org.Da.egPS <- dRDataLoader(RData=org.Da.egPS)
names(org.Da.egPS)</pre>
```

72 org.Da.string

org.Da.egSF	Annotations of Zebrafish Entrez Genes (EG) by domain superfamilies (SF).

### **Description**

An R object that contains domain superfamily information for Zebrafish Entrez Genes. This data is prepared based on SUPERFAMILY database, which provides SCOP domain architecture assignments to all completely sequenced genomes including eukaryotic genomes. The domain architecture for an Entrez gene is the protein product with the longest length of amino acids. Thus, domain superfamily information for Zebrafish Entrez gene is a list of domain superfamilies (excluding unknown gap) appearing in its domain architecture.

## Usage

```
org.Da.egSF <- dRDataLoader(RData=org.Da.egSF)</pre>
```

#### 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. SCOP domain superfamilies). The 4 columns are "setID" (i.e. "SCOP domain identifier"), "name" (i.e. "SCOP domain description"), "namespace" (i.e. "SCOP domain level") and "distance" (i.e. "SCOP domain classification").
- 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 al. (2011) SUPERFAMILY 1.75 including a domain-centric gene ontology method. *Nucleic Acids Res*, 39(Database issue):D427-34.

Andreeva et al. (2008) Data growth and its impact on the SCOP database: new developments. *Nucleic Acids Res*, 36(Database issue):D419-425

#### **Examples**

```
org.Da.egSF <- dRDataLoader(RData=org.Da.egSF)
names(org.Da.egSF)</pre>
```

org.Da.string

Zebrafish functional protein association network from STRING (version 9.1).

### **Description**

An igraph object that contains a functional protein association network in zebrafish. The network is extracted from the STRING database (version 9.1). Only those associations with medium confidence (score>=0.4) are retained.

org.Dm.eg

### Usage

```
org.Da.string <- dRDataLoader(RData=org.Da.string)</pre>
```

#### 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**

```
org.Da.string <- dRDataLoader(RData=org.Da.string)
org.Da.string</pre>
```

org.Dm.eg

Fruitfly Entrez Genes (EG).

#### **Description**

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

```
org.Dm.eg <- dRDataLoader(RData=org.Dm.eg)</pre>
```

74 org.Dm.egGOBP

#### 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**

```
org.Dm.eg <- dRDataLoader(RData=org.Dm.eg)
names(org.Dm.eg)
org.Dm.eg$gene_info[1:5,]</pre>
```

org.Dm.egGOBP

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

### **Description**

An R object that contains associations between Gene Ontology Biological Process terms and Fruitfly 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

```
org.Dm.egGOBP <- dRDataLoader(RData=org.Dm.egGOBP)</pre>
```

## 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.

```
org.Dm.egGOBP <- dRDataLoader(RData=org.Dm.egGOBP)
names(org.Dm.egGOBP)</pre>
```

org.Dm.egGOCC 75

org.Dm.egGOCC	Annotations of Fruitfly Entrez Genes (EG) by Gene Ontology Cellular Component (GOCC).

### **Description**

An R object that contains associations between Gene Ontology Cellular Component terms and Fruitfly 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

```
org.Dm.egGOCC <- dRDataLoader(RData=org.Dm.egGOCC)</pre>
```

### 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.

## **Examples**

```
org.Dm.egGOCC <- dRDataLoader(RData=org.Dm.egGOCC)
names(org.Dm.egGOCC)</pre>
```

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

## **Description**

An R object that contains associations between Gene Ontology Molecular Function terms and Fruitfly 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.

```
org.Dm.egGOMF <- dRDataLoader(RData=org.Dm.egGOMF)</pre>
```

76 org.Dm.egPS

#### 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**

```
org.Dm.egGOMF <- dRDataLoader(RData=org.Dm.egGOMF)
names(org.Dm.egGOMF)</pre>
```

org.Dm.egPS

Annotations of Fruitfly Entrez Genes (EG) by phylostratific age (PS).

### **Description**

An R object that contains phylostratific age information for Fruitfly Entrez Genes. This data is prepared based on 1) SUPERFAMILY database which provides 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 species tree of life (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 Fruitfly Entrez gene is the first appearance of its domain architecture along the branch from the eukaryotic ancestor to the fruitfly, and thus can be measured by: i) the most recent common ancestor, ii) 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.

## Usage

```
org.Dm.egPS <- dRDataLoader(RData=org.Dm.egPS)</pre>
```

### 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). 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

org.Dm.egSF

left empty (assumedly a hypothetical unknown ancestor). In the latter case, hypothetical unknown ancestor is filled with the information in its nearest descendant 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 al. (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**

```
org.Dm.egPS <- dRDataLoader(RData=org.Dm.egPS)
names(org.Dm.egPS)</pre>
```

org.Dm.egSF

Annotations of Fruitfly Entrez Genes (EG) by domain superfamilies (SF).

## Description

An R object that contains domain superfamily information for Fruitfly Entrez Genes. This data is prepared based on SUPERFAMILY database, which provides SCOP domain architecture assignments to all completely sequenced genomes including eukaryotic genomes. The domain architecture for an Entrez gene is the protein product with the longest length of amino acids. Thus, domain superfamily information for Fruitfly Entrez gene is a list of domain superfamilies (excluding unknown gap) appearing in its domain architecture.

## Usage

```
org.Dm.egSF <- dRDataLoader(RData=org.Dm.egSF)</pre>
```

# 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. SCOP domain superfamilies). The 4 columns are "setID" (i.e. "SCOP domain identifier"), "name" (i.e. "SCOP domain description"), "namespace" (i.e. "SCOP domain level") and "distance" (i.e. "SCOP domain classification").
- 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 al. (2011) SUPERFAMILY 1.75 including a domain-centric gene ontology method. *Nucleic Acids Res*, 39(Database issue):D427-34.

Andreeva et al. (2008) Data growth and its impact on the SCOP database: new developments. *Nucleic Acids Res*, 36(Database issue):D419-425

78 org.Dm.string

### **Examples**

```
org.Dm.egSF <- dRDataLoader(RData=org.Dm.egSF)
names(org.Dm.egSF)</pre>
```

org.Dm.string

Fruitfly functional protein association network from STRING (version 9.1).

## **Description**

An igraph object that contains a functional protein association network in fruitfly. The network is extracted from the STRING database (version 9.1). Only those associations with medium confidence (score>=0.4) are retained.

## Usage

```
org.Dm.string <- dRDataLoader(RData=org.Dm.string)</pre>
```

### 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.

