# Package 'BioNAR'

July 9, 2025

Title Biological Network Analysis in R

Version 1.10.0

**Description** the R package BioNAR, developed to step by step analysis of PPI network. The aim is to quantify and rank each protein's simultaneous impact into multiple complexes based on network topology and clustering. Package also enables estimating of co-occurrence of diseases across the network and specific clusters pointing towards shared/common mechanisms.

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addEdgeAtts

Copy edge attributes from one graph to another

### **Description**

Copy edge attributes from one graph to another

### Usage

```
addEdgeAtts(GG, gg)
```

### **Arguments**

igraph object, source of attributesigraph object, attributes recipient

#### Value

annotated version of gg igraph object

### **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
GG <- igraph::read_graph(file, format="gml")
gg<-findLCC(GG)
gg <- addEdgeAtts(GG, gg)
edge_attr_names(gg)</pre>
```

annotateGeneNames

Annotate Human Gene Names

#### **Description**

For the protein-protein interaction (PPI) or disease gene interaction (DGN) graphs that have EntrezID as a vertex name this function extract standard name from org.Hs.eg.db and annotate vertices.

### Usage

```
annotateGeneNames(gg, orgDB = org.Hs.eg.db, keytype = "ENTREZID")
```

#### **Arguments**

gg igraph object to annotate

orgDB ordDB object, by default human is assumed from org.Hs.eg.db

keytype type of IDs stored in the name vertex attribute, by default ENTREZID is assumed.

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#### **Details**

If vertex name attrubite stores not EntrezID or network is build not from human genes, other OrgDb-class object could be provided in orgDB and one of keytypes from that object that correspond to the nature of the vertex name attrubite could be provided in the keytype attribute.

If for some vertices name attrubite does not match keys with particular keytypes in the orgDB object, empty string is added as GeneName.

#### Value

igraph object with new vertex attribute GeneName

#### **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
agg<-annotateGeneNames(gg)
# due to error in org.Hs.eg.db we have to manually check annotation of one node
idx <- which(V(agg)$name == '80273')
paste(V(agg)$GeneName[idx], 'GRPEL1')</pre>
```

annotateGoBP

Add GO BP annotation to the graph vertices

### Description

The function loads an annotation data matrix called annoF, which contains three columns; the first containing gene Entrez IDs, the second gene GO BP ID terms, the third gene GO BP description terms. The function then performs a many-to-one mapping of each matrix row to a network vertex using matching Entrez IDs, filling the vertices attributes GO\_BP\_ID and GO\_BP.

### Usage

```
annotateGoBP(gg, annoF, idatt = "name")
```

#### **Arguments**

gg graph to update

annoF annotation matrix in Pair form

idatt optional name of the vertex attribute to map to the annotation data. frame first

column

#### Value

annotated igraph object

#### See Also

getAnnotationVertexList

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### **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
sfile<-system.file("extdata", "flatfile.go.BP.csv", package = "BioNAR")
goBP <- read.table(sfile, sep="\t", skip=1, header=FALSE,
strip.white=TRUE, quote="")
sgg <- annotateGoBP(gg, goBP)</pre>
```

annotateGoCC

Add GO CC annotation to the graph vertices

#### Description

The function loads an annotation data matrix called annoF, which contains three columns; the first containing gene Entrez IDs, the second gene GO ID terms, the third gene GO CC description terms. The function then performs a many-to-one mapping of each matrix row to a network vertex using matching Entrez IDs, filling the vertices attributes GO\_CC\_ID and GO\_CC.

#### Usage

```
annotateGoCC(gg, annoF, idatt = "name")
```

### **Arguments**

gg graph to update

annoF annotation matrix in Pair form

idatt optional name of the vertex attribute to map to the annotation data.frame first

column

### Value

annotated igraph object

#### See Also

getAnnotationVertexList

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
sfile<-system.file("extdata", "flatfile.go.CC.csv", package = "BioNAR")
goCC <- read.table(sfile, sep="\t", skip=1, header=FALSE,
strip.white=TRUE, quote="")
sgg <- annotateGoCC(gg, goCC)</pre>
```

annotateGoMF 7

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Add GO MF annotation to the graph vertices

#### **Description**

The function loads an annotation data matrix called annoF, which contains three columns; the first containing gene Entrez IDs, the second gene GO MF ID terms, the third gene GO MF description terms. The function then performs a many-to-one mapping of each matrix row to a network vertex using matching Entrez IDs, filling the vertices attributes GO\_MF\_ID and GO\_MF.

### Usage

```
annotateGoMF(gg, annoF, idatt = "name")
```

### **Arguments**

gg graph to update

annoF annotation matrix in Pair form

idatt optional name of the vertex attribute to map to the annotation data.frame first

column

#### Value

annotated igraph object

### See Also

getAnnotationVertexList

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
sfile<-system.file("extdata", "flatfile.go.MF.csv", package = "BioNAR")
goMF <- read.table(sfile, sep="\t", skip=1, header=FALSE,
strip.white=TRUE, quote="")
sgg <- annotateGoMF(gg, goMF)</pre>
```

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Annotate nodes with GO terms

### **Description**

For the protein-protein interaction (PPI) or disease gene interaction (DGN) graphs that have EntrezID as a vertex name this function extract GeneOntolgy annotation from orgDB, which should be OrgDb-class, split them into three ontology group (MF,BP,CC) and annotate vertices with .

#### Usage

```
annotateGOont(gg, orgDB = org.Hs.eg.db, keytype = "ENTREZID", idatt = "name")
```

#### **Arguments**

gg	igraph	object to	annotate
55	1514711	object to	amounce

orgDB ordDB object, by default human is assumed from org.Hs.eg.db

keytype type of IDs stored in the name vertex attribute, by default ENTREZID is assumed.

idatt optional name of the vertex attributes that contains IDs matching the keytype

#### **Details**

If vertex name attrubite stores not EntrezID or network is build not from human genes, other OrgDb-class object could be provided in orgDB and one of keytypes from that object that correspond to the nature of the vertex name attrubite could be provided in the keytype attribute.

If for some vertices name attrubite does not match keys with particular keytypes in the orgDB object, empty string is added as GeneName.

### Value

igraph object with new vertex attribute GeneName

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
ggGO <- annotateGOont(gg)</pre>
```

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annotateInterpro	Add InterPro Family and Domain annotation to the graph vertices

### **Description**

Function takes data from annoF matrix and add them to attributes InterPro\_Family for term and InterPro\_Family\_ID for IDs.

### Usage

```
annotateInterpro(gg, annoF, annoD, idatt = "name")
```

### **Arguments**

gg	graph to update
annoF	family annotation matrix in Pair form
annoD	domain annotation matrix in Pair form

idatt optional name of the vertex attributes that contains Entrez IDs

#### **Details**

Function takes data from annoD matrix and add them to attributes InterPro\_Domain for term and InterPro\_Domain\_ID for IDs.

#### Value

annotated igraph object

#### See Also

getAnnotationVertexList

```
annotate {\tt Presynaptic} \quad \textit{Add presynaptic functional groups}
```

### Description

Function takes from anno matrix manually curated presynaptic genes functional annotation derived from Boyken at al. (2013) doi:10.1016/j.neuron.2013.02.027 and add them to attributes PRESYNAPTIC.

#### Usage

```
annotatePresynaptic(gg, anno, idatt = "name")
```

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### **Arguments**

gg graph to update

anno annotation matrix in Pair form

idatt optional name of the vertex attributes that contains Entrez IDs

#### Value

annotated igraph object

#### See Also

getAnnotationVertexList

#### **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
sfile<-system.file("extdata", "PresynAn.csv", package = "BioNAR")
pres <- read.csv(sfile,skip=1,header=FALSE,strip.white=TRUE,quote="")
gg <- annotatePresynaptic(gg, pres)</pre>
```

annotateSCHanno

Add SCHanno synaptic functional groups

#### **Description**

The function loads an annotation data matrix of functional groups for schizopherina risk genes (1) called anno, which contains three columns; the first containing gene Entrez IDs, the second gene functional group ID terms, the third gene functional group description terms. The function then performs a many-to-one mapping of each matrix row to a network vertex using matching Entrez IDs, filling the SCHanno vertices attribute.

#### **Usage**

```
annotateSCHanno(gg, anno, idatt = "name")
```

### **Arguments**

gg igraph object to annotate

anno annotation matrix in Pairs form

idatt optional name of the vertex attributes that contains Entrez IDs

annotateTopOntoOVG

#### **Details**

#### References:

Lips E, Cornelisse L, Toonen R, Min J, Hultman C, the International Schizophernia Consortium, Holmans P, Donovan M, Purcell S, Smit A, Verhage M, Sullivan P, Visscher P, D P: Functional gene group analysis identifies synaptic gene groups as risk factor for schizophrenia. Molecular Psychiatry 2012,17:996–1006.

#### Value

annotated igraph object

#### See Also

getAnnotationVertexList

#### **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.csv", package = "BioNAR")
tbl <- read.csv(file, sep="\t")
gg <- buildNetwork(tbl)
afile<-system.file("extdata", "SCH_flatfile.csv", package = "BioNAR")
dis <- read.table(afile, sep="\t", skip=1, header=FALSE,
strip.white=TRUE, quote="")
agg<-annotateSCHanno(gg, dis)</pre>
```

annotateTopOntoOVG

Annotate graph with disease terms

#### **Description**

The function loads a human disease annotation matrix called dis, which contains three columns; the first containing gene Entrez IDs, the second gene Human Disease Ontology (HDO) ID terms, the third gene HDO description terms. For human protein-protein interaction (PPI) or disease-gene networks (DGN) that have human Entrez IDs for the igraph vertex name attribute. The function then performs a many-to-one mapping of each matrix row to a network vertex using matching Entrez IDs, filling the vertices attributes TopOnto\_OVG\_HDO\_ID and TopOnto\_OVG.

#### Usage

```
annotateTopOntoOVG(gg, dis, idatt = "name")
```

## Arguments

gg	igraph object to annotate
dis	annotation matrix in Pairs form
idatt	optional name of the vertex attributes that contains Entrez IDs

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#### Value

annotated igraph object

#### See Also

getAnnotationVertexList

### **Examples**

annotateVertex

Generic annotation function

### **Description**

Function to build and fill a vertex attribute given an igraph object. Where parameter 'name' is the new vertex attribute name and values are filled from a two column data.frame supplied to 'value' attribute. The first containing vertex name IDs, and the second the vertex annotation value.

#### Usage

```
annotateVertex(gg, name, values, idatt = "name")
```

#### **Arguments**

gg igraph object to annotate
name name of the attribute
values annotation data frame

idatt optional name of the vertex attribute to map to the annotation data.frame first

column

#### **Details**

As a first step all attributes with provided names will be removed.

