# Package 'geNetClassifier'

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Title classify diseases and build associated gene networks using gene expression profiles				
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Imports e1071, ipred, graphics				
Suggests leukemiasEset				
Enhances RColorBrewer, igraph				
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geNetClassifier-package

classify diseases and build associated gene networks using gene expression profiles

# Description

Comprehensive package to automatically train a multi-class SVM classifier based on gene expression data. Provides transparent selection of gene markers, their coexpression networks, and an interface to query the classifier.

# Details

Package: geNetClassifier

Type: Package
Version: 1.0
Date: 2013-02-28
License: GPL (>=2)

LazyLoad: yes

Depends: R (>= 2.10.1), Biobase (>= 2.5.5), EBarrays, minet, methods

Imports: e1071, ipred, graphics

Suggests: leukemiasEset

Enhances: RColorBrewer, igraph

# Author(s)

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# See Also

Main functions included in this package:

- geNetClassifier
- queryGeNetClassifier

# Query stats funtions:

- querySummary
- externalValidation.probMatrix
- externalValidation.stats
- plotAssignments

# Plots and genes info:

- calculateGenesRanking
- plotNetwork
- plotDiscriminantPower
- plotExpressionProfiles

# Classes:

- GenesRanking
- GenesNetwork
- GeNetClassifierReturn
- GeneralizationError

#### Related data sets:

• leukemiasEset

calculateGenesRanking

Calculate GenesRanking

## **Description**

Calculates the genes ranking and/or plots the posterior probability of the genes ordered by class ranking.

## Usage

```
calculateGenesRanking(eset=NULL, sampleLabels=NULL,
numGenesPlot=1000, plotTitle="Significant genes", plotLp=TRUE,
lpThreshold = 0.75, numSignificantGenesType="ranked",
returnRanking="full", nullHiphothesisFilter=0.95, nGenesExprDiff=1000,
geneLabels=NULL, precalcGenesRanking=NULL, IQRfilterPercentage= 0,
verbose=TRUE)
```

## **Arguments**

eset ExpressionSet or Matrix. Gene expression of the train samples.

sampleLabels Character. PhenoData variable (column name) containing the train samples class

labels.

Matrix or Factor. Class labels of the train samples.

numGenesPlot Integer. Number of genes to plot.

plotTitle Character. Plot title.

plotLp Logical. If FALSE no plot is drawn.

lpThreshold Numeric between 0 and 1. Required posterior probability value to consider a

gene 'significant'.

 ${\tt numSignificantGenesType}$ 

Character. Type of count for number of genes over lpThreshold.

- "global". Counts all genes of a class with posterior probability over lpThreshold, even if in the final ranking they were assigned to another class.
- "ranked". Counts only genes assigned to each class.

returnRanking

Character. Type of ranking to return:

- "full". Ranking of all available genes.
- "lp"/"significant"/"lpThreshold"/TRUE. Ranking of the significant genes (genes with posterior probability over lpThreshold).
- FALSE/NULL. No ranking is returned.

nullHiphothesisFilter

Numeric between 0 and 1. Genes with a Null Hipothesis with a posterior probability over this threshold will be removed from the ranking.

Null Hipothesis: They don't represent any class.

nGenesExprDiff

Numeric. Number of top genes to calculate the differencial expression for.

geneLabels Vector or Matrix. Gene name, ID or label which should be shown in the returned results and plots.

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```
IQRfilterPercentage
```

Integer. InterQuartile Range (IQR) filter applied to the initial data. Not recommended for more than two classes.

precalcGenesRanking

Allows providing a genesRanking provided by geNetClassifier or by a previous execution for the same data and parameters.

verbose

Logical. If TRUE, messages indicating the execution progress will be printed on screen.

#### **Details**

Significant genes: Genes with posterior probability over 'lpThreshold'. More significant genes may mean:

- · Very different class
- · More systemic disease

Plot lines represet the posterior probability of genes, sorted by rank from left to right.

In order to find genes that differentiate the classes from each other, the function ranks the genes bassed on their posterior probability for each class.

The posterior probability represents how well a gene differentiates samples from a class, from samples from other classes. Therefore, Genes with high posterior probability are good to differentiate a class from all the others.

This posterior probability is calculated by emfit (pkg:EBarrays), an expectation-maximization (EM) algorithm for gene expression mixture model.

# Value

- GenesRanking Optional. Requested genes ranking.
- Plot Optional. Plot of the posterior probability of the top genes.

## See Also

```
plot.GenesRanking is a shortcut to plotting a previously calculated genes ranking.
i.e. plot (genesRanking)
```

```
library(geNetClassifier)
# Load an expressionSet:
library(leukemiasEset)

data(leukemiasEset)

# Select the train samples:
trainSamples<- c(1:10, 13:22, 25:34, 37:46, 49:58)
# summary(leukemiasEset$LeukemiaType[trainSamples])

## Not run:
######
# Calculate/plot the significant genes (+ info) of a dataset
# without training classifier/calculating network</pre>
```

```
# Return only significant genes ranking (default)
signGenesRanking <- calculateGenesRanking(leukemiasEset[,trainSamples],</pre>
    sampleLabels="LeukemiaType")
numGenes(signGenesRanking)
# Return the full genes ranking:
fullRanking <- calculateGenesRanking(leukemiasEset[,trainSamples],</pre>
    sampleLabels="LeukemiaType", returnRanking="full")
numGenes (fullRanking)
numSignificantGenes(fullRanking)
# The significant genes can then be extracted from it:
signGenesRanking2 <- getTopRanking(fullRanking,
   numGenesClass=numSignificantGenes(fullRanking))
numGenes(signGenesRanking2)
# Changing the posterior probability required to consider genes significant:
signGenesRanking90 <- calculateGenesRanking(leukemiasEset[,trainSamples],</pre>
    sampleLabels="LeukemiaType", lpThreshold=0.9)
numGenes(signGenesRanking90)
## End(Not run)
######
# Ploting previously calculated rankings:
# Load or calculate a ranking (or a classifier with geNetClassifier)
data(leukemiasClassifier) # Sample trained classifier, @genesRanking
# Default plot:
# - equivalent to plot(leukemiasClassifier@genesRanking)
# - in this case, the previously calculated 'fullRanking'
  is equivalent to 'leukemiasClassifier@genesRanking'
calculateGenesRanking(precalcGenesRanking=leukemiasClassifier@genesRanking)
# Changing arguments:
numGenesPlot=5000, plotTitle="Leukemias", lpThreshold=0.9)
```