```
org.Dm.string <- dRDataLoader(RData=org.Dm.string)
org.Dm.string</pre>
```

org.Gg.eg

org.Gg.eg

Chicken Entrez Genes (EG).

# **Description**

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

### Usage

```
org.Gg.eg <- dRDataLoader(RData=org.Gg.eg)</pre>
```

### 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**

```
org.Gg.eg <- dRDataLoader(RData=org.Gg.eg)
names(org.Gg.eg)
org.Gg.eg$gene_info[1:5,]</pre>
```

org.Gg.egGOBP

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

## **Description**

An R object that contains associations between Gene Ontology Biological Process terms and Chicken 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.

```
org.Gg.egGOBP <- dRDataLoader(RData=org.Gg.egGOBP)</pre>
```

org.Gg.egGOCC

#### 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.

### **Examples**

```
org.Gg.egGOBP <- dRDataLoader(RData=org.Gg.egGOBP)
names(org.Gg.egGOBP)</pre>
```

org.Gg.egGOCC

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

## **Description**

An R object that contains associations between Gene Ontology Cellular Component terms and Chicken 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

```
\verb|org.Gg.egGOCC| <- dRDataLoader(RData=org.Gg.egGOCC)| \\
```

## 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.

org.Gg.egGOMF

### **Examples**

```
org.Gg.egGOCC <- dRDataLoader(RData=org.Gg.egGOCC)
names(org.Gg.egGOCC)</pre>
```

org.Gg.egGOMF

Annotations of Chicken Entrez Genes (EG) by Gene Ontology Molecular Function (GOMF).

## **Description**

An R object that contains associations between Gene Ontology Molecular Function terms and Chicken 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

```
org.Gg.egGOMF <- dRDataLoader(RData=org.Gg.egGOMF)</pre>
```

# 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.

```
org.Gg.egGOMF <- dRDataLoader(RData=org.Gg.egGOMF)
names(org.Gg.egGOMF)</pre>
```

82 org.Gg.egPS

org.Gg.egPS

Annotations of Chicken Entrez Genes (EG) by phylostratific age (PS).

### **Description**

An R object that contains phylostratific age information for Chicken Entrez Genes. This data is prepared based on 1) SUPERFAMILY database which provides 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 species tree of life (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 Chicken Entrez gene is the first appearance of its domain architecture along the branch from the eukaryotic ancestor to the chicken, and thus can be measured by: i) the most recent common ancestor, ii) 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.

### Usage

```
org.Gg.egPS <- dRDataLoader(RData=org.Gg.egPS)</pre>
```

#### 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). 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 nearest descendant 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 al. (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.

```
org.Gg.egPS <- dRDataLoader(RData=org.Gg.egPS)
names(org.Gg.egPS)</pre>
```

org.Gg.egSF

org.Gg.egSF	Annotations of Chicken Entrez Genes (EG) by domain superfamilies (SF).
	$(S\Gamma)$ .

### **Description**

An R object that contains domain superfamily information for Chicken Entrez Genes. This data is prepared based on SUPERFAMILY database, which provides SCOP domain architecture assignments to all completely sequenced genomes including eukaryotic genomes. The domain architecture for an Entrez gene is the protein product with the longest length of amino acids. Thus, domain superfamily information for Chicken Entrez gene is a list of domain superfamilies (excluding unknown gap) appearing in its domain architecture.

## Usage

```
org.Gg.egSF <- dRDataLoader(RData=org.Gg.egSF)</pre>
```

### 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. SCOP domain superfamilies). The 4 columns are "setID" (i.e. "SCOP domain identifier"), "name" (i.e. "SCOP domain description"), "namespace" (i.e. "SCOP domain level") and "distance" (i.e. "SCOP domain classification").
- 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 al. (2011) SUPERFAMILY 1.75 including a domain-centric gene ontology method. *Nucleic Acids Res*, 39(Database issue):D427-34.

Andreeva et al. (2008) Data growth and its impact on the SCOP database: new developments. *Nucleic Acids Res*, 36(Database issue):D419-425

### **Examples**

```
org.Gg.egSF <- dRDataLoader(RData=org.Gg.egSF)
names(org.Gg.egSF)</pre>
```

org.Gg.string

Chicken functional protein association network from STRING (version 9.1)

### **Description**

An igraph object that contains a functional protein association network in chicken. The network is extracted from the STRING database (version 9.1). Only those associations with medium confidence (score>=0.4) are retained.

84 org.Hs.eg

### Usage

```
org.Gg.string <- dRDataLoader(RData=org.Gg.string)</pre>
```

### 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**

```
org.Gg.string <- dRDataLoader(RData=org.Gg.string)
org.Gg.string</pre>
```

org.Hs.eg

Human Entrez Genes (EG).

#### **Description**

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

```
org.Hs.eg <- dRDataLoader(RData=org.Hs.eg)</pre>
```

org.Hs.egDGIdb 85

#### 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
org.Hs.eg <- dRDataLoader(RData=org.Hs.eg)
names(org.Hs.eg)
org.Hs.eg$gene_info[1:5,]</pre>
```

org.Hs.egDGIdb

Annotations of Human Entrez Genes (EG) by DGIdb categories.

### **Description**

An R object that contains associations between DGIdb categories and Human Entrez Genes. This data is prepared based on http://dgidb.genome.wustl.edu/downloads/categories.tsv.

## Usage

```
org.Hs.egDGIdb <- dRDataLoader(RData=org.Hs.egDGIdb)</pre>
```

### 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

Griffith et al. (2013) DGIdb: mining the druggable genome. Nature methods, 10(12):1209-10.

```
org.Hs.egDGIdb <- dRDataLoader(RData=org.Hs.egDGIdb)
names(org.Hs.egDGIdb)</pre>
```

86 org.Hs.egGOBP

org.Hs.egD0

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

# Description

An R object that contains associations between Disease Ontology terms and Human 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.

## Usage

```
org.Hs.egD0 <- dRDataLoader(RData=org.Hs.egD0)</pre>
```

#### 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**

```
org.Hs.egD0 <- dRDataLoader(RData=org.Hs.egD0)
names(org.Hs.egD0)</pre>
```

org.Hs.egGOBP

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

# Description

An R object that contains associations between Gene Ontology Biological Process terms and Human 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.

```
org.Hs.egGOBP <- dRDataLoader(RData=org.Hs.egGOBP)</pre>
```

org.Hs.egGOCC 87

#### 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.

### **Examples**

```
org.Hs.egGOBP <- dRDataLoader(RData=org.Hs.egGOBP)
names(org.Hs.egGOBP)</pre>
```

org.Hs.egGOCC

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

## **Description**

An R object that contains associations between Gene Ontology Cellular Component terms and Human 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

```
org.Hs.egGOCC <- dRDataLoader(RData=org.Hs.egGOCC)</pre>
```

## 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.

88 org.Hs.egGOMF

### **Examples**

```
org.Hs.egGOCC <- dRDataLoader(RData=org.Hs.egGOCC)
names(org.Hs.egGOCC)</pre>
```

org.Hs.egGOMF

Annotations of Human Entrez Genes (EG) by Gene Ontology Molecular Function (GOMF).

