# Package 'BiRewire'

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**Version** 3.36.0 **Date** 2022-02-10

**Title** High-performing routines for the randomization of a bipartite graph (or a binary event matrix), undirected and directed signed graph preserving degree distribution (or marginal totals)

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Description Fast functions for bipartite network rewiring through N consecutive switching steps (See References) and for the computation of the minimal number of switching steps to be performed in order to maximise the dissimilarity with respect to the original network. Includes functions for the analysis of the introduced randomness across the switching steps and several other routines to analyse the resulting networks and their natural projections. Extension to undirected networks and directed signed networks is also provided. Starting from version 1.9.7 a more precise bound (especially for small network) has been implemented. Starting from version 2.2.0 the analysis routine is more complete and a visual montioring of the underlying Markov Chain has been implemented. Starting from 3.6.0 the library can handle also matrices with NA (not for the directed signed graphs). Since version 3.27.1 it is possible to add a constraint for dsg generation: usually positive and negative arc between two nodes could be not accepted.

License GPL-3

Depends igraph, slam, Rtsne, Matrix

Suggests RUnit, BiocGenerics

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URL http://www.ebi.ac.uk/~iorio/BiRewire

biocViews Network

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#### **Description**

R package for computationally-efficient rewiring of bipartite graphs (or randomisation of 0-1 tables with prescribed marginal totals), undirected and directed signed graphs (dsg). The package provides useful functions for the analysis and the randomisation of large biological datasets that can be encoded as 0-1 tables, hence modeled as bipartite graphs by considering a 0-1 table as an incidence matrix, and for data that can be encoded as directed signed graphs such as patways and signaling networks. Large collections of such randomised tables can be used to approximate null models, preserving event-rates both across rows and columns, for statistical significance tests of combinatorial properties of the original dataset. The package provides an interface to a sampler routine useful for generating correctly such collections. Moreover a visual monitoring for the Markov Chain underlying the swithicing algorithm has been implemented. Since version 3.6.0 the SA can be performed also using matrices with NAs. In this case the positions of the NAs are preserved as the degree distribution. This extension is limited when the tables are provided instead of the graphs and does not work for the dsg.

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#### **Details**

Summary:

Package: BiRewire Version: 3.7.0

Date: 2017-02-27

Require: slam, igraph, Rtsne, Matrix, R>=2.10 URL: http://www.ebi.ac.uk/~iorio/BiRewire

License: GPL-3

### Author(s)

Andrea Gobbi [aut], Davide Albanese [cbt], Francesco Iorio [cbt], Giuseppe Jurman [cbt].

Maintainer: Andrea Gobbi <gobbi.andrea@mail.com>

#### References

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

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Jaccard, P. (1901), Étude comparative de la distribution florale dans une portion des Alpes et des *Jura*, Bulletin de la Société Vaudoise des Sciences Naturelles 37: 547–579.

R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), *On the uniform generation of random graphs with prescribed degree sequences*, eprint arXiv:cond-mat/0312028 Csardi, G. and Nepusz, T (2006)

Van der Maaten, L.J.P. and Hinton, G.E., *Visualizing High-Dimensional Data Using t-SNE*. Journal of Machine Learning Research 9(Nov):2579-2605, 2008 *The igraph software package for complex network research*, InterJournal, Complex Systems http://igraph.sf.net

birewire.analysis.bipartite

Analysis of Jaccard similarity trends across switching steps.

# Description

This function performs a sequence of *max.iter* switching steps on the input bipartite graph *g* and compute the Jaccard similarity between *g* (the initial network) and its rewired version each *step* switching steps. This procedure is performed *n.networks* times and a simple explorative plot, with mean and CI, is visualized if *display* is set to true.

# Usage

birewire.analysis.bipartite(incidence, step=10, max.iter="n",accuracy=0.00005, verbose=TRUE,MAXITER\_MUL=10,exact=FALSE,n.networks=50,display=TRUE)

# **Arguments**

incidence	Incidence matrix of the initial bipartite graph $g$ (can be extracted from an igraph bipartite graph using the get.incidence function). Since 3.6.0 this matrix can contain also NAs and the position of such entries will be preserved by the SA;
step	10 (default): the interval (in terms of switching steps) at which the Jaccard index between $g$ and the its current rewired version is computed;
max.iter	"n" (default) the number of switching steps to be performed (or if $exact==TRUE$ the number of successful switching steps). If equal to "n" then this number is considered equal to the analytically derived lower bound presented in $Gobbi$ $et$ $al$ . (see References): $N=e/2(1-d)\ln{((e-de)/\delta)}$ if exact is FALSE, $N=e(1-d)/2\ln{((e-de)/\delta)}$ otherwise , where $e$ is the number of edges of $g$ and $d$ its edge density . This bound is much lower than the empirical one proposed in $Milo$ $et$ $al$ . $2003$ (see References);
accuracy	0.00005 (default) is the desired level of accuracy reflecting the average distance between the Jaccard index at the N-th step and its analytically derived fixed point in terms of fracion of common edges;
verbose	TRUE (default). When TRUE a progression bar is printed during computation;
MAXITER_MUL	10 (default). If <i>exact==TRUE</i> in order to prevent a possible infinite loop the program stops anyway after MAXITER_MUL*max.iter iterations;
exact	FALSE (default). If TRUE the program performs <i>max.iter</i> swithcing steps, otherwise the program will count also the not-performed swithcing steps;
n.networks	50 (default), the number of independent rewiring process starting from the same inital graph from which the mean value and the CI is computed.
display	TRUE (default). If TRUE two explorative plots are displayed summarizing the trend of the Jaccard index in terms of mean and confidence interval.

#### **Details**

This function performs max.iter switching steps (see references). In particular, at each step two edges are randomly selected from the current version of g. Let these two edges be (a,b) and (c,d) (where a and c belong to the first class of nodes whereas b and d belong to the second one), with  $a \neq c$  and  $b \neq d$ .

If the (a, d) and (c, b) edges are not already present in the current current version of g then (a, d) and (c, b) replace (a, b) and (c, d).

At each *step* number of switching steps the function computes the **Jaccard index** between the original graph g and its current version.

This procedure is performed *n.networks* times and if *display* is set to TRUE, two explorative plots showing the mean value of the Jaccad Index over the SS and its CI are displayed.

#### Value

A list containing a data frame data collecting all the Jacard index computed (each row is a run of the SA), and the analytically derived lower bound N of switching steps to be performed by the switching algorithm in order to provide the revired version of g with the maximal level of achievable randomness (in terms of dissimilarity from the initial g).