```
externalValidation.probMatrix

Probability matrix.
```

## **Description**

Generates the probability matrix.

#### Usage

```
externalValidation.probMatrix(queryResult, realLabels, numDecimals=2)
```

#### **Arguments**

```
queryResult Object returned by queryGeNetClassifier realLabels Factor. Actual/real class of the samples.

numDecimals Integer. Number of decimals to return.
```

#### **Details**

A probability matrix contains the probabilities of assigning each assigned sample to each class. They help identifying where errors are likely to occur even though there were not actual errors in the external/cross validation.

#### Value

The probability matrix.

#### Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

#### See Also

```
Main package function and classifier training: geNetClassifier
```

Query the classifier: queryGeNetClassifier

Query summary: querySummary

External validation stats: external Validation.stats

```
###########################
## Classifier training
############################
library(geNetClassifier)
# Load an expressionSet:
library(leukemiasEset)
data(leukemiasEset)
# Select the train samples:
trainSamples<- c(1:10, 13:22, 25:34, 37:46, 49:58)
# summary(leukemiasEset$LeukemiaType[trainSamples])
# Train a classifier or load a trained one:
# leukemiasClassifier <- geNetClassifier(leukemiasEset[,trainSamples],</pre>
     sampleLabels="LeukemiaType", plotsName="leukemiasClassifier")
data(leukemiasClassifier) # Sample trained classifier
###########################
## External Validation
###########################
# Select the samples to query the classifier
  - External validation: samples not used for training
testSamples <- c(1:60) [-trainSamples]</pre>
# Make a query to the classifier:
queryResult <- queryGeNetClassifier(leukemiasClassifier, leukemiasEset[,testSamples])</pre>
```

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```
# Obtain the probability matrix for the assigned samples:
externalValidation.probMatrix(queryResult, leukemiasEset[,testSamples]$LeukemiaType)
```

```
externalValidation.stats
```

Statistics of the external validation.

## **Description**

Taking as input the confussion matrix resulting from external validation calculates the global Accuracy, Call Rate, Sensitivity, Specificity and Matthews Correlation Coefficient.

## Usage

```
externalValidation.stats(confussionMatrix, numDecimals = 2)
```

# **Arguments**

confussionMatrix

Confussion matrix containing the real class as rows and the assigned class as columns.

numDecimals Integer. Number of decimals to show on the statistics.

## Value

List:

• global General classifier stats.

Accuracy: Percentage of correctly assigned samples within all assigned samples. CallRate: Percentage of samples with were assigned to a class.

• byClass Stats by class.

Sensitivity: Percentage of samples of each class which were correctly identified (Rate of true positives)

Specificity: Percentage of samples assigned to a given class that really belonged to the class (Rate of true negatives)

MCC (Matthews Correlation Coefficient): Measure wich takes into account both, true and false positives and negatives. (100%: Perfect assignments) confMatrix Confussion matrix.

#### Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

# See Also

Main package function and classifier training: geNetClassifier

Querying the classifier: queryGeNetClassifier

Generating the probability matrix: externalValidation.probMatrix

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#### **Examples**

```
############################
## Classifier training
############################
library(geNetClassifier)
# Load an expressionSet:
library(leukemiasEset)
data(leukemiasEset)
# Select the train samples:
trainSamples<- c(1:10, 13:22, 25:34, 37:46, 49:58)
# summary(leukemiasEset$LeukemiaType[trainSamples])
# Train a classifier or load a trained one:
# leukemiasClassifier <- geNetClassifier(leukemiasEset[,trainSamples],</pre>
     sampleLabels="LeukemiaType", plotsName="leukemiasClassifier")
data(leukemiasClassifier) # Sample trained classifier
#############################
## External Validation:
# Select the samples to query the classifier
  - External validation: samples not used for training
testSamples <- c(1:60) [-trainSamples]</pre>
# Make a query to the classifier:
queryResult <- queryGeNetClassifier(leukemiasClassifier, leukemiasEset[,testSamples])</pre>
# Create the confusion matrix
confMatrix <- table(leukemiasEset[,testSamples]$LeukemiaType,queryResult$class)</pre>
# Calculate its accuracy, call rate, sensitivity and specificity:
externalValidation.stats(confMatrix)
```

gClasses-methods Classes in the ranking.

## **Description**

Returns the names of the classes in a GenesRanking

#### Methods

```
signature(object = "GenesRanking")
```

## See Also

Main package function and classifier training: geNetClassifier This method's class (GenesRanking) help page.

```
data(leukemiasClassifier)
gClasses(leukemiasClassifier@genesRanking)
```

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```
GeneralizationError-class
```

Class "GeneralizationError" (slot of GeNetClassifierReturn)

## **Description**

Contains the estimation of the Generalization Error and the gene stats for geNetClassifier executed with the given data and parameters. \ Calculated by 5-fold cross-validation.

#### **Slots**

```
accuracy: "Matrix". Accuracy and call rate.

sensitivitySpecificity: "Matrix". Sensitivity, Specificity, Matthews Correlation Coefficient and Call Rate for each of the classes.

confMatrix: "Matrix". Confussion matrix.

probMatrix: "Matrix". Probabilities of belonging to each class for the assigned samples. Helps identifying where errors are likely to occur even though there were not actual errors in the cross-validation.

querySummary: "List". Stats regarding the probability and number of assigned test samples to each class.

classificationGenes.stats: "List". Some basic statistics regarding the chosen genes. classificationGenes.num: "Matrix". Number of genes used for each of the 5 cross-validation classifiers.
```

# Methods

overview signature(object = "GeneralizationError"): Shows an overview of all
 the slots in the object.

#### Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

#### See Also

Main package function and classifier training: geNetClassifier

```
######
# Load data and train a classifier
######
library(geNetClassifier)
# Load an expressionSet:
library(leukemiasEset)
data(leukemiasEset)

# Select the train samples:
trainSamples<- c(1:10, 13:22, 25:34, 37:46, 49:58)</pre>
```

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```
# summary(leukemiasEset$LeukemiaType[trainSamples])
# Train a classifier or load a trained one:
# Note: Required 'estimateGError=TRUE'
# leukemiasClassifier <- geNetClassifier(leukemiasEset[,trainSamples],</pre>
     sampleLabels="LeukemiaType", plotsName="leukemiasClassifier",
     estimateGError=TRUE)
data(leukemiasClassifier) # Sample trained classifier
# Global view of the returned object and its structure:
leukemiasClassifier
names(leukemiasClassifier)
########
# Exploring the cross validation stats
# Note: Required 'estimateGError=TRUE' in geNetClassifier()
# Generalization Error estimated by cross-validation:
leukemiasClassifier@generalizationError
overview(leukemiasClassifier@generalizationError)
# i.e. probabilityMatrix:
leukemiasClassifier@generalizationError@probMatrix
# i.e. statistics of the genes chosen in any of the CV loops for for AML:
{\tt leukemiasClassifier@generalizationError@classificationGenes.stats\$ AML}
```

genesDetails-methods

Details of the genes in the network.