## **Description**

An R object that contains associations between Gene Ontology Molecular Function terms and Human 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

```
org.Hs.egGOMF <- dRDataLoader(RData=org.Hs.egGOMF)</pre>
```

# 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.

```
org.Hs.egGOMF <- dRDataLoader(RData=org.Hs.egGOMF)
names(org.Hs.egGOMF)</pre>
```

org.Hs.egHPMI 89

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

### **Description**

An R object that contains associations between HPMI terms and Human 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.

### Usage

```
org.Hs.egHPMI <- dRDataLoader(RData=org.Hs.egHPMI)</pre>
```

### 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.

### **Examples**

```
org.Hs.egHPMI <- dRDataLoader(RData=org.Hs.egHPMI)
names(org.Hs.egHPMI)</pre>
```

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

## **Description**

An R object that contains associations between HPON terms and Human 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.

```
org.Hs.egHPON <- dRDataLoader(RData=org.Hs.egHPON)</pre>
```

90 org.Hs.egHPPA

#### 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"

#### 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.

### **Examples**

```
org.Hs.egHPON <- dRDataLoader(RData=org.Hs.egHPON)
names(org.Hs.egHPON)</pre>
```

org.Hs.egHPPA

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

## **Description**

An R object that contains associations between Human Phenotype Phenotypic Abnormality terms and Human 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.

## Usage

```
data(org.Hs.egHPPA)
```

#### 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.

org.Hs.egMP

## **Examples**

```
#org.Hs.egHPPA <- dRDataLoader(RData=org.Hs.egHPPA)
data(org.Hs.egHPPA)
names(org.Hs.egHPPA)</pre>
```

org.Hs.egMP

Annotations of Human Entrez Genes (EG) by Mammalian Phenotype (MP).

# **Description**

An R object that contains associations between Mammalian Phenotype terms and Human 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, which results in annotations of Mouse Entrez Genes. Then, these annotations are transferred to Human Entrez Genes based on ftp://anonymous@ftp.ncbi.nih.gov/pub/HomoloGene/build67/homologene.data.

### Usage

```
org.Hs.egMP <- dRDataLoader(RData=org.Hs.egMP)</pre>
```

## 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.

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

```
org.Hs.egMP <- dRDataLoader(RData=org.Hs.egMP)
names(org.Hs.egMP)</pre>
```

org.Hs.egMsigdbC1

Annotations of Human Entrez Genes (EG) by C1 collections.

## **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C1 positional gene sets and Human Entrez Genes. C1 collections are about positional gene sets for each human chromosome and cytogenetic band, each gene set corresponding to each human chromosome and each cytogenetic band that has at least one gene. These gene sets are helpful in identifying effects related to chromosomal deletions or amplifications, dosage compensation, epigenetic silencing, and other regional effects.

### Usage

```
org.Hs.egMsigdbC1 <- dRDataLoader(RData=org.Hs.egMsigdbC1)</pre>
```

### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

## **Examples**

```
org.Hs.egMsigdbC1 <- dRDataLoader(RData=org.Hs.egMsigdbC1)
names(org.Hs.egMsigdbC1)</pre>
```

```
org.Hs.egMsigdbC2BIOCARTA
```

Annotations of Human Entrez Genes (EG) by C2:BIOCARTA collections.

## **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C2:BIOCARTA (BioCarta pathways) gene sets and Human Entrez Genes. C2:BIOCARTA gene sets are derived from the BioCarta pathway database http://www.biocarta.com/genes/index.asp.

## Usage

org.Hs.egMsigdbC2BIOCARTA <- dRDataLoader(RData=org.Hs.egMsigdbC2BIOCARTA)

#### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

# **Examples**

```
org.Hs.egMsigdbC2BIOCARTA <- dRDataLoader(RData=org.Hs.egMsigdbC2BIOCARTA)
names(org.Hs.egMsigdbC2BIOCARTA)</pre>
```

org. Hs. egMsigdbC2CGP Annotations of Human Entrez Genes (EG) by C2:CGP collections.

## Description

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C2:CGP (chemical and genetic perturbations) gene sets and Human Entrez Genes. C2:CGP gene sets are about expression signatures of genetic and chemical perturbations. A number of these gene sets come in pairs: an xxx\_UP (xxx\_DN) gene set representing genes induced (repressed) by the perturbation.

## Usage

```
org.Hs.egMsigdbC2CGP <- dRDataLoader(RData=org.Hs.egMsigdbC2CGP)</pre>
```

### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

## **Examples**

```
org.Hs.egMsigdbC2CGP <- dRDataLoader(RData=org.Hs.egMsigdbC2CGP)
names(org.Hs.egMsigdbC2CGP)</pre>
```

org.Hs.egMsigdbC2CP

Annotations of Human Entrez Genes (EG) by C2:CP collections.

## **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C2:CP (Canonical pathways) gene sets and Human Entrez Genes. C2:CP gene sets are from the pathway databases, and usually are canonical representations of a biological process compiled by domain experts.

### Usage

```
org.Hs.egMsigdbC2CP <- dRDataLoader(RData=org.Hs.egMsigdbC2CP)</pre>
```

### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

```
org.Hs.egMsigdbC2CP <- dRDataLoader(RData=org.Hs.egMsigdbC2CP)
names(org.Hs.egMsigdbC2CP)</pre>
```

org. Hs. egMsigdbC2KEGG Annotations of Human Entrez Genes (EG) by C2:KEGG collections.

## **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C2:KEGG (KEGG pathways) gene sets and Human Entrez Genes. C2:KEGG gene sets are derived from the KEGG pathway database http://www.genome.jp/kegg/pathway.html.

# Usage

```
org.Hs.egMsigdbC2KEGG <- dRDataLoader(RData=org.Hs.egMsigdbC2KEGG)</pre>
```

#### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

## **Examples**

```
org.Hs.egMsigdbC2KEGG <- dRDataLoader(RData=org.Hs.egMsigdbC2KEGG)
names(org.Hs.egMsigdbC2KEGG)</pre>
```

```
org.Hs.egMsigdbC2REACTOME
```

Annotations of Human Entrez Genes (EG) by C2:REACTOME collections.

## Description

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C2:REACTOME (Reactome pathways) gene sets and Human Entrez Genes. C2:REACTOME gene sets are derived from the Reactome pathway database http://www.reactome.org/.

```
org.Hs.egMsigdbC2REACTOME <- dRDataLoader(RData=org.Hs.egMsigdbC2REACTOME)</pre>
```

#### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

### **Examples**

```
\label{lem:condition} $$ \operatorname{dRDataLoader}(RData=\operatorname{org.Hs.egMsigdbC2REACTOME})$ $$ \operatorname{d
```

org. Hs. egMsigdbC3MIR Annotations of Human Entrez Genes (EG) by C3:MIR collections.