### Author(s)

Andrea Gobbi Maintainer: Andrea Gobbi <gobbi.andrea@mail.com> Special thanks to: Davide Albanese

#### References

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

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R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), On the uniform generation of random graphs with prescribed degree sequences, eprint arXiv:cond-mat/0312028

# **Examples**

```
library(BiRewire)
g <-graph.bipartite(rep(0:1,length=10), c(1:10))</pre>
```

birewire.analysis.dsg

```
##get the incidence matrix of g
m<-as.matrix(get.incidence(graph=g))</pre>
## set parameters
step=1
max=100*length(E(g))
## perform two different analysis using two different maximal number of switching steps
scores<-birewire.analysis.bipartite(m, step, max, n.networks=10)</pre>
scores2<-birewire.analysis.bipartite(m, step, "n", n.networks=10)</pre>
```

birewire.analysis.dsg Analysis of Jaccard similarity trends across switching steps.

### **Description**

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This function performs a sequence of max.iter.pos (and max.iter.pos) switching steps on the positive (and negative) part of the input dsg g and computes the Jaccard similarity between g (the initial network) and its rewired version each step switching steps. This procedure is pefromed n.networks times and a simple explorative plot, with mean and CI, is visualized if display is set to TRUE. The plot shows the trend of the Jaccad Index relative to the positve (and negative) part of g.

#### **Usage**

```
birewire.analysis.dsg(dsg, step=10, max.iter.pos='n',max.iter.neg='n',accuracy=0.00005,
     verbose=TRUE,MAXITER_MUL=10,exact=FALSE,n.networks=50,display=TRUE)
```

# **Arguments** dsg

	must contain a list of two incidence matrices and not igraph bipartite graphs.
step	10 (default): the interval (in terms of switching steps) at which the Jaccard index between <i>g</i> and the its current rewired version is computed;
max.iter.pos	"n" (default) the number of switching steps to be performed (or if $exact = TRUE$ the number of successful switching steps) for the positive part of $g$ . See birewire.rewire.bipartite for more details;
max.iter.neg	"n" (default) the same of max.iter.p but relative to the negative part;
accuracy	0.00005 (default) is the desired level of accuracy reflecting the average distance between the Jaccard index at the N-th step and its analytically derived fixed point

The initial dsg object (see birewire.induced.bipartite). Note that the dsg

in terms of fracion of common edges;

TRUE (default). When TRUE a progression bar is printed during computation; verbose 10 (default). If exact==TRUE in order to prevent a possible infinite loop the MAXITER\_MUL

program stops anyway after MAXITER\_MUL\*max.iter iterations;

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exact	FALSE (default). If TRUE the program performs <i>max.iter</i> swithcing steps, otherwise the program will count also the not-performed swithcing steps;
n.networks	50 (default), the number of independent rewiring process starting from the same inital graph from which the mean value and the CI is computed.
display	TRUE (default). If TRUE two explorative plots are displayed summarizing the trend of the Jaccard index in terms of mean and confidence interval.

#### **Details**

This procedure acts in the same way of birewire.analysis.bipartite but in the case of dsg. The similarity is measure using birewire.similarity.dsg.

#### Value

A list containing two lists: data that is a list collecting all the Jacard index computed (each row is a run of the SA) for the positive and negative part, and a list with the analytically derived lower bounds N for the positive and negative part of g.

#### Author(s)

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#### References

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

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R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), *On the uniform generation of random graphs with prescribed degree sequences*, eprint arXiv:cond-mat/0312028

# **Examples**

```
library(BiRewire)
data(test_dsg)
dsg <- birewire.induced.bipartite(test_dsg,sparse=FALSE)
a=birewire.analysis.dsg(dsg,verbose=FALSE,step=1,exact=TRUE,max.iter.pos=200,max.iter.neg=50)</pre>
```

birewire.analysis.undirected

Analysis of Jaccard similarity trends across switching steps.

# Description

This function performs a sequence of max.iter switching steps on the input undirected graph g and compute the Jaccard similarity between g (the initial network) and its rewired version each step switching steps. This procedure is performed n.networks times and a simple explorative plot, with mean and CI, is visualized if display is set to TRUE.

# Usage

birewire.analysis.undirected(adjacency, step=10, max.iter="n",accuracy=0.00005, verbose=TRUE,MAXITER\_MUL=10,exact=FALSE,n.networks=50,display=TRUE)

# **Arguments**

adjacency	Incidence matrix of the initial bipartite graph $g$ (can be extracted from an igraph undirected graph using the get.adjacency function). Since 3.6.0 this matrix can contain also NAs and the position of such entries will be preserved by the SA;
step	10 (default): the interval (in terms of switching steps) at which the Jaccard index between $g$ and the its current rewired version is computed;
max.iter	"n" (default) the number of switching steps to be performed (or if $exact==TRUE$ the number of successful switching steps). If equal to "n" then this number is considered equal to the analytically derived lower bound presented in $Gobbi\ et\ al.$ (see References): $N=e/(2d^3-dd^2+2d+2)\ln\left((e-de)/\delta\right)$ if exact is FALSE, $N=e(1-d)/2\ln\left((e-de)/\delta\right)$ otherwise , where $e$ is the number of edges of $g$ and $d$ its edge density . This bound is much lower than the empirical one proposed in $Milo\ et\ al.\ 2003$ (see References);
accuracy	0.00005 (default) is the desired level of accuracy reflecting the average distance between the Jaccard index at the N-th step and its analytically derived fixed point in terms of fracion of common edges;
verbose	TRUE (default). When TRUE a progression bar is printed during computation;
MAXITER_MUL	10 (default). If <i>exact==TRUE</i> in order to prevent a possible infinite loop the program stops anyway after MAXITER_MUL*max.iter iterations;
exact	FALSE (default). If TRUE the program performs <i>max.iter</i> swithcing steps, otherwise the program will count also the not-performed swithcing steps;
n.networks	50 (default), the number of independent rewiring process starting from the same inital graph from which the mean value and the CI is computed.
display	TRUE (default). If TRUE two explorative plots are displayed summarizing the trend of the Jaccard index in terms of mean and confidence interval.

#### **Details**

This function performs max.iter switching steps (see references). In particular, at each step two edges are randomly selected from the current version of g. Let these two edges be (a,b) and (c,d), with  $a \neq c$ ,  $b \neq d$ ,  $a \neq d$ ,  $b \neq c$ .

If the (a,d) and (c,b) (or (a,d) and (b,d)) edges are not already present in the current version of g then (a,d) and (c,b) replace (a,b) and (c,d) (or (a,b) and (c,d) replace (a,c) and (b,d)). If both of the configurations are allowed, then one of them is randomly selected.