# Description

Information of the genes in the ranking (table format).

## **Arguments**

object a GenesRanking

nGenes integer. Number of genes to show per class

numDecimals integer. Number of decimals to show in the numeric values

classes character. Classes of the genes to show

genes character. Genes to show

#### Value

A list containing a dataframe with the details of the genes of each class. For each gene, the following information is provided:

ranking Ranking of the gene.

gERankMean Mean rank the gene obtained in the cross-validation loops. Only available if

geNetClassifier() was called with option estimateGError=TRUE (False by

default).

class Class the gene was chosen for (the class the gene differentiates from the other

classes).

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postProb Posterior probability which the gene was assigned by the expectation-maximization

algorithm (emfit). Tied values are ranked based on the higher absolute value of exprsMeanDiff. Values are rounded. Several genes may look tied at posterior

probability '1' but may actually be i.e. 0.999998 and 0.999997.

exprsMeanDiff

exprsUpDw

Difference betwen the mean expression of the gene within its class and its mean

expression in the other classes.

Gene repressed (DOWN) or over-expressed(UP) for the current class (compared

to the other classes).

discriminantPower

Measure calculated based on the coordinates of the support vectors. Represents the weight that the classifier gives to each gene to separate the classes.

discrPwClass Class for which the Discriminant Power was calculated for.

isRedundant Does the gene have a high correlation or mutual information with other genes in

the list? The threshold to consider a gene redundant can be set through the arguments (by default: correlationsThreshold=0.8 and interactionsThreshold=0.5).

#### Methods

```
genesDetails(object, nGenes=NULL, numDecimals=4, classes=NULL, genes=NULL)
```

## See Also

Main package function and classifier training: geNetClassifier This method's class (GenesRanking) help page.

# **Examples**

```
data(leukemiasClassifier) # Sample geNetClassifier() return
options(width=200) # Optional, use in case the table rows are wrapped

genesDetails(leukemiasClassifier@classificationGenes)$CML
genesDetails(leukemiasClassifier@genesRanking, nGenes=5, numDecimals=2,
classes="AML")
genesDetails(leukemiasClassifier@genesRanking, genes=c("ENSG00000096006",
"ENSG00000168081","ENSG00000105699"))$CLL
```

GenesNetwork-class Class "GenesNetwork"

#### **Description**

Contains the network returned by geNetClassifier. (Slot: @genesNetwork)

#### Methods

```
getNodes signature(object = "GenesNetwork"): Returns the network nodes (genes).
getEdges signature(object = "GenesNetwork"): Returns the network edges (relationships).
```

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```
getNumNodes signature(object = "GenesNetwork"): Returns the number of nodes
    (genes) in the network.

getNumEdges signature(object = "GenesNetwork"): Returns the number of edges
    (relationships) in the network,

getSubNetwork signature(network = "GenesNetwork"): Returns a new network containing only the given genes.

network2txt signature(network = "GenesNetwork"): Exports the network as text file.
```

#### Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigación del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

#### See Also

Main package function and classifier training: geNetClassifier Plot network or export as iGraph: plotNetwork

```
######
# Load data and train a classifier
library(geNetClassifier)
# Load an expressionSet:
library(leukemiasEset)
data(leukemiasEset)
# Select the train samples:
trainSamples<- c(1:10, 13:22, 25:34, 37:46, 49:58)
# summary(leukemiasEset$LeukemiaType[trainSamples])
# Train a classifier or load a trained one:
# leukemiasClassifier <- geNetClassifier(leukemiasEset[,trainSamples],</pre>
    sampleLabels="LeukemiaType", plotsName="leukemiasClassifier")
data(leukemiasClassifier) # Sample trained classifier
######
# Explore the returned object
######
# Global view of the object and its structure:
leukemiasClassifier
names(leukemiasClassifier)
# List of Networks by classes:
{\tt leukemiasClassifier@genesNetwork}
# Access to the nodes or edges of each network:
getEdges(leukemiasClassifier@genesNetwork$AML)[1:5,]
getNodes(leukemiasClassifier@genesNetwork$AML)[1:50]
######
# Plotting
# Example: Plotting the sub-network of a class classificationGenes
```

14 GenesRanking-class

GenesRanking-class Class "GenesRanking"

## **Description**

Contains the genes ranking generated by geNetClassifier and the genes details. (Slots: @classificationGenes and @genesRanking)

## Methods

genesDetails signature(object = "GenesRanking"): Returns data.frames with information about the genes.

getRanking signature(object = "GenesRanking"): Returns a matrix containing the
 ranked genes.

getTopRanking signature(object = "GenesRanking", numGenesClass): Returns
 a new GenesRanking object containing only the top genes of each class.

**gClasses** signature (object = "GenesRanking"): Returns the classes for which the genes are ranked.

nGenes signature(object = "GenesRanking"): Returns the number of available ranked
genes per class.

numSignificantGenes signature(object = "GenesRanking"): Returns the number of significant genes per class (genes over the given posterior probability threshold).

plot signature(x = "GenesRanking", y = "missing"): Plots the genes' posterior
probability. Wrapper of calculateGenesRanking.

#### Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

#### See Also

Main package function and classifier training: geNetClassifier Plot the ranking genes's posterior probability: plot.GenesRanking

## **Examples**