### **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C3:MIR (microRNA targets) gene sets and Human Entrez Genes. C3 collections are about motif gene sets that contain genes that share a cis-regulatory motif that is conserved across the human, mouse, rat, and dog genomes, and represent known or likely regulatory elements in promoters and 3'-UTRs. C3:MIR gene sets contain genes that share a 3'-UTR microRNA binding motif.

### Usage

```
org.Hs.egMsigdbC3MIR <- dRDataLoader(RData=org.Hs.egMsigdbC3MIR)</pre>
```

## 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

```
org.Hs.egMsigdbC3MIR <- dRDataLoader(RData=org.Hs.egMsigdbC3MIR)
names(org.Hs.egMsigdbC3MIR)</pre>
```

org. Hs. egMsigdbC3TFT Annotations of Human Entrez Genes (EG) by C3:TFT collections.

# Description

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C3:TFT (transcription factor targets) gene sets and Human Entrez Genes. C3 collections are about motif gene sets that contain genes that share a cis-regulatory motif that is conserved across the human, mouse, rat, and dog genomes, and represent known or likely regulatory elements in promoters and 3'-UTRs. C3:TFT gene sets contain genes that share a transcription factor binding site defined in the TRANSFAC (version 7.4, http://www.gene-regulation.com/) database.

### **Usage**

```
org.Hs.egMsigdbC3TFT <- dRDataLoader(RData=org.Hs.egMsigdbC3TFT)</pre>
```

#### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

## **Examples**

```
org.Hs.egMsigdbC3TFT <- dRDataLoader(RData=org.Hs.egMsigdbC3TFT)
names(org.Hs.egMsigdbC3TFT)</pre>
```

 $\verb|org.Hs.egMsigdbC4CGN| Annotations of Human Entrez Genes (EG) by C4: CGN collections.$ 

# Description

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C4:CGN (cancer gene neighborhoods) gene sets and Human Entrez Genes. C4:CGN gene sets are defined by expression neighborhoods centered on 380 cancer-associated genes (see http://www.ncbi.nlm.nih.gov/pubmed/14593198).

```
org.Hs.egMsigdbC4CGN <- dRDataLoader(RData=org.Hs.egMsigdbC4CGN)</pre>
```

#### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

### **Examples**

```
org.Hs.egMsigdbC4CGN <- dRDataLoader(RData=org.Hs.egMsigdbC4CGN)
names(org.Hs.egMsigdbC4CGN)</pre>
```

org.Hs.egMsigdbC4CM

Annotations of Human Entrez Genes (EG) by C4:CM collections.

### **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C4:CM (cancer modules) gene sets and Human Entrez Genes. C4:CM gene sets are defined in http://www.ncbi.nlm.nih.gov/pubmed/15448693; the authors first compiled gene sets ('modules') from a variety of resources such as KEGG, GO, and others, and then by mining a large compendium of cancer-related microarray data, they identified 456 such modules as significantly changed in a variety of cancer conditions.

### Usage

```
org.Hs.egMsigdbC4CM <- dRDataLoader(RData=org.Hs.egMsigdbC4CM)</pre>
```

#### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

## **Examples**

```
org.Hs.egMsigdbC4CM <- dRDataLoader(RData=org.Hs.egMsigdbC4CM)
names(org.Hs.egMsigdbC4CM)</pre>
```

org.Hs.egMsigdbC5BP

Annotations of Human Entrez Genes (EG) by C5:BP collections.

## **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C5:BP (GO biological process) gene sets and Human Entrez Genes.

## Usage

```
org.Hs.egMsigdbC5BP <- dRDataLoader(RData=org.Hs.egMsigdbC5BP)</pre>
```

#### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

### **Examples**

```
org.Hs.egMsigdbC5BP <- dRDataLoader(RData=org.Hs.egMsigdbC5BP)
names(org.Hs.egMsigdbC5BP)</pre>
```

org.Hs.egMsigdbC5CC

Annotations of Human Entrez Genes (EG) by C5:CC collections.

### **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C5:CC (GO cellular component) gene sets and Human Entrez Genes.

```
org.Hs.egMsigdbC5CC <- dRDataLoader(RData=org.Hs.egMsigdbC5CC)</pre>
```

#### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

### **Examples**

```
org.Hs.egMsigdbC5CC <- dRDataLoader(RData=org.Hs.egMsigdbC5CC)
names(org.Hs.egMsigdbC5CC)</pre>
```

org.Hs.egMsigdbC5MF

Annotations of Human Entrez Genes (EG) by C5:MF collections.

### **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C5:MF (GO molecular function) gene sets and Human Entrez Genes.

## Usage

```
org.Hs.egMsigdbC5MF <- dRDataLoader(RData=org.Hs.egMsigdbC5MF)</pre>
```

## 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

```
org.Hs.egMsigdbC5MF <- dRDataLoader(RData=org.Hs.egMsigdbC5MF)
names(org.Hs.egMsigdbC5MF)</pre>
```

org.Hs.egMsigdbC6

org.Hs.egMsigdbC6

Annotations of Human Entrez Genes (EG) by C6 collections.

### **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C6 oncogenic signature gene sets and Human Entrez Genes. C6 collections contain gene sets that represent signatures of cellular pathways which are often dis-regulated in cancer.

### Usage

```
org.Hs.egMsigdbC6 <- dRDataLoader(RData=org.Hs.egMsigdbC6)</pre>
```

#### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)
- 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

## **Examples**

```
org.Hs.egMsigdbC6 <- dRDataLoader(RData=org.Hs.egMsigdbC6)
names(org.Hs.egMsigdbC6)</pre>
```

org.Hs.egMsigdbC7

Annotations of Human Entrez Genes (EG) by C7 collections.

## **Description**

This data is prepared based on the molecular signatures database (Msigdb; http://www.broadinstitute.org/gsea/msigdb/index.jsp). An R object that contains associations between Msigdb C7 immunologic signature gene sets and Human Entrez Genes. C7 collections contain gene sets that represent cell states and perturbations within the immune system.

```
org.Hs.egMsigdbC7 <- dRDataLoader(RData=org.Hs.egMsigdbC7)</pre>
```

102 org.Hs.egPS

#### 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, and the 4 columns are "setID" (i.e. "Geneset standard name"), "name" (i.e. "Geneset brief description"), "namespace" (i.e. Geneset category code) and "distance" (always empty)

• 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

Subramanian et al. (2005) Gene set enrichment analysis: a knowledge-based approach for interpreting genome-wide expression profiles. *Proc Natl Acad Sci U S A.*, 102(43):15545-50.

### **Examples**

```
org.Hs.egMsigdbC7 <- dRDataLoader(RData=org.Hs.egMsigdbC7)
names(org.Hs.egMsigdbC7)</pre>
```

org.Hs.egPS

Annotations of Human Entrez Genes (EG) by phylostratific age (PS).

