Package 'qpgraph'

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qpgraph-package

The q-order partial correlation graph learning software, apgraph.

Description

q-order partial correlation graphs, or qp-graphs for short, are undirected Gaussian graphical Markov models built from q-order partial correlations. They are useful for learning undirected graphical Gaussian Markov models from data sets where the number of random variables p exceeds the available sample size n as, for instance, in the case of microarray data where they can be employed to reverse engineer a molecular regulatory network.

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Functions

- qpNrr estimates non-rejection rates for every pair of variables.
- qpAvgNrr estimates average non-rejection rates for every pair of variables.
- qpGenNrr estimates generalized average non-rejection rates for every pair of variables.
- qpEdgeNrr estimate the non-rejection rate of one pair of variables.
- qpCItest performs a conditional independence test between two variables given a conditioning set.
- qpHist plots the distribution of non-rejection rates.
- qpGraph obtains a qp-graph from a matrix of non-rejection rates.
- qpAnyGraph obtains an undirected graph from a matrix of pairwise measurements.
- qpGraphDensity calculates and plots the graph density as function of the non-rejection rate.
- qpCliqueNumber calculates the size of the largest maximal clique (the so-called clique number or maximum clique size) in a given undirected graph.
- qpClique calculates and plots the size of the largest maximal clique (the so-called clique number or maximum clique size) as function of the non-rejection rate.
- qpGetCliques finds the set of (maximal) cliques of a given undirected graph.
- qpRndWishart random generation for the Wishart distribution.
- qpCov calculates the sample covariance matrix, just as the function cov() but returning a dspMatrix-class object which efficiently stores such a dense symmetric matrix.
- qpG2Sigma builds a random covariance matrix from an undrected graph. The inverse of the resulting matrix contains zeroes at the missing edges of the given undirected graph.
- qpUnifRndAssociation builds a matrix of uniformly random association values between 1 and +1 for all pairs of variables that follow from the number of variables given as input argument.
- qpK2ParCor obtains the partial correlation coefficients from a given concentration matrix.
- qpIPF performs maximum likelihood estimation of a sample covariance matrix given the independence constraints from an input list of (maximal) cliques.
- qpPAC estimates partial correlation coefficients and corresponding P-values for each edge in a given undirected graph, from an input data set.
- qpPCC estimates pairwise Pearson correlation coefficients and their corresponding P-values between all pairs of variables from an input data set.
- qpRndGraph builds a random undirected graph with a bounded maximum connectivity degree on every vertex.
- qpPrecisionRecall calculates the precision-recall curve for a given measure of association between all pairs of variables in a matrix.
- qpPRscoreThreshold calculates the score threshold at a given precision or recall level from a given precision-recall curve.
- qpFunctionalCoherence estimates functional coherence of a given transcriptional regulatory network using Gene Ontology annotations.
- qpTopPairs reports a top number of pairs of variables according to either an association measure and/or occurring in a given reference graph.

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• qpPlotNetwork plots a network using the Rgraphviz library.

This package provides an implementation of the procedures described in (Castelo and Roverato, 2006, 2009). An example of its use for reverse-engineering of transcriptional regulatory networks from microarray data is available in the vignette qpTxRegNet and, the same directory, contains a pre-print of a book chapter describing the basic functionality of the package which serves the purpose of a basic users's guide. This package is a contribution to the Bioconductor (Gentleman et al., 2004) and gR (Lauritzen, 2002) projects.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n. *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Castelo, R. and Roverato, A. Reverse engineering molecular regulatory networks from microarray data with qp-graphs. *J. Comput. Biol.* 16(2):213-227, 2009.

Gentleman, R.C., Carey, V.J., Bates, D.M., Bolstad, B., Dettling, M., Dudoit, S., Ellis, B., Gautier, L., Ge, Y., Gentry, J., Hornik, K. Hothorn, T., Huber, W., Iacus, S., Irizarry, R., Leisch, F., Li, C., Maechler, M. Rosinni, A.J., Sawitzki, G., Smith, C., Smyth, G., Tierney, L., Yang, T.Y.H. and Zhang, J. Bioconductor: open software development for computational biology and bioinformatics. *Genome Biol.*, 5:R80, 2004.

Lauritzen, S.L. (2002). gRaphical Models in R. R News, 3(2)39.

Ecoli0xygen

Preprocessed microarray oxygen deprivation data and filtered RegulonDB data

Description

The data consist of two classes of objects, one containing normalized gene expression microarray data from Escherichia coli (E. coli) and the other containing a subset of filtered RegulonDB transcription regulatory relationships on E. coli.

Usage

data(EcoliOxygen)

Format

gds680.eset
subset.gds680.eset
filtered.regulon6.1
subset.filtered.regulon6.1

ExpressionSet object containing n=43 experiments of various mutants under oxygen dep ExpressionSet object corresponding to a subset of gds680.eset defined by the transcript Data frame object containing a subset of the E. coli transcriptional network from RegulonI Subset of filtered.regulon6.1 containing the transcriptional regulatory relationships in

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Source

Covert, M.W., Knight, E.M., Reed, J.L., Herrgard, M.J., and Palsson, B.O. Integrating high-throughput and computational data elucidates bacterial networks. *Nature*, 429(6987):92-96, 2004.

Gama-Castro, S., Jimenez-Jacinto, V., Peralta-Gil, M., Santos-Zavaleta, A., Penaloza-Spinola, M.I., Contreras-Moreira, B., Segura-Salazar, J., Muniz-Rascado, L., Martinez-Flores, I., Salgado, H., Bonavides-Martinez, C., Abreu-Goodger, C., Rodriguez-Penagos, C., Miranda-Rios, J., Morett, E., Merino, E., Huerta, A.M., Trevino-Quintanilla, L., and Collado-Vides, J. RegulonDB (version 6.0): gene regulation model of Escherichia coli K-12 beyond transcription, active (experimental) annotated promoters and Textpresso navigation. *Nucleic Acids Res.*, 36(Database issue):D120-124, 2008.

References

Castelo, R. and Roverato, A. Reverse engineering molecular regulatory networks from microarray data with qp-graphs. *J. Comp. Biol.*, 16(2):213-227, 2009.

Examples

data(EcoliOxygen)
ls()

eQTLcross-class

eQTL experimental cross model class

Description

The expression quantitative trait loci (eQTL) experimental cross model class serves the purpose of holding all necessary information to simulate genetical genomics data from an experimental cross.

Author(s)

R. Castelo

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eQTLnetwork-class

eQTL network model class

Description

The expression quantitative trait loci (eQTL) network class serves the purpose of holding the result of estimating an eQTL network from genetical genomics data.

Author(s)

R. Castelo

See Also

 ${\tt eQTLnetworkEstimationParam-class} \ {\tt eQTLcross-class}$

 ${\it eQTL} network {\it Estimate}$

Estimation of a eQTL network from genetical genomics data

Description

The methods eQTLnetworkEstimate() perform the estimation of eQTL networks from genetical genomics data using as input an eQTLnetworkEstimationParam object and, eventually, a eQTLnetwork-class object.

Author(s)

R. Castelo

See Also

 ${\tt eQTLnetworkEstimationParam} \ {\tt eQTLnetwork-class}$

eQTLnetworkEstimationParam-class

eQTL network parameter model class

Description

The expression quantitative trait loci (eQTL) network parameter class serves the purpose of holding the parameter information to estimate an eQTL network from genetical genomics data with the function eQTLnetworkEstimate.

Author(s)

R. Castelo

See Also

eQTLnetworkEstimate eQTLnetwork-class

graphParam-class

Graph parameter classes

Description

Graph parameter classes are defined to ease the simulation of different types of graphs by using a single interface rgraphBAM().

Author(s)

R. Castelo

HMgmm-class

Homogeneous mixed graphical Markov model

Description

The "HMgmm" class is the class of homogeneous mixed graphical Markov models defined within the qpgraph package to store simulate and manipulate this type of graphical Markov models (GMMs).

An homogeneous mixed GMM is a family of multivariate conditional Gaussian distributions on mixed discrete and continuous variables sharing a set of conditional independences encoded by means of a marked graph. Further details can be found in the book of Lauritzen (1996).

Objects from the Class

Objects can be created by calls of the form HMgmm(g, ...) corresponding to constructor methods or rHMgmm(n, g, ...) corresponding to random simulation methods.

Slots

- pI: Object of class "integer" storing the number of discrete random variables.
- pY: Object of class "integer" storing the number of continuous random variables.
- g: Object of class graphBAM-class storing the associated marked graph.
- vtype: Object of class "factor" storing the type (discrete or continuous) of each random variable.
- dLevels: Object of class "integer" storing the number of levels of each discrete random variable.
- a: Object of class "numeric" storing the vector of additive linear effects on continuous variables connected to discrete ones.
- rho: Object of class "numeric" storing the value of the marginal correlation between two continuous random variables.
- sigma: Object of class dspMatrix-class storing the covariance matrix.
- mean: Object of class "numeric" storing the mean vector.
- eta2: Object of class "numeric" storing for each continuous variable connected to a discrete one, the fraction of variance of the continuous variable explained by the discrete one.

Methods

- HMgmm(g) Constructor method where g can be either an adjacency matrix or a graphBAM-class object.
- rHMgmm(n, g) Constructor simulation method that allows one to simulate homogeneous mixed GMMs where n is the number of GMMs to simulate and g can be either a markedGraphParam object, an adjacency matrix or a graphBAM-class object.
- names(x) Accessor method to obtain the names of the elements in the object x that can be retrieved with the \$ accessor operator.
- \$ Accessor operator to retrieve elements of the object in an analogous way to a list.
- dim(x) Dimension of the homogeneous mixed GMM corresponding to the number of discrete and continuous random variables.
- dimnames(x) Names of the discrete and continuous random variables in the homogeneous mixed GMM.
- show(object) Method to display some bits of information about the input homogeneous mixed GMM specified in object.
- summary(object) Method to display a sumarry of the main features of the input homogeneous mixed GMM specified in object.
- plot(x, ...) Method to plot the undirected graph associated to the the input homogeneous mixed GMM specified in x. It uses the plotting capabilities from the Rgraphviz library to which further arguments specified in ... are further passed.

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Author(s)

R. Castelo

References

Lauritzen, S.L. Graphical models. Oxford University Press, 1996.

See Also

UGgmm

qpAllCItests

Tests of conditional independence

Description

Performs a test of conditional independence for every pair of variables.

Usage

Arguments

Χ	data set from where to estimate the non-rejection rates. It can be an Expression-
	Set object, a data frame or a matrix.

I indexes or names of the variables in X that are discrete. See details below regarding this argument.

Q indexes or names of the variables in X forming the conditioning set.

pairup.i subset of vertices to pair up with subset pairup.j subset of vertices to pair up with subset pairup.i

long.dim.are.variables

logical; if TRUE it is assumed that when data are in a data frame or in a matrix, the longer dimension is the one defining the random variables (default); if FALSE, then random variables are assumed to be at the columns of the data frame or matrix

exact.test

logical; if FALSE an asymptotic conditional independence test is employed with mixed (i.e., continuous and discrete) data; if TRUE (default) then an exact conditional independence test with mixed data is employed. See details below regarding this argument.

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use a character string defining the way in which calculations are done in the presence

of missing values. It can be either "complete.obs" (default) or "em".

tol maximum tolerance controlling the convergence of the EM algorithm employed

when the argument use="em".

return.type type of value returned by this function. By default "p.value" indicates that a

list containing a matrix of p-values from all performed conditional independence (CI) tests will be returned. If return.type="statn" then a list containing the matrix of the statistics and the sample sizes on each CI test, will be returned. If return.type="all" then all previous matrices of values will be returned within

a list.

verbose show progress on the calculations.

R.code.only logical; if FALSE then the faster C implementation is used (default); if TRUE then

only R code is executed.

clusterSize size of the cluster of processors to employ if we wish to speed-up the calcula-

tions by performing them in parallel. A value of 1 (default) implies a single-processor execution. The use of a cluster of processors requires having previ-

ously loaded the packages snow and rlecuyer.

estimateTime logical; if TRUE then the time for carrying out the calculations with the given pa-

rameters is estimated by calculating for a limited number of adjacencies, specified by nAdj2estimateTime, and extrapolating the elapsed time; if FALSE (de-

fault) calculations are performed normally till they finish.

nAdj2estimateTime

number of adjacencies to employ when estimating the time of calculations (estimateTime=TRUE).

By default this has a default value of 10 adjacencies and larger values should provide more accurate estimates. This might be relevant when using a cluster

facility.

Details

When I is set different to NULL then mixed graphical model theory is employed and, concretely, it is assumed that the data comes from an homogeneous conditional Gaussian distribution. By default, with exact.test=TRUE, an exact test for conditional independence is employed, otherwise an asymptotic one will be used. Full details on these features can be found in Tur, Roverato and Castelo (2014).