### Value

igraph object where vertex attribute name contains annotation terms separated by semicolon.

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#### See Also

get Annotation Vertex List

### **Examples**

```
g1 <- make_star(10, mode="undirected")
V(g1)$name <- letters[1:10]
m<-rbind(data.frame(ID=letters[1:10], terms=letters[1:10]),
data.frame(ID=letters[1:10], terms=LETTERS[1:10]))
g2<-annotateVertex(g1, name='cap', values=m)
V(g2)$cap</pre>
```

applpMatrixToGraph

Add attributes to the vertex.

### **Description**

This function suits more for updating calculated vertex properties rather than node annotation. For the later case use annotateVertex.

### Usage

```
applpMatrixToGraph(gg, m)
```

#### **Arguments**

gg igraph object

m matrix of values to be applied as vertex attributes. matrix should contains col-

umn "ID" to map value to the vertex.

#### **Details**

Unlike annotateVertex, which is able to collapse multiple annotation terms, this function assume that vertex ID values are unique in the m matrix and corresponds to the name vertex attribute. If graph has no name vertex attribute error will be raised.

#### Value

modified igraph object

#### See Also

annotateVertex

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### **Examples**

```
g1 <- make_star(10, mode="undirected")
V(g1)$name <- letters[1:10]
m<-cbind(ID=letters[1:10], capital=LETTERS[1:10])
g1<-BioNAR::applpMatrixToGraph(g1,m)
V(g1)$capital</pre>
```

**BioNAR** 

BioNAR: Biological Network Analysis in R

### **Description**

The R package BioNAR, developed to step by step analysis of PPI network. The aim is to quantify and rank each protein's simultaneous impact into multiple complexes based on network topology and clustering. Package also enables estimating of co-occurrence of diseases across the network and specific clusters pointing towards shared/common mechanisms.

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#### See Also

Useful links:

• Report bugs at https://github.com/lptolik/BioNAR/issues/

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buildConsensusMatrix Build a consensus matrix from list of resampled clustering matrices outputted from the function sampleGraphClust

### **Description**

Build a consensus matrix from list of resampled clustering matrices outputted from the function sampleGraphClust

### Usage

buildConsensusMatrix(lcc)

### **Arguments**

1cc

list of membership matrices obtained from the sampleGraphClust

#### **Details**

Function build a consensus matrix from list of membership matrices, which are a three column matrix: the first column contains the vertex IDs of input network; the second column the vertex IDs of the subsampled network, or -1 if the vertex has been masked; the third column the cluster membership of subsampled network, or -1 if vertex has been masked. The randomised resampled membership matrices could be obtained from the function sampleGraphClust.

### Value

consensus matrix of Nvert X Nvert

buildNetwork

Build network from data.table

### **Description**

Wrapper for graph\_from\_data\_frame function which will always return the largest connect component for a given network ff. The function will also annotated the edges in ff with PubMed data from kw if provided.

#### Usage

```
buildNetwork(ff, kw = NA, LCC = TRUE, simplify = TRUE)
```

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### **Arguments**

ff network structure data frame with first two columns defining the network edge

nodes

kw pmid keyword annotation data.frame. If NA no annotation will be added

LCC if TRUE only largest connected component is returned simplify if TRUE loops and multiple edges will be removed

#### Value

igraph object of the largest connected component

### **Examples**

```
f<-data.frame(A=c('A', 'A', 'B', 'D'), B=c('B', 'C', 'C', 'E'))
gg<-buildNetwork(f)
V(gg)$name</pre>
```

calcAllClustering

Calculate memberships for all clustering algorithms and store them on the graph vertices.

#### **Description**

This function will call calcClustering for each clustering algorithm given in our predefined list. In the event no clustering could be performed, warnings will be issued and no new vertex attribute added to the graph.

#### Usage

```
calcAllClustering(gg, weights = NULL)
```

### **Arguments**

gg graph for analysis

weights The weights of the edges. It must be a positive numeric vector, NULL or NA. If

it is NULL and the input graph has a 'weight' edge attribute, then that attribute will be used. If NULL and no such attribute is present, then the edges will have equal weights. Set this to NA if the graph was a 'weight' edge attribute, but you don't want to use it for community detection. A larger edge weight means a stronger connection for this function. The weights value is ignored for the

spectral clustering.

#### Value

new graph object with all membership results stored as a vertex attribute.

calcBridgeness 17

#### See Also

calcClustering

#### **Examples**

```
g1 <- make_star(10, mode="undirected")
V(g1)$name <- letters[1:10]
g1<-calcAllClustering(g1)
clusteringSummary(g1)</pre>
```

calcBridgeness

Helper function that uses getBridgeness to calculate graph node bridgeness values for selected algorithm and consensus matrix and save them as a graph attribute BRIDGENESS. <alg> with <alg> replaced by the selected algorithm name.

### Description

Helper function that uses <code>getBridgeness</code> to calculate graph node bridgeness values for selected algorithm and consensus matrix and save them as a graph attribute <code>BRIDGENESS.<alg></code> with <code><alg></code> replaced by the selected algorithm name.

### Usage

```
calcBridgeness(gg, alg, conmat)
```

### **Arguments**

gg igraph object

alg clustering algorithm

conmat consensus matrix calculated with that algorithm

#### Value

graph with additional attributes to store Bridgeness value

### See Also

getBridgeness

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### **Examples**

```
library(BioNAR)
karate <- make_graph("Zachary")
# We need vertex ID in the 'name' attribute of the vertex
V(karate)$name<-c(LETTERS,letters)[1:vcount(karate)]
set.seed(100)
gg <- calcClustering(karate, 'louvain')
cnmat <- makeConsensusMatrix(gg, N=10, alg = 'louvain', type = 2, mask = 10)
gg<-calcBridgeness(gg, alg = 'louvain', cnmat)
hist(V(gg)$BRIDGENESS.louvain)</pre>
```

calcCentrality

Calculate the vertex centrality measures

#### **Description**

Calculate the vertex centrality measures (degree, betweenness, closeness, semi-local, etc....) for each graph vertex and store each result as new vertex attribute in the graph.

#### Usage

```
calcCentrality(gg, weights = NULL)
```

#### **Arguments**

gg igraph object

weights Possibly a numeric vector giving edge weights. If this is NULL and the graph

has a weight edge attribute, then the attribute is used. If this is NA then no

weights are used (even if the graph has a weight attribute).

#### **Details**

A wrapper function that first calls getCentralityMatrix, to calculate all vertex centrality measures, and then applpMatrixToGraph to store each centrality result as a new vertex attribute in the graph. The use of weights explained in details in getCentralityMatrix.

### Value

modified igraph object

### See Also

```
getCentralityMatrix()
```

```
data(karate,package='igraphdata')
ggm<-calcCentrality(karate)
V(ggm)$DEG</pre>
```

#### calcCentralityExternalDistances

Function to calculate a distance matrix between a list of permuted vertex centrality matrices and a unperturbed reference matrix.

### Description

Function to calculate a distance matrix between a list of permuted vertex centrality matrices and a unperturbed reference matrix.

### Usage

```
calcCentralityExternalDistances(m, 1, keepOrder = FALSE, dist = "euclidean")
```

### **Arguments**

m reference matrix, for example centrality obtained by invocation getCentralityMatrix

list of permuted matrix, for example centrality obtained by invocation getRandomGraphCentrality keepOrder if FALSE valuess will be sorted

methods available from dist function

### Value

matrix with seven columns containing distances between each element of 1 and reference matrix m

#### See Also

```
getRandomGraphCentrality
getCentralityMatrix
calcCentralityInternalDistances
```

```
data(karate,package='igraphdata')
m<-getCentralityMatrix(karate)
gnp<-list()
for(i in 1:10){
      gnp[[i]]<-getRandomGraphCentrality(karate,type = 'gnp')
}
gnpEDist<-calcCentralityExternalDistances(m,gnp)
summary(gnpEDist)</pre>
```

calcCentralityInternalDistances

Function calculates a set of distance metrics between each vertex pair given a list of vertex centrality matrices

### **Description**

Function calculates a set of distance metrics between each vertex pair given a list of vertex centrality matrices

#### Usage

```
calcCentralityInternalDistances(l, keepOrder = FALSE, dist = "euclidean")
```

### **Arguments**

list of matrices, for example centrality obtained by invocation getRandomGraphCentrality

keepOrder if FALSE values will be sorted before distance calculations

dist methods available from dist function

#### Value

matrix with seven columns containing distances between all pairs of 1 elements.

#### See Also

```
getRandomGraphCentrality
getCentralityMatrix
calcCentralityExternalDistances
```

```
data(karate,package='igraphdata')
m<-getCentralityMatrix(karate)
gnp<-list()
for(i in 1:10){
      gnp[[i]]<-getRandomGraphCentrality(karate,type = 'gnp')
}
gnpIDist<-calcCentralityInternalDistances(gnp)
summary(gnpIDist)</pre>
```

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calcClustering	Calculate community membership for given clustering algorithm and store the results as new vertex attributes in the graph
	store the results as her retrest annies in the graphin

### **Description**

When applying resampling the clustering results of a clustering algorithm applied to a graph can differ due to the stochastic nature of the resampling algorithm. To allow reproducible downstream analysis clustering results are stored as vertex attributes in the graph. This function call <code>getClustering</code> and stores community membership as new vertex attribute in the graph, and Modularity as a new graph attribute prefix with the alg name.

#### Usage

```
calcClustering(gg, alg, weights = NULL)
```

### **Arguments**

gg igraph object to cluster alg algorithm to apply

weights The weights of the edges. It must be a positive numeric vector, NULL or NA. If

it is NULL and the input graph has a 'weight' edge attribute, then that attribute will be used. If NULL and no such attribute is present, then the edges will have equal weights. Set this to NA if the graph was a 'weight' edge attribute, but you don't want to use it for community detection. A larger edge weight means a stronger connection for this function. The weights value is ignored for the

spectral clustering.

### **Details**

NOTE: getClustering verifies algorithm names with match.arg so correct membership will be calculated, but name of the attribute is taken from alg argument, so it is possible that vertex attribute name won't exactly match name of the algorithm from link{getClustering}.