At each *step* number of switching steps the function computes the **Jaccard index** between the original graph g and its current version.

This procedure is performed *n.networks* times and if *display* is set to TRUE, two explorative plots showing the mean value of the Jaccad Index over the SS and its CI are displayed.

#### Value

A list containing a data frame *data* collecting all the Jacard index computed (each row is a run of the SA), and the analytically derived lower bound N of switching steps to be performed by the switching algorithm in order to provide the revired version of g with the maximal level of achievable randomness (in terms of dissimilarity from the initial g).

#### Author(s)

Andrea Gobbi

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Special thanks to: Davide Albanese

# References

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

Gobbi, A. and Jurman, G. (2013) *Theoretical and algorithmic solutions for null models in network theory* (Doctoral dissertation) http://eprints-phd.biblio.unitn.it/1125/

Jaccard, P. (1901), Étude comparative de la distribution florale dans une portion des Alpes et des Jura, Bulletin de la Société Vaudoise des Sciences Naturelles 37: 547–579.

R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), On the uniform generation of random graphs with prescribed degree sequences, eprint arXiv:cond-mat/0312028

### **Examples**

```
library(BiRewire)
g <- erdos.renyi.game(1000,0.1)
##get the incidence matrix of g
m<-as.matrix(get.adjacency(graph=g,sparse=FALSE))

## set parameters
step=1000
max=100*length(E(g))

## perform two different analysis using two different numbers of switching steps
scores<-birewire.analysis.undirected(m,step,max,n.networks=10,verbose=FALSE)
scores2<-birewire.analysis.undirected(m,step,"n",n.networks=10,verbose=FALSE)</pre>
```

birewire.bipartite.from.incidence

Converts an incidence matrix into a bipartite graph.

### **Description**

This function creates an igraph bipartite graph from an incidence matrix.

### Usage

```
birewire.bipartite.from.incidence(matrix,directed=FALSE)
```

# **Arguments**

matrix incidence matrix: an (n-by-m) binary matrix where rows correspond to vertices

in the frist class while columns correspond to vertices in the second one;

directed Logical, if TRUE a directed graph is created.

### **Details**

The function calls graph.incidence of package igraph. See igraph documentation for more details.

### Value

Bipartite igraph graph.

### Author(s)

Andrea Gobbi

Maintainer: Andrea Gobbi <gobbi.andrea@mail.com>

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#### References

Csardi, G. and Nepusz, T (2006) *The igraph software package for complex network research*, Inter-Journal, Complex Systems url http://igraph.sf.net

#### **Examples**

```
library(igraph)
library(BiRewire)
g <- graph.bipartite( rep(0:1,length=10), c(1:10))
##gets the incidence matrix of g
m<-as.matrix(get.incidence(graph=g))
##rewire the current graph
m2=birewire.rewire.bipartite(m,100)
#create the rewired bipartite graph
g2<-birewire.bipartite.from.incidence(m2,TRUE)</pre>
```

birewire.build.dsg

Transform a dsg object in a SIF file.

# **Description**

The routine transforms the initial dsg (two bipartite graphs) into SIF dsg format.

# Usage

```
birewire.build.dsg(dsg,delimitators=list(negative='-',positive='+'))
```

# **Arguments**

dsg The dsg to be converted;

delimitators list(negative='-',positive='+') (default):a list with 'positive' and 'negative' names

identifying the character encoding the relation;

# **Details**

This fuction converts the dsg object into a SIF format that can be saved using birewire.write.dsg, an internal function, using the given delimitators for encoding the relations. It is the inverse function of birewire.induced.bipartite.

#### Value

A dsg in SIF format.

#### **Examples**

```
data(test_dsg)
dsg=birewire.induced.bipartite(test_dsg)
tmp= birewire.rewire.dsg(dsg,verbose=FALSE)
dsg2=birewire.build.dsg(tmp)
```

birewire.induced.bipartite

Transform a SIF data frame into a dsg object (a list of positive and negative incidence matrix).

### **Description**

The routine transforms the initial dsg graph in SIF format into a dsg object made of two bipartite graphs: one for positive edges and the other for negative edges.

# Usage

```
birewire.induced.bipartite(g,delimitators=list(negative='-',positive='+'),sparse=FALSE)
```

# **Arguments**

g	A dataframe in SIF format describing a dsg (for example the output of birewire.load.dsg;
delimitators	list(negative='-',positive='+') (default):a list with 'positive' and 'negative' names identifying the character encoding the relation;
sparse	FALSE (default): if TRUE the two bipartite graphs are saved as igraph bipartite graphs;

### **Details**

This fuction extract the positive and negative part of g and create a dsg object that can be used for example in the rewiring algorithm. Is is the inverse function of birewire.build.dsg.

#### Value

A list of two incidence matrix or bipartite igraph objects.

### References

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

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# **Examples**

```
data(test_dsg)
dsg=birewire.induced.bipartite(test_dsg)
```

birewire.load.dsg

Read a SIF file from a given path

# **Description**

The routine reads a SIF file and return a R table.

# Usage

```
birewire.load.dsg(path)
```

# **Arguments**

path

Path to the SIF file.

# Value

A R table that can be transformed into a dsg using birewire.induced.bipartite

birewire.rewire.bipartite

Efficient rewiring of bipartite graphs

# Description

Optimal implementation of the switching algorithm. It returns the rewired version of the initial bipartite graph or its incidence matrix.

# Usage

```
birewire.rewire.bipartite(incidence, max.iter="n",accuracy=0.00005,verbose=TRUE,
MAXITER_MUL=10,exact=FALSE)
```

#### **Arguments**

incidence Incidence matrix of the initial bipartite graph g (can be extracted from an igraph

bipartite graph using the get.incidence) function; or the entire bipartite igraph graph. Since 3.6.0, in the case the matrix is provided, such matrix can contain

also NAs and the position of such entries will be preserved by the SA

max.iter "n" (default) the number of switching steps to be performed (or if exact==TRUE

the number of **successful** switching steps). If equal to "n" then this number is considered equal to the analytically derived lower bound presented in *Gobbi et al.* (see References):  $N=e/2(1-d)\ln\left((e-de)/\delta\right)$  if exact is FALSE,  $N=e(1-d)/2\ln\left((e-de)/\delta\right)$  otherwise , where e is the number of edges of g and d its edge density . This bound is much lower than the empirical one

proposed in Milo et al. 2003 (see References);

accuracy 0.00005 (default) is the desired level of accuracy reflecting the average distance

between the Jaccard index at the N-th step and its analytically derived fixed point

in terms of fracion of common edges;

verbose TRUE (default). When TRUE a progression bar is printed during computation.

