

# Package ‘EGAD’

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**Type** Package

**Title** Extending guilt by association by degree

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**Description** The package implements a series of highly efficient tools to calculate functional properties of networks based on guilt by association methods.

**License** GPL-2

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|                      |  |
|----------------------|--|
| <i>assortativity</i> | <i>Calculating network assortativity</i> |
|----------------------|--|

### Description

The function calculates the assortativity of a network, that measures the preference of interactions between similar nodes. As in most literature, 'similarity' is here defined in terms of node degrees.

### Usage

```
assortativity(network)
```

### Arguments

network           matrix indicating network structure (symmetric)

### Value

Numeric value

### Examples

```
network <- matrix( sample(c(0,1),36, replace=TRUE), nrow=6,byrow=TRUE)
assort_value <- assortativity(network)
```

|                   |  |
|-------------------|--|
| <i>attr.human</i> | <i>Human GENCODE annotations (v22)</i> |
|-------------------|--|

### Description

A dataset containing identifiers for gene transcripts

### Format

A data frame with 60483 rows and 10 variables:

- chr** chromosome
- start** chromosomal start position, in base pairs
- end** chromosomal end position, in base pairs
- strand** chromosomal strand, + or -
- un** unknown
- ensemblID** ENSEMBL identifier
- type** type of transcript

**stat** status of transcript

**name** HUGO identifier

**entrezID** Entrez identifier

@source [ftp://ftp.sanger.ac.uk/pub/gencode/Gencode\\_human/release\\_22/](ftp://ftp.sanger.ac.uk/pub/gencode/Gencode_human/release_22/)

---

attr.mouse

*Mouse GENCODE annotations (M7)*

---

### Description

A dataset containing identifiers for gene transcripts

### Format

A data frame with 46517 rows and 10 variables:

**chr** chromosome

**start** chromosomal start position, in base pairs

**end** chromosomal end position, in base pairs

**strand** chromosomal strand, + or -

**un** unknown

**ensemblID** ENSEMBL identifier

**type** type of transcript

**stat** status of transcript

**name** HUGO identifier

**entrezID** Entrez identifier

@source [ftp://ftp.sanger.ac.uk/pub/gencode/Gencode\\_mouse/release\\_M7/](ftp://ftp.sanger.ac.uk/pub/gencode/Gencode_mouse/release_M7/)

---

auc\_multifunc

*Calculating AUC for functional groups from ranked lists*

---

### Description

The function calculates the AUC for a functional group analytically using an optimal ranked list of genes that indicates association between genes and groups.

### Usage

```
auc_multifunc(annotations, optimallist)
```

### Arguments

annotations      binary matrix indicating which list elements are in which functional groups.

optimallist      Ranked list (multifunctionality analysis, see [calculate\\_multifunc](#)).

**Value**

aucs array of aucs for each group in annotations

**Examples**

```
annotations <- c(rep(0,10))
annotations[c(1,3,5)] <- 1
optimallist <- 10:1
aurocs_mf <- auc_multifunc(annotations, optimallist)
```

---

auprc

*Area under the precision recall curve*

---

**Description**

The function calculates the area under the precision-recall curve

**Usage**

```
auprc(scores, labels)
```

**Arguments**

|        |               |
|--------|---------------|
| scores | numeric array |
| labels | binary array  |

**Value**

auprc Numeric value

**Examples**

```
labels <- c(rep(0,10))
labels[c(1,3,5)] <- 1
scores <- 10:1
auprc <- auprc(scores, labels)
```

---

|                |   |
|----------------|---|
| auroc_analytic | <i>Area under the receiver operating characteristic curve</i> |
|----------------|---|

---

**Description**

The function calculates the area under the receiver operating characteristic (ROC) curve analytically

**Usage**

```
auroc_analytic(scores, labels)
```

**Arguments**

|        |               |
|--------|---------------|
| scores | numeric array |
| labels | binary array  |

**Value**

auroc Numeric value

**Examples**

```
labels <- c(rep(0,10))
labels[c(1,3,5)] <- 1
scores <- 10:1
auroc <- auroc_analytic(scores, labels)
```

---

|         |                         |
|---------|-------------------------|
| biogrid | <i>BIOGRID v3.4.126</i> |
|---------|-------------------------|

---

**Description**

A data frame containing protein-protein interactions

**Format**

A data frame with 211506 rows and 2 variables:

**entrezID\_A** List of Entrez identifiers, interactor A

**entrezID\_B** List of Entrez identifiers, interactor B

@source <http://thebiogrid.org/>

---

build\_binary\_network *Builds a binary network*

---

### Description

The function creates a gene-by-gene matrix with binary entries indicating interaction (1) or no interaction (0) between the genes.