#### Value

modified igraph object with calculated membership stored as a vertex attribute and modularity as a graph attribute

#### See Also

getClustering

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#### **Examples**

```
karate <- make_graph("Zachary")
# We need vertex ID in the 'name' attribute of the vertex
V(karate)$name<-c(LETTERS,letters)[1:vcount(karate)]
g<-calcClustering(karate, 'louvain')
vertex_attr_names(g)
graph_attr(g, 'louvain')</pre>
```

calcDiseasePairs

Calculate each disease-disease pair overlap given a list of disease

### Description

Calculate each disease-disease pair overlap (or separation) on a given PPI network model, based on analysis described in Menche et al. 2015

### Usage

```
calcDiseasePairs(
   gg,
   name,
   diseases = NULL,
   permute = c("none", "random", "binned")
)
```

### **Arguments**

gg interactome network as igraph object

name of the attribute that stores disease annotation

diseases list of diseases to match

permute type of permutations. none – no permutation is applied, random – annotation

is randomly shuffled, binned – annotation is shuffled in a way to preserve node

degree-annotation relationship by degreeBinnedGDAs.

#### Value

list with three matrices:

- disease separation Ndisease X Ndisease matrix of separations
- gene\_disease\_separation Ngenes X Ndisease+2 matrix of gene-disease separation
- disease\_localisation matrix with diseases in rows and number of genes (N), average and standard deviation of gene-disease separation in columns

### References

Menche, J. et al. Uncovering disease-disease relationships through the incomplete interactome. Science, 347, (6224):1257601 (2015).

23 calcEntropy

#### See Also

```
degreeBinnedGDAs
sampleDegBinnedGDA
```

### **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")</pre>
gg <- igraph::read_graph(file, format="gml")</pre>
agg<-annotateGeneNames(gg)
# due to error in org.Hs.eg.db we have to manually check annotation of one node
idx \leftarrow which(V(agg)\normalfont{me} == '80273')
paste(V(agg)$GeneName[idx], 'GRPEL1')
p <- calcDiseasePairs(</pre>
agg,
name = "TopOntoOVGHDOID",
diseases = c("DOID:10652", "DOID:3312", "DOID:12849"),
permute = "n"
p$disease_separation
```

calcEntropy

Calculate the graph entropy for each perturbed vertex, and save the results as new vertex attributes in the graph.

#### **Description**

This function calculate the graph entropy for each perturbed vertex by calling getEntropy, and save the results as new vertex attributes SR\_UP and SR\_DOWN in the graph.

### Usage

```
calcEntropy(gg, maxSr = NULL, exVal = NULL)
```

### **Arguments**

igraph object gg

the maximum entropy rate maxSR, if NULL getEntropyRate will be called. maxSr exVal expression values boundaries. Two columns are expected: xx and lambda. If

NULL default values c(2,14) and c(-14,14) will be used for xx and lambda

respectively.

#### **Details**

According to Teschendorf et al., 2010, network entropy measure quantifies the degree of randomness in the local pattern information flux around single genes. For instance, in metastatic cancer this measure was found significantly higher than in non-metastatic and helped to identify genes and entire pathways involved on metastasis. However, for the assessment of scale-free structure we do not actually require gene expression data as it based solely on the network topology.

24 calcMembership

#### Value

graph with SR\_UP and SR\_DOWN vertex attributes storing the graph entropy values with over- or under-expressing each vertex.

#### See Also

```
getEntropy()
```

Other Entropy Functions: getEntropy(), getEntropyRate(), plotEntropy()

#### **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.csv", package = "BioNAR")
tbl <- read.csv(file, sep="\t")
gg <- buildNetwork(tbl)
gg<-annotateGeneNames(gg)
# due to error in org.Hs.eg.db we have to manually check annotation of one node
idx <- which(V(gg)$name == '80273')
paste(V(gg)$GeneName[idx], 'GRPEL1')
gg<- calcEntropy(gg)</pre>
```

calcMembership

Calculate cluster memberships for the graph.

#### **Description**

Calculates the clustering membership for one of the 10 clustering algorithms defined in function getClustering

#### Usage

```
calcMembership(
   gg,
   alg = c("lec", "wt", "fc", "infomap", "louvain", "sgG1", "sgG2", "sgG5", "spectral"),
   weights = NULL
)
```

### **Arguments**

gg igraph object to cluster alg algorithm name

weights

The weights of the edges. It must be a positive numeric vector, NULL or NA. If it is NULL and the input graph has a 'weight' edge attribute, then that attribute will be used. If it is NULL and no such attribute is present, then the edges will have equal weights. Set this to NA if the graph has a 'weight' edge attribute, but you don't want to use it for community detection. A larger edge weight means a stronger connection for this function. The weights value is ignored for the spectral clustering.

calcReclusterMatrix 25

#### Value

data.frame with columns names and membership

#### See Also

getClustering

#### **Examples**

```
karate <- make_graph("Zachary")
# We need vertex ID in the 'name' attribute of the vertex
V(karate)$name<-c(LETTERS,letters)[1:vcount(karate)]
m<-calcMembership(karate, 'lec')
head(m)</pre>
```

calcReclusterMatrix

Hierarchical graph clustering

### Description

This function takes in a gg and initial vertex community membership values mem as returned by calcMembership, and then performs a reclustering of the graph given the clustering algorithm alg to those clusters of size greater than CnMAX

### Usage

```
calcReclusterMatrix(
   gg,
   mem,
   alg,
   CnMAX = 10,
   weights = NULL,
   keepSplit = FALSE
)
```

#### **Arguments**

gg graph to cluster

mem data.frame with previous level clustering results

alg algorithm to apply

CnMAX maximus size of the cluster in mem that will not be processed

weights The weights of the edges. It must be a positive numeric vector, NULL or NA. If

it is NULL and the input graph has a 'weight' edge attribute, then that attribute will be used. If NULL and no such attribute is present, then the edges will have equal weights. Set this to NA if the graph was a 'weight' edge attribute, but you don't want to use it for community detection. A larger edge weight means a stronger connection for this function. The weights value is ignored for the

spectral clustering.

26 calcSparsness

keepSplit logical, wether to keep previous membership in the output matrix

### Value

remembership matrix, that contains vertex ID membership and result of reclustering

### **Examples**

```
data(karate,package='igraphdata')
alg<-'louvain'
mem<-calcMembership(karate,alg = alg)
remem<-calcReclusterMatrix(karate,mem,alg,10)</pre>
```

calcSparsness

Calculate sparsness of the graph.

### **Description**

For a simple unweighted, undirected graph G(N,E). Network sparseness is defined as the ratio of the actual number of graph edges (E) to the maximum number of edges possible in a graph with same number of vertices (N): E/binom(N,2)

#### Usage

```
calcSparsness(gg)
```

### **Arguments**

gg

graph to evaluate

#### Value

sparsness value

```
file <- system.file("extdata", "PPI_Presynaptic.csv", package = "BioNAR")
tbl <- read.csv(file, sep="\t")
gg <- buildNetwork(tbl)
calcSparsness(gg)</pre>
```

clusteringSummary 27

clusteringSummary

Matrix of cluster characteristics

#### **Description**

Function to calculate basic summary statistics after apply clustering algorithm:

- N number of vertices in the graph vcount
- mod clustering modularity modularity, the ratio of edges found within communities to the number of edges found between communities, relative to a randomised model
- C number of clusters
- Cn1 number of singletones (clusters of size 1)
- Cn100 number of clusters containing more than 100 nodes
- mu the ratio of edges found within communities to the number of edges found between communities
- Min. C minimum of the cluster size
- 1st Qu. C first quartile of the cluster size
- Median C median of the cluster size
- Mean C average cluster size
- 3rd Qu. C third quartile of the cluster size
- Max. C maximum of the cluster size

### Usage

```
clusteringSummary(
   gg,
   att = c("lec", "wt", "fc", "infomap", "louvain", "sgG1", "sgG2", "sgG5", "spectral")
)
```

### Arguments

```
gg graph to analyse
att vector of attribute names that contains membership data
```

#### Value

matrix of clustering characteristics

```
data(karate,package='igraphdata')
g<-calcAllClustering(karate)
clusteringSummary(g)</pre>
```

28 clusterORA

clusterORA

Calculate annotation enrichment for clusters in the graph

### **Description**

Calculate the cluster enrichment of a graph given a clustering algorithm alg and vertex annotation attribute 'name'. Function generates an enrichment table, one row for each cluster, containing: size of the cluster (Cn), number of annotated vertices in the graph  $F_n$  (Fn), number of annotated vertices in the cluster  $\mu$  (Mu), odds ratio (OR) and its 95% Confidence interval  $[CI_l, CI_u]$  (CI1 and CIu), two fold enrichment values  $F_e$  (Fe) and  $F_c$  (Fc). We also provide the list of vertices from the cluster that contribute to the annotation term, p.value of enrichment (pval) and depletion (palt) using the Hypergeometric test, adjusted p.values using Benjamini and Yekutieli correction (BY).

### Usage

```
clusterORA(g, alg, name, vid = "name", alpha = 1, col = COLLAPSE)
```

#### **Arguments**

g	graph to get annotation from
alg	cluster algorithm and membership attribute name
name	annotation attribute name
vid	attribute to be used as a vertex ID
alpha	probability threshold
col	list separation character in attribute, by default is ;

#### **Details**

Given the enrichment results, we can calculate the log of the Odds Ratio (OR) as:

$$\ln(OR) = \ln(\frac{\mu(N - F_n + \mu - C_n)}{(C_n - \mu)(F_n - \mu)})$$

and it's upper and lower 95% Confidence Interval:

$$CI(\ln(OR)) = \ln(OR) \pm 1.96\sqrt{\frac{1}{\mu} + \frac{1}{C_n - \mu} + \frac{1}{F_n - \mu} + \frac{1}{N - F_n + \mu - C_n}}$$

Using the odds ratio allows us to distinguish functionally enriched communities relative to functionally depleted communities.

Two types of fold enrichment values calculated as follow:

$$F_e = \frac{\left(\frac{\mu}{F_n}\right)}{\left(\frac{C_n}{N}\right)}$$

$$F_c = \frac{\left(\frac{\mu}{C_n}\right)}{\left(\frac{C_n}{N}\right)}$$

compMembership 29

#### Value

A table with overrepresentation results. Each row corresponds to a tested annotation in particular cluster. The columns are the following:

- alg name of the clustering algorithm;
- cl cluster ID;
- FL name of the enriched term;
- N number vertices in the network;
- Fn number of vertices in the graph annotated by term F1  $(F_n)$ ;
- Cn size of the cluster;
- Mu number of vertices in the cluster annotated by term F1 ( $\mu$ );
- OR odds ratio:
- CII odds ratio 95% confidence interval lower bound  $(CI_l)$ ;
- CIu odds ratio 95% confidence interval upper bound( $CI_u$ );
- Fe fold enrichment  $F_e$ ;
- Fc fold enrichment  $F_c$ ;
- pval an enrichment p-value from hypergeometric test;
- padj a BY-adjusted p-value;
- palt an depletion p-value from hypergeometric test;
- paltadj a BY-adjusted depletion p-value;
- overlapGenes vector with overlapping genes.