Value

A list with three entries called p.value, statistic and n corresponding to a dspMatrix-class symmetric matrix of p-values for the null hypothesis of coindtional independence with the diagonal set to NA values, an analogous matrix of the statistics of each test and of the sample sizes, respectively. These returned values, however, depend on the setting of argument return.type which, by default, enables only returning the matrix of p-values. If arguments pairup.i and pairup.j are employed, those cells outside the constrained pairs will get also a NA value.

Note, however, that when estimateTime=TRUE, then instead of the matrix of estimated non-rejection rates, a vector specifying the estimated number of days, hours, minutes and seconds for completion of the calculations is returned.

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Author(s)

R. Castelo, A. Roverato and I. Tur

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n, *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Tur, I., Roverato, A. and Castelo, R. Mapping eQTL networks with mixed graphical models. *Submitted*, http://arxiv.org/abs/1402.4547, 2014.

See Also

```
qpCItest
```

Examples

```
library(mvtnorm)

nVar <- 50  ## number of variables
maxCon <- 3  ## maximum connectivity per variable
nObs <- 30  ## number of observations to simulate

set.seed(123)

A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))

alltests <- qpAllCItests(X, verbose=FALSE)

## distribution of p-values for the present edges
summary(alltests$p.value[upper.tri(alltests$p.value) & A])

## distribution of p-values for the missing edges
summary(alltests$p.value[upper.tri(alltests$p.value) & !A])</pre>
```

qpAnyGraph

A graph

Description

Obtains an undirected graph from a matrix of pairwise measurements

Usage

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Arguments

measurementsMatrix

matrix of pairwise measurements.

threshold threshold on the measurements below or above which pairs of variables are as-

sumed to be disconnected in the resulting graph.

remove direction of the removal with the threshold. It should be either "below" (default)

or "above".

topPairs number of edges from the top of the ranking, defined by the pairwise measure-

ments in measurementsMatrix, to use to form the resulting graph. This param-

eter is incompatible with a value different from NULL in threshold.

decreasing logical, only applies when topPairs is set; if TRUE then the ranking is made in

decreasing order; if FALSE then is made in increasing order.

pairup.i subset of vertices to pair up with subset pairup.j subset of vertices to pair up with subset pairup.i

Details

This is a general purpose function for thresholding a matrix of pairwise measurements to select pairs of variables corresponding to selected edges in an undirected graph.

Value

The resulting undirected graph as a graphBAM object. Note that when some gold-standard graph is available for comparison, a value for the parameter threshold can be found by calculating a precision-recall curve with qpPrecisionRecall with respect to this gold-standard, and then using qpPRscoreThreshold. Parameters threshold and topPairs are mutually exclusive, that is, when we specify with topPairs=n that we want a graph with n edges then threshold cannot be used.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n, *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

See Also

qpNrr qpAvgNrr qpEdgeNrr qpGraph qpGraphDensity qpClique qpPrecisionRecall qpPRscoreThreshold

Examples

```
require(mvtnorm)
require(graph)

nVar <- 50 ## number of variables
maxCon <- 5 ## maximum connectivity per variable</pre>
```

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```
nObs <- 30 ## number of observations to simulate
set.seed(123)
A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)</pre>
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))</pre>
## estimate Pearson correlations
pcc.estimates <- qpPCC(X)</pre>
## the higher the threshold
g <- qpAnyGraph(abs(pcc.estimates$R), threshold=0.9,</pre>
                remove="below")
## the sparser the qp-graph
numEdges(g) / choose(numNodes(g), 2)
## the lower the threshold
g <- qpAnyGraph(abs(pcc.estimates$R), threshold=0.5,</pre>
                 remove="below")
# the denser the graph
numEdges(g) / choose(numNodes(g), 2)
```

qpAvgNrr

Average non-rejection rate estimation

Description

Estimates average non-rejection rates for every pair of variables.

Usage

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Arguments

tol

Χ data set from where to estimate the average non-rejection rates. It can be an ExpressionSet object, a data frame or a matrix. q0rders either a number of partial-correlation orders or a vector of vector of particular orders to be employed in the calculation. Ι indexes or names of the variables in X that are discrete. When X is an ExpressionSet then I may contain only names of the phenotypic variables in X. See details below regarding this argument. restrict.Q indexes or names of the variables in X that restrict the sample space of conditioning subsets Q. fix.Q indexes or names of the variables in X that should be fixed within every conditioning conditioning subsets Q. nTests number of tests to perform for each pair for variables. significance level of each test. alpha subset of vertices to pair up with subset pairup.j pairup.i pairup.j subset of vertices to pair up with subset pairup.i long.dim.are.variables logical; if TRUE it is assumed that when the data is a data frame or a matrix, the longer dimension is the one defining the random variables; if FALSE, then random variables are assumed to be at the columns of the data frame or matrix. type of average. By now only the arithmetic mean is available. type verbose show progress on the calculations. use identical conditioning subsets for every pair of vertices (default), otherwise identicalQs sample a new collection of nTests subsets for each pair of vertices. exact.test logical; if FALSE an asymptotic conditional independence test is employed with mixed (i.e., continuous and discrete) data; if TRUE (default) then an exact conditional independence test with mixed data is employed.

a character string defining the way in which calculations are done in the presence of missing values. It can be either "complete.obs" (default) or "em".

maximum tolerance controlling the convergence of the EM algorithm employed

when the argument use="em".

R.code.only logical; if FALSE then the faster C implementation is used (default); if TRUE then only R code is executed.

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clusterSize size of the cluster of processors to employ if we wish to speed-up the calcula-

tions by performing them in parallel. A value of 1 (default) implies a single-processor execution. The use of a cluster of processors requires having previ-

ously loaded the packages snow and rlecuyer.

estimateTime logical; if TRUE then the time for carrying out the calculations with the given pa-

rameters is estimated by calculating for a limited number of adjacencies, specified by nAdj2estimateTime, and extrapolating the elapsed time; if FALSE (de-

fault) calculations are performed normally till they finish.

nAdj2estimateTime

 $number\ of\ adjacencies\ to\ employ\ when\ estimating\ the\ time\ of\ calculations\ (estimate Time = TRUE).$

By default this has a default value of 10 adjacencies and larger values should provide more accurate estimates. This might be relevant when using a cluster

facility.

Details

Note that when specifying a vector of particular orders q, these values should be in the range 1 to min(p, n-3), where p is the number of variables and n the number of observations. The computational cost increases linearly within each q value and quadratically in p. When setting identicalQs to FALSE the computational cost may increase between 2 times and one order of magnitude (depending on p and q) while asymptotically the estimation of the non-rejection rate converges to the same value.

When I is set different to NULL then mixed graphical model theory is employed and, concretely, it is assumed that the data comes from an homogeneous conditional Gaussian distribution. In this setting further restrictions to the maximum value of q apply, concretely, it cannot be smaller than p plus the number of levels of the discrete variables involved in the marginal distributions employed by the algorithm. By default, with exact.test=TRUE, an exact test for conditional independence is employed, otherwise an asymptotic one will be used. Full details on these features can be found in Tur, Roverato and Castelo (2014).

Value

A dspMatrix-class symmetric matrix of estimated average non-rejection rates with the diagonal set to NA values. When using the arguments pairup.i and pairup.j, those cells outside the constraint pairs will get also a NA value.

Note, however, that when estimateTime=TRUE, then instead of the matrix of estimated average non-rejection rates, a vector specifying the estimated number of days, hours, minutes and seconds for completion of the calculations is returned.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. Reverse engineering molecular regulatory networks from microarray data with qp-graphs. *J. Comp. Biol.*, 16(2):213-227, 2009.

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Tur, I., Roverato, A. and Castelo, R. Mapping eQTL networks with mixed graphical models. *Submitted*, http://arxiv.org/abs/1402.4547, 2014.

See Also

```
qpNrr qpEdgeNrr qpHist qpGraphDensity qpClique
```

Examples

```
require(mvtnorm)
nVar <- 50 ## number of variables
maxCon <- 3 ## maximum connectivity per variable</pre>
nObs <- 30 ## number of observations to simulate
set.seed(123)
A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)</pre>
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))</pre>
avgnrr.estimates <- qpAvgNrr(X, verbose=FALSE)</pre>
## distribution of average non-rejection rates for the present edges
summary(avgnrr.estimates[upper.tri(avgnrr.estimates) & A])
## distribution of average non-rejection rates for the missing edges
summary(avgnrr.estimates[upper.tri(avgnrr.estimates) & !A])
## Not run:
library(snow)
library(rlecuyer)
## only for moderate and large numbers of variables the
## use of a cluster of processors speeds up the calculations
nVar <- 500
maxCon <- 3
A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)</pre>
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))</pre>
system.time(avgnrr.estimates <- qpAvgNrr(X, q=10, verbose=TRUE))</pre>
system.time(avgnrr.estimates <- qpAvgNrr(X, q=10, verbose=TRUE, clusterSize=4))</pre>
## End(Not run)
```

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Description

Calculates and plots the size of the largest vertex boundary as function of the non-rejection rate.

Usage

Arguments

nrrMatrix matrix of non-rejection rates.

n number of observations from where the non-rejection rates were estimated.

threshold.lim range of threshold values on the non-rejection rate.

breaks either a number of threshold bins or a vector of threshold breakpoints.

vertexSubset subset of vertices for which their maximum boundary size is calculated with

respect to all other vertices.

plot logical; if TRUE makes a plot of the result; if FALSE it does not.

qpBoundaryOutput

output from a previous call to qpBoundary. This allows one to plot the result changing some of the plotting parameters without having to do the calculation

again.

density.digits number of digits in the reported graph densities.

logscale.bdsize

logical; if TRUE then the scale for the maximum boundary size is logarithmic which is useful when working with more than 1000 variables; FALSE otherwise

(default).

titlebd main title to be shown in the plot.

verbose show progress on calculations.

Details

The maximum boundary is calculated as the largest degree among all vertices of a given qp-graph.

Value

A list with the maximum boundary size and graph density as function of threshold, the threshold on the non-rejection rate that provides a maximum boundary size strictly smaller than the sample size n and the resulting maximum boundary size.

Author(s)

R. Castelo and A. Roverato

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References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n. *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

See Also

```
qpHTF qpGraphDensity
```

Examples

```
require(mvtnorm)

nVar <- 50  ## number of variables
maxCon <- 5  ## maximum connectivity per variable
nObs <- 30  ## number of observations to simulate
set.seed(123)

A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))

## the higher the q the less complex the qp-graph
nrr.estimates <- qpNrr(X, q=1, verbose=FALSE)
qpBoundary(nrr.estimates, plot=FALSE)

qpBoundary(nrr.estimates, plot=FALSE)</pre>
```

qpCItest

Conditional independence test

Description

Performs a conditional independence test between two variables given a conditioning set.

Usage

qpCItest 19

Arguments

X	data set where the test should be performed. It can be either an ExpressionSet object, a qtl::cross object, a data frame, a matrix or an SsdMatrix-class object. In the latter case, the input matrix should correspond to a sample covariance matrix of data on which we want to test for conditional independence. The function qpCov() can be used to estimate such matrices.
i	index or name of one of the two variables in X to test.
j	index or name of the other variable in X to test.
Q	indexes or names of the variables in X forming the conditioning set.
I	indexes or names of the variables in X that are discrete. See details below regarding this argument.
long.dim.are.v	ariables
	logical; if TRUE it is assumed that when data are in a data frame or in a matrix, the longer dimension is the one defining the random variables (default); if FALSE, then random variables are assumed to be at the columns of the data frame or matrix.
exact.test	logical; if FALSE an asymptotic likelihood ratio test of conditional independence test is employed with mixed (i.e., continuous and discrete) data; if TRUE (default) then an exact likelihood ratio test of conditional independence with mixed data is employed. See details below regarding this argument.
use	a character string defining the way in which calculations are done in the presence of missing values. It can be either "complete.obs" (default) or "em".
tol	maximum tolerance controlling the convergence of the EM algorithm employed when the argument use=" em ".
R.code.only	logical; if FALSE then the faster C implementation is used (default); if TRUE then only R code is executed.

Details

When variables in i, j and Q are continuous and I=NULL, this function performs a conditional independence test using a t-test for zero partial regression coefficient (Lauritzen, 1996, pg. 150). Note that the size of possible Q sets should be in the range 1 to $\min(p, n-3)$, where p is the number of variables and n the number of observations. The computational cost increases linearly with the number of variables in Q.

When variables in i, j and Q are continuous and discrete (mixed data), indicated with the I argument when X is a matrix, then mixed graphical model theory (Lauritzen and Wermuth, 1989) is employed and, concretely, it is assumed that data come from an homogeneous conditional Gaussian

20 qpCItest

distribution. By default, with exact.test=TRUE, an exact likelihood ratio test for conditional independence is performed (Lauritzen, 1996, pg. 192-194; Tur, Roverato and Castelo, 2014), otherwise an asymptotic one is used.

In this setting further restrictions to the maximum value of q apply, concretely, it cannot be smaller than p plus the number of levels of the discrete variables involved in the marginal distributions employed by the algorithm.

Value

A list with class "htest" containing the following components:

statistic in case of pure continuous data and I=NULL, the t-statistic for zero partial re-

gression coefficient; when I!=NULL, the value Lambda of the likelihood ratio if

exact.test=TRUE and -n log Lambda otherwise.

parameter in case of pure continuous data and I=NULL, the degrees of freedom for the t-

statistic (n-q-2); when I!=NULL, the degrees of freedom for -n log Lambda of a chi-square distribution under the null hypothesis if exact.test=FALSE and the (a, b) parameters of a beta distribution under the null if exact.test=TRUE.

p.value the p-value for the test.

estimate in case of pure continuous data (I=NULL), the estimated partial regression coeffi-

cient. In case of mixed continuous and discrete data with I!=NULL, the estimated partial eta-squared: the fraction of variance from i or j explained by the other tested variable after excluding the variance explained by the variables in Q. If one of the tested variables i or j is discrete, then the partial eta-squared is calculated on the tested continuous variable. If both, i and j are continuous, then

the partial eta-squared is calculated on variable i.

alternative a character string describing the alternative hypothesis.

method a character string indicating what type of conditional independence test was

performed.

data.name a character string giving the name(s) of the random variables involved in the

conditional independence test.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n, *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Lauritzen, S.L. Graphical models. Oxford University Press, 1996.

Lauritzen, S.L and Wermuth, N. Graphical Models for associations between variables, some of which are qualitative and some quantitative. *Ann. Stat.*, 17(1):31-57, 1989.

Tur, I., Roverato, A. and Castelo, R. Mapping eQTL networks with mixed graphical models. *Submitted*, http://arxiv.org/abs/1402.4547, 2014.

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See Also