## Description

An R object that contains phylostratific age information for Human Entrez Genes. This data is prepared based on 1) SUPERFAMILY database which provides 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 species tree of life (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 Human Entrez gene is the first appearance of its domain architecture along the branch from the eukaryotic ancestor to the human, and thus can be measured by: i) the most recent common ancestor, ii) 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.

### Usage

```
org.Hs.egPS <- dRDataLoader(RData=org.Hs.egPS)</pre>
```

## 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). 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 nearest descendant with known taxonomic information.

org.Hs.egSF

 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 al. (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**

```
org.Hs.egPS <- dRDataLoader(RData=org.Hs.egPS)
names(org.Hs.egPS)</pre>
```

org.Hs.egSF

Annotations of Human Entrez Genes (EG) by domain superfamilies (SF).

## **Description**

An R object that contains domain superfamily information for Human Entrez Genes. This data is prepared based on SUPERFAMILY database, which provides SCOP domain architecture assignments to all completely sequenced genomes including eukaryotic genomes. The domain architecture for an Entrez gene is the protein product with the longest length of amino acids. Thus, domain superfamily information for Human Entrez gene is a list of domain superfamilies (excluding unknown gap) appearing in its domain architecture.

#### Usage

```
org.Hs.egSF <- dRDataLoader(RData=org.Hs.egSF)</pre>
```

## 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. SCOP domain superfamilies). The 4 columns are "setID" (i.e. "SCOP domain identifier"), "name" (i.e. "SCOP domain description"), "namespace" (i.e. "SCOP domain level") and "distance" (i.e. "SCOP domain classification").
- 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 al. (2011) SUPERFAMILY 1.75 including a domain-centric gene ontology method. *Nucleic Acids Res*, 39(Database issue):D427-34.

Andreeva et al. (2008) Data growth and its impact on the SCOP database: new developments. *Nucleic Acids Res*, 36(Database issue):D419-425

```
org.Hs.egSF <- dRDataLoader(RData=org.Hs.egSF)
names(org.Hs.egSF)</pre>
```

104 org.Hs.string

org.Hs.string	Human functional protein association network from STRING (version
	9.1).

### **Description**

An igraph object that contains a functional protein association network in human. The network is extracted from the STRING database (version 9.1). Only those associations with medium confidence (score>=0.4) are retained.

### Usage

```
org.Hs.string <- dRDataLoader(RData=org.Hs.string)</pre>
```

#### 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.

```
org.Hs.string <- dRDataLoader(RData=org.Hs.string)
org.Hs.string</pre>
```

org.Hs.string900

highest confidence (no less than 900).	org.Hs.string900	Human functional protein association network from STRING with highest confidence (no less than 900).
--	------------------	--

## Description

An igraph object that contains a functional protein association network in human. The network is extracted from the STRING database (version 9.1). Only those associations with medium confidence (score>=900) are retained.

## Usage

```
data(org.Hs.string900)
```

#### 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.

```
data(org.Hs.string900)
org.Hs.string900
```

106 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

```
org.Mm.eg <- dRDataLoader(RData=org.Mm.eg)</pre>
```

#### 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**

```
org.Mm.eg <- dRDataLoader(RData=org.Mm.eg)
names(org.Mm.eg)
org.Mm.eg$gene_info[1:5,]</pre>
```

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.

```
org.Mm.egD0 <- dRDataLoader(RData=org.Mm.egD0)</pre>
```

org.Mm.egGOBP

#### 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**

```
org.Mm.egD0 <- dRDataLoader(RData=org.Mm.egD0)
names(org.Mm.egD0)</pre>
```

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

```
org.Mm.egGOBP <- dRDataLoader(RData=org.Mm.egGOBP)</pre>
```

## 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.

108 org.Mm.egGOCC

### **Examples**

```
org.Mm.egGOBP <- dRDataLoader(RData=org.Mm.egGOBP)
names(org.Mm.egGOBP)</pre>
```

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

```
org.Mm.egGOCC <- dRDataLoader(RData=org.Mm.egGOCC)</pre>
```

## 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.

```
org.Mm.egGOCC <- dRDataLoader(RData=org.Mm.egGOCC)
names(org.Mm.egGOCC)</pre>
```

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

```
org.Mm.egGOMF <- dRDataLoader(RData=org.Mm.egGOMF)</pre>
```

#### 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**

```
org.Mm.egGOMF <- dRDataLoader(RData=org.Mm.egGOMF)
names(org.Mm.egGOMF)</pre>
```

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.

org.Mm.egHPON

#### Usage

```
org.Mm.egHPMI <- dRDataLoader(RData=org.Mm.egHPMI)</pre>
```

#### 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**

```
org.Mm.egHPMI <- dRDataLoader(RData=org.Mm.egHPMI)
names(org.Mm.egHPMI)</pre>
```

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

```
org.Mm.egHPON <- dRDataLoader(RData=org.Mm.egHPON)</pre>
```

# 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

#### 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**

```
org.Mm.egHPON <- dRDataLoader(RData=org.Mm.egHPON)
names(org.Mm.egHPON)</pre>
```

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

```
org.Mm.egHPPA <- dRDataLoader(RData=org.Mm.egHPPA)</pre>
```

#### 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.

## **Examples**

```
org.Mm.egHPPA <- dRDataLoader(RData=org.Mm.egHPPA)
names(org.Mm.egHPPA)</pre>
```

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org.Mm.egMP	Annotations of 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

```
org.Mm.egMP <- dRDataLoader(RData=org.Mm.egMP)</pre>
```

## 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**

```
org.Mm.egMP <- dRDataLoader(RData=org.Mm.egMP)
names(org.Mm.egMP)</pre>
```

org.Mm.egPS

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

# Description

An R object that contains phylostratific age information for Mouse Entrez Genes. This data is prepared based on 1) SUPERFAMILY database which provides 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 species tree of life (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, and thus can be measured by: i) the most recent common ancestor, ii) 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.egSF

#### Usage

```
org.Mm.egPS <- dRDataLoader(RData=org.Mm.egPS)</pre>
```

#### 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). 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 nearest descendant 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 al. (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**

```
org.Mm.egPS <- dRDataLoader(RData=org.Mm.egPS)
names(org.Mm.egPS)</pre>
```

org.Mm.egSF

Annotations of Mouse Entrez Genes (EG) by domain superfamilies (SF).