```
######
# Load data and obtain a genesRanking
library(geNetClassifier)
# Load an expressionSet:
library(leukemiasEset)
data(leukemiasEset)
# Select the train samples:
trainSamples<- c(1:10, 13:22, 25:34, 37:46, 49:58)
# summary(leukemiasEset$LeukemiaType[trainSamples])
# Train a classifier or load a trained one:
# leukemiasClassifier <- geNetClassifier(leukemiasEset[,trainSamples],</pre>
      sampleLabels="LeukemiaType", plotsName="leukemiasClassifier")
data(leukemiasClassifier) # Sample trained classifier
# Or directly obtain a genesRanking with calculateGenesRanking()
## Not run: genesRanking <- calculateGenesRanking(leukemiasEset[,trainSamples],
    sampleLabels="LeukemiaType", returnRanking="full")
## End(Not run)
######
# Exploring the rankings
######
genesRanking <- leukemiasClassifier@genesRanking</pre>
# Number of available genes in the ranking:
numGenes(genesRanking)
# Number of significant genes (genes with posterior probability over the threshold.
# Default: lpThreshold=0.75):
numSignificantGenes(genesRanking)
# Top 10 genes of CML:
genesDetails(genesRanking)$CML[1:10,]
# To get a sub ranking with the top 10 genes:
getTopRanking(genesRanking, 10)
# Genes details of the top 10 genes:
genesDetails(getTopRanking(genesRanking, 10))
###
# Genes used for training the classifier:
###
numGenes(leukemiasClassifier@classificationGenes)
leukemiasClassifier@classificationGenes
#genesDetails(leukemiasClassifier@classificationGenes)
                                                          # List by classes
genesDetails(leukemiasClassifier@classificationGenes)$AML # Show a class genes
# If your R console wraps the table rows, try widening your display width:
# options(width=200)
```

geNetClassifier

Main function of the geNetClassifier package.

Trains the multi-class SVM classifier based on the given gene expression data through transparent detection of gene markers and their associated networks.

#### **Description**

Allows to train the classifier, calculate the genes network...

#### **Usage**

```
geNetClassifier(eset, sampleLabels, plotsName = NULL,
buildClassifier = TRUE, estimateGError = FALSE,
calculateNetwork = TRUE, labelsOrder = NULL, geneLabels = NULL,
numGenesNetworkPlot = 100,
minGenesTrain = 1, maxGenesTrain = 100, continueZeroError = FALSE,
numIters = 6, lpThreshold = 0.75, numDecimals = 3,
removeCorrelations = FALSE, correlationsThreshold = 0.8,
removeInteractions = FALSE, interactionsThreshold = 0.5,
minProbAssignCoeff = 1, minDiffAssignCoeff = 0.8,
IQRfilterPercentage = 0, skipInteractions = FALSE,
precalcGenesNetwork = NULL, precalcGenesRanking = NULL,
returnAllGenesRanking = TRUE, verbose = TRUE)
```

#### **Arguments**

eset ExpressionSet or matrix. Gene expression of the train samples.

sampleLabels Character. PhenoData variable (column name) containing the train samples class

labels.

Matrix or Factor. Class labels of the train samples.

labelsOrder Vector or Factor. Order in which the labels should be shown in the returned

results and plots.

plotsName Character. File name with which the plots should be saved. If not provided, no

plots will be drawn.

buildClassifier

Logical. If TRUE trains a classifier with the given samples.

estimateGError

Logical. If TRUE uses cross-validation to estimate the Generalization Error of a classiffier trained with the given samples.

calculateNetwork

Logical. If TRUE calculates the coexpression network between the best genes.

geneLabels Vector or Matrix. Gene name, ID or label which should be shown in the returned results and plots.

numGenesNetworkPlot

Integer. Number of genes to show in the coexpression network for each class.

minGenesTrain

Integer. Maximum number of genes per class to train the classifier with.

maxGenesTrain

Integer. Maximum number of genes per class to train the classifier with.

continueZeroError

Logical. If TRUE, the program will continue testing combinations with more genes even if error 0 has been reached.

numIters Integer. Number of iterations to determine the optimum number of genes (between minGenesTrain and maxGenesTrain).

lpThreshold Numeric between 0 and 1. Required posterior probability value to consider a gene 'significant'.

removeCorrelations

Logical. If TRUE, no correlated genes will be chosen to train the classifier.

correlationsThreshold

Numeric between 0 and 1. Minimum Pearson's correlation coefficient to consider genes correlated.

removeInteractions

Logical. If TRUE, genes with Mutual Information coefficient over the threshold will not be chosen to train the classifier.

interactionsThreshold

Numeric between 0 and 1. Minimum Mutual Information coefficient to consider two genes equivalent.

numDecimals Integer. Number of decimals to show in the statistics.

minProbAssignCoeff

Numeric. Allows modifying the required probability to assign a sample to a class in the internal crossvalidation. For details see: queryGeNetClassifier

minDiffAssignCoeff

Numeric. Allows modifying the difference of probabilities required between the most likely class and second most likely class to assign a sample. For details see: queryGeNetClassifier

IQRfilterPercentage

Integer. InterQuartile Range (IQR) filter applied to the initial data. Not recommended for more than two classes.

skipInteractions

Logical. If TRUE, the interactions between genes are not calculated (they will not appear on the genes network). Saves some execution time. Only available if removeInteractions=FALSE.

precalcGenesNetwork

GenesNetwork from a previous execution with the same expression data and parameters.

precalcGenesRanking

GenesRanking from a previous execution with the same expression data and parameters.

returnAllGenesRanking

Logical. If TRUE, returns the whole genes ranking. If FALSE the returned ranking contains only the significant genes (genes over lpThreshold).

verbose Logical. If TRUE, messages indicating the execution progress will be shown.

#### Value

A GeNetClassifierReturn object containing the classifier and the genes chosen to train it (classificationGenes), Cross-Validation statistics, the whole GenesRanking and each class' GenesNetwork (if requested). Several plots saved as 'plotsName\_...pdf' in the working directory.

#### Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

#### References

Packages used by this function:

EBarrays: emfit (Implements EM algorithm for gene expression mixture model) and ebPatterns, for calculating the gene ranking.

Ming Yuan, Michael Newton, Deepayan Sarkar and Christina Kendziorski (2007). EBarrays: Unified Approach for Simultaneous Gene Clustering and Differential Expression Identification. R package.