```
qpCov qpNrr qpEdgeNrr
```

Examples

qpClique

Complexity of the resulting qp-graphs

Description

Calculates and plots the size of the largest maximal clique (the so-called clique number or maximum clique size) as function of the non-rejection rate.

Usage

Arguments

nrrMatrix matrix of non-rejection rates.

n number of observations from where the non-rejection rates were estimated.

threshold.lim range of threshold values on the non-rejection rate.

breaks either a number of threshold bins or a vector of threshold breakpoints.

plot logical; if TRUE makes a plot of the result; if FALSE it does not.

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exact.calculation

logical; if TRUE then the exact clique number is calculated; if FALSE then a

lower bound is given instead.

approx.iter number of iterations to be employed in the calculation of the lower bound (i.e.,

only applies when exact.calculation=FALSE).

qpCliqueOutput output from a previous call to qpClique. This allows one to plot the result

changing some of the plotting parameters without having to do the calculation

again.

density.digits number of digits in the reported graph densities.

logscale.clqsize

logical; if TRUE then the scale for the maximum clique size is logarithmic which is useful when working with more than 1000 variables; FALSE otherwise (de-

fault).

titleclq main title to be shown in the plot.
verbose show progress on calculations.

Details

The estimate of the complexity of the resulting qp-graphs is calculated as the area enclosed under the curve of maximum clique sizes.

The maximum clique size, or clique number, is obtained by calling the function qpCliqueNumber The calculation of the clique number of an undirected graph is an NP-complete problem which means that its computational cost is bounded by an exponential running time (Pardalos and Xue, 1994). Therefore, giving breakpoints between 0.95 and 1.0 may result into very dense graphs which can lead to extremely long execution times. If it is necessary to look at that range of breakpoints it is recommended either to use the lower bound on the clique number (exact.calculation=FALSE) or to look at qpGraphDensity.

Value

A list with the maximum clique size and graph density as function of threshold, an estimate of the complexity of the resulting qp-graphs across the thresholds, the threshold on the non-rejection rate that provides a maximum clique size strictly smaller than the sample size n and the resulting maximum clique size.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n. *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Pardalos, P.M. and Xue, J. The maximum clique problem. J. Global Optim., 4:301-328, 1994.

See Also

qpCliqueNumber qpGraphDensity

qpCliqueNumber 23

Examples

```
require(mvtnorm)

nVar <- 50  ## number of variables
maxCon <- 5  ## maximum connectivity per variable
nObs <- 30  ## number of observations to simulate
set.seed(123)

A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))

## the higher the q the less complex the qp-graph
nrr.estimates <- qpNrr(X, q=1, verbose=FALSE)
qpClique(nrr.estimates, plot=FALSE)$complexity
nrr.estimates <- qpNrr(X, q=5, verbose=FALSE)
qpClique(nrr.estimates, plot=FALSE)$complexity</pre>
```

qpCliqueNumber

Clique number

Description

Calculates the size of the largest maximal clique (the so-called clique number or maximum clique size) in a given undirected graph.

Usage

Arguments

g either a graphNEL object or an adjacency matrix of the given undirected graph.

exact.calculation

logical; if TRUE then the exact clique number is calculated; if FALSE then a lower bound is given instead.

return.vertices

logical; if TRUE a set of vertices forming a maximal clique of maximum size is returned; if FALSE only the maximum clique size is returned.

approx.iter

number of iterations to be employed in the calculation of the lower bound (i.e., only applies when exact.calculation=FALSE.

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verbose show progress on calculations.

R.code.only logical; if FALSE then the faster C implementation is used (default); if TRUE

then only R code is executed.

Details

The calculation of the clique number of an undirected graph is one of the basic NP-complete problems (Karp, 1972) which means that its computational cost is bounded by an exponential running time (Pardalos and Xue, 1994). The current implementation uses C code from the GNU GPL Cliquer library by Niskanen and Ostergard (2003) based on the, probably the fastest to date, algorithm by Ostergard (2002).

The lower bound on the maximum clique size is calculated by ranking the vertices by their connectivity degree, put the first vertex in a set and go through the rest of the ranking adding those vertices to the set that form a clique with the vertices currently within the set. Once the entire ranking has been examined a large clique should have been built and eventually one of the largests ones. This process is repeated a number of times (approx.iter) each of which the ranking is altered with increasing levels of randomness acyclically (altering 1 to \$p\$ vertices and again). Larger values of approx.iter should provide tighter lower bounds although it has been proven that no polynomial time algorithm can approximate the maximum clique size within a factor of n^{ϵ} ($\epsilon > 0$), unless P=NP (Feige et al, 1991; Pardalos and Xue, 1994).

Value

a lower bound of the size of the largest maximal clique in the given graph, also known as its clique number.

Author(s)

R. Castelo

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n. *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Feige, U., Goldwasser, S., Lov\'asz, L., Safra, S. and Szegedy, M. Approximating the maximum clique is almost NP-Complete. *Proc. 32nd IEEE Symp. on Foundations of Computer Science*, 2-12, 1991.

Karp, R.M. Reducibility among combinatorial problems. *Complexity of computer computations*, 43:85-103, 1972.

Niskanen, S. Ostergard, P. Cliquer User's Guide, Version 1.0. Communications Laboratory, Helsinki University of Technology, Espoo, Finland, Tech. Rep. T48, 2003. (http://users.tkk.fi/~pat/cliquer.html)

Ostergard, P. A fast algorithm for the maximum clique problem. Discrete Appl. Math. 120:197-207, 2002.

Pardalos, P.M. and Xue, J. The maximum clique problem. J. Global Optim., 4:301-328, 1994.

qpCov 25

See Also

```
qpClique
```

Examples

```
require(graph)
nVar <- 50
set.seed(123)
g1 <- randomEGraph(V=as.character(1:nVar), p=0.3)
qpCliqueNumber(g1, verbose=FALSE)
g2 <- randomEGraph(V=as.character(1:nVar), p=0.7)
qpCliqueNumber(g2, verbose=FALSE)</pre>
```

qpCov

Calculation of the sample covariance matrix

Description

Calculates the sample covariance matrix, just as the function cov() but returning a dspMatrix-class object which efficiently stores such a dense symmetric matrix.

Usage

```
qpCov(X, corrected=TRUE)
```

Arguments

X data set from where to calculate the sample covariance matrix. As the cov()

function, it assumes the columns correspond to random variables and the rows

to multivariate observations.

corrected flag set to TRUE when calculating the sample covariance matrix (default; and set

to FALSE when calculating the uncorrected sum of squares and deviations.

Details

This function makes the same calculation as the cov function but returns a sample covariance matrix stored in the space-efficient class dspMatrix-class and, moreover, allows one for calculating the uncorrected sum of squares and deviations which equals (n-1) * cov().

Value

A sample covariance matrix stored as a dspMatrix-class object. See the Matrix package for full details on this object class.

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Author(s)

R. Castelo

See Also

qpPCC

Examples

```
require(graph)
require(mvtnorm)
nVar <- 50 ## number of variables
nObs <- 10 ## number of observations to simulate
set.seed(123)
g <- randomEGraph(as.character(1:nVar), p=0.15)</pre>
Sigma <- qpG2Sigma(g, rho=0.5)</pre>
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))</pre>
S \leftarrow qpCov(X)
## estimate Pearson correlation coefficients by scaling the sample covariance matrix
R <- cov2cor(as(S, "matrix"))</pre>
## get the corresponding boolean adjacency matrix
A \leftarrow as(g, "matrix") == 1
## Pearson correlation coefficients of the present edges
summary(abs(R[upper.tri(R) & A]))
## Pearson correlation coefficients of the missing edges
summary(abs(R[upper.tri(R) & !A]))
```

qpEdgeNrr

Non-rejection rate estimation for a pair of variables

Description

Estimates the non-rejection rate for one pair of variables.

Usage

Arguments

X	data set from where the non-rejection rate should be estimated. It can be either an ExpressionSet object a data frame, a matrix or an SsdMatrix-class object. In the latter case, the input matrix should correspond to a sample covariance matrix of data from which we want to estimate the non-rejection rate for a pair of variables. The function qpCov() can be used to estimate such matrices.	
i	index or name of one of the two variables in X to test.	
j	index or name of the other variable in X to test.	
q	order of the conditioning subsets employed in the calculation.	
I	indexes or names of the variables in \boldsymbol{X} that are discrete when \boldsymbol{X} is a matrix or a data frame.	
restrict.Q	indexes or names of the variables in \boldsymbol{X} that restrict the sample space of conditioning subsets \boldsymbol{Q} .	
fix.Q	indexes or names of the variables in \boldsymbol{X} that should be fixed within every conditioning conditioning subsets \boldsymbol{Q} .	
nTests	number of tests to perform for each pair for variables.	
alpha	significance level of each test.	
long.dim.are.variables		

logical; if TRUE it is assumed that when data are in a data frame or in a matrix, the longer dimension is the one defining the random variables (default); if FALSE, then random variables are assumed to be at the columns of the data frame or matrix.

frame or matrix.

logical; if FALSE an asymptotic conditional independence test is employed with mixed (i.e., continuous and discrete) data; if TRUE (default) then an exact conditional independence test with mixed data is employed. See details below regarding this argument.

a character string defining the way in which calculations are done in the presence of missing values. It can be either "complete.obs" (default) or "em".

exact.test

use

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tol maximum tolerance controlling the convergence of the EM algorithm employed

when the argument use="em".

R.code.only logical; if FALSE then the faster C implementation is used (default); if TRUE

then only R code is executed.

Details

The estimation of the non-rejection rate for a pair of variables is calculated as the fraction of tests that accept the null hypothesis of conditional independence given a set of randomly sampled q-order conditionals.

Note that the possible values of q should be in the range 1 to min(p,n-3), where p is the number of variables and n the number of observations. The computational cost increases linearly with q.

When I is set different to NULL then mixed graphical model theory is employed and, concretely, it is assumed that the data comes from an homogeneous conditional Gaussian distribution. In this setting further restrictions to the maximum value of q apply, concretely, it cannot be smaller than p plus the number of levels of the discrete variables involved in the marginal distributions employed by the algorithm. By default, with exact.test=TRUE, an exact test for conditional independence is employed, otherwise an asymptotic one will be used. Full details on these features can be found in Tur, Roverato and Castelo (2014).

The argument I specifying what variables are discrete actually applies only when X is a matrix object since in the other cases data types are specified for each data columns or slot.

Value

An estimate of the non-rejection rate for the particular given pair of variables.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n, *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Tur, I., Roverato, A. and Castelo, R. Mapping eQTL networks with mixed graphical models. *Submitted*, http://arxiv.org/abs/1402.4547, 2014.

See Also

qpNrr qpAvgNrr qpHist qpGraphDensity qpClique qpCov

Examples