### Usage

```
build_binary_network(data, list)
```

### Arguments

|      |   |
|------|---|
| data | 2-column matrix, each row a pair indicating a relationship or interaction |
| list | string array of genes/labels/ids  |

### Value

net matrix binary characterizing interactions

### Examples

```
data <- cbind(edgeA=c('gene1', 'gene2'), edgeB=c('gene3', 'gene3'))  
list <- c('gene1', 'gene2', 'gene3')  
network <- build_binary_network(data, list)
```

---

build\_coexp\_expressionSet

*Builds a coexpression network from an expressionSet*

---

### Description

The function generates a dense coexpression network from expression data stored in the expressionSet data type. Correlation coefficients are used as to weight the edges of the nodes (genes). Calls [build\\_coexp\\_network](#).

### Usage

```
build_coexp_expressionSet(exprsSet, gene.list, method = "spearman",  
  flag = "rank")
```

### Arguments

|           |  |
|-----------|--|
| exprsSet  | data class ExpressionSet                           |
| gene.list | array of gene labels                               |
| method    | correlation method to use, default Spearman's rho  |
| flag      | string to indicate if the network should be ranked |

**Value**

net Matrix symmetric

**Examples**

```
exprs <- matrix( rnorm(1000), ncol=10,byrow=TRUE)
gene.list <- paste('gene',1:100, sep='')
sample.list <- paste('sample',1:10, sep='')
rownames(exprs) <- gene.list
colnames(exprs) <- sample.list
network <- build_coexp_expressionSet(exprs, gene.list, method='pearson')
```

---

|                   |   |
|-------------------|---|
| build_coexp_GEOID | <i>Builds a coexpression network given a GEO ID</i> |
|-------------------|---|

---

**Description**

The function generates a dense coexpression network from expression data stored in GEO. The expression data is downloaded from GEO. Correlation coefficients are used as to weight the edges of the nodes (genes). Calls [get\\_expression\\_matrix\\_from\\_GEO](#) and [build\\_coexp\\_network](#).

**Usage**

```
build_coexp_GEOID(gseid, gene.list, method = "spearman", flag = "rank")
```

**Arguments**

|           |  |
|-----------|--|
| gseid     | string GEO ID of expression experiment             |
| gene.list | array of gene labels                               |
| method    | correlation method to use, default Spearman's rho  |
| flag      | string to indicate if the network should be ranked |

**Value**

net Matrix symmetric

---

|                     |  |
|---------------------|--|
| build_coexp_network | <i>Builds a coexpression network from an expressionSet</i> |
|---------------------|--|

---

**Description**

The function generates a dense coexpression network from expression data stored as a matrix, with the genes as row labels, and samples as column labels. Correlation coefficients are used as to weight the edges of the nodes (genes). Calls [cor](#).

**Usage**

```
build_coexp_network(exprs, gene.list, method = "spearman", flag = "rank")
```

**Arguments**

|           |  |
|-----------|--|
| exprs     | matrix of expression data                          |
| gene.list | array of gene labels                               |
| method    | correlation method to use, default Spearman's rho  |
| flag      | string to indicate if the network should be ranked |

**Value**

net Matrix symmetric

**Examples**

```
exprs <- matrix( rnorm(1000), ncol=10,byrow=TRUE)
gene.list <- paste('gene',1:100, sep='')
sample.list <- paste('sample',1:10, sep='')
rownames(exprs) <- gene.list
colnames(exprs) <- sample.list
network <- build_coexp_network(exprs, gene.list)
```

---

build\_semantic\_similarity\_network

*Builds a semantic similarity network*

---

**Description**

The function builds a semantic similarity network given a data and labels

**Usage**

```
build_semantic_similarity_network(genes.labels, genes)
```

**Arguments**

|              |   |
|--------------|---|
| genes.labels | matrix with rows as genes and columns as a function/label |
| genes        | array of gene IDs   |

**Value**

net Numeric value

**Examples**

```
genes.labels <- matrix( sample(c(0,1), 100, replace=TRUE),ncol=10,nrow=10)
rownames(genes.labels) <- 1:10
genes <- 1:10
net <- build_semantic_similarity_network(genes.labels, genes)
```

---

build\_weighted\_network  
*Builds a weighted network*

---

**Description**

The function creates a gene-by-gene matrix with binary entries indicating interaction (1) or no interaction (0) between the genes.