# **Description**

An R object that contains domain superfamily information for Mouse Entrez Genes. This data is prepared based on SUPERFAMILY database, which provides SCOP domain architecture assignments to all completely sequenced genomes including eukaryotic genomes. The domain architecture for an Entrez gene is the protein product with the longest length of amino acids. Thus, domain superfamily information for Mouse Entrez gene is a list of domain superfamilies (excluding unknown gap) appearing in its domain architecture.

```
org.Mm.egSF <- dRDataLoader(RData=org.Mm.egSF)</pre>
```

114 org.Mm.string

#### 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. SCOP domain superfamilies). The 4 columns are "setID" (i.e. "SCOP domain identifier"), "name" (i.e. "SCOP domain description"), "namespace" (i.e. "SCOP domain level") and "distance" (i.e. "SCOP domain classification").

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

Andreeva et al. (2008) Data growth and its impact on the SCOP database: new developments. *Nucleic Acids Res*, 36(Database issue):D419-425

## **Examples**

```
org.Mm.egSF <- dRDataLoader(RData=org.Mm.egSF)
names(org.Mm.egSF)</pre>
```

org.Mm.string

Mouse functional protein association network from STRING (version 9.1).

# Description

An igraph object that contains a functional protein association network in mouse. The network is extracted from the STRING database (version 9.1). Only those associations with medium confidence (score>=0.4) are retained.

# Usage

```
org.Mm.string <- dRDataLoader(RData=org.Mm.string)</pre>
```

#### 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

org.Rn.eg

- 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**

```
org.Mm.string \leftarrow dRDataLoader(RData=org.Mm.string) org.Mm.string
```

org.Rn.eg

Rat Entrez Genes (EG).

# **Description**

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

# Usage

```
org.Rn.eg <- dRDataLoader(RData=org.Rn.eg)</pre>
```

# 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**

```
org.Rn.eg <- dRDataLoader(RData=org.Rn.eg)
names(org.Rn.eg)
org.Rn.eg$gene_info[1:5,]</pre>
```

org.Rn.egGOCC

org.Rn.egGOBP	Annotations of Rat Entrez Genes (EG) by Gene Ontology Biological Process (GOBP).

## **Description**

An R object that contains associations between Gene Ontology Biological Process terms and Rat 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

```
org.Rn.egGOBP <- dRDataLoader(RData=org.Rn.egGOBP)</pre>
```

#### 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.

# **Examples**

```
org.Rn.egGOBP <- dRDataLoader(RData=org.Rn.egGOBP)
names(org.Rn.egGOBP)</pre>
```

org.Rn.egGOCC Annotations of Rat Entrez Genes (EG) by Gene Ontology Cellular Component (GOCC).

# **Description**

An R object that contains associations between Gene Ontology Cellular Component terms and Rat 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.

```
org.Rn.egGOCC <- dRDataLoader(RData=org.Rn.egGOCC)</pre>
```

org.Rn.egGOMF

#### 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.

#### **Examples**

```
org.Rn.egGOCC <- dRDataLoader(RData=org.Rn.egGOCC)
names(org.Rn.egGOCC)</pre>
```

org.Rn.egGOMF

Annotations of Rat Entrez Genes (EG) by Gene Ontology Molecular Function (GOMF).

## **Description**

An R object that contains associations between Gene Ontology Molecular Function terms and Rat 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

```
org.Rn.egGOMF <- dRDataLoader(RData=org.Rn.egGOMF)</pre>
```

#### 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**

```
org.Rn.egGOMF <- dRDataLoader(RData=org.Rn.egGOMF)
names(org.Rn.egGOMF)</pre>
```

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org.Rn.egPS

Annotations of Rat Entrez Genes (EG) by phylostratific age (PS).

#### **Description**

An R object that contains phylostratific age information for Rat Entrez Genes. This data is prepared based on 1) SUPERFAMILY database which provides 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 species tree of life (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 Rat Entrez gene is the first appearance of its domain architecture along the branch from the eukaryotic ancestor to the rat, and thus can be measured by: i) the most recent common ancestor, ii) 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.

## Usage

```
org.Rn.egPS <- dRDataLoader(RData=org.Rn.egPS)</pre>
```

#### 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). 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 nearest descendant 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 al. (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**

```
org.Rn.egPS <- dRDataLoader(RData=org.Rn.egPS)
names(org.Rn.egPS)</pre>
```

org.Rn.egSF

org.Rn.egSF

Annotations of Rat Entrez Genes (EG) by domain superfamilies (SF).

# Description

An R object that contains domain superfamily information for Rat Entrez Genes. This data is prepared based on SUPERFAMILY database, which provides SCOP domain architecture assignments to all completely sequenced genomes including eukaryotic genomes. The domain architecture for an Entrez gene is the protein product with the longest length of amino acids. Thus, domain superfamily information for Rat Entrez gene is a list of domain superfamilies (excluding unknown gap) appearing in its domain architecture.

## Usage

```
org.Rn.egSF <- dRDataLoader(RData=org.Rn.egSF)</pre>
```

#### 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. SCOP domain superfamilies). The 4 columns are "setID" (i.e. "SCOP domain identifier"), "name" (i.e. "SCOP domain description"), "namespace" (i.e. "SCOP domain level") and "distance" (i.e. "SCOP domain classification").
- 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 al. (2011) SUPERFAMILY 1.75 including a domain-centric gene ontology method. *Nucleic Acids Res*, 39(Database issue):D427-34.

Andreeva et al. (2008) Data growth and its impact on the SCOP database: new developments. *Nucleic Acids Res*, 36(Database issue):D419-425

## **Examples**

```
org.Rn.egSF <- dRDataLoader(RData=org.Rn.egSF)
names(org.Rn.egSF)</pre>
```

org.Rn.string

Rat functional protein association network from STRING (version 9.1).

# Description

An igraph object that contains a functional protein association network in rat. The network is extracted from the STRING database (version 9.1). Only those associations with medium confidence (score>=0.4) are retained.

```
org.Rn.string <- dRDataLoader(RData=org.Rn.string)</pre>
```

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#### 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**

```
org.Rn.string <- dRDataLoader(RData=org.Rn.string)
org.Rn.string</pre>
```

TCGA\_mutations

TCGA mutational profiles across 12 major cancer types from Kandoth et al. (2013)

# Description

This dataset is available from TCGA, containing somatic mutational profiles for 3096 cancer samples with survival data. These cancer samples belong to one of 12 major cancer types, including breast adenocarcinoma (BRCA), lung adenocarcinoma (LUAD), lung squamous cell carcinoma (LUSC), uterine corpus endometrial carcinoma (UCEC), glioblastoma multiforme (GBM), head and neck squamous cell carcinoma (HNSC), colon and rectal carcinoma (COAD/READ), bladder urothelial carcinoma (BLCA), kidney renal clear cell carcinoma (KIRC), ovarian serous carcinoma (OV) and acute myeloid leukaemia (LAML). For each patient sample, somatic mutations are represented as a profile of states on genes, where non-zero entry indicates a gene for which how many mutations have occurred in the tumor relative to germ line. The dataset is provided as an 'ExpressionSet' object.