#### **Examples**

```
options("show.error.messages"=TRUE)
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
g <- igraph::read_graph(file, format="gml")
anL<-getAnnotationVertexList(g, 'TopOntoOVGHDOID')
res<-clusterORA(g, alg='louvain', name='TopOntoOVGHDOID', vid='name')
andf<-unique(data.frame(ID=vertex_attr(g, 'TopOntoOVGHDOID'),
Term=vertex_attr(g, 'TopOntoOVG')))
rr<-merge(andf, res, by.y='FL', by.x='ID')
rr[order(rr$cl), ]</pre>
```

compMembership

Calculate cluster memberships for one of the graph component.

### **Description**

Calculates the clustering membership for one of the 10 clustering algorithms defined in function getClustering for selected graph component

30 degreeBinnedGDAs

### Usage

```
compMembership(
   gg,
   alg = c("lec", "wt", "fc", "infomap", "louvain", "sgG1", "sgG2", "sgG5", "spectral"),
   compnum = 0,
   weights = NULL
)
```

### **Arguments**

gg igraph object to cluster

alg algorithm name

compnum number of the componet to cluster

weights The weights of the edges. It must be a positive numeric vector, NULL or NA. If

it is NULL and the input graph has a 'weight' edge attribute, then that attribute will be used. If it is NULL and no such attribute is present, then the edges will have equal weights. Set this to NA if the graph has a 'weight' edge attribute, but you don't want to use it for community detection. A larger edge weight means a stronger connection for this function. The weights value is ignored for the

spectral clustering.

#### Value

data.frame with columns names and membership

#### See Also

getClustering

degreeBinnedGDAs

Prepare mapping for degree-aware annotation shuffling.

### Description

Function to randomly shuffle vertex annotation terms, whilst preserving the vertex degree originally found with that annotation term.

#### Usage

```
degreeBinnedGDAs(gg, GDA, dtype)
```

### Arguments

gg graph to analyse

GDA vertex annotations returned by prepareGDA dtype list of unique annotation terms to analyze

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#### Value

mapping matrix between vertices, vertex-degree groups and annotation terms.

#### See Also

```
prepareGDA
getAnnotationList
sampleDegBinnedGDA
```

#### **Examples**

```
options("show.error.messages"=TRUE)
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
agg<-annotateGeneNames(gg)
# due to error in org.Hs.eg.db we have to manually check annotation of one node
idx <- which(V(agg)$name == '80273')
paste(V(agg)$GeneName[idx], 'GRPEL1')
gda<-prepareGDA(agg, 'TopOntoOVGHDOID')
m<-degreeBinnedGDAs(agg, gda, getAnnotationList(gda))
c(dim(m), vcount(agg), length(getAnnotationList(gda)))
head(m)</pre>
```

diseasome

Barabasi's Diseasome Network

### **Description**

In the paper Goh.t al. (2007) doi:10.1073/pnas.0701361104 Barabasi with colleagues published Diseasome: a network of disorders and disease genes linked by known disorder–gene associations. We extract definition of the genes, disorders and interactions from papers supplementary materials and store it as graph object.

### Usage

diseasome

#### **Format**

A bipartite graph as graph object.

Vertex attributes: 'name' for the node ID, 'Name' for the human readable node name, 'Disorder.class', 'Type' for the human readable node type, 'label' and 'shape' for plotting the graph, 'type' the node type for bipartite graph representation.

#### **Details**

Diseasesome is a bipartite graph that have nodes of two types gene and disease and links are allowed only between nodes of different types. It could be projected to Human Disease Network (HDN) and Disease Gene Network (DGN).

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#### **Source**

Goh, K.-I. et al. The human disease network. Proc. Natl. Acad. Sci. U.S.A. 104, 8685–8690 (2007). https://pnas.org/doi/full/10.1073/pnas.0701361104

escapeAnnotation

Escapes elements of list in annotation.

### Description

In situations when a given list of annotation ID terms may not be well formatted, and therefore not be interoperated as unique. For example, given a list of HDO IDs: HDO:14, HDO:143, HDO:1433, and HDO:14330, a grep for the term HDO:14 could return: HDO:143, HDO:1433, HDO:14330. To avoid this all terms should be enclosed in escape characters, which unlikely to find within annotation itself.

### Usage

```
escapeAnnotation(annVec, col = COLLAPSE, esc = ESC)
```

### **Arguments**

annVec vector of annotation strings
col term list separator character
esc escape character

# Details

NOTE: spaces are treated as regular characters, no trimming is applied before or after escaping.

#### Value

vector of annotation strings with elements escaped

#### See Also

unescapeAnnotation

```
annVec<-apply(matrix(letters, ncol=13), 2, paste, collapse=';')
cbind(annVec, escapeAnnotation(annVec, ';', '|'))</pre>
```

evalCentralitySignificance

Compare distance distributions of internal and external distances

#### **Description**

Function to compare two distance distributions using the Kolmogorov-Smirnov test. Where the first distance distribution is generated internally and calculates the distance between random graph centralities. The second distance distribution is generated externally, and measures the distance between random and the original graph centralities.

### Usage

```
evalCentralitySignificance(dmi, dme)
```

### **Arguments**

dmi distribution of internal distances between random graph centralities

dme distribution of external distances between random and original graph centralities

#### Value

list of lists for each centrality value in the input matrix three element list is created where ks contains Kolmogorov-Smirnov test result from class ks.test; pval contains Kolmogorov-Smirnov test pvalue; and dt contains input distribution.

### See Also

ks.test

```
data(karate,package='igraphdata')
m<-getCentralityMatrix(karate)
gnp<-list()
for(i in 1:10){
          gnp[[i]]<-getRandomGraphCentrality(karate,type = 'gnp')
}
gnpIDist<-calcCentralityInternalDistances(gnp)
gnpEDist<-calcCentralityExternalDistances(m,gnp)

simSig<-evalCentralitySignificance(gnpIDist,gnpEDist)
sapply(simSig,function(.x).x$ks$p.value)</pre>
```

34 fitDegree

 ${\tt findLCC}$ 

Find Largest Connected Component of the graph

### Description

Find Largest Connected Component of the graph

### Usage

```
findLCC(GG)
```

### **Arguments**

GG

igraph object to analyze

### Value

igraph representation LCC

### **Examples**

```
g1 <- make_star(10, mode="undirected") %du% make_ring(7) %du% make_ring(5)
lcc<-findLCC(g1)
summary(lcc)</pre>
```

fitDegree

Fit Power Law to degree distribution.

### Description

Fit a Powerlaw distribution to graph's degree distribution using the R "PoweRlaw" package (version 0.50.0) (Gillespie, 2015)

### Usage

```
fitDegree(
  DEG,
  Nsim = 100,
  plot = FALSE,
  DATAleg = "Fit power-law",
  threads = 4,
  WIDTH = 480,
  HEIGHT = 480,
  legpos = "bottomleft",
  showErr = TRUE
)
```

fitSigmoid 35

#### **Arguments**

DEG	degree distribution
Nsim	number of bootstrap iterations
plot	logical, do you want plot to be drawn
DATAleg	legend string for degree data
threads	number of parallel computational threads
WIDTH	width of the plot in ptx
HEIGHT	heigth of the plot in ptx
legpos	position of the legend @seealso legend()
showErr	logical, do you want error on the plot legend

#### Value

an object of class law-class with results of fitting

### **Examples**

```
##No: of bootstrap iterations use nsim > 100 for reliable result
nsim <- 10

##Legend Titles
Legend <- "Presynaptic PPI"

file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
pFit <- fitDegree( as.vector(igraph::degree(graph=gg)),
DATAleg=Legend,threads=1, Nsim=nsim)</pre>
```

fitSigmoid

Fit Fold-enrichment distribution to sigmoid function

### **Description**

This function calculates fit of the Fold-Enrichment distribution to the sigmoid function with the levels of noise specidied in SDV for all clustering algorithms, which have non-zero SUM3\$`Psig&ORsig` in the enrichment table summary results. The function returns the list in which each element contains result for one of the noise level.

### Usage

```
fitSigmoid(stat, SDv = c(0, 0.05, 0.1, 0.5))
```

### Arguments

stat enrichment results obtained from summaryStats

SDv vector of noise SD values

36 flatfile.go.CC.csv

#### **Details**

Results are repersented as a list with five elements:

- gridplot that allow comparison of fitting for different clustering algorithms;
- plots the list of individual plots from gridplot;
- fitInfo the data.frame that contains results of fitting, such as message, number of iterations and exit code;
- parInfo values and standard deviations for all sigmoid parameters;
- ks table of Kolmogorov-Smirnov test p-values.

Grid plot is designed in a way to be viewed in the device at least 12 inches in width and 12 inches in height.

#### Value

list of fitted functions tables and plots

#### See Also

summaryStats()

flatfile.go.BP.csv

Annotation from Gene Ontology Biological Process (GO\_BP)

#### **Description**

Annotation, downloaded from Gene Ontology for Biological Proceess domain. The table has columns: the first containing gene gene functional group ID terms, the second gene functional group description terms, the third - Human gene Entrez IDs; in csv format

#### See Also

annotateGoBP

flatfile.go.CC.csv

Annotation from Gene Ontology Cellular Compartment (GO\_CC)

### **Description**

Annotation, downloaded from Gene Ontology for Cellular Compartment domain. The table has columns: the first containing gene gene functional group ID terms, the second gene functional group description terms, the third - Human gene Entrez IDs; in csv format

#### See Also

annotateGoCC

flatfile.go.MF.csv 37

flatfile.go.MF.csv

Annotation from Gene Ontology Molecular Function (GO\_MF)

# **Description**

Annotation, downloaded from Gene Ontology for Molecular Function domain. The table has columns: the first containing gene gene functional group ID terms, the second gene functional group description terms, the third - Human gene Entrez IDs; in csv format

#### See Also

annotateGoMF

```
flatfile_human_gene2HDO.csv
```

Human Gene Disease Associations (GDA)

# Description

Annotation derived from Human Disease Ontology database (HDO). The table contains three columns; the first containing gene Entrez IDs, the second gene Human Disease Ontology (HDO) ID terms, the third gene HDO description terms; in csv format

# See Also

annotateTopOntoOVG

getAnnotationList

Extract unique values from annotations.

#### **Description**

It is not uncommon to find both duplicated vertex annotation terms, and vertices annotated with multiple terms, in a given annotation list. This function creates a vector of unique annotation terms for each vertex given an input annotation list.

