## Package 'BiRewire'

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**Title** High-performing routines for the randomization of a bipartite graph (or a binary event matrix) 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 function for the analysis of the introduced randomness across the switching and several other routines to analyse the resulting networks and their natural projections.
Extension to undirected networks (not bipartite) 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

useful and a visual montioring of the underlying Markov Chain has been implemented.

License GPL-3

Depends igraph, slam, tsne

Suggests RUnit, BiocGenerics

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

biocViews Network

NeedsCompilation yes

## **R** topics documented:

BiRewire-package																 •							2
birewire.analysis.bipartite			•	•			•	 •	•			•		•		 •		•	•			•	3
birewire.analysis.undirected	•	•	•	•	•	•	•		•			•	•	•	•	 •	•	•	•	•	•	•	5

birewire.bipartite.from.incidence	7
birewire.rewire.bipartite	8
birewire.rewire.bipartite.and.projections	10
birewire.rewire.undirected	12
birewire.sampler.bipartite	13
birewire.sampler.undirected	15
birewire.similarity	16
birewire.visual.monitoring.bipartite	17
birewire.visual.monitoring.undirected	19
BRCA_binary_matrix	21
	22

## Index

BiRewire-package The BiRewire package

## Description

R package for computationally-efficient rewiring of bipartite graphs (or randomisation of 0-1 tables with prescribed marginal totals) and undirected. 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. 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 swithicng algorithm has been implemented.

#### Details

Summary:

Package:BiRewireVersion:2.2.3Date: 2015-07-072.2.0Require: slam, igraph,tsne, R>=2.10URL: http://www.ebi.ac.uk/~iorio/BiRewireLicense:GPL-3

#### Author(s)

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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.

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 <i>g</i> (can be extracted from an igraph bipartite graph using the get.incidence function);
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 <i>exact==TRUE</i> the number of successful switching steps). If equal to "n" then this number is considered equal to the analytically derived lower bound presented in <i>Gobbi et al.</i> (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 <i>e</i> is the number of edges of <i>g</i> and <i>d</i> its edge density. This bound is much lower than the empirical one proposed in <i>Milo et al. 2003</i> (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 initial 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 perfomed *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.

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 <-graph.bipartite( rep(0:1,length=10), c(1:10))
##get the incidence matrix of g
m<-as.matrix(get.incidence(graph=g))
## set parameters
step=1
max=100*length(E(g))
## perform two different evaluate uping two different</pre>
```

## perform two different analysis using two different maximal number of switching steps scores<-birewire.analysis.bipartite(m,step,max,n.networks=10) scores2<-birewire.analysis.bipartite(m,step,"n",n.networks=10)</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 <i>g</i> (can be extracted from an igraph undirected graph using the get.adjacency function);
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;

"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 <i>Gobbi et</i> <i>al.</i> (see References): $N = e/(2d^3 - dd^2 + 2d + 2) \ln ((e - de)/\delta)$ if exact is FALSE, $N = e(1 - d)/2 \ln ((e - de)/\delta)$ otherwise, where <i>e</i> is the number of edges of <i>g</i> and <i>d</i> its edge density. This bound is much lower than the empirical one proposed in <i>Milo et al. 2003</i> (see References);
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;
TRUE (default). When TRUE a progression bar is printed during computation;
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;
FALSE (default). If TRUE the program performs <i>max.iter</i> swithcing steps, otherwise the program will count also the not-performed swithcing steps;
50 (default), the number of independent rewiring process starting from the same initial graph from which the mean value and the CI is computed.
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 perfomed *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.

Gobbi, A. and Jurman, G. (in preparation), *Number of required Switching Steps in the Switching Algorithm for undirected graphs*.

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))</pre>
```

```
## 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)
scores2<-birewire.analysis.undirected(m,step,"n",n.networks=10)</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>

#### 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)</pre>

```
#create the rewired bipartite graph
g2<-birewire.bipartite.from.incidence(m2,TRUE)</pre>
```

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
max.iter	"n" (default) the number of switching steps to be performed (or if <i>exact==TRUE</i> 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 <i>Gobbi et al.</i> (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 <i>e</i> is the number of edges of <i>g</i> and <i>d</i> its edge density. This bound is much lower than the empirical one proposed in <i>Milo et al. 2003</i> (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;

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

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)))</pre>
```

```
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 $N = e(1-d)/2\ln((e-de)/\delta)$ presented in <i>Gobbi et al.</i> (see References);

10

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;
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 order 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.

#### Examples