```
require(mvtnorm)

nObs <- 100 ## number of observations to simulate

## the following adjacency matrix describes an undirected graph

## where vertex 3 is conditionally independent of 4 given 1 AND 2
```

qpFunctionalCoherence 29

qpFunctionalCoherence Functional coherence estimation

Description

Estimates functional coherence for a given transcriptional regulatory network specified either as an adjacency matrix with a list of transcription factor gene identifiers or as a list of transcriptional regulatory modules, whose element names determine which genes encode for transcription factor proteins.

Usage

```
## S4 method for signature lsCMatrix
qpFunctionalCoherence(object, TFgenes, geneUniverse=rownames(object),
                               chip, minRMsize=5, removeGOterm="transcription",
                                            verbose=FALSE, clusterSize=1)
## S4 method for signature lspMatrix
qpFunctionalCoherence(object, TFgenes, geneUniverse=rownames(object),
                               chip, minRMsize=5, removeGOterm="transcription",
                                            verbose=FALSE, clusterSize=1)
## S4 method for signature lsyMatrix
qpFunctionalCoherence(object, TFgenes, geneUniverse=rownames(object),
                               chip, minRMsize=5, removeGOterm="transcription",
                                            verbose=FALSE, clusterSize=1)
## S4 method for signature matrix
qpFunctionalCoherence(object, TFgenes, geneUniverse=rownames(object),
                               chip, minRMsize=5, removeGOterm="transcription",
                                         verbose=FALSE, clusterSize=1)
## S4 method for signature list
qpFunctionalCoherence(object, geneUniverse=unique(c(names(object), unlist(object, use.names=FALSE)))
                               chip, minRMsize=5, removeGOterm="transcription",
                                       verbose=FALSE, clusterSize=1)
```

Arguments

object containing the transcriptional regulatory modules for which we want to

estimate their functional coherence. It can be an adjacency matrix of the undirected graph representing the transcriptional regulatory network or a list of gene target sets where the name of the entry should be the transcription factor gene

identifier.

TFgenes when the input object is a matrix, it is required to provide a vector of tran-

scription factor gene identifiers (which should match somewhere in the row and

column names of the matrix.

geneUniverse vector of all genes considered in the analysis. By default it equals the rows and

column names of object when it is a matrix, or the set of all different gene

identifiers occuring in object when it is a list.

chip name of the .db package containing the Gene Ontology (GO) annotations.

minRMsize minimum size of the target gene set in each regulatory module where functional

enrichment will be calculated and thus where functional coherence will be esti-

mated.

removeGOterm word, or regular pattern, matching GO terms that should be excluded in the

transcription factor gene GO annotations, and in the target gene if the regulatory module has only one gene, prior to the calculation of functional coherence.

verbose logical; if TRUE the function will show progress on the calculations; if FALSE

the function will remain quiet (default).

clusterSize size of the cluster of processors to employ if we wish to speed-up the calcula-

tions by performing them in parallel. A value of 1 (default) implies a single-processor execution. The use of a cluster of processors requires having previ-

ously loaded the packages snow and rlecuyer.

Details

This function estimates the functional coherence of a transcriptional regulatory network represented by means of an undirected graph encoded by either an adjacency matrix and a vector of transcription factor genes, or a list of regulatory modules each of them defined by a transcription factor gene and its targets. The functional coherence of a transcriptional regulatory network is calculated as specified by Castelo and Roverato (2009) and corresponds to the distribution of individual functional coherence values of every of the regulatory modules of the network each of them defined as a transcription factor and its set of putatively regulated target genes. In the calculation of the functional coherence value of a regulatory module, Gene Ontology (GO) annotations are employed through the given annotation .db package and the conditional hyper-geometric test implemented in the GOstats package from Bioconductor.

When a regulatory module has only one target gene, then no functional enrichment is calculated and, instead, the GO trees, grown from the GO annotations of the transcription factor gene and its target, are directly compared.

Value

A list with the following elements: the transcriptional regulatory network as a list of regulatory modules and their targets; the previous list of regulatory modules but excluding those with no

enriched GO BP terms. When the regulatory module has only one target, then instead the GO BP annotations of the target gene are included; a vector of functional coherence values.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. Reverse engineering molecular regulatory networks from microarray data with qp-graphs. *J. Comp. Biol.*, 16(2):213-227, 2009.

See Also

qpAvgNrr qpGraph

Examples

```
## example below takes about minute and a half to execute and for
## that reason it is not executed by default
## Not run:
library(GOstats)
library(org.EcK12.eg.db)
## load RegulonDB data from this package
data(EcoliOxygen)
## pick two TFs from the RegulonDB data in this package
TFgenes <- c("mhpR", "iscR")</pre>
## get their Entrez Gene Identifiers
TFgenesEgIDs <- unlist(mget(TFgenes, AnnotationDbi::revmap(org.EcK12.egSYMBOL)))</pre>
## get all genes involved in their regulatory modules from
## the RegulonDB data in this package
mt <- match(filtered.regulon6.1[,"EgID_TF"], TFgenesEgIDs)</pre>
allGenes <- as.character(unique(as.vector(</pre>
            as.matrix(filtered.regulon6.1[!is.na(mt),
                                            c("EgID_TF", "EgID_TG")]))))
mtTF <- match(filtered.regulon6.1[,"EgID_TF"],allGenes)</pre>
mtTG <- match(filtered.regulon6.1[,"EgID_TG"],allGenes)</pre>
## select the corresponding subset of the RegulonDB data in this package
subset.filtered.regulon6.1 <- filtered.regulon6.1[!is.na(mtTF) & !is.na(mtTG),]</pre>
TFi <- match(subset.filtered.regulon6.1[,"EgID_TF"], allGenes)</pre>
TGi <- match(subset.filtered.regulon6.1[,"EgID_TG"], allGenes)
subset.filtered.regulon6.1 <- cbind(subset.filtered.regulon6.1,</pre>
                                      idx_TF=TFi, idx_TG=TGi)
```

32 qpG2Sigma

qpG2Sigma

Random covariance matrix

Description

Builds a positive definite matrix from an undirected graph G that can be used as a covariance matrix for a Gaussian graphical model with graph G. The inverse of the resulting matrix contains zeroes at the missing edges of the given undirected graph G.

Usage

Arguments

g undirected graph specified either as a graphNEL object or as an adjacency matrix.

rho real number between -1/(n.var-1) and 1 corresponding to the mean marginal correlation

matrix.completion
 algorithm to employ in the matrix completion operations employed to construct a positive definite matrix with the zero pattern specified in g

tol tolerance under which the matrix completion algorithm stops.

verbose show progress on the calculations.

R.code.only logical; if FALSE then the faster C implementation is used in the internal call to

the HTF, or IPF, algorithm (default); if TRUE then only R code is executed.

qpGenNrr 33

Details

The random covariance matrix is built by first generating a random matrix with the function qpRndWishart from a Wishart distribution whose expected value is a matrix with unit diagonal and constant off-diagonal entries equal to rho.

Value

A random positive definite matrix that can be used as a covariance matrix for a Gaussian graphical model with graph G.

Author(s)

A. Roverato

References

Tur, I., Roverato, A. and Castelo, R. Mapping eQTL networks with mixed graphical Markov models. *Genetics*, 198(4):1377-1393, 2014.

See Also

qpRndGraph qpGetCliques qpIPF qpRndWishart rmvnorm

Examples

```
set.seed(123)
G <- qpRndGraph(p=5, d=2)
Sigma <- qpG2Sigma(G, rho=0.5)
round(solve(Sigma), digits=2)
as(G, "matrix")</pre>
```

qpGenNrr

Generalized non-rejection rate estimation

Description

Estimates generalized non-rejection rates for every pair of variables from two or more data sets.

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Usage

```
## S4 method for signature ExpressionSet
qpGenNrr(X, datasetIdx=1, qOrders=NULL, I=NULL, restrict.Q=NULL,
                           fix.Q=NULL, return.all=FALSE, nTests=100, alpha=0.05,
                       pairup.i=NULL, pairup.j=NULL, verbose=TRUE, identicalQs=TRUE,
                         exact.test=TRUE, use=c("complete.obs", "em"), tol=0.01,
                          R.code.only=FALSE, clusterSize=1, estimateTime=FALSE,
                                   nAdj2estimateTime=10)
## S4 method for signature data.frame
qpGenNrr(X,\ datasetIdx=1,\ qOrders=NULL,\ I=NULL,\ restrict.Q=NULL,
                           fix.Q=NULL, return.all=FALSE, nTests=100, alpha=0.05,
                      pairup.i=NULL, pairup.j=NULL, long.dim.are.variables=TRUE,
                               verbose=TRUE, identicalQs=TRUE, exact.test=TRUE,
                       use=c("complete.obs", "em"), tol=0.01, R.code.only=FALSE,
                        clusterSize=1, estimateTime=FALSE, nAdj2estimateTime=10)
## S4 method for signature matrix
qpGenNrr(X, datasetIdx=1, qOrders=NULL, I=NULL, restrict.Q=NULL,
                          fix.Q=NULL, return.all=FALSE, nTests=100, alpha=0.05,
                     pairup.i=NULL, pairup.j=NULL, long.dim.are.variables=TRUE,
                            verbose=TRUE, identicalQs=TRUE, exact.test=TRUE,
                       use=c("complete.obs", "em"), tol=0.01, R.code.only=FALSE,
                       clusterSize=1, estimateTime=FALSE, nAdj2estimateTime=10)
```

Arguments

X	data set from where to estimate the average non-rejection rates. It can be an ExpressionSet object, a data frame or a matrix.
datasetIdx	either a single number, or a character string, indicating the column in the phenotypic data of the ExpressionSet object, or in the input matrix or data frame, containing the indexes to the data sets. Alternatively, it can be a vector of these indexes with as many positions as samples.
q0rders	either a NULL value (default) indicating that a default guess on the q-order will be employed for each data set or a vector of particular orders with one for each data set. The default guess corresponds to the floor of the median value among the valid q orders of the data set.
I	indexes or names of the variables in X that are discrete. When X is an ExpressionSet then I may contain only names of the phenotypic variables in X. See details below regarding this argument.
restrict.Q	indexes or names of the variables in X that restrict the sample space of conditioning subsets Q.
fix.Q	indexes or names of the variables in X that should be fixed within every conditioning conditioning subsets Q.
return.all	logical; if TRUE all intervining non-rejection rates will be return in a matrix per dataset within a list; FALSE (default) if only generalized non-rejection rates should be returned.
nTests	number of tests to perform for each pair for variables.

qpGenNrr 35

alpha significance level of each test.

pairup.i subset of vertices to pair up with subset pairup.j subset of vertices to pair up with subset pairup.i

long.dim.are.variables

logical; if TRUE it is assumed that when the data is a data frame or a matrix, the longer dimension is the one defining the random variables; if FALSE, then random variables are assumed to be at the columns of the data frame or matrix.

verbose show progress on the calculations.

identicalQs use identical conditioning subsets for every pair of vertices (default), otherwise

sample a new collection of nTests subsets for each pair of vertices.

exact.test logical; if FALSE an asymptotic conditional independence test is employed with

mixed (i.e., continuous and discrete) data; if TRUE (default) then an exact condi-

tional independence test with mixed data is employed.

use a character string defining the way in which calculations are done in the presence

of missing values. It can be either "complete.obs" (default) or "em".

tol maximum tolerance controlling the convergence of the EM algorithm employed

when the argument use="em".

R.code.only logical; if FALSE then the faster C implementation is used (default); if TRUE

then only R code is executed.

clusterSize size of the cluster of processors to employ if we wish to speed-up the calcula-

tions by performing them in parallel. A value of 1 (default) implies a single-processor execution. The use of a cluster of processors requires having previ-

ously loaded the packages snow and rlecuyer.

estimateTime logical; if TRUE then the time for carrying out the calculations with the given pa-

rameters is estimated by calculating for a limited number of adjacencies, specified by nAdj2estimateTime, and extrapolating the elapsed time; if FALSE (de-

fault) calculations are performed normally till they finish.

nAdj2estimateTime

number of adjacencies to employ when estimating the time of calculations (estimateTime=TRUE).

By default this has a default value of 10 adjacencies and larger values should provide more accurate estimates. This might be relevant when using a cluster

facility.

Details

Note that when specifying a vector of particular orders q, these values should be in the range 1 to $\min(p,n-3)$, where p is the number of variables and n the number of observations for the corresponding data set. The computational cost increases linearly within each q value and quadratically in p. When setting identicalQs to FALSE the computational cost may increase between 2 times and one order of magnitude (depending on p and q) while asymptotically the estimation of the non-rejection rate converges to the same value.

When I is set different to NULL then mixed graphical model theory is employed and, concretely, it is assumed that the data comes from an homogeneous conditional Gaussian distribution. In this setting further restrictions to the maximum value of q apply, concretely, it cannot be smaller than p plus the number of levels of the discrete variables involved in the marginal distributions employed

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by the algorithm. By default, with exact.test=TRUE, an exact test for conditional independence is employed, otherwise an asymptotic one will be used. Full details on these features can be found in Tur, Roverato and Castelo (2014).

Value

A list containing the following two or more entries: a first one with name genNrr with a dspMatrix-class symmetric matrix of estimated generalized non-rejection rates with the diagonal set to NA values. When using the arguments pairup.i and pairup.j, those cells outside the constraint pairs will get also a NA value; a second one with name qOrders with the q-orders employed in the calculation for each data set; if return.all=TRUE then there will be one additional entry for each data set containing the matrix of the non-rejection rates estimated from that data set with the corresponding q-order, using the indexing value of the data set as entry name.

Note, however, that when estimateTime=TRUE, then instead of the list with matrices of estimated (generalized) non-rejection rates, a vector specifying the estimated number of days, hours, minutes and seconds for completion of the calculations is returned.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. Reverse engineering molecular regulatory networks from microarray data with qp-graphs. *J. Comp. Biol.*, 16(2):213-227, 2009.

Tur, I., Roverato, A. and Castelo, R. Mapping eQTL networks with mixed graphical models. *Submitted*, http://arxiv.org/abs/1402.4547, 2014.

See Also

qpNrr qpAvgNrr qpEdgeNrr qpHist qpGraphDensity qpClique

Examples