# Usage

```
getAnnotationList(
  annVec,
  col = COLLAPSE,
  sort = c("none", "string", "frequency")
)
```

## **Arguments**

annVec vector of annotation strings
col list separator character
sort how to sort the result list

#### Value

vector of unique annotation terms

#### See Also

getAnnotationVertexList

## **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
annVec<-V(gg)$TopOntoOVG
al<-getAnnotationList(annVec)
al</pre>
```

getAnnotationVertexList

Return vertex list for each term in annotation attribute

# Description

For different purposes annotation of graph vertices could be represented in three forms:

Pairs dataframe with vertex ID and annotation terms

Vertex Annotation list named with vertex ID and containing terms annotating each vertex

Annotation Vertices list named with term and containing vertex IDs

## Usage

```
getAnnotationVertexList(g, name, vid = "name", col = COLLAPSE)
```

# Arguments

g	graph to get annotation from
name	annotation attribute name
vid	attribute to be used as a vertex ID

col list separation character in attribute, by default is ;

getBridgeness 39

#### **Details**

This function takes Vertex Annotation from vertex attribute and convert it to Annotation Vertices form

#### Value

named list with annotation in Annotation Vertices form

# **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
avl<-getAnnotationVertexList(gg, 'TopOntoOVGHDOID')
head(avl)</pre>
```

getBridgeness

Calculate bridginess from consensus matrix

# **Description**

Bridginess takes into account a vertices shared community membership together with its local neighbourhood. It was proposed in Nepusz et al., 2008 doi:10.1103/PhysRevE.77.016107.

#### Usage

```
getBridgeness(gg, alg, conmat)
```

#### **Arguments**

gg igraph object

alg clustering algorithm

conmat consensus matrix calculated with that algorithm

#### **Details**

Function assumes clustering already been performed by the clustering algorithm, and its membership values stored in vertex attributes. If clustering algorithm vertex alg attribute is not found an error will be issued.

#### Value

data.frame with first column contains vertex ID, if GeneName attribute assigned to the vertices its value will be stored as a second column, the last column contains bridginess values for the

40 getCentralityMatrix

#### **Examples**

```
library(BioNAR)
karate <- make_graph("Zachary")
# We need vertex ID in the 'name' attribute of the vertex
V(karate)$name<-c(LETTERS,letters)[1:vcount(karate)]
gg <- calcClustering(karate, 'louvain')
cnmat <- makeConsensusMatrix(gg, N=10, alg = 'louvain', type = 2, mask = 10)
br<-getBridgeness(gg, alg = 'louvain', cnmat)</pre>
```

getCentralityMatrix

Calculate centrality measures for graph nodes.

#### **Description**

Calculate centrality measures for graph nodes.

## Usage

```
getCentralityMatrix(gg, weights = NULL)
```

## Arguments

gg igraph object

weights Possibly a numeric vector giving edge weights. If this is NULL and the graph

has a weight edge attribute, then the attribute is used. If this is NA then no

weights are used (even if the graph has a weight attribute).

## **Details**

The edge attribute weights treated differently by different functions calculating centrality measures. For example, betweenness use weights as an edge length, while in page\_rank "an edge with a larger weight is more likely to be selected by the surfer", which infer the opposite meaning. Taking into account that all methods in getClustering treat edge weights in the same way as page\_rank, we calculate the distance=l/weights as edge weights for BET, dBET, mnSP, and sdSP values. So we treat weights in the package consistently as the strength and closiness of vertices, rather the distance between them.

#### Value

data.frame with following columns:

- ID vertex ID
- DEG degree
- iDEG in-degree (directed graph only)
- oDEG out-degree (directed graph only)
- BET betweenness for undirected graph

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- dBET betweenness when directionality is taken into account (directed graph only)
- CC clustering coefficient
- SL semilocal centrality
- mnSP mean shortest path
- PR page rank for undirected graph
- dPR page rank when directionality is taken into account (directed graph only)
- sdSP standard deviation of the shortest path

# **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.csv", package = "BioNAR")
tbl <- read.csv(file, sep="\t")
gg <- buildNetwork(tbl)
m<-getCentralityMatrix(gg)</pre>
```

getClustering

Get clustering results for the graph.

## **Description**

Wrapper function for calculation of clustering for predefined set of ten algorithms:

- lec leading eigenvector community (version of cluster\_leading\_eigen), directed graph will be converted to undirected by as\_undirected with mode collapse;
- wt walktrap community cluster\_walktrap;
- fc fastgreedy community cluster\_fast\_greedy, directed graph will be converted to undirected by as\_undirected with mode collapse;
- infomap infomap community cluster\_infomap;
- louvain cluster\_louvain cluster\_louvain, directed graph will be converted to undirected by as\_undirected with mode collapse;
- sgG1 spin-glass model and simulated annealing clustering (version of cluster\_spinglass with spins=500 and gamma=1);
- sgG2 spin-glass model and simulated annealing clustering (version of cluster\_spinglass with spins=500 and gamma=2);
- sgG5 spin-glass model and simulated annealing clustering (version of cluster\_spinglass with spins=500 and gamma=7);
- spectral spectral modularity clustering spectral\_igraph\_communities;

# Usage

```
getClustering(
   gg,
   alg = c("lec", "wt", "fc", "infomap", "louvain", "sgG1", "sgG2", "sgG5", "spectral"),
   weights = NULL
)
```

## **Arguments**

gg igraph object to cluster
alg clustering algorithm name

weights The weights of the edges. It must be a positive numeric vector, NULL or NA. If

it is NULL and the input graph has a 'weight' edge attribute, then that attribute will be used. If NULL and no such attribute is present, then the edges will have equal weights. Set this to NA if the graph was a 'weight' edge attribute, but you don't want to use it for community detection. A larger edge weight means a stronger connection for this function. The weights value is ignored for the

spectral clustering.

#### **Details**

graph suppose to be undirected. If algorithm failed warning will be issued and function returned NULL.

Algorithm names are verified with match.arg.

#### Value

communities object or NULL if algorithm failed.

## **Examples**

```
data(karate,package='igraphdata')
c<-getClustering(karate,'lec')
c$modularity</pre>
```

```
getClusterSubgraphByID
```

Return induced subgraph for cluster

# Description

Function reads in a graph gg, vertex cluster membership vector mem, and returns an induced subgraph given a cluster membership number 'clID'.

#### Usage

```
getClusterSubgraphByID(clID, gg, mem)
```

#### **Arguments**

clID cluster ID to extracte
gg graph to analyze
mem membership vector

getCommunityGraph 43

# Value

induced subgraph as igraph object

# **Examples**

```
data(karate,package='igraphdata')
alg<-'louvain'
c<-getClustering(karate,alg = alg)
gc3<-getClusterSubgraphByID(3,karate,membership(c))
#plot(gc3,vertex.label=V(gc3)$name)</pre>
```

 ${\tt getCommunityGraph}$ 

Create new graph with communities as a nodes.

# Description

The idea based upon this StackOverflow answer

# Usage

```
getCommunityGraph(gg, membership)
```

# **Arguments**

gg graph to convert

membership participation list for new graph

# Value

community graph

```
data(karate,package='igraphdata')
alg<-'louvain'
mem<-calcMembership(karate,alg = alg)
cg<-getCommunityGraph(karate,mem$membership)</pre>
```

getDType

getDiseases

Get HDO disease IDs

# Description

Return vector of HDO disease IDs for synaptic PPI analysis.

# Usage

```
getDiseases()
```

# Value

vector of disease IDs of interest

# See Also

getDType

# **Examples**

getDiseases()

getDType

Get DiseaseTypes

# Description

Return vector of disease abbreviations for synaptic PPI analysis.

# Usage

```
getDType()
```

#### Value

vector of disease abbreviations for synaptic PPI analysis.

# See Also

getDiseases

```
getDType()
```

getDYNAMO 45

getDYNAMO	Calculate DYNAMO sensitivity matrix.

# Description

This function calculates sensitivity matrix that represents perturbation patterns defined by topology and edge weights of the network. If weights are signed value sensitivity matrix is able to reproduce not only activation but inhibition relationships in the network.

# Usage

```
getDYNAMO(g, attr = NULL, vid = "name", alpha = 0.9)
```

# Arguments

g	igraph object
attr	NULL or the name of edge attribute containing numerical weight values
vid	name of the vertex attribute to be used as row and column names
alpha	parameter characterizing the propagation strength, default value 0.9 taken from Santolini paper.

#### **Details**

Algorithm proposed in:

Santolini,M. and Barabasi,A.-L. (2018) Predicting perturbation patterns from the topology of biological networks. Proc Natl Acad Sci USA, 169, 201720589.

#### Value

sparce sensitivity matrix defined by the network topology and edge values

```
data(karate, package='igraphdata')
upgrade_graph(karate)
d<-getDYNAMO(karate,attr='weight')
df<-metlMatrix(d)
head(df)</pre>
```

46 getEntropy

getEntropy	ge	tEr	ntr	on	v
------------	----	-----	-----	----	---

Calculates vertex perturbation graph entropy.

#### Description

According to Teschendorf et al., 2010, network entropy measure quantifies the degree of randomness in the local pattern information flux around single genes. For instance, in metastatic cancer this measure was found significantly higher than in non-metastatic and helped to identify genes and entire pathways involved on metastasis. However, for the assessment of scale-free structure we do not actually require gene expression data as it based solely on the network topology.

#### Usage

```
getEntropy(gg, maxSr = NULL, exVal = NULL)
```

#### **Arguments**

gg igraph object

maxSr the maximum entropy rate maxSR, if NULL getEntropyRate will be called.

exval expression values boundaries. Two columns are expected: xx and lambda. If

NULL default values c(2,14) and c(-14,14) will be used for xx and lambda

respectively.

#### **Details**

In this function, following procedure described in (Teschendorff et al., 2015), all vertexes are artificially assigned a uniform weight then sequentially perturbed with the global entropy rate (SR) after each protein's perturbation being calculated and plotted against the log of the protein's degree. In case of scale-free or approximate scale-free topologies, we see a clear bi-modal response between over-weighted vertices and their degree and an opposing bi-phasic response in under-weighted vertices and their degrees.

#### Value

matrix containing for each Gene:

- Entrez ID,
- · Name,
- Degree,
- UP Graph Entropy values when gene is expressed up,
- DOWN Graph Entropy values when gene is expressed down.

#### Note

Entropy is calculated with respect to GeneName property, if there is no such vertex attribute in the graph vertex name will be copied to the GeneName attribute. If any NA is found in GeneNames error will be thrown.

getEntropyRate 47

#### See Also

Other Entropy Functions: calcEntropy(), getEntropyRate(), plotEntropy()

#### **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.csv", package = "BioNAR")
tbl <- read.csv(file, sep="\t")
gg <- buildNetwork(tbl)
gg<-annotateGeneNames(gg)
any(is.na(V(gg)$GeneName))
# due to error in org.Hs.eg.db we have to manually check annotation of one node
idx <- which(V(gg)$name == '80273')
paste(V(gg)$GeneName[idx], 'GRPEL1')
e<- getEntropy(gg)</pre>
```

getEntropyRate

Calculate the maximum entropy rate and initial entropy rate.