**Usage**

```
build_weighted_network(data, list)
```

**Arguments**

|      |   |
|------|---|
| data | 3-column matrix, each row a pair indicating a relationship or interaction, and the last column the weight |
| list | string array of genes/labels/ids  |

**Value**

net matrix characterizing interactions

**Examples**

```
data <- cbind(edgeA=c('gene1','gene2'),edgeB=c('gene3','gene3'), weight=c(0.5, 0.9))
list <- c('gene1','gene2','gene3')
network <- build_weighted_network(data,list)
```

---

calculate\_multifunc *Performing multifunctionality analysis*

---

**Description**

The function performs multifunctionality analysis ([1]) for a set of annotated genes and creates a rank based optimallist. For annotations use an ontology that is large enough to serve as a prior (e.g. GO, Phenocarta).

**Usage**

```
calculate_multifunc(genes.labels)
```

**Arguments**

|              |                   |
|--------------|-------------------|
| genes.labels | Annotation matrix |
|--------------|-------------------|

**Value**

gene.mfs Returns matrix with evaluation of gene function prediction by given labels:

**Examples**

```
genes.labels <- matrix( sample(c(0,1), 100, replace=TRUE),ncol=10,nrow=10)
rownames(genes.labels) = paste('gene', 1:10, sep='')
colnames(genes.labels) = paste('label', 1:10, sep='')
mf <- calculate_multifunc(genes.labels)
```

---

|               |                            |
|---------------|----------------------------|
| conv_smoother | <i>Plot smoothed curve</i> |
|---------------|----------------------------|

---

**Description**

The function plots a smoothed curve using the `convolve` function.

**Usage**

```
conv_smoother(X, Y, window, xlab = "", ylab = "", raw = FALSE)
```

**Arguments**

|        |  |
|--------|--|
| X      | numeric array                                  |
| Y      | numeric array                                  |
| window | numeric value indicating size of window to use |
| xlab   | string of x-axis label                         |
| ylab   | string of y-axis label                         |
| raw    | boolean  |

**Value**

smoothed X,Y and std Y matrix

**Examples**

```
x <- 1:1000
y <- rnorm(1000)
conv <- conv_smoother(x,y,10)
```

---

|                     |                               |
|---------------------|-------------------------------|
| example_annotations | <i>Example of annotations</i> |
|---------------------|-------------------------------|

---

**Description**

This dataset includes

---

example\_binary\_network

*Example of binary network*

---

**Description**

This dataset includes

**Format**

Matrices and vectors

---

example\_coexpression *Example of binary network*

---

**Description**

This dataset includes

**Format**

Matrices and vectors

---

example\_neighbor\_voting

*Example of binary network*

---

**Description**

This dataset includes

**Format**

**entrezID** chromosomal start position, in base pairs

**name** HUGO gene identifier

**species** species

**disease** disease

---

|                |   |
|----------------|---|
| extend_network | <i>Builds an extended network from a binary network</i> |
|----------------|---|

---

**Description**

The function extends a binary network by using the inverse of the path length between nodes as a weighted edge

**Usage**

```
extend_network(net, max = 6)
```

**Arguments**

|     |                                 |
|-----|---------------------------------|
| net | matrix binary and symmetric     |
| max | numeric maximum number of jumps |

**Value**

ext\_net matrix dense and symmetric

**Examples**

```
net <- matrix( sample(c(0,1),36, replace=TRUE), nrow=6,byrow=TRUE)
ext_net <- extend_network(net)
```

---

|                |                         |
|----------------|-------------------------|
| filter_network | <i>Filter on matrix</i> |
|----------------|-------------------------|

---

**Description**

The function filters out the rows or columns of a matrix such that the size of the group is exclusively between given min and max values

**Usage**

```
filter_network(network, flag = 1, min = 0, max = 1, ids = NA)
```

**Arguments**

|         |   |
|---------|---|
| network | numeric matrix                                      |
| flag    | numeric 1 for row filtering, 2 for column filtering |
| min     | numeric value                                       |
| max     | numeric value                                       |
| ids     | array to filter on                                  |

**Value**

network numeric matrix

**Examples**

```
net <- matrix( rnorm(10000), nrow=100)
filt_net <- filter_network(net,1,10,100)
```

---

filter\_network\_cols     *Filter on columns*

---

**Description**

The function filters out the columns of a matrix such that the size of the group is exclusively between given min and max values

**Usage**

```
filter_network_cols(network, min = 0, max = 1, ids = NA)
```

**Arguments**

|         |                |
|---------|----------------|
| network | numeric matrix |
| min     | numeric value  |
| max     | numeric value  |
| ids     | array          |