MAXITER\_MUL 10 (default). If exact==TRUE in order to prevent a possible infinite loop the

program stops anyway after MAXITER\_MUL\*max.iter iterations;

exact FALSE (default). If TRUE the program performs max.iter swithcing steps, oth-

erwise the program will count also the not-performed swithcing steps;

#### **Details**

Main function of the package. It performs at most max.iter switching steps producing a rewired version of an initial bipartite graph.

#### Value

Incidence matrix of the rewired graphn or the *igraph* corresponding object depending on the input type.

### Author(s)

Andrea Gobbi

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# References

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), *On the uniform generation of random graphs with prescribed degree sequences*, eprint arXiv:cond-mat/0312028

# **Examples**

```
library(igraph)
library(BiRewire)
g <-graph.bipartite( rep(0:1,length=10), c(1:10))

##gets the incidence matrix of g
m<-as.matrix(get.incidence(graph=g))

##rewiring
m2=birewire.rewire.bipartite(m,100*length(E(g)))
##creates the corresponding bipartite graph
g2<-birewire.bipartite.from.incidence(m2,directed=TRUE)</pre>
```

birewire.rewire.bipartite.and.projections

Analysis and rewiring function processing a bipartite graphs and its two projections

#### **Description**

This function performs the same analysis of birewire.analysis.bipartite but additionally it provides in output a rewired version of the two networks resulting from the natural projections of the initial graph, together with the corresponding Jaccard index trends.

# Usage

```
birewire.rewire.bipartite.and.projections(graph, step=10, max.iter="n",
accuracy=0.00005, verbose=TRUE, MAXITER_MUL=10)
```

# Arguments

graph	A bipartite graph g;
max.iter	"n" (default) the number of successful switching steps to be performed. If equal to "n" then this number is considered equal to the analytically derived lower bound $N=e(1-d)/2\ln\left((e-de)/\delta\right)$ presented in <i>Gobbi et al.</i> (see References);
step	10 (default): the interval (in terms of switching steps) at which the Jaccard index between <i>g</i> and the its current rewired version is computed;

accuracy 0.00005 (default) is the desired level of accuracy reflecting the average distance

between the Jaccard index at the N-th step and its analytically derived fixed point

in terms of fracion of common edges;

verbose TRUE (default) boolean value. If TRUE print a processing bar during the rewiring

algorithm.

MAXITER\_MUL 10 (default). Since N indicates the number of successful switching steps, in or-

der to prevent a possible infinite loop the program stops anyway after MAX-

ITER\_MUL\*max.iter iterations;

### **Details**

See birewire.analysis.bipartite for details.

#### Value

A list containing the three sequences of Jaccard index values (similarity\_scores, similarity\_scores.proj1, similarity\_scores.proj2) for the three resulting graphs respectively (rewired, rewired.proj1, rewired.proj2). The first one is the rewired version of the initial graph g, while the second and the third one are rewired versions of its natural projections.

#### Author(s)

Andrea Gobbi

Maintainer: Andrea Gobbi <gobbi.andrea@mail.com>

# References

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

# **Examples**

birewire.rewire.dsg 17

birewire.rewire.dsg Efficient rewiring of directed signed graphs

# Description

Optimal implementation of the switching algorithm. It returns the rewired version of the initial directed signed graph (dsg).

# Usage

```
birewire.rewire.dsg(dsg,exact=FALSE,verbose=1,max.iter.pos='n',max.iter.neg='n',
    accuracy=0.00005,MAXITER_MUL=10,path=NULL,delimitators=list(positive='+',negative='-'),check_pos_
```

# Arguments

dsg	A dsg object: is a list of two incidence matrices (see References), "positive" and "negative", encoding the positive edges and negative edges. This list can be obtained reading a SIF file using birewire.load.dsg function and converting the resulting dataframe using birewire.induced.bipartite;
exact	FALSE (default). If TRUE the program performs <i>max.iter</i> <b>successful</b> swithcing steps, otherwise the program will count also the not-performed swithcing steps;
verbose	TRUE (default). When TRUE a progression bar is printed during computation;
max.iter.pos	"n" (default) the number of switching steps to be performed on the positive part of $dsg$ (or if $exact==TRUE$ the number of <b>successful</b> switching steps). If equal to "n" then this number is considered equal to the analytically derived lower bound presented in $Gobbi\ et\ al.$ (see References): $N=e/2(1-d)\ln{((e-de)/\delta)}$ if exact is FALSE, $N=e(1-d)/2\ln{((e-de)/\delta)}$ otherwise , where $e$ is the number of edges of $g$ and $d$ its edge density . This bound is much lower than the empirical one proposed in $Milo\ et\ al.\ 2003$ (see References);
max.iter.neg	"n" (default) the number of switching steps to be performed on the negative part of $dsg$ (or if $exact==TRUE$ the number of $successful$ switching steps). If equal to "n" then this number is considered equal to the analytically derived lower bound presented in $Gobbi$ et al. (see References): $N=e/2(1-d)\ln\left((e-de)/\delta\right)$ if exact is FALSE, $N=e(1-d)/2\ln\left((e-de)/\delta\right)$ otherwise , where $e$ is the number of edges of $g$ and $d$ its edge density . This bound is much lower than the empirical one proposed in $Milo$ et al. $2003$ (see References);
accuracy	0.00005 (default) is the desired level of accuracy reflecting the average distance between the Jaccard index at the N-th step and its analytically derived fixed point in terms of fracion of common edges;
MAXITER_MUL	10 (default). If <i>exact</i> == <i>TRUE</i> in order to prevent a possible infinite loop the program stops anyway after MAXITER_MUL*max.iter iterations;
path	NULL (default). If not NULL, the dsg is saved in path in SIF format;

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delimitators list(positive='+',negative='-') (default). If save.file is true, the dsg is saved us-

ing delimitators as characters encoding the relations. See birewire.build.dsg

for more details.

check\_pos\_neg FALSE (default). if true check if there are positive and negative loops in the

same arc, if so the routine will not save the dsg but return it anyway with a

warining

in\_sampler FALSE (default).if TRUE does not print the warining due to the positive-negative

check

#### **Details**

This fuction runs birewire.rewire.bipartite on the positive and negative part of *dsg*. See references for more details.

#### Value

Rewired dsg.