```
nVar <- 50 ## number of variables
maxCon <- 5 ## maximum connectivity per variable
nObs <- 30 ## number of observations to simulate

set.seed(123)

## simulate two independent Gaussian graphical models determined
## by two undirected d-regular graphs
model1 <- rUGgmm(dRegularGraphParam(p=nVar, d=maxCon), rho=0.5)
model2 <- rUGgmm(dRegularGraphParam(p=nVar, d=maxCon), rho=0.5)

## simulate two independent data sets from the previous graphical models
X1 <- rmvnorm(nObs, model1)
dim(X1)
X2 <- rmvnorm(nObs, model2)
dim(X2)</pre>
```

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```
## estimate generalized non-rejection rates from the joint data
nrr.estimates <- qpGenNrr(rbind(X1, X2),</pre>
                          datasetIdx=rep(1:2, each=n0bs),
                          q0rders=c("1"=5, "2"=5),
                          long.dim.are.variables=FALSE, verbose=FALSE)
## create adjacency matrices from the undirected graphs
## determining the two Gaussian graphical models
A1 <- as(model1$g, "matrix") == 1
A2 <- as(model2$g, "matrix") == 1
## distribution of generalized non-rejection rates for the common present edges
summary(nrr.estimates$genNrr[upper.tri(nrr.estimates$genNrr) & A1 & A2])
## distribution of generalized non-rejection rates for the present edges specific to A1
summary(nrr.estimates$genNrr[upper.tri(nrr.estimates$genNrr) & A1 & !A2])
## distribution of generalized non-rejection rates for the present edges specific to A2
summary(nrr.estimates$genNrr[upper.tri(nrr.estimates$genNrr) & !A1 & A2])
## distribution of generalized non-rejection rates for the common missing edges
summary(nrr.estimates$genNrr[upper.tri(nrr.estimates$genNrr) & !A1 & !A2])
## compare with the average non-rejection rate on the pooled data set
avgnrr.estimates <- qpNrr(rbind(X1, X2), q=5, long.dim.are.variables=FALSE, verbose=FALSE)
## distribution of average non-rejection rates for the common present edges
summary(avgnrr.estimates[upper.tri(avgnrr.estimates) & A1 & A2])
## distribution of average non-rejection rates for the present edges specific to A1
summary(avgnrr.estimates[upper.tri(avgnrr.estimates) & A1 & !A2])
## distribution of average non-rejection rates for the present edges specific to A2
summary(avgnrr.estimates[upper.tri(avgnrr.estimates) & !A1 & A2])
## distribution of average non-rejection rates for the common missing edges
summary(avgnrr.estimates[upper.tri(avgnrr.estimates) & !A1 & !A2])
```

qpGetCliques

Clique list

Description

Finds the set of (maximal) cliques of a given undirected graph.

Usage

```
qpGetCliques(g, clqspervtx=FALSE, verbose=TRUE)
```

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Arguments

g either a graphNEL object or an adjacency matrix of the given undirected graph.

clqspervtx logical; if TRUE then the resulting list returned by the function includes addi-

tionally p entries at the beginning (p=number of variables) each corresponding to a vertex in the graph and containing the indices of the cliques where that vertex belongs to; if FALSE these additional entries are not included (default).

verbose show progress on calculations.

Details

To find the list of all (maximal) cliques in an undirected graph is an NP-hard problem which means that its computational cost is bounded by an exponential running time (Garey and Johnson, 1979). For this reason, this is an extremely time and memory consuming computation for large dense graphs. The current implementation uses C code from the GNU GPL Cliquer library by Niskanen and Ostergard (2003).

Value

A list of maximal cliques. When clqspervtx=TRUE the first p entries (p=number of variables) contain, each of them, the indices of the cliques where that particular vertex belongs to.

Author(s)

R. Castelo

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n. *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Garey, M.R. and Johnson D.S. *Computers and intractability: a guide to the theory of NP-completeness*. W.H. Freeman, San Francisco, 1979.

Niskanen, S. Ostergard, P. Cliquer User's Guide, Version 1.0. Communications Laboratory, Helsinki University of Technology, Espoo, Finland, Tech. Rep. T48, 2003. (http://users.tkk.fi/~pat/cliquer.html)

See Also

```
qpCliqueNumber qpIPF
```

```
require(graph)
set.seed(123)
nVar <- 50
g1 <- randomEGraph(V=as.character(1:nVar), p=0.3)
clqs1 <- qpGetCliques(g1, verbose=FALSE)
length(clqs1)</pre>
```

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```
summary(sapply(clqs1, length))
g2 <- randomEGraph(V=as.character(1:nVar), p=0.7)
clqs2 <- qpGetCliques(g2, verbose=FALSE)
length(clqs2)
clqs2 <- qpGetCliques(g2, verbose=FALSE)
summary(sapply(clqs2, length))</pre>
```

qpGraph-class

The qp-graph

Description

The "qpGraph" class is the class to store and manipulate q-order (partial) correlation graphs, or qp-graphs for short. See Castelo and Roverato (2006, 2009) for a mathematical and statistical definition of a qp-graph.

In earlier versions 1.x of the qpgraph package there was a function called qpGraph() to obtain a qp-graph from a matrix of non-rejection rates. This function, as it was written, has been deprecated and replaced by this class and corresponding constructor methods of the same name. The main difference with respect to earlier 1.x versions is that the argument threshold is now called epsilon, the argument return. type has been removed and the current version returns an object of this class qpGraph described in this manual page.

Arguments

epsilon	maximum cutoff value met by the edges present in the qp-graph.
topPairs	number of edges from the top of the ranking, defined by the non-rejection rates in nrrMatrix, to use to form the resulting qp-graph. This parameter is incompatible with a value different from NULL in epsilon.
pairup.i	subset of vertices to pair up with subset pairup.j
pairup.j	subset of vertices to pair up with subset pairup.i
q	q-order employed to derive the input matrix of non-rejection rates nrrMatrix.
n	when the input matrix of non-rejection rates nrrMatrix has been estimated from data, this is the number of observations in the data set.

Objects from the Class

Objects can be created by calls of the form qpGraph(nrrMatrix, ...) corresponding to constructor methods that take as input a matrix of non-rejection rates, calculated with qpNrr.

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Slots

- p: number of vertices, in one-to-one correspondence with random variables.
- q: order of the qp-graph, always smaller than p-2.
- n: when the qp-graph has been estimated from data, this is the number of observations in the data set, which must be larger than q+2.
- epsilon: maximum cutoff value of the non-rejection rate met by the edges that are present in the qp-graph.
- g: undirected graph structure of the qp-graph stored as a graphBAM-class object.

Methods

qpGraph(nrrMatrix, ...) Constructor method where nrrMatrix is a matrix of non-rejection
rates.

show(object) Method to display some bits of information about the qp-graph stored in the input argument object.

Author(s)

R. Castelo

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n, *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Castelo, R. and Roverato, A. Reverse engineering molecular regulatory networks from microarray data with qp-graphs. *J. Comp. Biol.*, 16(2):213-227, 2009.

See Also

qpNrr

qpGraphDensity

Densities of resulting qp-graphs

Description

Calculates and plots the graph density as function of the non-rejection rate.

Usage

qpGraphDensity 41

Arguments

nrrMatrix matrix of non-rejection rates.

threshold.lim range of threshold values on the non-rejection rate.

breaks either a number of threshold bins or a vector of threshold breakpoints.

plot logical; if TRUE makes a plot of the result; if FALSE it does not.

qpGraphDensityOutput

output from a previous call to qpGraphDensity. This allows one to plot the result changing some of the plotting parameters without having to do the calcu-

lation again.

density.digits number of digits in the reported graph densities.

titlegd main title to be shown in the plot.

Details

The estimate of the sparseness of the resulting qp-graphs is calculated as one minus the area enclosed under the curve of graph densities.

Value

A list with the graph density as function of threshold and an estimate of the sparseness of the resulting qp-graphs across the thresholds.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n, *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

See Also

```
qpNrr qpAvgNrr qpEdgeNrr qpClique
```

```
require(mvtnorm)

nVar <- 50  ## number of variables
maxCon <- 5  ## maximum connectivity per variable
nObs <- 30  ## number of observations to simulate
set.seed(123)

A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))</pre>
```

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```
## the higher the q the sparser the qp-graph
nrr.estimates <- qpNrr(X, q=1, verbose=FALSE)
qpGraphDensity(nrr.estimates, plot=FALSE)$sparseness
nrr.estimates <- qpNrr(X, q=5, verbose=FALSE)
qpGraphDensity(nrr.estimates, plot=FALSE)$sparseness</pre>
```

qpHist

Histograms of non-rejection rates

Description

Plots the distribution of non-rejection rates.

Usage

Arguments

nrrMatrix matrix of non-rejection rates.

A adjacency matrix of an undirected graph whose present and missing edges will

be employed to show separately the distribution of non-rejection rates.

titlehist main title of the histogram(s).

freq logical; if TRUE, the histograms show frequencies (counts) of occurrence of

the different non-rejection rate values; if FALSE, then probability densities are

plotted

Details

This function plots histograms using the R-function hist and therefore the way they are displayed follows that of this R-function.

Value

None

Author(s)

R. Castelo and A. Roverato

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References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n, *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

See Also

```
qpNrr qpAvgNrr qpEdgeNrr qpGraphDensity qpClique
```

Examples

```
require(mvtnorm)

nVar <- 50  ## number of variables
maxCon <- 5  ## maximum connectivity per variable
nObs <- 30  ## number of observations to simulate

A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))

nrr.estimates <- qpNrr(X, q=5, verbose=FALSE)

qpHist(nrr.estimates, A)</pre>
```

qpHTF

Hastie Tibshirani Friedman iterative regression algorithm

Description

Performs maximum likelihood estimation of a covariance matrix given the independence constraints from an input undirected graph.

Usage

```
qpHTF(S, g, tol = 0.001, verbose = FALSE, R.code.only = FALSE)
```

Arguments

S input matrix, in the context of this package, the	he sample covariance matrix.
---	------------------------------

g input undirected graph.

tol tolerance under which the iterative algorithm stops.

verbose show progress on calculations.

R.code.only logical; if FALSE then the faster C implementation is used (default); if TRUE

then only R code is executed.

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Details

This is an alternative to the Iterative Proportional Fitting (IPF) algorithm (see, Whittaker, 1990, pp. 182-185 and qpIPF) which also adjusts the input matrix to the independence constraints in the input undirected graph. However, differently to the IPF, it works by going through each of the vertices fitting the marginal distribution over the corresponding vertex boundary. It stops when the adjusted matrix at the current iteration differs from the matrix at the previous iteration in less or equal than a given tolerance value. This algorithm is described by Hastie, Tibshirani and Friedman (2009, pg. 634), hence we name it here HTF, and it has the advantage over the IPF that it does not require the list of maximal cliques of the graph which may be exponentially large. In contrast, it requires that the maximum boundary size of the graph is below the number of samples where the input sample covariance matrix S was estimated. For the purpose of exploring qp-graphs that meet such a requirement, one can use the function qpBoundary.

Value

The input matrix adjusted to the constraints imposed by the input undirected graph, i.e., a maximum likelihood estimate of the sample covariance matrix that includes the independence constraints encoded in the undirected graph.

Note

Thanks to Giovanni Marchetti for bringing us our attention to this algorithm and sharing an early version of its implementation on the R package ggm.

Author(s)

R. Castelo

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n. *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Hastie, T., Tibshirani, R. and Friedman, J.H. The Elements of Statistical Learning, Springer, 2009.