**Value**

network numeric matrix

**Examples**

```
genes.labels <- matrix( sample( c(0,1), 10000, replace=TRUE), nrow=100)
rownames(genes.labels) = paste('gene', 1:100, sep='')
colnames(genes.labels) = paste('function', 1:100, sep='')
genes.labels <- filter_network_cols(genes.labels,50,200)
```

```
genes.labels <- matrix( sample( c(0,1), 10000, replace=TRUE), nrow=100)
rownames(genes.labels) = paste('gene', 1:100, sep='')
colnames(genes.labels) = paste('function', 1:100, sep='')
genes.labels <- filter_network_cols(genes.labels,ids = paste('function', 1:20, sep=''))
```

---

filter\_network\_rows     *Filter on rows*

---

### Description

The function filters out the rows of a matrix such that the size of the group is exclusively between given min and max values

### Usage

```
filter_network_rows(network, min = 0, max = 1, ids = NA)
```

### Arguments

|         |                    |
|---------|--------------------|
| network | numeric matrix     |
| min     | numeric value      |
| max     | numeric value      |
| ids     | array to filter on |

### Value

network numeric matrix

### Examples

```
genes.labels <- matrix( sample( c(0,1), 10000, replace=TRUE), nrow=100)
rownames(genes.labels) = paste('gene', 1:100, sep='')
colnames(genes.labels) = paste('function', 1:100, sep='')
genes.labels <- filter_network_rows(genes.labels,50,200)

genes.labels <- matrix( sample( c(0,1), 10000, replace=TRUE), nrow=100)
rownames(genes.labels) = paste('gene', 1:100, sep='')
colnames(genes.labels) = paste('function', 1:100, sep='')
genes.labels <- filter_network_rows(genes.labels,ids = paste('gene', 1:20, sep=''))
```

---

filter\_orthologs     *Filter on orthologs*

---

### Description

The function filters away the labels for the genes that are not in the orthologs list

### Usage

```
filter_orthologs(annotations, genelist, orthologs)
```

**Arguments**

|             |                    |
|-------------|--------------------|
| annotations | binary matrix      |
| genelist    | array of gene ids  |
| orthologs   | array to filter on |

**Value**

annotations\_filtered binary matrix

**Examples**

```
genes.labels <- matrix( sample( c(0,1), 1000, replace=TRUE), nrow=100)
rownames(genes.labels) = paste('gene', 1:100, sep='')
colnames(genes.labels) = paste('function', 1:10, sep='')
gene.list <- paste('gene', 1:100, sep='')
orthologs <- paste('gene', (1:50)*2, sep='')
genes.labels.filt <- filter_orthologs(genes.labels, gene.list, orthologs)
```

---

fmeasure

*Fmeasure of precision-recall*

---

**Description**

The function calculates fmeasure for a given beta of a precision-recall curve

**Usage**

```
fmeasure(recall, precis, beta = 1)
```

**Arguments**

|        |                             |
|--------|-----------------------------|
| recall | numeric array               |
| precis | numeric array               |
| beta   | numeric value, default is 1 |

**Value**

fmeasure Numeric value

**Examples**

```
labels <- c(rep(0,10))
labels[c(1,3,5)] <- 1
scores <- 10:1
prc <- get_prc(scores, labels)
fm <- fmeasure(prc[,1], prc[,2])
```

---

|       |                                    |
|-------|------------------------------------|
| genes | <i>Genes from BIOGRID v3.4.126</i> |
|-------|------------------------------------|

---

**Description**

An array containing identifiers for genes in biogrid

**Format**

Array

**genes** List of Entrez identifiers

@source <http://thebiogrid.org/>

---

|         |  |
|---------|--|
| get_auc | <i>Calculates the area under a curve</i> |
|---------|--|

---

**Description**

The function calculates the area under the curve defined by x and y

**Usage**

```
get_auc(x, y)
```

**Arguments**

x                numeric array

y                numeric array

**Value**

auc numeric value

**Examples**

```
x <- 1:100  
y <- 1:100  
auc <- get_auc(x,y)
```

---

`get_biogrid`*Downloading and filtering BIOGRID*

---

**Description**

The function downloads the specified version of biogrid for a particular taxon

**Usage**

```
get_biogrid(species = "9606", version = "3.4.131",
            interactions = "physical")
```

**Arguments**

|                           |  |
|---------------------------|--|
| <code>species</code>      | numeric taxon of species                               |
| <code>version</code>      | string of biogrid version                              |
| <code>interactions</code> | string stating either physical or genetic interactions |

**Value**

biogrid data.frame with interactions

---

`get_counts`*Get counts*

---

**Description**

The function formats the count distribution from the histogram function

**Usage**

```
get_counts(hist)
```

**Arguments**

|                   |           |
|-------------------|-----------|
| <code>hist</code> | histogram |
|-------------------|-----------|