#### Author(s)

Andrea Gobbi: <gobbi.andrea@mail.com>

# References

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), *On the uniform generation of random graphs with prescribed degree sequences*, eprint arXiv:cond-mat/0312028

#### **Examples**

library(BiRewire)
data(test\_dsg)
dsg=birewire.induced.bipartite(test\_dsg)
tmp= birewire.rewire.dsg(dsg,verbose=FALSE)

birewire.rewire.undirected 19

birewire.rewire.undirected

Efficient rewiring of undirected graphs

# **Description**

Optimal implementation of the switching algorithm. It returns the rewired version of the initial undirected graph or its adjacency matrix.

# Usage

```
birewire.rewire.undirected(adjacency, max.iter="n",accuracy=0.00005,
verbose=TRUE,MAXITER_MUL=10,exact=FALSE)
```

# Arguments

adjacency	An igraph undirected graph $g$ or its adjacency matrix (can be extracted from $g$ using get.adjacency). Since 3.6.0, if the matrix is provided, such matrix can contain also NAs and the position of such entries will be preserved by the SA
max.iter	"n" (default) the number of switching steps to be performed (or if $exact==TRUE$ the number of <b>successful</b> switching steps). If equal to "n" then this number is considered equal to the analytically derived lower bound presented in $Gobbi$ $et$ $al$ . (see References): $N=e/(2d^3-6d^2+2d+2)\ln{(e-de)}$ if exact is FALSE, $N=e(1-d)/2\ln{((e-de)/\delta)}$ otherwise , where $e$ is the number of edges of $g$ and $d$ its edge density . This bound is much lower than the empirical one proposed in $Milo$ $et$ $al$ . $2003$ (see References);
accuracy	0.00005 (default) is the desired level of accuracy reflecting the average distance between the Jaccard index at the N-th step and its analytically derived fixed point in terms of fracion of common edges;
verbose	TRUE (default) boolean value. If TRUE print a processing bar during the rewiring algorithm.
MAXITER_MUL	10 (default). If <i>exact</i> == <i>TRUE</i> in order to prevent a possible infinite loop the program stops anyway after MAXITER_MUL*max.iter iterations;
exact	FALSE (default). If TRUE the program performs <i>max.iter</i> swithcing steps, otherwise the program will count also the not-performed swithcing steps;

#### **Details**

Performs at most max.iter number of rewiring steps producing a rewired version of an initial undirected graph.

#### Value

Adjacency matrix of the rewired graph or the relative *igraph* object depending on the input type.

#### Author(s)

Andrea Gobbi

Maintainer: Andrea Gobbi <gobbi.andrea@mail.com>

Special thanks to:Davide Albanese

#### References

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T.and Jurman, G.and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

Gobbi, A. and Jurman, G. (2013) Theoretical and algorithmic solutions for null models in network theory (Doctoral dissertation) http://eprints-phd.biblio.unitn.it/1125/
R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), On the uniform generation of random graphs with prescribed degree sequences, eprint arXiv:cond-mat/0312028

# **Examples**

```
library(igraph)
library(BiRewire)
g <- erdos.renyi.game(1000,0.1)
##gets the incidence matrix of g
m<-as.matrix(get.adjacency(graph=g,sparse=FALSE))

## sets parameters
step=1000
max=100*length(E(g))

##rewiring
m2=birewire.rewire.undirected(m,100*length(E(g)))
##creates the corresponding bipartite graph
g2<-graph.adjacency(m2,mode="undirected")</pre>
```

birewire.sampler.bipartite

Efficient generation of a null model for a given bipartite graph

# **Description**

The routine samples correctly from the null model of a given bipartite graph creating a set of randomized version of the initial bipartite graph.

# Usage

birewire.sampler.bipartite(incidence,K,path,max.iter="n", accuracy=0.00005, verbose=TRUE,MAXITER\_MUL=10,exact=FALSE,write.sparse=TRUE)

#### **Arguments**

incidence	Incidence matrix of the initial bipartite graph. Since 3.6.0 this matrix can contain also NAs and the position of such entries will be preserved by the SA;
K	The number of networks that has to be generated;
path	The directory in which the routine stores the outputs;
max.iter	"n" (default) the number of switching steps to be performed (or if $exact = TRUE$ the number of <b>successful</b> switching steps). If equal to "n" then this number is considered equal to the analytically derived lower bound presented in $Gobbi$ $et$ $al$ . (see References): $N = e/2(1-d)\ln\left((e-de)/\delta\right)$ if exact is FALSE, $N = e(1-d)/2\ln\left((e-de)/\delta\right)$ otherwise , where $e$ is the number of edges of $g$ and $d$ its edge density . This bound is much lower than the empirical one proposed in $Milo$ $et$ $al$ . $2003$ (see References);
accuracy	0.00005 (default) is the desired level of accuracy reflecting the average distance between the Jaccard index at the N-th step and its analytically derived fixed point in terms of fracion of common edges;
verbose	TRUE (default). When TRUE a progression bar is printed during computation.
MAXITER_MUL	10 (default). If <i>exact</i> == <i>TRUE</i> in order to prevent a possible infinite loop the program stops anyway after MAXITER_MUL*max.iter iterations;
exact	FALSE (default). If TRUE the program performs <i>max.iter</i> swithcing steps, otherwise the program will count also the not-performed swithcing steps;
write.sparse	TRUE (default). If FALSE the table is written as an R data.frame (long time and

# **Details**

The routine creates, starting from the given path, different subfolders in order to have maximum 1000 files for folder. Moreover the incidence matrices are saved using write\_stm\_CLUTO (sparse matrices) that can be loaded using read\_stm\_CLUTO. The set is generated calling birewire.rewire.bipartite on the last generated graph starting from the input graph.

### Author(s)

Andrea Gobbi: <gobbi.andrea@mail.com>

more space needed)

22 birewire.sampler.dsg

#### References

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), *On the uniform generation of random graphs with prescribed degree sequences*, eprint arXiv:cond-mat/0312028

birewire.sampler.dsg Efficient generation of a null model for a given dsg.

# Description

Efficient generation of a null model for a given dsg. The routine samples correctly from the null model of a given dsg creating a set of randomized dsgs.