```
e1071: svm.
```

Evgenia Dimitriadou, Kurt Hornik, Friedrich Leisch, David Meyer and Andreas Weingessel (2011). e1071: Misc Functions of the Department of Statistics (e1071), TU Wien. R package. http://CRAN.R-project.org/package=e1071

ipred: kfoldev (computes feasible sample sizes for the k groups in k-fold ev) for the cross-validations.

Andrea Peters and Torsten Hothorn (2012). ipred: Improved Predictors. R package. http://CRAN.R-project.org/package=ipred

minet for the Mutual Information network.

Patrick E. Meyer, Frederic Lafitte and Gianluca Bontempi (2008). MINET: An open source R/Bioconductor Package for Mutual Information based Network Inference. BMC Bioinformatics. http://www.biomedcentral.com/1471-2105/9/461

RColorBrewer for palettes in some of the plots.

Erich Neuwirth (2011). RColorBrewer: ColorBrewer palettes. R package.

http://CRAN.R-project.org/package=RColorBrewer

igraph for the graphical representation of the networks.

Csardi G, Nepusz T: The igraph software package for complex network research, InterJournal, Complex Systems 1695. 2006. http://igraph.sf.net

#### See Also

To query the classifier: queryGeNetClassifier
All functions in the package: geNetClassifier-package

```
#######
# NOTE: Training the classifier takes a while...
# Choose ONE of the followings, or modify to suit your needs:
## Not run:
# "Basic" execution: All default parameters
leukemiasClassifier <- qeNetClassifier(eset=leukemiasEset[,trainSamples],</pre>
    sampleLabels="LeukemiaType", plotsName="leukemiasClassifier")
# All default parameters also estimatings the classiffier's Generalization Error:
# ( by default: buildClassifier=TRUE, calculateNetwork=TRUE)
# Takes longer time than the basic execution
leukemiasClassifier <- geNetClassifier(eset=leukemiasEset[,trainSamples],</pre>
    sampleLabels="LeukemiaType", plotsName="leukemiasClassifier",
    estimateGError=TRUE)
# Faster execution (few minutes - depending on the computer):
# By skipping the calculation of the interactions (MI) betwen the genes,
# and reducing the number of genes to explore when training the classifier
# (100 by default), the execution time can be sightly reduced
leukemiasClassifier <- qeNetClassifier(eset=leukemiasEset[,trainSamples],</pre>
sampleLabels="LeukemiaType", plotsName="leukemiasClassifier",
skipInteractions= TRUE, maxGenesTrain=20)
# To any of these examples, you can add/remove the argument geneLabels,
# in order to show/remove the gene name in the rankings and plots:
# The argument labelsOrder allows showing the classes in a specific order
# i.e.: labelsOrder=c("ALL","CLL","AML",CML","NoL")
save(leukemiasClassifier, file="leukemiasClassifier.RData") # Save execution result
# For loading the saved object in the future...
# (If it doesn't find it, use getwd() to make sure you are in the right directory)
#load("leukemiasClassifier.RData")
# To avoid having to train a classifier to continue learning to use the package,
# you can load the package's pre-executed example:
data(leukemiasClassifier)
#This example classifier was trained with the following code:
#leukemiasClassifier <- geNetClassifier(leukemiasEset[,trainSamples],
     "LeukemiaType", plotsName="leukemiasClassifier", buildClassifier=TRUE,
     estimateGError=TRUE, calculateNetwork=TRUE, geneLabels=geneSymbols)
#
#######
# Explore the returned object:
#######
names(leukemiasClassifier)
# More details on the class' help file:
?GeNetClassifierReturn
# Further options:
# The trained classifier can be used to find the class of new samples:
?queryGeNetClassifier
# The default plots can be modified and presonalized to fit the user needs:
?calculateGenesRanking
```

```
?plotNetwork
?plotDiscriminantPower
?plotExpressionProfiles
## End(Not run)
```

GeNetClassifierReturn-class

Class "GeNetClassifierReturn"

#### **Description**

Object wich wraps all the items returned by geNetClassifier. It usually contains the classifier, the genes ranking and information, the network and any other requested statistics.

#### Methods

```
names signature(x = "GeNetClassifierReturn"): Shows the available slots in the object.
```

overview signature(object = "GeNetClassifierReturn"): Shows an overview of
 all the slots in the object.

#### **Slots**

Available slots deppends on the arguments used to call geNetClassifier():

call: call. Always available.

classifier: list. SVM classifier. Only available if geNetClassifier() was called with option buildClassifier=TRUE (default settings).

classificationGenes: GenesRanking. Genes used to train the classifier. Only available if geNetClassifier() was called with option buildClassifier=TRUE (default settings).

generalizationError: GeneralizationError. Statistics calculated for the current training set and options.

Only available if geNetClassifier() was called with option estimateGError=TRUE (False by default).

genesRanking: GenesRanking. Whole genes ranking (if returnAllGenesRanking=TRUE) or significant genes ranking (if returnAllGenesRanking=FALSE, includes only the genes with posterior probability over lpThreshold)

genesRankingType: character. "all", "significant" or "significantNonRedundant"

genesNetwork: List of GenesNetwork. Only available if geNetClassifier() was called with option calculateNetwork=TRUE (default settings).

genesNetworkType: character. At the moment, only "topGenes" available.

# Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

#### See Also

Main package function and classifier training: geNetClassifier plot.GeNetClassifierReturn

```
######
# Load data and train a classifier
library(geNetClassifier)
# Load an expressionSet:
library(leukemiasEset)
data(leukemiasEset)
# Select the train samples:
trainSamples<- c(1:10, 13:22, 25:34, 37:46, 49:58)
# summary(leukemiasEset$LeukemiaType[trainSamples])
# Train a classifier or load a trained one:
# leukemiasClassifier <- geNetClassifier(leukemiasEset[,trainSamples],</pre>
     sampleLabels="LeukemiaType", plotsName="leukemiasClassifier")
data(leukemiasClassifier) # Sample trained classifier
######
# Explore the returned object
######
# Global view of the object and its structure:
leukemiasClassifier
names(leukemiasClassifier)
### Depending on the available slots:
# Call and acess to the classifier:
leukemiasClassifier@call
leukemiasClassifier@classifier
# Genes used for training the classifier:
numGenes(leukemiasClassifier@classificationGenes)
{\tt leukemiasClassifier@classificationGenes}
# Show de tetails of the genes of a class
genesDetails(leukemiasClassifier@classificationGenes)$AML
# If your R console wraps the table rows, try widening your display width:
# options(width=200)
# Generalization Error estimated by cross-validation:
leukemiasClassifier@generalizationError
overview(leukemiasClassifier@generalizationError)
# i.e. probabilityMatrix:
leukemiasClassifier@generalizationError@probMatrix
# i.e. statistics of the genes chosen in any of the CV loops for for AML:
leukemias Classifier \tt @generalization Error \tt @classification Genes.stats \tt \$AML
# List of Networks by classes:
leukemiasClassifier@genesNetwork
# Access to the nodes or edges of each network:
getEdges(leukemiasClassifier@genesNetwork$AML)
getNodes(leukemiasClassifier@genesNetwork$AML)
```

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```
# Genes ranking:
leukemiasClassifier@genesRanking
# Number of available genes in the ranking:
numGenes(leukemiasClassifier@genesRanking)
# Number of significant genes
# (genes with posterior probability over lpThreshold, default=0.75)
numSignificantGenes(leukemiasClassifier@genesRanking)
# Top 10 genes of CML:
genesDetails(leukemiasClassifier@genesRanking)$CML[1:10,]
# To get a sub ranking with the top 10 genes:
getTopRanking(leukemiasClassifier@genesRanking, 10)
# Genes details of the top 10 genes:
genesDetails(getTopRanking(leukemiasClassifier@genesRanking, 10))
```