Tur, I., Roverato, A. and Castelo, R. Mapping eQTL networks with mixed graphical Markov models. *Genetics*, 198(4):1377-1393, 2014.

Whittaker, J. Graphical Models in Applied Multivariate Statistics. Wiley, 1990.

See Also

```
qpBoundary qpIPF qpPAC
```

```
require(graph)
require(mvtnorm)

nVar <- 50  ## number of variables
nObs <- 100  ## number of observations to simulate</pre>
```

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```
set.seed(123)
g <- randomEGraph(as.character(1:nVar), p=0.15)
Sigma <- qpG2Sigma(g, rho=0.5)
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))
## MLE of the sample covariance matrix
S <- cov(X)
## more efficient MLE of the sample covariance matrix using HTF
S_htf <- qpHTF(S, g)
## get the adjacency matrix and put the diagonal to one
A <- as(g, "matrix")
diag(A) <- 1
## entries in S and S_htf for present edges in g should coincide
max(abs(S_htf[A==1] - S[A==1]))
## entries in the inverse of S_htf for missing edges in g should be zero
max(solve(S_htf)[A==0])</pre>
```

qpImportNrr

Import non-rejection rates

Description

Imports non-rejection rates from an external flat file.

Usage

```
qpImportNrr(filename, nTests)
```

Arguments

filename name of the flat file with the data on the non-rejection rates.

nTests number of tests performed in the estimation of these non-rejection rates.

Details

This function expects a flat file with three tab-separated columns corresponding to, respectively, 0-based index of one of the variables, 0-based index of the other variable, number of non-rejected tests for the pair of variables of that row in the text file. An example of a few lines of that file would be:

6	3	95
6	4	98
6	5	23

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7	0	94
7	1	94

After reading the file the function builds a matrix of non-rejection rates by dividing the number of non-rejected tests by nTests. Note that if the flat file to be imported would eventually have directly the rates instead of the number of tests, these can be also imported by setting nTests=1.

This function is thought to be used to read files obtained from the standalone parallel version of qpNrr which can be downloaded from http://functionalgenomics.upf.edu/qp.

Value

A symmetric matrix of non-rejection rates with the diagonal set to the NA value.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n, *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

See Also

qpNrr

qpIPF	Iterative proportional fitting algorithm

Description

Performs maximum likelihood estimation of a covariance matrix given the independence constraints from an input list of (maximal) cliques.

Usage

```
qpIPF(vv, clqlst, tol = 0.001, verbose = FALSE, R.code.only = FALSE)
```

Arguments

VV	input matrix, in the context of this package, the sample covariance matrix.
clqlst	list of maximal cliques obtained from an undirected graph by using the function qpGetCliques.
tol	tolerance under which the iterative algorithm stops.
verbose	show progress on calculations.
R.code.only	logical; if FALSE then the faster C implementation is used (default); if TRUE then only R code is executed.

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Details

The Iterative proportional fitting algorithm (see, Whittaker, 1990, pp. 182-185) adjusts the input matrix to the independence constraints in the undirected graph from where the input list of cliques belongs to, by going through each of the cliques fitting the marginal distribution over the clique for the fixed conditional distribution of the clique. It stops when the adjusted matrix at the current iteration differs from the matrix at the previous iteration in less or equal than a given tolerance value.

Value

The input matrix adjusted to the constraints imposed by the list of cliques, i.e., a maximum likelihood estimate of the sample covariance matrix that includes the independence constraints encoded in the undirected graph formed by the given list of cliques.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n. *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Tur, I., Roverato, A. and Castelo, R. Mapping eQTL networks with mixed graphical Markov models. *Genetics*, 198(4):1377-1393, 2014.

Whittaker, J. Graphical models in applied multivariate statistics. Wiley, 1990.

See Also

```
qpGetCliques qpPAC
```

```
require(graph)
require(mvtnorm)

nVar <- 50  ## number of variables
nObs <- 100  ## number of observations to simulate

set.seed(123)

g <- randomEGraph(as.character(1:nVar), p=0.15)

Sigma <- qpG2Sigma(g, rho=0.5)
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))

## MLE of the sample covariance matrix
S <- cov(X)

## more efficient MLE of the sample covariance matrix using IPF
clqs <- qpGetCliques(g, verbose=FALSE)</pre>
```

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```
S_ipf <- qpIPF(S, clqs)

## get the adjacency matrix and put the diagonal to one
A <- as(g, "matrix")
diag(A) <- 1

## entries in S and S_ipf for present edges in g should coincide
max(abs(S_ipf[A==1] - S[A==1]))

## entries in the inverse of S_ipf for missing edges in g should be zero
max(solve(S_ipf)[A==0])</pre>
```

qpK2ParCor

Partial correlation coefficients

Description

Obtains partial correlation coefficients from a given concentration matrix.

Usage

```
qpK2ParCor(K)
```

Arguments

Κ

positive definite matrix, typically a concentration matrix.

Details

This function applies cov2cor to the given concentration matrix and then changes the sign of the off-diagonal entries in order to obtain a partial correlation matrix.

Value

A partial correlation matrix.

Author(s)

R. Castelo and A. Roverato

References

Lauritzen, S.L. Graphical models. Oxford University Press, 1996.

See Also

```
qpG2Sigma
```

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Examples

```
require(graph)
n.var <- 5 # number of variables
set.seed(123)
g <- randomEGraph(as.character(1:n.var), p=0.15)
Sigma <- qpG2Sigma(g, rho=0.5)
K <- solve(Sigma)
round(qpK2ParCor(K), digits=2)
as(g, "matrix")</pre>
```

qpNrr

Non-rejection rate estimation

Description

Estimates non-rejection rates for every pair of variables.

Usage

```
## S4 method for signature ExpressionSet
qpNrr(X, q=1, restrict.Q=NULL, fix.Q=NULL, nTests=100,
                                alpha=0.05, pairup.i=NULL, pairup.j=NULL,
                               verbose=TRUE, identicalQs=TRUE, exact.test=TRUE,
                       use=c("complete.obs", "em"), tol=0.01, R.code.only=FALSE,
                       clusterSize=1, estimateTime=FALSE, nAdj2estimateTime=10)
## S4 method for signature cross
qpNrr(X, q=1, restrict.Q=NULL, fix.Q=NULL, nTests=100,
                        alpha=0.05, pairup.i=NULL, pairup.j=NULL, verbose=TRUE,
                 identicalQs=TRUE, exact.test=TRUE, use=c("complete.obs", "em"),
                 tol=0.01, R.code.only=FALSE, clusterSize=1, estimateTime=FALSE,
                         nAdj2estimateTime=10)
## S4 method for signature data.frame
qpNrr(X, q=1, I=NULL, restrict.Q=NULL, fix.Q=NULL, nTests=100,
                             alpha=0.05, pairup.i=NULL, pairup.j=NULL,
                             long.dim.are.variables=TRUE, verbose=TRUE,
                   identicalQs=TRUE, exact.test=TRUE, use=c("complete.obs", "em"),
                             tol=0.01, R.code.only=FALSE, clusterSize=1,
                             estimateTime=FALSE, nAdj2estimateTime=10)
## S4 method for signature matrix
qpNrr(X, q=1, I=NULL, restrict.Q=NULL, fix.Q=NULL, nTests=100,
                         alpha=0.05, pairup.i=NULL, pairup.j=NULL,
                   long.dim.are.variables=TRUE, verbose=TRUE, identicalQs=TRUE,
                        exact.test=TRUE, use=c("complete.obs", "em"), tol=0.01,
                         R.code.only=FALSE, clusterSize=1, estimateTime=FALSE,
                         nAdj2estimateTime=10)
```

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Arguments

X data set from where to estimate the non-rejection rates. It can be an ExpressionSet

object, a qtl/cross object, a data.frame object or a matrix object.

q partial-correlation order to be employed.

I indexes or names of the variables in X that are discrete. See details below re-

garding this argument.

restrict.Q indexes or names of the variables in X that restrict the sample space of condi-

tioning subsets Q.

fix.0 indexes or names of the variables in X that should be fixed within every condi-

tioning conditioning subsets Q.

nTests number of tests to perform for each pair for variables.

alpha significance level of each test.

pairup.i subset of vertices to pair up with subset pairup.j subset of vertices to pair up with subset pairup.i

long.dim.are.variables

logical; if TRUE it is assumed that when data are in a data frame or in a matrix, the longer dimension is the one defining the random variables (default); if FALSE, then random variables are assumed to be at the columns of the data frame or

matrix.

verbose show progress on the calculations.

identicalQs use identical conditioning subsets for every pair of vertices (default), otherwise

sample a new collection of nTests subsets for each pair of vertices.

exact.test logical; if FALSE an asymptotic conditional independence test is employed with

mixed (i.e., continuous and discrete) data; if TRUE (default) then an exact conditional independence test with mixed data is employed. See details below re-

garding this argument.

use a character string defining the way in which calculations are done in the presence

of missing values. It can be either "complete.obs" (default) or "em".

tol maximum tolerance controlling the convergence of the EM algorithm employed

when the argument use="em".

R.code.only logical; if FALSE then the faster C implementation is used (default); if TRUE then

only R code is executed.

clusterSize size of the cluster of processors to employ if we wish to speed-up the calcula-

tions by performing them in parallel. A value of 1 (default) implies a single-processor execution. The use of a cluster of processors requires having previ-

ously loaded the packages snow and rlecuyer.

estimateTime logical; if TRUE then the time for carrying out the calculations with the given pa-

rameters is estimated by calculating for a limited number of adjacencies, specified by nAdj2estimateTime, and extrapolating the elapsed time; if FALSE (de-

fault) calculations are performed normally till they finish.

nAdj2estimateTime

number of adjacencies to employ when estimating the time of calculations (estimateTime=TRUE).

By default this has a default value of 10 adjacencies and larger values should provide more accurate estimates. This might be relevant when using a cluster

facility.

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Details

Note that for pure continuous data the possible values of q should be in the range 1 to min(p, n-3), where p is the number of variables and n the number of observations. The computational cost increases linearly with q and quadratically in p. When setting identicalQs to FALSE the computational cost may increase between 2 times and one order of magnitude (depending on p and q) while asymptotically the estimation of the non-rejection rate converges to the same value. Full details on the calculation of the non-rejection rate can be found in Castelo and Roverato (2006).

When I is set different to NULL then mixed graphical model theory is employed and, concretely, it is assumed that the data comes from an homogeneous conditional Gaussian distribution. In this setting further restrictions to the maximum value of q apply, concretely, it cannot be smaller than p plus the number of levels of the discrete variables involved in the marginal distributions employed by the algorithm. By default, with exact.test=TRUE, an exact test for conditional independence is employed, otherwise an asymptotic one will be used. Full details on these features can be found in Tur, Roverato and Castelo (2014).

The argument I specifying what variables are discrete actually applies only when X is a matrix object since in the other cases data types are specified for each data columns or slot.

In the case that X is a qtl/cross object, the default NULL values in arguments pairup.i and pairup.j actually imply pairing all markers and phenotypes with numerical phenotypes only (including integer phenotypes). Likewise, the default argument restrict.Q=NULL implies setting restrict.Q to all numeric phenotypes. Setting these arguments to values other than NULL allows the user to use those particular values being set.

Value

A dspMatrix-class symmetric matrix of estimated non-rejection rates with the diagonal set to NA values. If arguments pairup.i and pairup.j are employed, those cells outside the constrained pairs will get also a NA value.

Note, however, that when estimateTime=TRUE, then instead of the matrix of estimated non-rejection rates, a vector specifying the estimated number of days, hours, minutes and seconds for completion of the calculations is returned.

Author(s)

R. Castelo, A. Roverato and I. Tur

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n, *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Tur, I., Roverato, A. and Castelo, R. Mapping eQTL networks with mixed graphical Markov models. *Genetics*, 198(4):1377-1393, 2014.