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## Usage

```
data(TCGA_mutations)
```

#### Value

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

- assayData: a matrix of 19171 genes X 3096 samples
- phenoData: variables describing sample phenotypes (i.e. columns in assayData), including clinical/survival information about samples: "time" (i.e. survival time in days), "status" (i.e., survival status: 0=alive; 1=dead), "Age" (the patient age in years), "Gender" (the patient gender: male/female), "TCGA\_tumor\_type", "Tumor\_stage", "Tumor\_grade"
- featureData: variables describing features (i.e. rows in assayData), including information about features/genes: "EntrezID" for gene EntrezID, "Symbol" for gene symbol, "Desc" for gene description, "Synonyms" for gene symbol alias

#### References

Kandoth et al. (2013). Mutational landscape and significance across 12 major cancer types. *Nature*, 502(7471):333-9.

## **Examples**

```
#TCGA_mutations <- dRDataLoader(RData=TCGA_mutations)
data(TCGA_mutations)
TCGA_mutations
library(Biobase)
# extract information about the first 5 samples
pData(TCGA_mutations)[1:5,]
# extract information about the first 5 features
fData(TCGA_mutations)[1:5,]
# number of samples for each cancer type
table(pData(TCGA_mutations)$TCGA_tumor_type)</pre>
```

visBoxplotAdv

Function to visualise a data frame using advanced boxplot

#### **Description**

visBoxplotAdv is supposed to visualise a data frame using advanced boxplot. In addition to boxplot, a scatter plot is also drawn with various methods to avoid co-incident points so that each point is visible (with fine-controling the color and plotting character). Also, these points can be pies or thermometers, which allows an additional proportation data to be visualised as well.

```
visBoxplotAdv(formula, data, orientation = c("vertical", "horizontal"),
method = c("center", "hex", "square", "swarm"), corral = c("none",
   "gutter", "wrap", "random", "omit"), corralWidth, cex = 1, spacing = 1,
   breaks = NULL, labels, at = NULL, add = FALSE, log = FALSE,
   xlim = NULL, ylim = NULL, xlab = NULL, ylab = NULL,
   pch = c("circles", "thermometers", "pies")[1], col = par("col"),
```

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```
bg = NA, pwpch = NULL, pwcol = NULL, pwbg = NULL, pwpie = NULL,
do.plot = TRUE, do.boxplot = TRUE, boxplot.notch = FALSE,
boxplot.border = "#888888CO", boxplot.col = "transparent", ...)
```

## **Arguments**

formula a formula, such as 'y ~ grp', where 'y' is a numeric vector of data values to be

split into groups according to the grouping variable 'grp' (usually a factor)

data a data.frame (or list) from which the variables in 'formula' should be taken.

orientation the orientation. It can be one of "vertical" for the vertical orientation, "horizon-

tal" for the horizontal orientation

method the method for arranging the points. It can be one of "swarm" for arranging

points in increasing order (if a point would overlap an existing point, it is shifted sideways (along the group axis) by a minimal amount sufficient to avoid overlap), "center" for first discretizing the values along the data axis (in order to create more efficient packing) and then using a square grid to produce a symmetric swarm, "hex" for first discretization and then arranging points in a hexagonal grid, and "square" for first discretization and then arranging points in a square

grid

corral the method to adjust points that would be placed outside their own group region.

It can be one of "none" for not adjusting runaway points, "gutter" for collecting runaway points along the boundary between groups, "wrap" for wrapping runaway points to produce periodic boundaries, "random" for placing runaway

points randomly in the region, and "omit" for omitting runaway points

corralWidth the width of the "corral" in user coordinates

cex size of points relative to the default. This must be a single value

breaks breakpoints (optional). If NULL, breakpoints are chosen automatically

spacing relative spacing between points

labels labels for each group. Recycled if necessary. By default, these are inferred from

the data

at numeric vector giving the locations where the swarms should be drawn; defaults

to '1:n' where n is the number of groups

add whether to add to an existing plot

log whether to use a logarithmic scale on the data axis

xlim limits for x-axis
ylim limits for y-axis
xlab labels for x-aixs
ylab labels for y-aixs

pch plotting characters, specified by group and recycled if necessary. In additon to

the convertional pch values, it can also be "circles", "thermometers", or "pies". For "pies" (or "thermometers"), users can also specify the proportional values (see below "pwpie") to visualise another information in the pie (or themometer)

chart

col plotting colors, specified by group and recycled if necessary

bg plotting background, specified by group and recycled if necessary

pwpch point-wise version of pch

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pwcol	point-wise version of col
pwbg	point-wise version of bg
pwpie	point-wise proportion used when drawing pies or themometers
do.plot	whether to draw main plot
do.boxplot	whether to draw boxplot. It only works when the main plot is drawn
	whether to draw a notch in the boxplot. If the notches of two plots do not overlap this is 'strong evidence' that the two medians differ
boxplot.border	the color for the outlines of the boxplots
boxplot.col	the color for the bodies of the boxplots
•••	additional graphic parameters for the plot

## Value

A data frame with plotting information. It has the same row names as the input data

## Note

none

## See Also

visBoxplotAdv

# **Examples**

```
data(TCGA_mutations)
pd <- Biobase::pData(TCGA_mutations)
# only tumor types "LAML" or "BLCA"
data <- pd[pd$TCGA_tumor_type=="LAML" | pd$TCGA_tumor_type=="BLCA",]
labels <- levels(as.factor(data$TCGA_tumor_type))
# colors for gender
pwcol <- as.numeric((data$Gender))
# pie for relative age
pwpie <- data$Age/(max(data$Age))
out <- visBoxplotAdv(formula=time ~ TCGA_tumor_type, data=data,
pch="pies", pwcol=pwcol, pwpie=pwpie)
legend("topright", legend=levels(data$Gender), box.col="transparent",
pch=19, col=unique(pwcol))</pre>
```

visDAG Function to visualise a direct acyclic graph (DAG) with node colorings according to a named input data vector (if provided)

# Description

visDAG is supposed to visualise a direct acyclic graph (DAG) with node colorings according to a named input data vector (if provided)

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#### Usage