# **Description**

This function calculates the maximum entropy rate maxSR (maxSr) and initial entropy rate  $SR_0$  (SRo) given a connected network.

#### Usage

```
getEntropyRate(gg)
```

## **Arguments**

gg

igroph object

#### **Details**

The maximum entropy rate being calculated from the network's adjacency matrix:

$$maxSR = \sum_{i,j} p_{ij} = \frac{A_{ij}\nu_j}{\lambda\nu_i}$$

where  $\nu$  and  $\lambda$  are the leading eigenvector and eigenvalue of the network adjacency matrix A respectively.

The initial configuration occurs when the entropy for each node is maximal. This can be calculated by setting the expression value for each gene/node in the network to be the same, and thus the maximal node entropy is dependent only on the node's degree k:

$$SR_0 = \frac{1}{N\bar{k}} \sum_{j} k_j \log k_i$$

where N here is the number of nodes and  $\bar{k}$  the average node degree found in the network.

48 getGNP

# Value

list with values of maxSr and SRo

# See Also

```
Other Entropy Functions: calcEntropy(), getEntropy(), plotEntropy()
```

# **Examples**

```
karate <- make_graph("Zachary")
# We need vertex ID in the 'name' attribute of the vertex
V(karate)$name<-c(LETTERS,letters)[1:vcount(karate)]
ent <- getEntropyRate(karate)</pre>
```

getGNP

Generate random graph from reference

# **Description**

Function generates random G(n,p) Erdos-Renyi graph (sample\_gnp) with the same number of vertices and edges as in in the reference graph gg.

# Usage

```
getGNP(gg, ...)
```

# **Arguments**

```
gg reference graph
... additional arguments to be passed to sample_gnp
```

# Value

new instance of the random graph.

```
data(karate,package='igraphdata')
vcount(karate)
ecount(karate)
rg<- getGNP(karate)
vcount(rg)
ecount(rg)</pre>
```

getGraphCentralityECDF

Convert centrality matrix into ECDF

# **Description**

Convert centrality matrix into ECDF

# Usage

```
getGraphCentralityECDF(m)
```

## **Arguments**

m

centrality matrix from getCentralityMatrix invocation.

#### Value

list of several ecdf objects, corresponding to values in centrality matrix from getCentralityMatrix invocation.

# See Also

getCentralityMatrix

# **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.csv", package = "BioNAR")
tbl <- read.csv(file, sep="\t")
gg <- buildNetwork(tbl)
m<-getCentralityMatrix(gg)
ecdfL<-getGraphCentralityECDF(m)</pre>
```

getIDs

Utility function to get vertex ids from vertex attributes The function obtain attribute values and check duplicates in it. It fails if any duplicate found.

# Description

Utility function to get vertex ids from vertex attributes The function obtain attribute values and check duplicates in it. It fails if any duplicate found.

## Usage

```
getIDs(gg, idatt)
```

50 getPA

#### **Arguments**

gg graph

idatt attribute name

#### Value

idatt attribute values

getPA

Generate random graph from reference

# Description

The function generates random Barabasi-Albert graph (sample\_pa) with the same vertex number as in the reference graph gg and the power specified by parameter pwr. If pwr is missing, we are trying to estimate pwr from the reference graph gg.

# Usage

```
getPA(gg, pwr, ...)
```

## **Arguments**

gg reference graph

the power parameter for

pwr the power parameter for the sample\_pa

... additional parameters to be passed to the sample\_pa

#### Value

new instance of the random graph.

```
data(karate,package='igraphdata')
vcount(karate)
ecount(karate)
rg<- getPA(karate,pwr=1.25)
vcount(rg)
ecount(rg)</pre>
```

getRandomGraphCentrality

Centrality measures for random graphs induced by input one

#### **Description**

Generate a random graph that mimics the properties of the input graph and calls getCentralityMatrix to calculate all available vertex centrality measures. There are four different types of random graph to generate

#### Usage

```
getRandomGraphCentrality(
  gg,
  type = c("gnp", "pa", "cgnp", "rw"),
  power = NULL,
  weights = NULL,
  ...
)
```

## **Arguments**

gg template graph to mimic

type type of random graph to generate:

- gnp G(n,p) Erdos-Renyi model (sample\_gnp)
- pa Barabasi-Albert model (sample\_pa)
- cgnp new random graph from a given graph by randomly a dding/removing edges (sample\_correlated\_gnp)
- rw new random graph from a given graph by rewiring 25% of edges preserving the degree distribution sample\_gnp, sample\_correlated\_gnp, and sample\_pa

power

optional argument of the power of the preferential attachment to be passed to sample\_pa. If power is NULL the power of the preferential attachment will be estimated from fitDegree function.

weights

Possibly a numeric vector giving edge weights. If this is NULL and the graph has a weight edge attribute, then the attribute is used. If this is NA then no weights are used (even if the graph has a weight attribute).

... other parameters passed to random graph generation functions

#### Value

matrix of random graph vertices centrality measure.

#### See Also

getCentralityMatrix() for explanation of the use of weights.

52 getRobustness

# **Examples**

getRobustness

Calculate cluster robustness from consensus matrix

#### **Description**

This function takes as argument a network (gg), the name of a clustering algorithm (alg) which can be found in the network, and a consensus matrix (conmat) generated from the clustering network. The function uses the consensus matrix to generate a measure of cluster robustness  $C_{rob}$  (Crob) for each cluster (C) using the R function clrob. Briefly, this is done by summing elements of the consensus matrix that are found in the same cluster, and dividing this by the total number of entries in the matrix:

$$C_{rob} = \frac{2}{C_n(C_n - 1)} \sum_{\substack{i,j \in I_C \\ i \le j}} conmat_{i,j}$$

where  $I_C$  – indices of vertices of the cluster C,  $C_n$  is the number of nodes found inside the cluster C

#### Usage

```
getRobustness(gg, alg, conmat)
```

#### **Arguments**

gg igroph object
alg clustering algorithm
conmat consensus matrix

#### Value

data.frame that for each cluster C shows

- its size  $C_n$  (Cn),
- robustness  $C_{rob}$  (Crob) and
- robustness scaled to range between 0 and 1 (CrobScaled).

gofs 53

#### See Also

Other Robustness Functions: makeConsensusMatrix()

# **Examples**

```
karate <- make_graph("Zachary")
# We need vertex ID in the 'name' attribute of the vertex
V(karate)$name<-c(LETTERS,letters)[1:vcount(karate)]
alg<-'louvain'
gg<-calcClustering(karate, alg = alg)
conmat<-makeConsensusMatrix(gg, N=100, mask = 10, alg = alg, type = 2)
clrob<-getRobustness(gg, alg = alg, conmat)
clrob</pre>
```

gofs

Goodnes of fit KS test

# Description

This is internal function and do not suppose to be called by user.

#### Usage

```
gofs(x, rate, model, sigma2 = NULL, countDATA = TRUE)
```

# Arguments

X	steps along the Fe
rate	parameters of the sigmoid
model	fitted model
sigma2	noise strength
countDATA	should points to be counted

#### Value

list of ks. test values for each value in rate

54 layoutByCluster

law-class

Result of PawerLaw fit

# Description

Result of PawerLaw fit

#### **Slots**

```
fit displ-class result of power law fit.
p numeric.
alpha numeric degree of power-law.
SDxmin numeric bootstrap sd of Xmin.
SDalpha numeric bootstrap sd of alpha.
```

layoutByCluster

Calculate layout based upon membership

# Description

Function to split graph into clusters and layout each cluster independently..

# Usage

```
layoutByCluster(gg, mem, layout = layout_with_kk)
```

# **Arguments**

gg graph to layout

mem membership data.frame from calcMembership

layout algorithm to use for layout

#### Value

Layout in a form of 2D matrix.

#### See Also

```
igraph::layout_
```

```
data(karate,package='igraphdata')
alg<-'louvain'
mem<-calcMembership(karate,alg = alg)
lay<-layoutByCluster(karate,mem)
#plot(karate,layout=lay)</pre>
```

layoutByRecluster 55

layoutByRecluster

Calculate two-level layout from recluster matrix

# **Description**

Takes results of recluster and apply layoutByCluster to each

## Usage

```
layoutByRecluster(gg, remem, layout = layout_with_kk)
```

## **Arguments**

gg graph to layout

remem recluster result obtained by calcReclusterMatrix invocation

layout one of the layout algorithms from layout\_

#### Value

Layout in a form of 2D matrx.

# **Examples**

```
data(karate,package='igraphdata')
alg<-'louvain'
mem<-calcMembership(karate,alg = alg)
remem<-calcReclusterMatrix(karate,mem,alg,10)
lay<-layoutByRecluster(karate,remem)
#plot(karate,layout=lay)</pre>
```

makeConsensusMatrix

Function to make random resampling consensus matrix in memory

#### **Description**

Function to make random resampling consensus matrix in memory

# Usage

```
makeConsensusMatrix(
   gg,
   N = 500,
   mask = 20,
   alg,
   type,
   weights = NULL,
```

56 makeConsensusMatrix

```
reclust = FALSE,
Cnmax = 10
)
```

#### **Arguments**

gg graph to perturb

N number of perturbation steps
mask percentage of elements to perturbe

alg clustering alg.

type edges (1) or nodes (2) to mask

weights The weights of the edges. It must be a positive numeric vector, NULL or NA. If

it is NULL and the input graph has a 'weight' edge attribute, then that attribute will be used. If NULL and no such attribute is present, then the edges will have equal weights. Set this to NA if the graph was a 'weight' edge attribute, but you don't want to use it for community detection. A larger edge weight means a stronger connection for this function. The weights value is ignored for the

spectral clustering.

reclust logical to decide wether to invoke reclustering via recluster

Cnmax maximum size of the cluster in mem that will not be processed if reclustering is

invoked

#### **Details**

Function to assess the robustness of network clustering. A randomisation study is performed apply the same clustering algorithm to N perturbed networks, and which returns the consensus matrix where each vertex pair is assigned the probability of belong to the same cluster. The inputted network is perturbed by randomly removing a mask percentage of edges (type=1) or vertices (type=2) from the network before clustering.

#### Value

consensus matrix of Nvert X Nvert

#### See Also

Other Robustness Functions: getRobustness()