```
library(igraph)
library(BiRewire)
g <- simplify(graph.bipartite( rep(0:1,length=100),
c(c(1:100),seq(1,100,3),seq(1,100,7),100,seq(1,100,13),
seq(1,100,17),seq(1,100,19),seq(1,100,23),100
)))
##gets the incidence matrix of g
m<-as.matrix(get.incidence(graph=g))
## rewires g and its projections</pre>
```

result=birewire.rewire.bipartite.and.projections(g,step=10,max.iter="n",accuracy=0.00005)

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, exactly accuracy=0.00005, verbose=TRUE, maxit accuracy=0.00005, verbose=TRUE, verbose=TRU

## Arguments

adjacency	An igraph undirected graph $g$ or its adjacency matrix (can be extracted from $g$ using get.adjacency(g));
max.iter	"n" (default) the number of switching steps to be performed (or if <i>exact==TRUE</i> 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 <i>Gobbi et al.</i> (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 <i>e</i> is the number of edges of <i>g</i> and <i>d</i> its edge density. This bound is much lower than the empirical one proposed in <i>Milo et al. 2003</i> (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==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 Special thanks to: Maintainer: Andrea Gobbi <gobbi.andrea@mail.com> 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.

Gobbi, A. and Jurman, G. (in preparation), *Number of required Switching Steps in the Switching Algorithm for undirected graphs*.

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))</pre>
```

```
## 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;
К	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 <i>exact==TRUE</i> 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 <i>Gobbi et al.</i> (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 <i>e</i> is the number of edges of <i>g</i> and <i>d</i> its edge density. This bound is much lower than the empirical one proposed in <i>Milo et al. 2003</i> (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 <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.bipartite on the last generated graph starting from the input graph.

#### Author(s)

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.

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;
К	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\_CLUT0 (sparse matrices) that can be loaded using read\_stm\_CLUT0. 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> 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.

Gobbi, A. and Jurman, G. (in preparation), *Number of required Switching Steps in the Switching Algorithm for undirected graphs*.

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

<pre>birewire.similarity</pre>	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 nonnull 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.

#### Value

Returns the Jaccard similarity index between the objects.

#### 16

birewire.visual.monitoring.bipartite

#### 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.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 tsne). If *display* is 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;
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 <i>sequence</i> ;
perplexity	15 (default): the value of perplexity passed to the function tsne;
sequence	c(1,5,100,"n")(default) the sequence of step for wich generating a sampler (seebirewire.rewire.sample
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.

birewire.visual.monitoring.bipartite

#### 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 tsne. 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 *tsne*.

#### 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.

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 <- graph.bipartite( rep(0:1,length=100), c(1:100))
birewire.visual.monitoring.bipartite(g,display=FALSE,n.networks=10)</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 tsne). If *display* is 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;
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 <i>se-quence</i> ;
perplexity	15 (default): the value of perplexity passed to the function tsne;
sequence	c(1,5,100,"n")(default) the sequence of step for wich generating a sampler (seebirewire.rewire.sample
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 tsne. 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 *tsne*.

#### 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.

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)
birewire.visual.monitoring.undirected(g,display=FALSE,n.networks=10)</pre>
```

20

#### 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/

#### 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.

# Index

```
*Topic bipartite graph, incidece
                                              birewire.sampler.bipartite, 13
        matrix
                                              birewire.sampler.undirected, 15
    birewire.bipartite.from.incidence,
                                              birewire.similarity, 16
                                              birewire.visual.monitoring.bipartite,
*Topic bipartite graph, projection,
                                                       17
        rewire
                                              birewire.visual.monitoring.undirected,
    birewire.rewire.bipartite.and.projections,
                                                      19
        10
                                              BRCA_binary_matrix, 21
*Topic bipartite graph, rewire
    birewire.rewire.bipartite, 8
*Topic datasets
    BRCA_binary_matrix, 21
*Topic directed graph, rewire,
        pathway, signaling
    birewire.sampler.bipartite, 13
*Topic package
    BiRewire-package, 2
*Topic rewire, bipartite graph
    birewire.analysis.bipartite,3
    birewire.visual.monitoring.bipartite,
        17
    birewire.visual.monitoring.undirected,
        19
*Topic rewire, undirected graph
    birewire.analysis.undirected, 5
*Topic similarity, jaccard
    birewire.similarity, 16
*Topic undirected graph, rewire
    birewire.rewire.undirected.12
    birewire.sampler.undirected, 15
BiRewire (BiRewire-package), 2
BiRewire-package, 2
birewire.analysis.bipartite, 3, 10, 11
birewire.analysis.undirected, 5
birewire.bipartite.from.incidence,7
birewire.rewire.bipartite, 8
birewire.rewire.bipartite.and.projections,
        10
birewire.rewire.undirected, 12
```