**Value**

`x,y`

**Examples**

```
x <- runif(1000)
counts <- get_counts( hist(x, plot=FALSE))
```

---

|             |                    |
|-------------|--------------------|
| get_density | <i>Get density</i> |
|-------------|--------------------|

---

**Description**

The function formats the density distribution from the histogram function

**Usage**

```
get_density(hist)
```

**Arguments**

|      |           |
|------|-----------|
| hist | histogram |
|------|-----------|

**Value**

array

**Examples**

```
x <- runif(1000)
density <- get_density( hist(x, plot=FALSE))
```

---

|                           |   |
|---------------------------|---|
| get_expression_data_gemma | <i>Obtain expression matrix from the GEMMA database</i> |
|---------------------------|---|

---

**Description**

The function downloads and parses the expression matrix from the GEMMA database, specified by the GEO ID

**Usage**

```
get_expression_data_gemma(gseid, filtered = "true")
```

**Arguments**

|          |  |
|----------|--|
| gseid    | GEO ID of the expression experiment            |
| filtered | flag to indicate whether or not the data is QC |

**Value**

list of genes and the expression matrix

---

`get_expression_matrix_from_GEO`*Obtain expression matrix from GEO database*

---

**Description**

The function downloads and parses the expression matrix from the GEO file specified by the GEO ID

**Usage**

```
get_expression_matrix_from_GEO(gseid)
```

**Arguments**

`gseid`                    GEO ID of the expression experiment

**Value**

list of genes and the expression matrix

---

`get_phenocarta`*Downloading and filtering Phenocarta*

---

**Description**

The function downloads the latest version of phenocarta

**Usage**

```
get_phenocarta(species = "human", type = "all")
```

**Arguments**

`species`                string

`type`                    string

**Value**

data data.frame with phenocarta data

---

|         |                                     |
|---------|-------------------------------------|
| get_prc | <i>Build precision-recall curve</i> |
|---------|-------------------------------------|

---

**Description**

The function calculates the recall and precision

**Usage**

```
get_prc(scores, labels)
```

**Arguments**

|        |               |
|--------|---------------|
| scores | numeric array |
| labels | binary array  |

**Value**

recall,precision numeric arrays

**Examples**

```
labels <- c(rep(0,10))
labels[c(1,3,5)] <- 1
scores <- 10:1
prc <- get_prc(scores, labels)
```

---

|         |  |
|---------|--|
| get_roc | <i>Build receiver operating characteristic curve</i> |
|---------|--|

---

**Description**

The function calculates the FPR and TRPR for the receiver operating characteristic (ROC)

**Usage**

```
get_roc(scores, labels)
```

**Arguments**

|        |               |
|--------|---------------|
| scores | numeric array |
| labels | binary array  |

**Value**

FPR,TPR numeric arrays

**Examples**

```

labels <- c(rep(0,10))
labels[c(1,3,5)] <- 1
scores <- 10:1
roc <- get_roc(scores, labels)

```

---

GO.human

*GO - human*


---

**Description**

A dataset of the gene GO associations

**Format**

A data frame with 2511938 rows and 4 variables:

**name** gene symbol

**entrezID** entrez identifier

**GO** gene ontology term ID

**evidence** evidence code

@source <http://geneontology.org/>

---

GO.mouse

*GO - mouse*


---

**Description**

A dataset of the gene GO associations

**Format**

A data frame with 2086086 rows and 4 variables:

**name** gene symbol

**entrezID** entrez identifier

**GO** gene ontology term ID

**evidence** evidence code

@source <http://geneontology.org/>

---

|        |                                 |
|--------|---------------------------------|
| GO.voc | <i>Gene ontology vocabulary</i> |
|--------|---------------------------------|

---

**Description**

A dataset of the gene ontology vocabulary

**Format**

A data frame with 42266 rows and 3 variables:

**GOID** GO identifier

**term** GO description

**domain** GO domain

@source <http://geneontology.org/>

---

|                    |                                  |
|--------------------|----------------------------------|
| make_annotatations | <i>Creating gene annotations</i> |
|--------------------|----------------------------------|

---

**Description**

The function annotates a list of genes according to a given ontology. It creates a binary matrix associating genes (rows) with labels (columns).