```
karate <- make_graph("Zachary")
# We need vertex ID in the 'name' attribute of the vertex
V(karate)$name<-c(LETTERS,letters)[1:vcount(karate)]
alg<-'louvain'
gg<-calcClustering(karate, alg = alg)
conmat<-makeConsensusMatrix(gg, N=100, mask = 10, alg = alg, type = 2)
dim(conmat)</pre>
```

makeMembership 57

makeMembership

Create membership data. frame from graph for arbitrary annotation

#### Description

Create membership data.frame from graph vertex attribute or vector of cluster names, IDs or indices. This function is simular to calcMembership but do not linked to clustering algorithm.

# Usage

```
makeMembership(gg, membership)
```

# **Arguments**

gg igraph object to assign membership

membership either name of the vertex attribute or vector of membership

# **Details**

Any annotation coercible to factor could be converted to the membership data.frame. This function is useful, for example, to make layout with layoutByCluster.

#### Value

data. frame with two columns names and membership

```
karate <- make_graph("Zachary")
# We need vertex ID in the 'name' attribute of the vertex
V(karate)$name<-c(LETTERS,letters)[1:vcount(karate)]
m<-makeMembership(karate,rep(c(1,2),length.out=vcount(karate)))
head(m)</pre>
```

58 metlMatrix

#### markBowTie

Calculates bow-tie decomposition and marks vertices with one of the following in the BowTie attribute:

- *SCC maximal strong connected component*;
- *IN vertices not in SCC, but SCC is reachable from them;*
- OUT vertices not in SCC, but reachable from SCC;
- TU vertices not in all three above, but reachable from IN and OUT is reachable from them (TUBES);
- IDR vertices not in SCC, but they are reachable from IN and OUT is NOT reachable from them (INTENDRILS);
- ODR vertices not is SCC, but they are NOT reachable from IN and OUT is reachable from them (OUTTENDRILS);
- *OTR all other vertices*.

# Description

Algorithm proposed in:

#### Usage

markBowTie(g)

# **Arguments**

g

graph to analyse

#### **Details**

"Bow-tie Decomposition in Directed Graphs" - Yang et al. IEEE (2011)

## Value

graph with BowTie vertex attribute

metlMatrix

Convert sparce matrix into triplet data. frame.

# **Description**

For very large graphs handling adjacency-like matrices is difficult due to its sparse nature. This function convert sparse matrix into triplet data.frame with row and column indices and names, and cell value.

normModularity 59

# Usage

```
metlMatrix(sparceM)
```

#### **Arguments**

sparceM

sparce matrix to convert into triplet data. frame

# Value

data.frame with three colums:

- i row index;
- j column index;
- x cell value;
- Rname i-th row name;
- Cname j-th column name.

# **Examples**

```
data(karate, package='igraphdata')
upgrade_graph(karate)
Ws <- as_adjacency_matrix(karate,type='both',attr='weight',sparse = TRUE)
mdf<-metlMatrix(Ws)
head(mdf)</pre>
```

 ${\tt normModularity}$ 

Calculates the normalised network modularity value.

# **Description**

Function to compare network Modularity of input network with networks of different size and connectivity.

# Usage

```
normModularity(
   gg,
   alg = c("lec", "wt", "fc", "infomap", "louvain", "sgG1", "sgG2", "sgG5"),
   Nint = 1000,
   weights = NULL
)
```

60 normModularity

#### **Arguments**

gg graph object to analyze
alg clustering algorithm
Nint number of iterations

weights The weights of the edges. It must be a positive numeric vector, NULL or NA. If

it is NULL and the input graph has a 'weight' edge attribute, then that attribute will be used. If NULL and no such attribute is present, then the edges will have equal weights. Set this to NA if the graph was a 'weight' edge attribute, but you don't want to use it for community detection. A larger edge weight means a stronger connection for this function. The weights value is ignored for the

spectral clustering.

#### **Details**

Used the normalised network modularity value Qm based on the previous studies by Parter et al., 2007, Takemoto, 2012, Takemoto, 2013, Takemoto and Borjigin, 2011, which was defined as:

$$Q_m = \frac{Q_{real} - Q_{rand}}{Q_{max} - Q_{rand}}$$

Where  $Q_{real}$  is the network modularity of a real-world signalling network and,  $Q_{rand}$  is the average network modularity value obtained from 10,000 randomised networks constructed from its real-world network.  $Q_{max}$  was estimated as: 1 - 1/M, where M is the number of modules in the real network.

Randomised networks were generated from a real-world network using the edge-rewiring algorithm (Maslov and Sneppen, 2002).

#### Value

normalized modularity value

#### References

Takemoto, K. & Kihara, K. Modular organization of cancer signaling networks is associated with patient survivability. Biosystems 113, 149–154 (2013).

```
file <- system.file("extdata", "PPI_Presynaptic.csv", package = "BioNAR")
tbl <- read.csv(file, sep="\t")
gg <- buildNetwork(tbl)
nm<-normModularity(gg, alg='louvain',Nint=10)</pre>
```

permute 61

permute

Randomly shuffle annotations

#### **Description**

This function is a convinience wrapper to sample with replace= FALSE

# Usage

```
permute(GNS, N)
```

# **Arguments**

GNS annotation list to take data from

N size of the sample

#### Value

random list of GNS values

## **Examples**

```
permute(LETTERS, 15)
```

plotBridgeness

Plot Bridgeness values

# Description

Semi-local centrality measure (Chen et al., 2011) lies between 0 and 1 indicating whether protein is important globally or locally. By plotting Bridgeness against semi-local centrality we can categorises the influence each protein found in our network has on the overall network structure:

- Region 1, proteins having a 'global' rather than 'local' influence in the network (also been called bottle-neck bridges, connector or kinless hubs (0<Sl<0.5; 0.5<Br<1).
- Region 2, proteins having 'global' and 'local' influence (0.5<Sl<1, 0.5<Br<1).
- Region 3, proteins centred within the community they belong to, but also communicating with a few other specific communities (0<Sl<0.5; 0.1<Br<0.5).
- Region 4, proteins with 'local' impact, primarily within one or two communities (local or party hubs, 0.5<Sl<1, 0<Br<0.5).

62 plotBridgeness

# Usage

```
plotBridgeness(
 gg,
 alg,
 VIPs,
 Xatt = "SL",
 Xlab = "Semilocal Centrality (SL)",
 Ylab = "Bridgeness (B)",
 bsize = 3,
 spsize = 7,
 MainDivSize = 0.8,
 xmin = 0,
 xmax = 1,
 ymin = 0,
 ymax = 1,
 baseColor = "royalblue2",
 SPColor = "royalblue2"
)
```

# **Arguments**

gg	igraph object with bridgenes values stored as attributes, after call to calcBridgeness
alg	clustering algorithm that was used to calculate bridgeness values
VIPs	list of 'specical' genes to be marked on the plot
Xatt	name of the attribute that stores values to be used as X-axis values. By default SL for semi-local centrality
Xlab	label for the X-axis
Ylab	label for the Y-axis
bsize	point size for genes
spsize	point size for 'specical' genes
MainDivSize	size of the line for the region separation lines
xmin	low limit for X-axis
xmax	upper limit for X-axis
ymin	low limit for Y-axis
ymax	upper limit for Y-axis
baseColor	basic color for genes
SPColor	colour highlighting any 'specical' genes

# Value

```
ggplot object with plot
```

plotEntropy 63

## **Examples**

```
karate <- make_graph("Zachary")
# We need vertex ID in the 'name' attribute of the vertex
V(karate)$name<-c(LETTERS,letters)[1:vcount(karate)]
set.seed(100)
gg <- calcClustering(karate, 'louvain')
gg <- calcCentrality(gg)
cnmat <- makeConsensusMatrix(gg, N=10, alg = 'louvain', type = 2, mask = 10)
gg<-calcBridgeness(gg, alg = 'louvain', cnmat)
plotBridgeness(gg,alg = 'louvain', VIPs=c("Mr Hi", "John A"))</pre>
```

plotEntropy

Plot graph entropy values versus vertex degree for each perturbed vertex value.

## Description

Following procedure described in (Teschendorff et al., 2015), all vertexes are artificially assigned a uniform weight then sequentially perturbed with the global entropy rate (SRprime) after each protein's perturbation being calculated by getEntropy function.

## Usage

```
plotEntropy(SRprime, subTIT = "Entropy", SRo = NULL, maxSr = NULL)
```

# **Arguments**

SRprime results of getEntropy invocation

subTIT entropy axis label

SRo initial entropy rate  $SR_0$ , results of getEntropyRate invocation

maxSr the maximum entropy rate maxSR, results of getEntropyRate invocation

## **Details**

This function plot SRprime against the log of the protein's degree. In case of scale-free or approximate scale-free topologies, we see a clear bi-modal response between over-weighted vertices and their degree and an opposing bi-phasic response in under-weighted vertices and their degrees.

If maxSr or SRo is set to their default value NULL getEntropyRate will be called and returned values will be used in the following calculations. As maxSr is required for SRprime calculation by getEntropy using explicit values could save some time in the case of large network.

#### Value

ggplot2 object with diagram

64 plotRatio

# See Also

```
getEntropy()
```

```
Other Entropy Functions: calcEntropy(), getEntropy(), getEntropyRate()
```

# Examples

```
file <- system.file("extdata", "PPI_Presynaptic.csv", package = "BioNAR")
tbl <- read.csv(file, sep="\t")
gg <- buildNetwork(tbl)
gg<-annotateGeneNames(gg)
# due to error in org.Hs.eg.db we have to manually check annotation of one node
idx <- which(V(gg)$name == '80273')
paste(V(gg)$GeneName[idx], 'GRPEL1')
ent <- getEntropyRate(gg)
SRprime <- getEntropy(gg, maxSr = NULL)
plotEntropy(SRprime, subTIT = "Entropy", SRo = ent$SRo, maxSr = ent$maxSr)</pre>
```

plotRatio

Plot fraction of enriched communities

## **Description**

Plot fraction of enriched communities

# Usage

```
plotRatio(
    x,
    desc = "",
    anno = "",
    LEGtextSize = 1.5,
    LEGlineSize = 4,
    type = NULL
)
```

#### **Arguments**

x enrichment statistics
desc plot subtitle
anno name of annotation used
LEGtextSize size of the text
LEGlineSize width of the line
type type of the plot

#### Value

ggplot object

plotSigmoid 65

# Description

Plot results of the sigmoid fit

# Usage