#### Usage

birewire.sampler.dsg(dsg,K,path,delimitators=list(negative='-',positive='+'),exact=FALSE,
 verbose=TRUE, max.iter.pos='n',max.iter.neg='n', accuracy=0.00005,MAXITER\_MUL=10,check\_pos\_neg=FALSE

#### **Arguments**

dsg

A dsg object: is a list of two incidence matrices (see References), "positive" and "negative", encoding the positive edges and negative edges. This list can be obtained reading a SIF file using birewire.load.dsg function and converting the resulting dataframe using birewire.induced.bipartite.

max.iter.pos

"n" (default) the number of switching steps to be performed on the positive part of dsg (or if exact==TRUE the number of **successful** switching steps). If equal to "n" then this number is considered equal to the analytically derived lower bound presented in  $Gobbi\ et\ al.$  (see References):  $N=e/2(1-d)\ln\left((e-de)/\delta\right)$  if exact is FALSE,  $N=e(1-d)/2\ln\left((e-de)/\delta\right)$  otherwise, where e is the number of edges of g and d its edge density. This bound is much lower than the empirical one proposed in  $Milo\ et\ al.\ 2003$  (see References);

max.iter.neg

"n" (default) the number of switching steps to be performed on the negative part of dsg (or if exact==TRUE the number of **successful** switching steps). If equal to "n" then this number is considered equal to the analytically derived lower bound presented in  $Gobbi\ et\ al.$  (see References):  $N=e/2(1-d)\ln{((e-de)/\delta)}$  if exact is FALSE,  $N=e(1-d)/2\ln{((e-de)/\delta)}$  otherwise , where e is the number of edges of g and d its edge density . This bound is much lower than the empirical one proposed in  $Milo\ et\ al.\ 2003$  (see References);

birewire.sampler.dsg 23

accuracy 0.00005 (default) is the desired level of accuracy reflecting the average distance

between the Jaccard index at the N-th step and its analytically derived fixed point

in terms of fracion of common edges;

verbose TRUE (default). When TRUE a progression bar is printed during computation.

MAXITER\_MUL 10 (default). If exact==TRUE in order to prevent a possible infinite loop the

program stops anyway after MAXITER\_MUL\*max.iter iterations;

exact FALSE (default). If TRUE the program performs max.iter swithcing steps, oth-

erwise the program will count also the not-performed swithcing steps;

path The directory in which the routine stores the outputs;

K The number of network that has to be generated;

delimitators list(negative='-',positive='+') (default):a list with 'positive' and 'negative' names

identifying the character encoding the relation used for writing the ouput with

birewire.build.dsg;

check\_pos\_neg FALSE (default): if true check if there are positive and negative loops in the

same arc, if so the routine will not save the dsg and goes on with the marcov

chain (it can be slower due to the chekcs);

#### **Details**

The routine creates, starting from a given path, different subfolders in order to have maximum 1000 files for folder; the SIF files are saved using birewire.write.dsg, an internal routine. The set is generated calling birewire.rewire.dsg on the last generated dsg starting from the input dsg.

#### Author(s)

Andrea Gobbi: <gobbi.andrea@mail.com>

#### References

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), *On the uniform generation of random graphs with prescribed degree sequences*, eprint arXiv:cond-mat/0312028

birewire.sampler.undirected

Efficient generation of a null model for a given undirected graph

# **Description**

The routine samples correctly from the null model of a given undirected graph creating a set of randomized version of the initial undirected graph.

#### Usage

```
birewire.sampler.undirected(adjacency,K,path,max.iter="n", accuracy=0.00005,
verbose=TRUE,MAXITER_MUL=10,exact=FALSE,write.sparse=TRUE)
```

#### **Arguments**

adjacency	Adjacency matrix of the initial undirected graph. Since 3.6.0 this matrix can contain also NAs and the position of such entries will be preserved by the SA;
K	The number of networks that has to be generated;
path	The directory in which the routine stores the outputs;
max.iter	"n" (default) see birewire.rewire.undirected for references
accuracy	0.00005 (default) is the desired level of accuracy reflecting the average distance between the Jaccard index at the N-th step and its analytically derived fixed point in terms of fracion of common edges;
verbose	TRUE (default). When TRUE a progression bar is printed during computation.
MAXITER_MUL	10 (default). If <i>exact==TRUE</i> in order to prevent a possible infinite loop the program stops anyway after MAXITER_MUL*max.iter iterations;
exact	FALSE (default). If TRUE the program performs <i>max.iter</i> swithcing steps, otherwise the program will count also the not-performed swithcing steps;
write.sparse	TRUE (default). If FALSE the table is written as an R data.frame (long time and more space needed)

#### **Details**

The routine creates, starting from the given path, different subfolders in order to have maximum 1000 files for folder. Moreover the incidence matrices are saved using write\_stm\_CLUTO (sparse matrices) that can be loaded using read\_stm\_CLUTO. The set is generated calling birewire.rewire.undirected on the last generated graph starting from the input graph.

#### Author(s)

Andrea Gobbi: <gobbi.andrea@mail.com>

birewire.similarity 25

#### References

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T.and Jurman, G.and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

Gobbi, A. and Jurman, G. (2013) *Theoretical and algorithmic solutions for null models in network theory* (Doctoral dissertation) http://eprints-phd.biblio.unitn.it/1125/

R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), *On the uniform generation of random graphs with prescribed degree sequences*, eprint arXiv:cond-mat/0312028

birewire.similarity

Compute the Jaccard similarity index between two binary matrices with the same number of non-null entries and the sam row- and column-wise sums.

# Description

Compute the Jaccard similarity index between two binary matrices with the same number of non-null entries and the sam row- and column-wise sums. The function accept also two *igraph* objects.

#### **Usage**

birewire.similarity( m1,m2)

### Arguments

m1 First matrix or graph; m2 Second matrix or graph.

#### Details

The **Jaccard** index between two sets *M* and *N* is defined as:

$$|M \cup N|/|M \cap N|$$

With M and N binary matrices, the Jaccard index is computed as:

$$\frac{\sum N_{i,j} \wedge M_{i,j}}{\sum N_{i,j} \vee M_{i,j}}.$$

The Jaccard index ranges between 0 and 1 and since 3.6.0 can be computed also among matrix with NAs.

#### Value

Returns the Jaccard similarity index between the objects.

### Author(s)

Andrea Gobbi

Maintainer: Andrea Gobbi <gobbi.andrea@mail.com>

# **Examples**

```
library(igraph)
library(BiRewire)
g <- graph.bipartite( rep(0:1,length=10), c(1:10))
g2=birewire.rewire.bipartite(g)
birewire.similarity(get.incidence(g,sparse=FALSE),get.incidence(g2,sparse=FALSE))
birewire.similarity(g,g2)</pre>
```

birewire.similarity.dsg

Compute the Jaccard similarity index between dsg.

# **Description**

Compute the Jaccard similarity index between dsg objects described in the same way (matrices of graphs).