getEdges-methods Edges in the network.

#### **Description**

Returns the network's edges (relations between genes).

#### Methods

```
signature(object = "GenesNetwork")
```

#### See Also

Main package function and classifier training: geNetClassifier This method's class (GenesNetwork) help page.

# Examples

```
data(leukemiasClassifier)
getEdges(leukemiasClassifier@genesNetwork$AML)[1:5,]
```

getNodes-methods Nodes in the network.

#### **Description**

Returns the network's nodes (genes).

#### Methods

```
signature(object = "GenesNetwork")
```

# See Also

Main package function and classifier training: geNetClassifier This method's class (GenesNetwork) help page.

getNumEdges-methods 23

#### **Examples**

```
data(leukemiasClassifier)
getNodes(leukemiasClassifier@genesNetwork$AML)[1:5]
```

```
getNumEdges-methods
```

Number of edges in the network.

#### **Description**

Returns the number of edges (relationships) in the network.

#### Methods

```
signature(object = "GenesNetwork")
```

#### See Also

Main package function and classifier training: geNetClassifier This method's class (GenesNetwork) help page.

# **Examples**

```
data(leukemiasClassifier)
getNumEdges(leukemiasClassifier@genesNetwork$AML)
```

```
getNumNodes-methods
```

Number of nodes in the network.

## **Description**

Returns the number of nodes (genes) in the network.

## Methods

```
signature(object = "GenesNetwork")
```

#### See Also

Main package function and classifier training: geNetClassifier This method's class (GenesNetwork) help page.

```
data(leukemiasClassifier)
getNumNodes(leukemiasClassifier@genesNetwork$AML)
```

getRanking-methods Shows the genes ranking.

# **Description**

Shows the ranking as matrix: Ranked genes by classes.

#### **Arguments**

object a GenesRanking
showGeneID boolean. If TRUE, the genes will be shown with the gene IDs used in the expressionSet. This matrix will be ...\$geneID in the returned list.
showGeneLabels

boolean. If TRUE, and if the ranking contains gene labels, the ranking matrix will use them. This matrix will be . . . \$geneLabels in the returned list.

#### Value

The method returns a list with one or two matrices: ...\$geneLabels and ...\$geneID.

#### See Also

Main package function and classifier training: geNetClassifier This method's class (GenesRanking) help page.

# **Examples**

```
data(leukemiasClassifier)
getRanking(leukemiasClassifier@classificationGenes)

# Top 7 genes (two ways):
getRanking(leukemiasClassifier@genesRanking)$geneLabels[1:7,]
getRanking(getTopRanking(leukemiasClassifier@genesRanking, 7))

# Show gene ID and select a class:
getRanking(leukemiasClassifier@classificationGenes, showGeneID=TRUE)$geneID[,"CML", drop=FALSE]
```

```
getSubNetwork-methods
```

Get a sub-network.

## **Description**

Returns the sub-network formed by the given genes.

## Usage

```
getSubNetwork(network, genes, showWarnings=TRUE)
```

#### **Arguments**

network GenesNetwork or GenesNetwork list containing the whole network.

genes GenesRanking or character vector. Genes in the new network.

showWarnings Logical. If true, shows warnings if the given genes are not in the network.

#### Value

A GenesNetwork or list of networks between the given genes.

#### See Also

Main package function and classifier training: geNetClassifier This method's class (GenesNetwork) help page.

#### **Examples**

```
data(leukemiasClassifier)
clGenesSubNet <- getSubNetwork(leukemiasClassifier@genesNetwork,
leukemiasClassifier@classificationGenes)
getSubNetwork(leukemiasClassifier@genesNetwork, getTopRanking(leukemiasClassifier@genesRanking)</pre>
```

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```
getTopRanking-methods
```

Gets a new ranking with the given top genes.

## **Description**

Returns a new ranking containing only the top genes of each class.

# **Arguments**

```
object a GenesRanking
numGenesClass
integer. Number of genes per class.
```

## Methods

```
getTopRanking(object, numGenesClass)
```

## See Also

Main package function and classifier training: geNetClassifier This method's class (GenesRanking) help page.

```
data(leukemiasClassifier) # Sample classifier
# Sub-ranking with the top 10 genes:
getTopRanking(leukemiasClassifier@genesRanking, 10)
```

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```
leukemiasClassifier
```

Sample leukemias classifier

# **Description**

A sample of the object returned by geNetClassifier. Containins the classifier, the network, and the gene statistics.

## Usage

```
data(leukemiasClassifier)
```

#### **Format**

```
GeNetClassifierReturn object
```

```
data(leukemiasClassifier)
# Global view of the object and its structure:
leukemiasClassifier
names(leukemiasClassifier)
# Call and acess to the classifier:
leukemiasClassifier@call
leukemiasClassifier@classifier
# Genes used for training the classifier:
numGenes(leukemiasClassifier@classificationGenes)
leukemiasClassifier@classificationGenes
genesDetails(leukemiasClassifier@classificationGenes)
# Generalization Error estimated by cross-validation:
  leukemiasClassifier@generalizationError
# overview(leukemiasClassifier@generalizationError)
# List of Networks by classes:
leukemiasClassifier@genesNetwork
# Access to the nodes or edges of each network:
getEdges(leukemiasClassifier@genesNetwork$AML)[1:5,]
getNodes(leukemiasClassifier@genesNetwork$AML)[1:50]
# Global genes ranking:
leukemiasClassifier@genesRanking
numGenes(leukemiasClassifier@genesRanking)
numSignificantGenes(leukemiasClassifier@genesRanking)
# getTopRanking(leukemiasClassifier@genesRanking, 10)
```

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network2txt network2txt

## **Description**

Saves the GenesNetwork as text file.