See Also

qpAvgNrr qpEdgeNrr qpHist qpGraphDensity qpClique

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```
nVar <- 50 ## number of variables
maxCon <- 3 ## maximum connectivity per variable</pre>
nObs <- 30 ## number of observations to simulate
set.seed(123)
## simulate an undirected Gaussian graphical model
## determined by some random undirected d-regular graph
model <- rUGgmm(dRegularGraphParam(p=nVar, d=maxCon), rho=0.5)</pre>
## simulate data from this model
X <- rmvnorm(nObs, model)</pre>
dim(X)
## estimate non-rejection rates with q=3
nrr.estimates <- qpNrr(X, q=3, verbose=FALSE)</pre>
## create an adjacency matrix of the undirected graph
## determining the undirected Gaussian graphical model
A <- as(model$g, "matrix") == 1
## distribution of non-rejection rates for the present edges
summary(nrr.estimates[upper.tri(nrr.estimates) & A])
## distribution of non-rejection rates for the missing edges
summary(nrr.estimates[upper.tri(nrr.estimates) & !A])
## Not run:
## using R code only this would take much more time
qpNrr(X, q=3, R.code.only=TRUE, estimateTime=TRUE)
## only for moderate and large numbers of variables the
## use of a cluster of processors speeds up the calculations
library(snow)
library(rlecuyer)
nVar <- 500
maxCon <- 3
model <- rUGgmm(dRegularGraphParam(p=nVar, d=maxCon), rho=0.5)</pre>
X <- rmvnorm(nObs, model)</pre>
system.time(nrr.estimates <- qpNrr(X, q=10, verbose=TRUE))</pre>
system.time(nrr.estimates <- qpNrr(X, q=10, verbose=TRUE, clusterSize=4))</pre>
## End(Not run)
```

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Description

Estimates partial correlation coefficients (PACs) for a Gaussian graphical model with undirected graph G and their corresponding P-values for the hypothesis of zero partial correlations.

Usage

Arguments

Χ	data set from where to estimate the partial correlation coefficients. It can be an
	ExpressionSet object, a data frame or a matrix.

g either a graphNEL object or an adjacency matrix of the given undirected graph.

return.K logical; if TRUE this function also returns the concentration matrix K; if FALSE

it does not return it (default).

long.dim.are.variables

logical; if TRUE it is assumed that when X is a data frame or a matrix, the longer dimension is the one defining the random variables (default); if FALSE, then random variables are assumed to be at the columns of the data frame or matrix.

tol maximum tolerance in the application of the IPF algorithm.

matrix.completion

algorithm to employ in the matrix completion operations employed to construct

a positive definite matrix with the zero pattern specified in g

verbose show progress on the calculations.

R.code.only logical; if FALSE then the faster C implementation is used (default); if TRUE

then only R code is executed.

Details

In the context of maximum likelihood estimation (MLE) of PACs it is a necessary condition for the existence of MLEs that the sample size n is larger than the clique number w(G) of the graph G.

The PAC estimation is done by first obtaining a MLE of the covariance matrix using the qpIPF function and the P-values are calculated based on the estimation of the standard errors (see Roverato and Whittaker, 1996).

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Value

A list with two matrices, one with the estimates of the PACs and the other with their P-values.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n. *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Castelo, R. and Roverato, A. Reverse engineering molecular regulatory networks from microarray data with qp-graphs. *J. Comp. Biol.*, 16(2):213-227, 2009.

Roverato, A. and Whittaker, J. Standard errors for the parameters of graphical Gaussian models. *Stat. Comput.*, 6:297-302, 1996.

See Also

qpGraph qpCliqueNumber qpClique qpGetCliques qpIPF

```
require(mvtnorm)
nVar <- 50 ## number of variables
maxCon <- 5 ## maximum connectivity per variable</pre>
nObs <- 30 ## number of observations to simulate
set.seed(123)
A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)</pre>
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))</pre>
nrr.estimates <- qpNrr(X, verbose=FALSE)</pre>
g <- qpGraph(nrr.estimates, threshold=0.5)</pre>
pac.estimates <- qpPAC(X, g=g, verbose=FALSE)</pre>
## distribution absolute values of the estimated
## partial correlation coefficients of the present edges
summary(abs(pac.estimates$R[upper.tri(pac.estimates$R) & A]))
## distribution absolute values of the estimated
## partial correlation coefficients of the missing edges
summary(abs(pac.estimates$R[upper.tri(pac.estimates$R) & !A]))
```

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qpPCC

Estimation of Pearson correlation coefficients

Description

Estimates Pearson correlation coefficients (PCCs) and their corresponding P-values between all pairs of variables from an input data set.

Usage

```
## S4 method for signature ExpressionSet
qpPCC(X)
## S4 method for signature data.frame
qpPCC(X, long.dim.are.variables=TRUE)
## S4 method for signature matrix
qpPCC(X, long.dim.are.variables=TRUE)
```

Arguments

Χ

data set from where to estimate the Pearson correlation coefficients. It can be an ExpressionSet object, a data frame or a matrix.

```
long.dim.are.variables
```

logical; if TRUE it is assumed that when X is a data frame or a matrix, the longer dimension is the one defining the random variables (default); if FALSE, then random variables are assumed to be at the columns of the data frame or matrix.

Details

The calculations made by this function are the same as the ones made for a single pair of variables by the function cor.test but for all the pairs of variables in the data set.

Value

A list with two matrices, one with the estimates of the PCCs and the other with their P-values.

Author(s)

R. Castelo and A. Roverato

See Also

qpPAC

56 qpPlotMap

Examples

```
require(graph)
require(mvtnorm)

nVar <- 50 ## number of variables
nObs <- 10 ## number of observations to simulate

set.seed(123)

g <- randomEGraph(as.character(1:nVar), p=0.15)

Sigma <- qpG2Sigma(g, rho=0.5)

X <- rmvnorm(nObs, sigma=as.matrix(Sigma))

pcc.estimates <- qpPCC(X)

## get the corresponding boolean adjacency matrix
A <- as(g, "matrix") == 1

## Pearson correlation coefficients of the present edges
summary(abs(pcc.estimates$R[upper.tri(pcc.estimates$R) & A]))

## Pearson correlation coefficients of the missing edges
summary(abs(pcc.estimates$R[upper.tri(pcc.estimates$R) & !A]))</pre>
```

qpPlotMap

Plots a map of associated pairs

Description

Plots a map of associated pairs defined by adjusted p-values

Usage

Arguments

p.valueMatrix squared symmetric matrix with raw p-values for all pairs.

markerPos two-column matrix containing chromosome and position of each genetic marker.

genePos two-column matrix containing chromosome and position of each gene.

chrLen named vector with chromosome lengths. Vector names should correspond to

chromosome names, which are displayed in the axes of the plot. This vector should be ordered following the same convention for chromosomes in arguments

markerPos and genePos.

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p.value	adjusted p-value cutoff.
adjust.method	method employed to adjust the raw p-values. It is passed in a call to p.adjust() in its method argument.
xlab	label for the x-axis.
ylab	label for the y-axis.
main	main title of the plot, set to the empty string by default.
	further arguments passed to the plot() function.

Details

This function plots a map of present associations, typically between genetic markers and gene expression profiles (i.e., eQTL associations), according to the chromosomal locations of both the genetic markers and the genes. The input argument p.valueMatrix should contain the raw p-values of these associations. Present associations are selected by a cutoff given in the p.value argument applied to the adjusted p-values.

The input raw p-values can be obtained with the function qpAllCItests.

Value

The selected present associations are invisibly returned.

Author(s)

R. Castelo

See Also

```
qpAllCItests
```

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```
## make matrix symmetric
rndpvalues <- rndpvalues + t(rndpvalues)
stopifnot(isSymmetric(rndpvalues))
rndpvalues[1:m, 1:m] <- rndpvalues[(m+1):p,(m+1):p] <- NA

## create chromosomal map
chrlen <- c("chr1"=1000)
posmarkers <- matrix(c(rep(1, m), seq(1, chrlen, length.out=m)), nrow=m)
posgenes <- matrix(c(rep(1, g), seq(1, chrlen, length.out=g)), nrow=g)
rownames(posmarkers) <- paste0("m", 1:m)
rownames(posgenes) <- paste0("g", 1:g)

qpPlotMap(rndpvalues, posmarkers, posgenes, chrlen, cex=3)</pre>
```

qpPlotNetwork

Plots a graph

Description

Plots a graph using the Rgraphviz library

Usage

Arguments

g graph to plot provided as a graphNEL-class object.

vertexSubset subset of vertices that define the induced subgraph to be plotted.

boundary flag set to TRUE when we wish that the subset specified in vertexSubset also

includes the vertices connected to them; FALSE otherwise.

minimumSizeConnComp

minimum size of the connected components to be plotted.

pairup.i subset of vertices to pair up with subset pairup.j.
pairup.j subset of vertices to pair up with subset pairup.i.

highlight subset of vertices to highlight by setting the color font to red.

annotation name of an annotation package to transform gene identifiers into gene symbols

when vertices correspond to genes.

layout argument for the Rgraphviz library that plots the network. Possible values

are twopi (default), dot, neato, circo, fdp.

Details

This function acts as a wrapper for the functionality provided by the Rgraphviz package to plot graphs in R. It should help to plot networks obtained with methods from theqpgraph package.

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Value

The plotted graph is invisibly returned as a graphNEL-class object.

Author(s)

R. Castelo

See Also

```
qpGraph qpAnyGraph
```

Examples

```
## Not run:
require(Rgraphviz)

rndassociations <- qpUnifRndAssociation(10)
g <- qpAnyGraph(abs(rndassociations), threshold=0.7, remove="below")
qpPlotNetwork(g) ## this does not work at the moment and should be fixed
## End(Not run)</pre>
```

qpPrecisionRecall

Calculation of precision-recall curves

Description

Calculates the precision-recall curve (see Fawcett, 2006) for a given measure of association between all pairs of variables in a matrix.

Usage

Arguments

 ${\tt measurementsMatrix}$

matrix containing the measure of association between all pairs of variables.

refGraph a reference graph from which to calculate the precision-recall curve provided

either as an adjacency matrix, a two-column matrix of edges, a graphNEL-class

object or a graphAM-class object.

decreasing logical; if TRUE then the measurements are ordered in decreasing order; if

FALSE then in increasing order.

pairup.i subset of vertices to pair up with subset pairup.j.
pairup.j subset of vertices to pair up with subset pairup.i.
recallSteps steps of the recall on which to calculate precision.

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Details

The measurementsMatrix should be symmetric and may have also contain NA values which will not be taken into account. That is an alternative way to restricting the variable pairs with the parameters pairup.i and pairup.j.

Value

A matrix where rows correspond to recall steps and columns correspond, respetively, to the actual recall, the precision, the number of true positives at that recall rate and the threshold score that yields that recall rate.

Author(s)

R. Castelo and A. Roverato

References

Fawcett, T. An introduction to ROC analysis. Pattern Recogn. Lett., 27:861-874, 2006.

See Also

qpPRscoreThreshold qpGraph qpAvgNrr qpPCC

```
require(mvtnorm)
nVar <- 50 ## number of variables
maxCon <- 5 ## maximum connectivity per variable</pre>
nObs <- 30 ## number of observations to simulate
set.seed(123)
A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)</pre>
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))</pre>
## estimate non-rejection rates
nrr.estimates <- qpNrr(X, q=5, verbose=FALSE)</pre>
## estimate Pearson correlation coefficients
pcc.estimates <- qpPCC(X)</pre>
## calculate area under the precision-recall curve
## for both sets of estimated values of association
nrr.prerec <- qpPrecisionRecall(nrr.estimates, refGraph=A, decreasing=FALSE,</pre>
                                  recallSteps=seq(0, 1, 0.1))
f <- approxfun(nrr.prerec[, c("Recall", "Precision")])</pre>
integrate(f, 0, 1)$value
pcc.prerec <- qpPrecisionRecall(abs(pcc.estimates$R), refGraph=A,</pre>
```

qpPRscoreThreshold 61

```
recall Steps = seq(0, 1, 0.1)) \\ f \leftarrow approxfun(pcc.prerec[, c("Recall", "Precision")]) \\ integrate(f, 0, 1) \\ state = (f, 0,
```

qpPRscoreThreshold

Calculation of scores thresholds attaining nominal precision or recall levels

Description

Calculates the score threshold at a given precision or recall level from a given precision-recall curve.

Usage

```
qpPRscoreThreshold(preRecFun, level, recall.level=TRUE, max.score=9999999)
```

Arguments

preRecFun precision-recall function (output from qpPrecisionRecall).

level recall or precision level.

recall.level logical; if TRUE then it is assumed that the value given in the level parameter

corresponds to a desired level of recall; if FALSE then it is assumed a desired

level of precision.

max. score maximum score given by the method that produced the precision-recall function

to an association.

Value

The score threshold at which a given level of precision or recall is attained by the given precision-recall function. For levels that do not form part of the given function their score is calculated by linear interpolation and for this reason is important to carefully specify a proper value for the max.score parameter.

Author(s)

R. Castelo and A. Roverato

References

Fawcett, T. An introduction to ROC analysis. Pattern Recogn. Lett., 27:861-874, 2006.