# Usage

```
birewire.similarity.dsg( m1,m2)
```

# Arguments

m1 First dsg; m2 Second dsg.

# **Details**

See birewire.similarity for more details.

#### Value

Returns the Jaccard similarity index between the objects.

#### Author(s)

Andrea Gobbi

Maintainer: Andrea Gobbi <gobbi.andrea@mail.com>

### **Examples**

```
library(BiRewire)
data(test_dsg)
dsg <- birewire.induced.bipartite(test_dsg,sparse=FALSE)
birewire.similarity.dsg(dsg,birewire.rewire.dsg(dsg))
dsg <- birewire.induced.bipartite(test_dsg,sparse=TRUE)
birewire.similarity.dsg(dsg,birewire.rewire.dsg(dsg))</pre>
```

birewire.slum.to.sparseMatrix

The function transforms a triplet sparse matrix from slum package to a Matrix sparse matrix.

# **Description**

Transform a triplet sparse matrix from *slum* package to a *Matrix* sparse matrix that can be used by *igraph* for creating a network. This function could be used in order to analyze graphs obtained from samplers routines (birewire.sampler.undirected,birewire.sampler.dsg and birewire.sampler.bipartite.)

### Usage

```
birewire.slum.to.sparseMatrix( simple_triplet_matrix_sparse)
```

# **Arguments**

```
simple_triplet_matrix_sparse
```

A triplet sparse matrix, usually the object coming from read\_stm\_CLUTO.

#### Value

Returns an Matrix sparse matrix that could be used for building an igraph graph using graph.adjacency.

#### Author(s)

Andrea Gobbi

Maintainer: Andrea Gobbi <gobbi.andrea@mail.com>

birewire.visual.monitoring.bipartite

Visual monitoring of the Markov chain underlying the SA for directed graphs.

#### **Description**

This function generates a cascade-sampling from the model at different switching steps given in *sequence*. For each step the routine computes the pairwise Jaccard distance (1-JI) among the samples and perfroms, on the resulting matix, a dimentional scaling reduction (using Rtsne). If *display* is set to *TRUE* the relative plot is displayed.

# Usage

```
birewire.visual.monitoring.bipartite(data,accuracy=0.00005,verbose=FALSE,MAXITER_MUL=10,
    exact=FALSE,n.networks=100,perplexity=15,sequence=c(1,5,100,"n"),ncol=2,
    nrow=length(sequence)/ncol,display=TRUE)
```

# **Arguments**

data	The initial bipartite graph, either an incidence matrix or an <i>igraph</i> bipartite graph object. Since 3.6.0, if the matrix is provided, such matrix can contain also NAs and the position of such entries will be preserved by the SA;
accuracy	0.00005 (default) is the desired level of accuracy reflecting the average distance between the Jaccard index at the N-th step and its analytically derived fixed point in terms of fracion of common edges;
verbose	TRUE (default). When TRUE a progression bar is printed during computation.
MAXITER_MUL	10 (default). If <i>exact==TRUE</i> in order to prevent a possible infinite loop the program stops anyway after MAXITER_MUL*max.iter iterations;
exact	FALSE (default). If TRUE the program performs <i>max.iter</i> swithcing steps, otherwise the program will count also the not-performed swithcing steps;
n.networks	100 (default): the number of network generated for each step defined in sequence;
perplexity	15 (default): the value of perplexity passed to the function Rtsne;
sequence	c(1,5,100,"n")(default) the sequence of step for wich generating a sampler (seebirewire.sampler.bipar
ncol	2 (default). The number of column in the plot;
nrow	length(sequence)/ncol (default). The number of row in the plot;
display	TRUE (default). If TRUE the result is displayed.

#### **Details**

For each value p in sequence (it that can also contain the special character "n", see birewire.rewire.bipartite), the routine generates n.networks sampled each p SS from the SA initialized with the given data. Pariwise distance are computed using the Jaccard distance and the resulting matrix is the input for the dimensional scaling performed by the function Rtsne. An explorative plot is displayed if display is set to TRUE.

#### Value

A list containing the list containing the distance matrices *dist* and the list containing the tsne results *Rtsne* 

#### Author(s)

Andrea Gobbi

Maintainer: Andrea Gobbi <gobbi.andrea@mail.com>

#### References

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

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R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), *On the uniform generation of random graphs with prescribed degree sequences*, eprint arXiv:cond-mat/0312028

Van der Maaten, L.J.P. and Hinton, G.E. Visualizing High-Dimensional Data Using t-SNE. Journal of Machine Learning Research 9(Nov):2579-2605, 2008

#### **Examples**

```
library(BiRewire)
g <- graph.bipartite( rep(0:1,length=100), c(1:100))
tsne = birewire.visual.monitoring.bipartite(g,display=FALSE,n.networks=10,perplexity=1)</pre>
```

birewire.visual.monitoring.dsg

Visual monitoring of the Markov chain underlying the SA for dsgs.

### Description

This function generates a cascade-sampling from the model at different switching steps given in *sequence*. For each step the routine computes the pairwise Jaccard distance (1-JI) among the samples and perfroms, on the resulting matix, a dimentional scaling reduction (using Rtsne). If *display* is set to *TRUE* the relative plot is displayed.

#### Usage

```
birewire.visual.monitoring.dsg(data,accuracy=0.00005,verbose=FALSE,MAXITER\_MUL=10,exact=FALSE,n.net sequence.pos=c(1,5,100,"n"), sequence.neg=c(1,5,100,"n"),ncol=2,nrow=length(sequence.pos)/ncol,display=TRUE)
```

# **Arguments**

data	The initial dsg either in matrix or graph formulation 9see birewire.induced.bipartite.
accuracy	0.00005 (default) is the desired level of accuracy reflecting the average distance between the Jaccard index at the N-th step and its analytically derived fixed point in terms of fracion of common edges;
verbose	TRUE (default). When TRUE a progression bar is printed during computation.
MAXITER_MUL	10 (default). If <i>exact==TRUE</i> in order to prevent a possible infinite loop the program stops anyway after MAXITER_MUL*max.iter iterations;
exact	FALSE (default). If TRUE the program performs <i>max.iter</i> swithcing steps, otherwise the program will count also the not-performed swithcing steps;
n.networks	100 (default): the number of network generated for each step defined in sequence;
perplexity	15 (default): the value of perplexity passed to the function Rtsne;
sequence.pos	c(1,5,100,"n") (default) the sequence of step for wich generating a sampler (seebirewire.sampler.dsg) for the positive part of $data$
sequence.neg	same as sequence.pos but for the negative part
ncol	2 (default). The number of column in the plot;
nrow	length(sequence)/ncol (default). The number of row in the plot;
display	TRUE (default). If TRUE the result of tsne is displayed.