**Usage**

```
make_annotatations(data, listA, listB)
```

**Arguments**

|       |   |
|-------|---|
| data  | 2-column matrix, each row a pair indicating a relationship or interaction |
| listA | string array of genes   |
| listB | string array of labels/functions  |

**Value**

net matrix binary

**Examples**

```
gene.list <- paste('gene', 1:100, sep='')
labels.list <- paste('labels', 1:10, sep='')
data <- matrix(0,nrow=100, ncol=2)
data[,1] <- sample(gene.list, 100, replace=TRUE)
data[,2] <- sample(labels.list, 100, replace=TRUE)
net <- make_annotatations(data, gene.list, labels.list)
```

---

make\_genelist                    *Creating list of all genes in the data set.*

---

**Description**

The function extracts the list of all genes in the data set

**Usage**

```
make_genelist(gene_data_interacting)
```

**Arguments**

gene\_data\_interacting  
2-column matrix, each row a pair indicating a relationship or interaction

**Value**

list array of data labels

**Examples**

```
gene.list <- paste('gene', 1:100, sep='')  
data <- matrix(0,nrow=100, ncol=2)  
data[,1] <- sample(gene.list, 50, replace=TRUE)  
data[,2] <- sample(gene.list, 50, replace=TRUE)  
genes <- make_genelist(data)
```

---

make\_gene\_network                *Creating gene-by-gene network*

---

**Description**

The function creates a gene-by-gene matrix with binary entries indicating interaction (1) or no interaction (0) between the genes.

**Usage**

```
make_gene_network(data, list)
```

**Arguments**

data                    2-column matrix, each row a pair indicating a relationship or interaction  
list                    string array of genes

**Value**

net matrix binary characterizing interactions

**Examples**

```
gene.list <- paste('gene', 1:100, sep='')
data <- matrix(0,nrow=100, ncol=2)
data[,1] <- sample(gene.list, 100)
data[,2] <- sample(gene.list, 100)
net <- make_gene_network(data, gene.list)
```

---

|                  |   |
|------------------|---|
| make_transparent | <i>Make a color transparent (Taken from an answer on StackOverflow by Nick Sabbe)</i> |
|------------------|---|

---

**Description**

Make a color transparent (Taken from an answer on StackOverflow by Nick Sabbe)

**Usage**

```
make_transparent(someColor, alpha = 100)
```

**Arguments**

|           |  |
|-----------|--|
| someColor | color number, string or hexadecimal code |
| alpha     | numeric transparency                     |

**Value**

someColor rgb

---

|                 |  |
|-----------------|--|
| neighbor_voting | <i>Evaluating Gene Function Prediction</i> |
|-----------------|--|

---

**Description**

The function performs gene function prediction based on 'guilt by association' using cross validation ([1]). Performance and significance are evaluated by calculating the AUROC or AUPRC of each functional group.

**Usage**

```
neighbor_voting(genes.labels, network, nFold = 3, output = "AUROC",
  FLAG_DRAW = FALSE)
```

**Arguments**

|              |  |
|--------------|--|
| genes.labels | numeric array                                |
| network      | numeric array symmetric, gene-by-gene matrix |
| nFold        | numeric value, default is 3                  |
| output       | string, default is AUROC                     |
| FLAG_DRAW    | binary flag to draw roc plot                 |

**Value**

scores numeric matrix

**Examples**

```
genes.labels <- matrix( sample( c(0,1), 1000, replace=TRUE), nrow=100)
rownames(genes.labels) = paste('gene', 1:100, sep='')
colnames(genes.labels) = paste('function', 1:10, sep='')
net <- cor( matrix( rnorm(10000), ncol=100), method='spearman')
rownames(net) <- paste('gene', 1:100, sep='')
colnames(net) <- paste('gene', 1:100, sep='')

aurocs <- neighbor_voting(genes.labels, net, output = 'AUROC')

auprcs <- neighbor_voting(genes.labels, net, output = 'PR')
```

---

node\_degree

*Calculate node degree*

---

**Description**

The function calculates the node degree of a network

**Usage**

```
node_degree(net)
```

**Arguments**

net                    numeric matrix

**Value**

node\_degree numeric array

**Examples**

```
net <- cor( matrix(rnorm(1000), ncol=10))
n <- 10
net <- matrix(rank(net, na.last = 'keep', ties.method = 'average'), nrow = n, ncol = n)
net <- net/max(net, na.rm=TRUE)
nd <- node_degree(net)
```

---

|       |                       |
|-------|-----------------------|
| ortho | <i>Gene orthologs</i> |
|-------|-----------------------|

---

**Description**

A list containing identifiers for the subsets of gene orthologs

**Format**

List orthologs for 5 species

**dros** List of Entrez identifiers, Drosophila

**celeg** List of Entrez identifiers, C. elegans

**yeast** List of Entrez identifiers, Yeast

**mouse** List of Entrez identifiers, Mouse

**zf** List of Entrez identifiers, Zebrafish

@source <http://useast.ensembl.org/index.html/>

---

|       |                   |
|-------|-------------------|
| pheno | <i>Phenocarta</i> |
|-------|-------------------|