See Also

```
qpPrecisionRecall qpGraph
```

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Examples

```
require(mvtnorm)

nVar <- 50  ## number of variables
maxCon <- 5  ## maximum connectivity per variable
nObs <- 30  ## number of observations to simulate

set.seed(123)

A <- qpRndGraph(p=nVar, d=maxCon)
Sigma <- qpG2Sigma(A, rho=0.5)
X <- rmvnorm(nObs, sigma=as.matrix(Sigma))

nrr.estimates <- qpNrr(X, q=1, verbose=FALSE)

nrr.prerec <- qpPrecisionRecall(nrr.estimates, A, decreasing=FALSE, recallSteps=seq(0, 1, by=0.1))

qpPRscoreThreshold(nrr.prerec, level=0.5, recall.level=FALSE, max.score=0)

qpPRscoreThreshold(nrr.prerec, level=0.5, recall.level=FALSE, max.score=0)</pre>
```

qpRndGraph

Undirected random d-regular graphs

Description

Samples an undirected d-regular graph approximately uniformly at random.

Usage

Arguments

p number of vertices.d degree of every vertex.

labels vertex labels.

exclude vector of vertices inducing edges that should be excluded from the sampled d-

regular graph.

verbose show progress on the calculations.

return. type class of object to be returned by the function

R.code.only logical; if FALSE then the faster C implementation is used (default); if TRUE then

only R code is executed.

qpRndGraph 63

Details

This function implements the algorithm from Steger and Wormald (1999) for sampling undirected d-regular graphs from a probability distribution of all d-regular graphs on p vertices which is approximately uniform. More concretely, for all vertex degree values d that grow as a small power of p, all d-regular graphs on p vertices will have in the limit the same probability as p grows large. Steger and Wormald (1999, pg. 396) believe that for d » sqrt(p) the resulting probability distribution will no longer be approximately uniform.

This function is provided in order to generate a random undirected graph as input to the function qpG2Sigma which samples a random covariance matrix whose inverse (aka, precision matrix) has zeroes on those cells corresponding to the missing edges in the input graph. d-regular graphs are useful for working with synthetic graphical models for two reasons: one is that d-regular graph density is a linear function of d and the other is that the minimum connectivity degree of two disconnected vertices is an upper bound of their outer connectivity (see Castelo and Roverato, 2006, pg. 2646).

Value

The adjacency matrix of the resulting graph.

Author(s)

R. Castelo and A. Roverato

References

Castelo, R. and Roverato, A. A robust procedure for Gaussian graphical model search from microarray data with p larger than n, *J. Mach. Learn. Res.*, 7:2621-2650, 2006.

Steger, A. and Wormald, N.C. Generating random regular graphs quickly, *Combinatorics, Probab. and Comput.*, 8:377-396.

See Also

```
qpG2Sigma
```

```
set.seed(123)
A <- qpRndGraph(p=50, d=3)
summary(apply(A, 1, sum))</pre>
```

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qpRndWishart Random Wishart distribution
--

Description

Random generation for the (n.var * n.var) Wishart distribution (see Press, 1972) with matrix parameter A=diag(delta)%*%P%*%diag(delta) and degrees of freedom df.

Usage

```
qpRndWishart(delta=1, P=0, df=NULL, n.var=NULL)
```

Arguments

delta	a numeric vector of n.var positive values. If a scalar is provided then this is extended to form a vector.
Р	a (n.var * n.var) positive definite matrix with unit diagonal. If a scalar is provided then this number is used as constant off-diagonal entry for P.
df	degrees of freedom.
n.var	dimension of the Wishart matrix. It is required only when both delata and P are scalar.

Details

The degrees of freedom are df > n.var-1 and the expected value of the distribution is equal to df * A. The random generator is based on the algorithm of Odell and Feiveson (1966).

Value

A list of two n.var * n.var matrices rW and meanW where rW is a random value from the Wishart and meanW is the expected value of the distribution.

Author(s)

A. Roverato

References

Odell, P.L. and Feiveson, A.G. A numerical procedure to generate a sample covariance matrix. *J. Am. Statist. Assoc.* 61, 199-203, 1966.

Press, S.J. Applied Multivariate Analysis: Using Bayesian and Frequentist Methods of Inference. New York: Holt, Rinehalt and Winston, 1972.

Tur, I., Roverato, A. and Castelo, R. Mapping eQTL networks with mixed graphical Markov models. *Genetics*, 198(4):1377-1393, 2014.

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See Also

```
qpG2Sigma
```

Examples

```
## Construct an adjacency matrix for a graph on 6 vertices

nVar <- 6
A <- matrix(0, nVar, nVar)
A[1,2] <- A[2,3] <- A[3,4] <- A[3,5] <- A[4,6] <- A[5,6] <- 1
A=A + t(A)
A
set.seed(123)
M <- qpRndWishart(delta=sqrt(1/nVar), P=0.5, n.var=nVar)
M
set.seed(123)
d=1:6
M <- qpRndWishart(delta=d, P=0.7, df=20)
M</pre>
```

qpTopPairs

Report pairs of variables

Description

Report a top number of pairs of variables according to either some association measure and/or occurring in a given reference graph.

Usage

Arguments

measurementsMatrix

matrix containing the measure of association between all pairs of variables.

refGraph a reference graph containing the pairs that should be reported and provided ei-

ther as an adjacency matrix, a graphNEL-class object or a graphAM-class

object

n number of pairs to report, 6 by default, use Inf for reporting all of them.

file file name to dump the pairs information as tab-separated column text.

decreasing logical; if TRUE then the measurements are employed to be ordered in decreas-

ing order; if FALSE then in increasing order.

pairup.i subset of vertices to pair up with subset pairup.j.

pairup.j subset of vertices to pair up with subset pairup.i.

annotation name of an annotation package to transform gene identifiers into gene symbols

when variables correspond to genes.

fcOutput output of qpFunctionalCoherence.

fcOutput.na.rm flag set to TRUE when pairs with NA values from fcOutput should not be re-

ported; FALSE (default) otherwise.

digits number of decimal digits reported in the values of measurementsMatrix and

functional coherence values. By default digits=NULL, and therefore, no round-

ing is performed.

Details

The measurementsMatrix should be symmetric and may have also contain NA values which will not be taken into account. That is an alternative way to restricting the variable pairs with the parameters pairup. i and pairup. j. The same holds for refGraph. One of these two, should be specified.

Value

The ranking of pairs is invisibly returned.

Author(s)

R. Castelo

See Also

qpGraph qpPrecisionRecall qpFunctionalCoherence

Examples

```
qpTopPairs(matrix(runif(100), nrow=10, dimnames=list(1:10,1:10)))
```

qpUnifRndAssociation Uniformly random association values

Description

Builds a matrix of uniformly random association values between -1 and +1 for all pairs of variables that follow from the number of variables given as input argument.

Usage

```
qpUnifRndAssociation(n.var, var.names=1:n.var)
```

Arguments

n.var number of variables.

var.names names of the variables to use as row and column names in the resulting matrix.

Details

This function simply generates uniformly random association values with no independence pattern associated to them. For generating a random covariance matrix that reflects such a pattern use the function qpG2Sigma.

Value

A symmetric matrix of uniformly random association values between -1 and +1.

Author(s)

R. Castelo

See Also

qpG2Sigma

Examples

```
rndassociation <- qpUnifRndAssociation(100)
summary(rndassociation[upper.tri(rndassociation)])</pre>
```

qpUpdateCliquesRemoving

Update clique list when removing one edge

Description

Updates the set of (maximal) cliques of a given undirected graph when removing one edge.

Usage

```
qpUpdateCliquesRemoving(g, clqlst, v, w, verbose=TRUE)
```

show progress on calculations.

Arguments

verbose

g	either a graphNEL object or an adjacency matrix of the given undirected graph.
clqlst	list of cliques of the graph encoded in g. this list should start on element n+1 (for n vertices) while between elements 1 to n there should be references to the cliques to which each of the 1 to n vertices belong to (i.e., the output of qpGetCliques) with parameter clqspervtx=TRUE.
V	vertex of the edge being removed.
W	vertex of the edge being removed.

Details

To find the list of all (maximal) cliques in an undirected graph is an NP-hard problem which means that its computational cost is bounded by an exponential running time (Garey and Johnson, 1979). For this reason, this is an extremely time and memory consuming computation for large dense graphs. If we spend the time to obtain one such list of cliques and we remove one edge of the graph with this function we may be able to update the set of maximal cliques instead of having to generate it again entirely with qpGetCliques but it requires that in the first call to qpGetCliques we set clqspervtx=TRUE. It calls a C implementation of the algorithm from Stix (2004).

Value

The updated list of maximal cliques after removing one edge from the input graph. Note that because the corresponding input clique list had to be generated with the argument clqspervtx=TRUE in the call to qpGetCliques, the resulting updated list of cliques also includes in its first p entries (p=number of variables) the indices of the cliques where that particular vertex belongs to. Notice also that although this strategy might be in general more efficient than generating again the entire list of cliques, when removing one edge from the graph, the clique enumeration problem remains NP-hard (see Garey and Johnson, 1979) and therefore depending on the input graph its computation may become unfeasible.

Author(s)

R. Castelo

References

Garey, M.R. and Johnson D.S. *Computers and intractability: a guide to the theory of NP-completeness*. W.H. Freeman, San Francisco, 1979.

Stix, V. Finding all maximal cliques in dynamic graphs *Comput. Optimization and Appl.*, 27:173-186, 2004.

See Also

qpCliqueNumber qpGetCliques qpIPF

```
## the example below takes about 30 seconds to execute and for that reason
## it is not executed by default
## Not run:
require(graph)

set.seed(123)
nVar <- 1000
g1 <- randomEGraph(V=as.character(1:nVar), p=0.1)
g1
clqs1 <- qpGetCliques(g1, clqspervtx=TRUE, verbose=FALSE)
length(clqs1)</pre>
```

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```
g2 <- removeEdge(from="1", to=edges(g1)[["1"]][1], g1)
g2

system.time(clqs2a <- qpGetCliques(g2, verbose=FALSE))

system.time(clqs2b <- qpUpdateCliquesRemoving(g1, clqs1, "1", edges(g1)[["1"]][1], verbose=FALSE))

length(clqs2a)

length(clqs2b)-nVar

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

SsdMatrix-class

Sum of squares and deviations Matrices

Description

The "SsdMatrix" class is the class of symmetric, dense matrices in packed storage (just as a dspMatrix-class, i.e., only the upper triangle is stored) defined within the qpgraph package to store corrected, or uncorrected, matrices of the sum of squares and deviations (SSD) of pairs of random variables. A corrected SSD matrix corresponds to a sample covariance matrix.

Objects from the Class

Objects can be created by calls of the form new("SsdMatrix", ...) or by using qpCov() which estimates a sample covariance matrix from data returning an object of this class.

Slots

ssd: Object of class dspMatrix-class storing the SSD matrix.

n: Object of class "numeric" storing the sample size employed to estimate the SSD matrix stored in the slot ssd. This is specially relevant when the SSD matrix was estimated from data with missing values by using complete observations only, which is the default mode of operation of qpCov().

Extends

"SsdMatrix" extends class "dspMatrix", directly.

Methods

```
dim signature(x = "SsdMatrix")
dimnames signature(x = "SsdMatrix")
show signature(object = "SsdMatrix")
determinant signature(object = "SsdMatrix", logarithm = "missing")
```

UGgmm-class

Undirected Gaussian graphical Markov model

Description

The "UGgmm" class is the class of undirected Gaussian graphical Markov models defined within the qpgraph package to store simulate and manipulate this type of graphical Markov models (GMMs).

An undirected Gaussian GMM is a family of multivariate normal distributions sharing a set of conditional independences encoded by means of an undirected graph. Further details can be found in the book of Lauritzen (1996).

Objects from the Class

Objects can be created by calls of the form UGgmm(g, ...) corresponding to constructor methods or rUGgmm(n, g, ...) corresponding to random simulation methods.

Slots

- p: Object of class "integer" storing the dimension of the undirected Gaussian GMM corresponding to the number of random variables.
- g: Object of class graphBAM-class storing the associated undirected labeled graph.

mean: Object of class "numeric" storing the mean vector.

sigma: Object of class dspMatrix-class storing the covariance matrix.

Methods

- UGgmm(g) Constructor method where g can be either an adjacency matrix or a graphBAM-class object.
- rUGgmm(n, g) Constructor simulation method that allows one to simulate undirected Gaussian GMMs where n is the number of GMMs to simulate and g can be either a graphParam object, an adjacency matrix or a graphBAM-class object.
- names(x) Accessor method to obtain the names of the elements in the object x that can be retrieved with the \$ accessor operator.
- \$ Accessor operator to retrieve elements of the object in an analogous way to a list.
- dim(x) Dimension of the undirected Gaussian GMM corresponding to the total number of random variables.
- dimnames(x) Names of the random variables in the undirected Gaussian GMM.
- show(object) Method to display some bits of information about the input undirected Gaussian GMM specified in object.
- summary(object) Method to display a sumarry of the main features of the input undirected Gaussian GMM specified in object.
- plot(x, ...) Method to plot the undirected graph associated to the the input undirected Gaussian GMM specified in x. It uses the plotting capabilities from the Rgraphviz library to which further arguments specified in ... are further passed.

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Author(s)

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