# **Details**

See birewire.visual.monitoring.bipartite for more details.

### Value

A list containing the list containing the distance matrices *dist* and the list containing the tsne results *Rtsne*.

# Author(s)

Andrea Gobbi

Maintainer: Andrea Gobbi <gobbi.andrea@mail.com>

#### References

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

Jaccard, P. (1901), Étude comparative de la distribution florale dans une portion des Alpes et des *Jura*, Bulletin de la Société Vaudoise des Sciences Naturelles 37: 547–579.

R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), *On the uniform generation of random graphs with prescribed degree sequences*, eprint arXiv:cond-mat/0312028

Van der Maaten, L.J.P. and Hinton, G.E. Visualizing High-Dimensional Data Using t-SNE. Journal of Machine Learning Research 9(Nov):2579-2605, 2008

# **Examples**

```
library(BiRewire)
data(test_dsg)
##bigger dsg
test_dsg_2=test_dsg
test_dsg_2[,1]=paste(test_dsg_2[,1],"_",sep="")
test_dsg_2[,3]=paste(test_dsg_2[,3],"_",sep="")

dsg <- birewire.induced.bipartite(rbind(test_dsg,test_dsg_2),sparse=FALSE)

tsne = birewire.visual.monitoring.dsg(dsg,exact=TRUE,sequence.pos=c(1,2,"n",100),sequence.neg=c(1,2,"n",60),n.networks=50,perplexity=1)</pre>
```

birewire.visual.monitoring.undirected

Visual monitoring of the Markov chain underlying the SA for undirected graphs.

# **Description**

This function generates a cascade-sampling from the model at different switching steps given in *sequence*. For each step the routine computes the pairwise Jaccard distance (1-JI) among the samples and perfroms, on the resulting matix, a dimentional scaling reduction (using Rtsne). If *display* is set to *TRUE* the relative plot is displayed.

### Usage

birewire.visual.monitoring.undirected(data,accuracy=0.00005,verbose=FALSE,MAXITER\_MUL=10,
exact=FALSE,n.networks=100,perplexity=15,sequence=c(1,5,100,"n"),ncol=2,
nrow=length(sequence)/ncol,display=TRUE)

# Arguments

data	The initial undirected graph, either an adjacency matrix or an <i>igraph</i> undirected graph object. Since 3.6.0, if the matrix is provided, such matrix can contain also NAs and the position of such entries will be preserved by the SA;
accuracy	0.00005 (default) is the desired level of accuracy reflecting the average distance between the Jaccard index at the N-th step and its analytically derived fixed point in terms of fracion of common edges;
verbose	TRUE (default). When TRUE a progression bar is printed during computation.
MAXITER_MUL	10 (default). If <i>exact==TRUE</i> in order to prevent a possible infinite loop the program stops anyway after MAXITER_MUL*max.iter iterations;
exact	FALSE (default). If TRUE the program performs <i>max.iter</i> swithcing steps, otherwise the program will count also the not-performed swithcing steps;
n.networks	100 (default): the number of network generated for each step defined in $sequence$ ;
perplexity	15 (default): the value of perplexity passed to the function Rtsne;
sequence	$c(1,5,100,"n") (default) \ the \ sequence \ of \ step \ for \ wich \ generating \ a \ sampler \ (see \ birewire.sampler.undirected)$
ncol	2 (default). The number of column in the plot;
nrow	length(sequence)/ncol (default). The number of row in the plot;
display	TRUE (default). If TRUE the result of tsne is displayed.

#### **Details**

For each value *p* in *sequence* (it that can also contain the special character "n", see birewire.rewire.bipartite), the routine generates *n.networks* sampled each *p* SS from the SA initialized with the given *data*. Pariwise distance are computed using the Jaccard distance and the resulting matrix is the input for the dimensional scaling performed by the function Rtsne. An explorative plot is displayed if *display* is set to TRUE.

#### Value

A list containing the list containing the distance matrices *dist* and the list containing the tsne results *Rtsne*.

### Author(s)

Andrea Gobbi

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#### References

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

Iorio, F. and and Bernardo-Faura, M. and Gobbi, A. and Cokelaer, T. and Jurman, G. and Saez-Rodriguez, J. (2016) *Efficient randomization of biological networks while preserving functional characterization of individual nodes* Bioinformatics 2016 1 (17):542 doi: 10.1186/s12859-016-1402-1.

Jaccard, P. (1901), Étude comparative de la distribution florale dans une portion des Alpes et des *Jura*, Bulletin de la Société Vaudoise des Sciences Naturelles 37: 547–579.

R. Milo, N. Kashtan, S. Itzkovitz, M. E. J. Newman, U. Alon (2003), *On the uniform generation of random graphs with prescribed degree sequences*, eprint arXiv:cond-mat/0312028

Van der Maaten, L.J.P. and Hinton, G.E. Visualizing High-Dimensional Data Using t-SNE. Journal of Machine Learning Research 9(Nov):2579-2605, 2008

# **Examples**

```
library(BiRewire)
g <- erdos.renyi.game(1000,0.1)
tsne = birewire.visual.monitoring.undirected(g,display=FALSE,n.networks=10,perplexity=1)</pre>
```

BRCA\_binary\_matrix

TCGA Brest Cancer data

### **Description**

Breast cancer samples and their respective mutations downloaded from the Cancer Cancer Genome Atlas (TCGA), used in *Gobbi et al.*. Germline mutations were filtered out of the list of reported mutations; synonymous mutations and mutations identified as benign and tolerated were also removed from the dataset. The bipartite graph resulting when considering this matrix as an incidence matrix has  $n_r = 757, n_c = 9757, e = 19758$  for an edge density equal to 0.27%.

#### **Usage**

```
data(BRCA_binary_matrix)
```

#### Source

http://tcga.cancer.gov/dataportal/

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# References

Gobbi, A. and Iorio, F. and Dawson, K. J. and Wedge, D. C. and Tamborero, D. and Alexandrov, L. B. and Lopez-Bigas, N. and Garnett, M. J. and Jurman, G. and Saez-Rodriguez, J. (2014) *Fast randomization of large genomic datasets while preserving alteration counts* Bioinformatics 2014 30 (17): i617-i623 doi: 10.1093/bioinformatics/btu474.

test\_dsg

Tool example of dsg

# Description

A simple dsg for testing routines.

# Usage

data(test\_dsg)

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