---

**Description**

A dataset of gene disease associations

**Format**

A data frame with 142272 rows and 4 variables:

**entrezID** chromosomal start position, in base pairs

**name** HUGO gene identifier

**species** species

**disease** disease

@source <http://www.chibi.ubc.ca/Gemma/phenotypes.html>

---

plot\_densities      *Plot densities*

---

### Description

The function plots multiple density curves and compares their modes

### Usage

```
plot_densities(hists, id, col = c("lightgrey"), xlab = "",
  ylab = "Density", mode = "hist")
```

### Arguments

|       |  |
|-------|--|
| hists | list of histogram objects or density objects |
| id    | string                                       |
| col   | color for shading                            |
| xlab  | string x-axis label                          |
| ylab  | string y-axis label                          |
| mode  | flag indicating histogram or density         |

### Value

null

### Examples

```
aurocsA <- density((runif(1000)+runif(1000)+runif(1000)+runif(1000))/4)
aurocsB <- density((runif(1000)+runif(1000)+runif(1000))/3)
aurocsC <- density(runif(1000))
hists <- list(aurocsA, aurocsB, aurocsC)
temp <- plot_densities(hists, '', mode='density')
```

---

plot\_density\_compare      *Plot density comparisons*

---

### Description

The function plots two density curves and compares their modes

### Usage

```
plot_density_compare(aucA, aucB, col = "lightgrey",
  xlab = "AUROC (neighbor voting)", ylab = "Density", mode = TRUE)
```

**Arguments**

|      |                              |
|------|------------------------------|
| aucA | numeric array of aurops      |
| aucB | numeric array of aurops      |
| col  | color of lines               |
| xlab | string label                 |
| ylab | string label                 |
| mode | boolean to plot mode or mean |

**Value**

null

**Examples**

```

auropsA <- (runif(1000)+runif(1000)+runif(1000)+runif(1000))/4
auropsB <- runif(1000)
plot_density_compare(auropsA, auropsB)

```

---

plot\_distribution      *Plot distribution histogram*

---

**Description**

The function plots a the distribution of AUROCs

**Usage**

```

plot_distribution(auc, b = 20, col = "lightgrey", xlab = "",
  ylab = "Density", xlim = c(0.4, 1), ylim = c(0, 5), med = TRUE,
  avg = TRUE, density = TRUE, bars = FALSE)

```

**Arguments**

|         |                             |
|---------|-----------------------------|
| auc     | numeric aucs                |
| b       | array of breaks             |
| col     | color of line               |
| xlab    | string label                |
| ylab    | string label                |
| xlim    | range of values for xaxis   |
| ylim    | range of values for yaxis   |
| med     | boolean to plot median auc  |
| avg     | boolean to plot average auc |
| density | boolean                     |
| bars    | boolean for barplot         |

**Value**

auc list and quartiles

**Examples**

```
aurocs <- (runif(1000)+runif(1000)+runif(1000)+runif(1000))/4
d <- plot_distribution(aurocs)
```

---

plot\_network\_heatmap *Plot network heatmap*

---

**Description**

The function draws a heatmap to visualize a network

**Usage**

```
plot_network_heatmap(net, colrs)
```

**Arguments**

|       |                                       |
|-------|---------------------------------------|
| net   | a numeric matrix of edge weights      |
| colrs | a range of colors to plot the network |

**Value**

null

**Examples**

```
network <- cor(matrix( rnorm(10000), nrow=100))
plot_network_heatmap(network)
```

---

plot\_prc *Plot precision recall curve*

---

**Description**

The function calculates the precision and recall and plots the curve

**Usage**

```
plot_prc(scores, labels)
```

**Arguments**

|        |               |
|--------|---------------|
| scores | numeric array |
| labels | binary array  |

**Value**

prc numeric arrays

**Examples**

```
labels <- c(rep(0,10))
labels[c(1,3,5)] <- 1
scores <- 10:1
roc <- plot_prc(scores, labels)
```

---

plot\_roc

*Plot receiver operating characteristic curve*

---

**Description**

The function calculates the FPR and TRPR for the receiver operating characteristic (ROC) and plots the curve

**Usage**

```
plot_roc(scores, labels)
```

**Arguments**

|        |               |
|--------|---------------|
| scores | numeric array |
| labels | binary array  |