## Usage

```
network2txt(network, filePrefix = NULL, nwClass = NULL)
```

## **Arguments**

network GenesNetwork or list of GenesNetworks.

filePrefix Character. File name prefix. nwClass Character. Network class.

#### Value

Saves the networks as text (.txt) files. The files will be saved in the current working directory as filePrefix\_className.txt.

#### See Also

Main package function and classifier training: geNetClassifier This method's class (GenesNetwork) help page.

## **Examples**

```
## Load or calculate a network:
library(geNetClassifier)
data(leukemiasClassifier)

## Export as text:
network2txt(leukemiasClassifier@genesNetwork, filePrefix="leukemiasNetwork")
```

numGenes-methods Number of genes in the genesRanking.

# Description

Provides the number of genes in the genesRanking.

#### Methods

```
signature(object = "GenesRanking")
```

# See Also

Main package function and classifier training: geNetClassifier This method's class (GenesRanking) help page.

#### **Examples**

```
data(leukemiasClassifier)
numGenes(leukemiasClassifier@genesRanking)
```

numSignificantGenes-methods

Number of ranked genes over the posterior probability threshold.

#### **Description**

Provides the number of ranked genes over the posterior probability threshold

## **Arguments**

```
object a GenesRanking

lpThreshold Posterior probability threshold

numSignificantGenesType

"ranked" or "global".
```

Ranked will show the count of genes on the ranking of each class. Each gene will be counted only once, since it is only kept in the class for which it had better ranking.

Global counts the genes over the threshold before assigning them to a class. i.e. a gene might have 0.3 for one class, and 0.25 for another, if we are taking a thershold of 0.20, it will be counted on both classes.

## Methods

numSignificantGenes(object, lpThreshold=0.75, numSignificantGenesType="ranke")

#### See Also

Main package function and classifier training: geNetClassifier This method's class (GenesRanking) help page.

```
data(leukemiasClassifier)
# Total number of genes in the ranking:
numGenes(leukemiasClassifier@genesRanking)
# Number of genes owher the posterior probability threshold
# Default: lpThreshold=0.75, numSignificantGenesType="ranked"
numSignificantGenes(leukemiasClassifier@genesRanking)
numSignificantGenes(leukemiasClassifier@genesRanking, numSignificantGenesType="global")
numSignificantGenes(leukemiasClassifier@genesRanking, lpThreshold=0.90)
```

overview-methods 29

```
overview-methods Overview
```

## **Description**

Provides an overview of all the slots in the object.

#### Methods

It can be applied to the following classes:

```
signature(object = "GenesNetwork")
signature(object = "GenesRanking")
signature(object = "GeNetClassifierReturn")
signature(object = "GeneralizationError")
```

## See Also

Main package function and classifier training: geNetClassifier This method's classes help pages:

```
GenesRanking
GenesNetwork
GeNetClassifierReturn
GeneralizationError
```

# **Examples**

```
library(geNetClassifier)
data(leukemiasClassifier)
# geNetClassifier return:
overview(leukemiasClassifier)
# Generalization Error and stats estimated by cross-validation:
overview(leukemiasClassifier@generalizationError)
# A GenesNetwork:
# (a class has to be selected, otherwise it is a list)
overview(leukemiasClassifier@genesNetwork$ALL)
# For a GenesRanking, we recommend to use genesDetails() instead:
genesDetails(leukemiasClassifier@classificationGenes)$AML
```

```
plot.GenesRanking Plot GenesRanking
```

#### **Description**

Plots the posterior probability of the genes ordered by class ranking.

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#### Usage

#### **Arguments**

```
    x GenesRanking.
    numGenesPlot Numeric. Number of genes to plot.
    plotTitle Character. Plot main title.
    lpThreshold Numeric between 0 and 1. Required posterior probability value to consider a gene 'significant'.
    y Not required.
    ... Not required
```

#### **Details**

Significant genes: Genes with posterior probability over 'lpThreshold'. More significant genes may mean:

- · Very different class
- More systemic disease

Plot lines represet the posterior probability of genes, sorted by rank from left to right.

In order to find genes that differentiate the classes from each other, the function ranks the genes bassed on their posterior probability for each class.

The posterior probability represents how well a gene differentiates samples from a class, from samples from other classes. Therefore, Genes with high posterior probability are good to differentiate a class from all the others.

This posterior probability is calculated by emfit (pkg:EBarrays), an expectation-maximization (EM) algorithm for gene expression mixture model.

#### Value

Posterior probability plot of the top genes.

```
# Load or calculate a ranking (or a classifier with geNetClassifier)
data(leukemiasClassifier) # Sample trained classifier, @genesRanking
# Default plot:
plot(leukemiasClassifier@genesRanking)
# Changing options:
plot(leukemiasClassifier@genesRanking,
    numGenesPlot=5000, plotTitle="Leukemias", lpThreshold=0.9)
```

```
plot.GeNetClassifierReturn

Plot GeNetClassifierReturn
```

# Description

Allows generating the plots from the objet created by geNetClassifier.

## Usage

```
## S3 method for class 'GeNetClassifierReturn'
plot(x, y="missing", fileName = NULL, lpThreshold = 0.75,
    numGenesLpPlot = 1000, numGenesNetworkPlot = 100,
geneLabels = NULL, verbose = TRUE, ...)
```

## **Arguments**

X	$GeNet Classifier Return. \ Object \ returned \ by \ the \ main \ function \ "geNetClassifier".$				
fileName	Character. File name to save the plots.				
lpThreshold	Numeric between 0 and 1. Required posterior probability value to consider a gene 'significant'.				
numGenesLpPlot					
	Integer. Number of genes to show in the significant genes plot.				
numGenesNetworkPlot					
	Integer. Number of genes (nodes) to plot in the network.				
geneLabels	Vector or Matrix. Gene name, ID or label which should be shown in the returned results and plots.				
verbose	Logical. If TRUE, messages indicating the execution progress will be printed on screen.				
У	Not required.				
	Not required				

#### **Details**

The plots are generated by default by <code>geNetClassifier</code>. This function allows re-plotting them with different parameters.