```
plotSigmoid(x, rates, model, alg = "", pv = 0)
```

# **Arguments**

X	steps along the Fe
rates	parameters of the sigmoid
model	fitted model
alg	name of the clustering algorithm

pv Kolmogorov-Smirnov test's p-value

#### Value

ggplot object with sigmoid fit plot

PPI\_Presynaptic.csv Table of protein protein interactions for presynaptic compartment

# Description

Protein-protein interactions (PPIS) for presynaptic compartment, extracted from Synaptome.db, in a csv form. Columns A and B correspond to Entrez IDs for interacting proteins A and B (node names); column We contains the edge weights, if available.

#### See Also

buildNetwork

66 prepareGDA

PPI\_Presynaptic.gml

PPI graph for presynaptic compartment

## **Description**

Protein-protein interactions (PPIS) for presynaptic compartment, extracted from Synaptome.db, and saved in a graph format. Graph contains node attributes, such as names (Entrez IDs), Gene Names, disease association (TopOntoOVG, TopOntoOVGHDOID), annotation with schizophrenia-related genes (Schanno (v/c), function annotation from GO (GOBPID, GOBP, GOMFID, GOMF, GOCCID, GOCC), centrality measures (DEG - degree, BET - betweenness, CC - clustering coefficient, SL - semilocal centrality, mnSP - mean shortest path, PR - page rank, sdSP - standard deviation of the shortest path), and clustering memberships for 8 clustering algorithms (lec, wt, fc, infomap, louvain, sgG1, sgG2, sgG5)

prepareGDA

Function to return vertex annotation from a graph in the Vertex Annotation form and format it for further analysis.

#### **Description**

Function to return vertex annotation from a graph in the Vertex Annotation form and format it for further analysis.

#### Usage

```
prepareGDA(gg, name)
```

#### **Arguments**

gg igraph object to take annotation from

name of the vertex attribute that contains annotation. If graph has no such vertex

attribute an error is thrown..

# Value

escaped annotation in Vertex Annotation form

## See Also

getAnnotationVertexList escapeAnnotation

PresynAn.csv 67

## **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
agg<-annotateGeneNames(gg)
# due to error in org.Hs.eg.db we have to manually check annotation of one node
idx <- which(V(agg)$name == '80273')
paste(V(agg)$GeneName[idx], 'GRPEL1')
gda<-prepareGDA(agg, 'TopOntoOVGHDOID')
gda<-prepareGDA(agg, 'TopOntoOVGHDOID')
head(gda)</pre>
```

PresynAn.csv

Presynaptic genes specific functional annotation

#### **Description**

Presynaptic genes functional annotation derived from Boyken at al. (2013) doi:10.1016/j.neuron. 2013.02.027. The table has columns: the first containing functional group ID terms, the second - gene functional group description terms, third - gene Human Entrez Ids; in csv format

#### See Also

annotatePresynaptic

recluster

Hierarchical graph clustering

## Description

Function reads in a graph GG with cluster membership stored in vertex attribute ALGN, and reapplies the clustering algorithm ALGN to all clusters larger than CnMAX

## Usage

```
recluster(GG, ALGN, CnMAX, weights = NULL)
```

## **Arguments**

GG graph to cluster
ALGN algorithm to apply

CnMAX maximum size of the cluster in mem that will not be processed

weights The weights of the edges. It must be a positive numeric vector, NULL or NA. If

it is NULL and the input graph has a 'weight' edge attribute, then that attribute will be used. If NULL and no such attribute is present, then the edges will have equal weights. Set this to NA if the graph was a 'weight' edge attribute, but you don't want to use it for community detection. A larger edge weight means a stronger connection for this function. The weights value is ignored for the

spectral clustering.

68 remove Vertex Term

# Value

remembership matrix, that contains vertex ID membership and result of reclustering

# **Examples**

```
data(karate,package='igraphdata')
alg<-'louvain'
mem<-calcMembership(karate,alg = alg)
remem<-calcReclusterMatrix(karate,mem,alg,10)</pre>
```

removeVertexTerm

Remove vertex property.

# **Description**

Remove vertex property.

## Usage

```
removeVertexTerm(GG, NAME)
```

# Arguments

GG igraph object

NAME name of the vertex property to remove

#### Value

igraph object with attribute removed

```
data(karate, package='igraphdata')
upgrade_graph(karate)
vertex_attr_names(karate)
m<-removeVertexTerm(karate, 'color')
vertex_attr_names(m)</pre>
```

runPermDisease 69

runPermDisease	Calculate disease-disease pair overlaps on permuted network to esti-
	mate its statistical significance

## **Description**

Function to calculate the disease-pair overlap characteristics of an inputted network, before applying Nperm permutations on the disease annotations of #' type "random" or "binned" permute. From the permuted networks the function estimates the significance of disease overlap: p-value, Bonferoni-adjusted p-value, and q-value in the Disease\_overlap\_sig. The function also compares the average disease separation between inputted and permuted networks, and calculates its significance using the Wilcox test and store. Significance of disease-pair overlap and disease separation results are stored in the matrix Disease\_location\_sig.

# Usage

```
runPermDisease(
   gg,
   name,
   diseases = NULL,
   Nperm = 100,
   permute = c("random", "binned"),
   alpha = c(0.05, 0.01, 0.001)
)
```

## Arguments

gg interactome network as igraph object

name of the attribute that stores disease annotation

diseases list of diseases to match

Nperm number of permutations to apply

permute type of permutations. random – annotation is randomly shuffled, binned – an-

notation is shuffled in a way to preserve node degree-annotation relationship by

degreeBinnedGDAs.

alpha statistical significance levels

# Details

Run with care, as large number of permutations could require a lot of memory and be timeconsuming.

# Value

list of two matrices: Disease\_overlap\_sig gives statistics for each pair of disease, and Disease\_location\_sig gives intra-disease statistics

## **Examples**

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
agg<-annotateGeneNames(gg)
# due to error in org.Hs.eg.db we have to manually check annotation of one node
idx <- which(V(agg)$name == '80273')
paste(V(agg)$GeneName[idx], 'GRPEL1')
r <- runPermDisease(
agg,
name = "TopOntoOVGHDOID",
diseases = c("DOID:10652", "DOID:3312", "DOID:12849", "DOID:1826"),
Nperm = 10,
alpha = c(0.05, 0.01, 0.001))
r$Disease_location_sig</pre>
```

sampleDegBinnedGDA

Function to randomly shuffle vertex annotation terms, whilst preserving the vertex degree originally found with that annotation term.

# Description

Function to randomly shuffle vertex annotation terms, whilst preserving the vertex degree originally found with that annotation term..

#### Usage

```
sampleDegBinnedGDA(org.map, term)
```

#### **Arguments**

org.map degree-annotation mapping returned by degreeBinnedGDAs term annotation term to shuffle

#### Value

vertex IDs to assign term in shuffled annotation

#### See Also

degreeBinnedGDAs

```
file <- system.file("extdata", "PPI_Presynaptic.gml", package = "BioNAR")
gg <- igraph::read_graph(file, format="gml")
agg<-annotateGeneNames(gg)
# due to error in org.Hs.eg.db we have to manually check annotation of one node
idx <- which(V(agg)$name == '80273')</pre>
```

sampleGraphClust 71

```
paste(V(agg)$GeneName[idx], 'GRPEL1')
gda<-prepareGDA(agg, 'TopOntoOVGHDOID')
diseases<-getAnnotationList(gda)
m<-degreeBinnedGDAs(agg, gda, diseases)
sampleDegBinnedGDA(m, diseases[1])</pre>
```

sampleGraphClust

Perturbe graph and calculate its clustering

# **Description**

Function will mask mask a percentage of edges (type=1) or vertices (type=2) from the network, find the largest connected component of the masked network and cluster it. The clustering results are stored in a three column matrix: the first column contains the vertex IDs of input network; the second column the vertex IDs of the subsampled network, or -1 if the vertex has been masked; the third column the cluster membership of subsampled network, or -1 if vertex has been masked.

# Usage

```
sampleGraphClust(
   gg,
   mask = 20,
   alg,
   type,
   weights = NULL,
   reclust = FALSE,
   Cnmax = 10
)
```

#### **Arguments**

gg	graph

mask percentage of elements to perturbe

alg clustering alg.

type edges=>1 or nodes=>2 to mask

weights The weights of the edges. It must be a positive numeric vector, NULL or NA. If

it is NULL and the input graph has a 'weight' edge attribute, then that attribute will be used. If NULL and no such attribute is present, then the edges will have equal weights. Set this to NA if the graph was a 'weight' edge attribute, but you don't want to use it for community detection. A larger edge weight means a stronger connection for this function. The weights value is ignored for the

spectral clustering.

reclust logical to decide whether to invoke reclustering via recluster

Cnmax maximum size of the cluster in mem that will not be processed if reclustering is

invoked

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#### **Details**

This is internal function and not supposed to be calle by end user.

#### Value

list of Nx3 matrices

# **Examples**

```
data(karate,package='igraphdata')
alg<-'louvain'
mem<-calcMembership(karate,alg = alg)
smpl<-BioNAR:::sampleGraphClust(karate,mask=10,alg,type=2)</pre>
```

SCH\_flatfile.csv

Schizopherina related synaptic gene functional annotation.

## Description

Annotation, manually curated from an external file: Lips et al., (2012) doi:10.1038/mp.2011.117.The table has columns: the first containing gene Human Entrez IDs, the second gene functional group ID terms, the third gene functional group description terms; in csv format

## See Also

annotateSCHanno

summaryStats

Calculate summary statistics from enrichment table

# Description

Calculate summary statistics from enrichment table

#### Usage

```
summaryStats(RES, ALPHA, usePadj = FALSE, FeMAX = 0, FcMAX = 0)
```

## **Arguments**

RES enrichment results data.frame

ALPHA p-value cut-off

usePadj logical, wether to use plain or adjusted p-value

 $\begin{array}{ll} \text{FeMAX} & \text{max of the FE} \\ \text{FcMAX} & \text{max of the FC} \end{array}$ 

unescapeAnnotation 73

# Value

list of data.frame

unescape Annotation

Unescape annotation strings

# **Description**

Function to remove all escape characters from annotation strings (opposite to escapeAnnotation).

# Usage

```
unescapeAnnotation(annVec, col = COLLAPSE, esc = ESC)
```

# **Arguments**

annVec vector of annotation strings

col list separator character within annotation string

esc escape character

## **Details**

NOTE: spaces are treated as regular characters, no trimming is applied before or after escaping.

#### Value

vector of annotation strings with removed escape characters

# See Also

escapeAnnotation

```
annVec<-apply(matrix(letters, ncol=13), 2, paste, collapse=';')
escVec<-escapeAnnotation(annVec, ';', '|')
cbind(annVec, escVec, unescapeAnnotation(escVec, ';', '|'))</pre>
```

74 zeroNA

zeroNA

Auxiliary function to replace NAs with zeros.

# Description

Auxiliary function to replace NAs with zeros.

# Usage

```
zeroNA(x)
```

# Arguments

Χ

matrix or vector to process

# Value

matrix or vector with NAs replaced by zero.

```
x<-matrix(NA,nrow = 3,ncol = 3)
zeroNA(x)</pre>
```

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