```
visDAG(g, data = NULL, height = 7, width = 7, margin = rep(0.1, 4),
colormap = c("yr", "bwr", "jet", "gbr", "wyr", "br", "rainbow", "wb",
"lightyellow-orange"), ncolors = 40, zlim = NULL, colorbar = T,
colorbar.fraction = 0.1, newpage = T,
layout.orientation = c("left_right", "top_bottom", "bottom_top",
"right_left"), node.info = c("none", "term_id", "term_name", "both",
"full_term_name"), graph.node.attrs = NULL, graph.edge.attrs = NULL,
node.attrs = NULL)
```

# **Arguments**

g an object of class "igraph"

data a named input data verctor used to color-code vertices/nodes. The input data

vector must have names, and these names should include all node names of input graph, i.e. V(g)\$name, since there is a mapping operation. The way of how to color-code is to map values in the data onto the whole colormap (see the

next arguments: colormap, ncolors, zlim and colorbar)

height a numeric value specifying the height of device width a numeric value specifying the width of device

margin margins as units of length 4 or 1

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. "lightyellow-orange" (by default), "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/data 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 data is null, it always sets to

false

colorbar.fraction

the relative fraction of colorbar block against the device size

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

opening a new page

layout.orientation

the orientation of the DAG layout. It can be one of "left\_right" for the left-right layout (viewed from the DAG root point), "top\_bottom" for the top-bottom layout, "bottom\_top" for the bottom-top layout, and "right\_left" for the right-left

layout

node.info tells the ontology term information used to label nodes. It can be one of "none"

for no node labeling, "term\_id" for using Term ID, "term\_name" for using Term Name (the first 15 characters), "both" for using both of Term ID and Name (the first 15 characters), and "full\_term\_name" for using the full Term Name

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graph.node.attrs

a list of global node attributes. These node attributes will be changed globally. See 'Note' below for details on the attributes

graph.edge.attrs

a list of global edge attributes. These edge attributes will be changed globally. See 'Note' below for details on the attributes

node.attrs

a list of local edge attributes. These node attributes will be changed locally; as such, for each attribute, the input value must be a named vector (i.e. using Term ID as names). See 'Note' below for details on the attributes

#### Value

An object of class 'Ragraph'

#### Note

A list of global node attributes used in "graph.node.attrs":

- "shape": the shape of the node: "circle", "rectangle", "rect", "box" and "ellipse"
- "fixedsize": the logical to use only width and height attributes. By default, it sets to true for not expanding for the width of the label
- "fillcolor": the background color of the node
- "color": the color for the node, corresponding to the outside edge of the node
- "fontcolor": the color for the node text/labelings
- "fontsize": the font size for the node text/labelings
- "height": the height (in inches) of the node: 0.5 by default
- "width": the width (in inches) of the node: 0.75 by default
- "style": the line style for the node: "solid", "dashed", "dotted", "invis" and "bold"

A list of global edge attributes used in "graph.edge.attrs":

- "color": the color of the edge: gray by default
- "weight": the weight of the edge: 1 by default
- "style": the line style for the edge: "solid", "dashed", "dotted", "invis" and "bold"

A list of local node attributes used in "node.attrs" (only those named Term IDs will be changed locally!):

- "label": a named vector specifying the node text/labelings
- "shape": a named vector specifying the shape of the node: "circle", "rectangle", "rect", "box" and "ellipse"
- "fixedsize": a named vector specifying whether it sets to true for not expanding for the width of the label
- "fillcolor": a named vector specifying the background color of the node
- "color": a named vector specifying the color for the node, corresponding to the outside edge
  of the node
- "fontcolor": a named vector specifying the color for the node text/labelings
- "fontsize": a named vector specifying the font size for the node text/labelings
- "height": a named vector specifying the height (in inches) of the node: 0.5 by default
- "width": a named vector specifying the width (in inches) of the node: 0.75 by default
- "style": a named vector specifying the line style for the node: "solid", "dashed", "dotted", "invis" and "bold"

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#### See Also

dDAGreverse, dDAGroot, dDAGinduce, dDAGlevel

### **Examples**

```
# 1) load HPPA as igraph object
data(ig.HPPA)
g <- ig.HPPA
# 2) randomly select vertices as the query nodes
# the more common, the query nodes can be term id
nodes\_query \leftarrow V(g)[sample(V(g),5)]$name
# 3) obtain the induced subgraph based on all possible paths
subg <- dDAGinduce(g, nodes_query, path.mode="all_paths")</pre>
# 4) just visualise the induced subgraph
visDAG(g=subg, node.info="both")
# 5) color-code nodes/terms according to its level
data <- dDAGlevel(subg)</pre>
visDAG(g=subg, data=data, node.info="both")
# 5a) globally change the node and edge attributes
visDAG(g=subg, data=data, layout.orientation="top_bottom",
node.info="both",
graph.node.attrs=list(fixedsize=FALSE, shape="box", color="transparent"),
graph.edge.attrs=list(color="black"))
# 5b) locally highlight the root by changing its shape into "box"
root <- dDAGroot(subg)</pre>
root.shape <- "box"
names(root.shape) <- V(subg)[root]$name</pre>
visDAG(g=subg, data=data, node.info="both",
node.attrs=list(shape=root.shape))
# 5c) further locally remove the root labelling
root.label <- ""
names(root.label) <- V(subg)[root]$name</pre>
visDAG(g=subg, data=data, node.info="both",
node.attrs=list(shape=root.shape,label=root.label))
```

visGSEA

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

# **Description**

visGSEA 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).

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

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#### **Arguments**

an object of class "eTerm" eTerm which sample will be used. It can be index or sample names which\_sample which\_term which term will be used. It can be index or term ID or term names type of score weigth. It can be "0" for unweighted (an equivalent to Kolmogorovweight 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

```
dGSEA, dGSEAview
```

# **Examples**

```
## Not run:
visGSEA(eTerm, which_sample=1, which_term=1)
## End(Not run)
```

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",
"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", ...)
```

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## **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 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.

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```
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, ...)
```

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## **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) edge.lty line type for the edges (default 1) 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>
```

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```
vcolors <- palette.name(length(com))[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),]$ind</pre>
# 6) visualise graph using 1-dimensional arc diagram
visNetArc(g, ordering=ordering, labels=V(g)$name,
vertex.label.color=vcolors,
vertex.color=vcolors, vertex.frame.color=vcolors,
vertex.size=log(vdegrees)+0.1)
# 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.frame.color=vcolors,
vertex.shape="sphere")
```

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	<pre>an object of class "communities" (see http://igraph.sourceforge.net/doc/ R/communities.html)</pre>
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

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ordering a numeric vector about the ordering of vertices. It is optional. It is highly recommend to order vertices according to communities and degrees short name for the colormap. It can be one of "jet" (jet colormap), "bwr" (bluecolormap white-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 vertex.size the size of each vertex. By default, it is decided according to node degrees 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", "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". 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) mark.expand a numeric scalar or vector, the size of the border around the marked vertex 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

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## **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=vcolors, vertex.shape="sphere",
mark.groups=mark.groups, mark.col=mark.col,
mark.border=mark.border, mark.shape=1, mark.expand=10,
edge.color=edge.color)
```

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

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#### Usage

```
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, 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

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 http:

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

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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	
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>
```

 $\verb|visNetReorder|$ 

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

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#### Usage

```
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", ...)
```

## **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

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>

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

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## 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>
```

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