**Value**

FPR,TPR numeric arrays

**Examples**

```
labels <- c(rep(0,10))
labels[c(1,3,5)] <- 1
scores <- 10:1
roc <- plot_roc(scores, labels)
```

---

plot\_roc\_overlay      *Plot ROC overlay*

---

### Description

The function plots a density overlay of ROCs given the scores and labels

### Usage

```
plot_roc_overlay(scores.mat, labels.mat, nbins = 100)
```

### Arguments

|            |               |
|------------|---------------|
| scores.mat | numeric array |
| labels.mat | numeric array |
| nbins      | numeric value |

### Value

list of Z(matrix) and roc\_sum (average ROC curve)

### Examples

```
genes.labels <- matrix( c(rep(1, 1000), rep(0,9000)), nrow=1000, byrow=TRUE)
rownames(genes.labels) = paste('gene', 1:1000, sep='')
colnames(genes.labels) = paste('function', 1:10, sep='')

scores <- matrix( rnorm(10000), nrow=1000)
scores <- apply(scores, 2, rank)
rownames(scores) = paste('gene', 1:1000, sep='')
colnames(scores) = paste('function', 1:10, sep='')

z <- plot_roc_overlay(scores, genes.labels)
```

---

plot\_value\_compare      *Plot value comparisons*

---

### Description

The function plots a scatter

### Usage

```
plot_value_compare(aucA, aucB, xlab = "AUROC", ylab = "AUROC", xlim = c(0,
1), ylim = c(0, 1))
```

**Arguments**

|      |                           |
|------|---------------------------|
| aucA | numeric array of aucs     |
| aucB | numeric array of aucs     |
| xlab | string label              |
| ylab | string label              |
| xlim | range of values for xaxis |
| ylim | range of values for yaxis |

**Value**

null

---

predictions *Performing Gene Function Prediction*

---

**Description**

The function performs gene function prediction on the whole data set using the 'guilt by association' principle ([1]).

**Usage**

```
predictions(genes.labels, network)
```

**Arguments**

|              |  |
|--------------|--|
| genes.labels | numeric array                                |
| network      | numeric array symmetric, gene-by-gene matrix |

**Value**

scores numeric matrix

**Examples**

```
genes.labels <- matrix( sample( c(0,1), 1000, replace=TRUE), nrow=100)
rownames(genes.labels) = paste('gene', 1:100, sep='')
colnames(genes.labels) = paste('function', 1:10, sep='')
net <- cor( matrix( rnorm(10000), ncol=100), method='spearman')
rownames(net) <- paste('gene', 1:100, sep='')
colnames(net) <- paste('gene', 1:100, sep='')

preds <- predictions(genes.labels, net)
```

---

|        |                                  |
|--------|----------------------------------|
| repmat | <i>Rep function for matrices</i> |
|--------|----------------------------------|

---

**Description**

The function generates a matrix by binding the columns and rows

**Usage**

```
repmat(X, m, n)
```

**Arguments**

|   |                                       |
|---|---------------------------------------|
| X | numeric matrix                        |
| m | numeric value, repeat rows m times    |
| n | numeric value, repeat columns n times |

**Value**

list of genes and the expression matrix

**Examples**

```
genes.labels <- matrix( sample( c(0,1), 1000, replace=TRUE), nrow=100)
expand <- repmat( genes.labels, 1,2)
```

---

|         |   |
|---------|---|
| run_GBA | <i>Performing 'Guilt by Association' Analysis</i> |
|---------|---|

---

**Description**

The function runs and evaluates gene function prediction based on the 'guilt by association'-principle using neighbor voting ([neighbor\\_voting](#)) [1]. As a measure of performance and significance of results, AUCs of all evaluated functional groups are calculated.

**Usage**

```
run_GBA(network, labels, min = 20, max = 1000, nfold = 3)
```

**Arguments**

|         |  |
|---------|--|
| network | numeric array symmetric, gene-by-gene matrix |
| labels  | numeric array                                |
| min     | numeric value to limit gene function size    |
| max     | numeric value to limit gene function size    |
| nfold   | numeric value, default is 3                  |

**Value**

list roc.sub, genes, auroc

**Examples**

```
genes.labels <- matrix( sample( c(0,1), 1000, replace=TRUE), nrow=100)
rownames(genes.labels) = paste('gene', 1:100, sep='')
colnames(genes.labels) = paste('function', 1:10, sep='')
net <- cor( matrix( rnorm(10000), ncol=100), method='spearman')
rownames(net) <- paste('gene', 1:100, sep='')
colnames(net) <- paste('gene', 1:100, sep='')

gba <- run_GBA(net, genes.labels, min=10)
```

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