# Value

Plots (depending on the available info):

- Significant genes
- Classification genes' Discriminant Power
- Top ranked genes network (for each class)

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#### See Also

```
Main package function and classifier training: geNetClassifier Class GeNetClassifierReturn
Other plotting functions:
-plotDiscriminantPower
-plot.GenesRanking
-plotNetwork
```

#### **Examples**

```
library(geNetClassifier)

# Train or load an already trained classifier
data(leukemiasClassifier)

# Plot default plots on-screen
plot(leukemiasClassifier)

# Save plots on file
# (includes Discriminant Power of all genes, but the networks will not be interactive)
plot(leukemiasClassifier, fileName="newPlots")
```

plotAssignments

Plot assignment probabilities

# **Description**

Plots the assignment probabilities of a previous query.

#### Usage

# **Arguments**

Numeric. Total number of classes the classifier was trained with. The assignment probability is determined bassed on it. It is not needed if there are samples of all the training classes.

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#### Value

Plot. If the plot is shown on the usual X11 or quartz devices, clicking on a point will identify the sample the point represents. Press ESC or right-click on the plot screen to finish.

#### See Also

```
Main package function and classifier training: geNetClassifier Querying the classifier: queryGeNetClassifier
```

## **Examples**

```
###########################
## Classifier training
#############################
library(geNetClassifier)
# Load an expressionSet:
library(leukemiasEset)
data(leukemiasEset)
# Select the train samples:
trainSamples<- c(1:10, 13:22, 25:34, 37:46, 49:58)
# summary(leukemiasEset$LeukemiaType[trainSamples])
# Train a classifier or load a trained one:
 leukemiasClassifier <- geNetClassifier(leukemiasEset[,trainSamples],</pre>
     sampleLabels="LeukemiaType", plotsName="leukemiasClassifier")
data(leukemiasClassifier) # Sample trained classifier
#############################
## External Validation:
###########################
# Select the samples to query the classifier
  - External validation: samples not used for training
testSamples <- c(1:60) [-trainSamples]</pre>
# Make a query to the classifier:
queryResult <- queryGeNetClassifier(leukemiasClassifier, leukemiasEset[,testSamples])</pre>
## Plot:
##############################
plotAssignments(queryResult, realLabels=leukemiasEset[,testSamples]$LeukemiaType)
```

```
plotDiscriminantPower
```

Plots the genes' Discriminant Power.

## **Description**

Calculates and plots the Discriminant Power of the genes in the given classifier.

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#### Usage

#### **Arguments**

classifier Classifier returned by geNetClassifier. (@classifier) classificationGenes Vector or Matrix. IDs of the genes to plot. If matrix: genes should be ordered by classes. Columns should be named after the classes. Vector or Matrix. Gene name, ID or label which should be shown in the returned geneLabels results and plots. Named vector. Short version of the class names if different from the ones used classNames to train the classifier. Logical. If TRUE, plots the discriminant power of the given genes. plotDP fileName Character. File name to save the plot with. If not provided, the plots will be shown through the standard output device. Logical. If TRUE, returns a table with the genes discriminant power. returnTable verbose Logical. If TRUE, messages indicating the execution progress will be printed on screen.

#### **Details**

The Discriminant Power represents the weight the (SVM) classifier gives each gene to separate the classes. It is calculated based on the coordinates of the support vectors. Genes with a high Discriminant Power are better for identifying samples from the class.

## Value

- Data frame Optional. Data.frame containing the genes and their Discriminant Power.
- Discriminant Power plot Optional. Shown throught the standard output devide or saved in the working directory as 'fileName.pdf' if fileName was provided.

## Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

## See Also

Main package function and classifier training: geNetClassifier

```
######
# Load data and train a classifier
######
library(geNetClassifier)
# Load an expressionSet:
```

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```
library(leukemiasEset)
data(leukemiasEset)
# Select the train samples:
trainSamples<- c(1:10, 13:22, 25:34, 37:46, 49:58)
# summary(leukemiasEset$LeukemiaType[trainSamples])
# Train a classifier or load a trained one:
# leukemiasClassifier <- geNetClassifier(leukemiasEset[,trainSamples],</pre>
     sampleLabels="LeukemiaType", plotsName="leukemiasClassifier")
data(leukemiasClassifier) # Sample trained classifier
######
# Discriminant Power
######
# Default (plots up to 20 genes)
plotDiscriminantPower(leukemiasClassifier)
# Plot a specific gene:
plotDiscriminantPower(leukemiasClassifier, classificationGenes="ENSG00000169575")
# Plot top5 genes of a class, and return their discriminant power:
# Note: The discriminant Power can only be calculated for 'classificationGenes'
             (genes chosen for training the classifier)
genes <- getRanking(leukemiasClassifier@classificationGenes,</pre>
    showGeneID=TRUE) $geneID[1:5, "AML", drop=FALSE] # Top 5 genes of AML
discPowerTable2 <- plotDiscriminantPower(leukemiasClassifier,</pre>
    classificationGenes=genes, returnTable=TRUE)
# For plotting more than 20 genes or saving the plots as .pdf, provide a fileName
plotDiscriminantPower(leukemiasClassifier,
     fileName="leukemiasClassifier_DiscriminantPower.pdf")
```

plotExpressionProfiles

Expression profiles plot.

# Description

Plots the expression profiles of the given genes.

#### Usage

# Arguments

eset ExpressionSet or Matrix. Gene expression of the samples.

genes Vector or Matrix. IDs of the genes to plot.

If matrix: genes should be ordered by classes. Columns should be named after

the classes. If not provided, all available genes will be plot.

fileName Character. File name to save the plots. If not provided, up to 20 genes will be

shown on screen.

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geneLabels Vector or Matrix. Gene name, ID or label which should be shown in the returned results and plots. sampleLabels Character. PhenoData variable (column name) containing the train samples class labels. Matrix or Factor. Class labels of the train samples. Vector or Factor. Order in which the labels should be shown in the returned labelsOrder results and plots. showSampleNames Logical. If TRUE, the sample names are shown in the plot. Not recommended for big datasets. showMean Logical. If TRUE, plots the class expression mean.

Logical. If TRUE, plots all the genes in the same expression scale. sameScale

sameScale

verbose Logical. If TRUE, messages indicating the execution progress will be printed

on screen.

#### Value

The expression profiles plot, saved in the working directory as 'fileName.pdf'.

# summary(leukemiasEset\$LeukemiaType[trainSamples])

## Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

```
######
# Load libraries and expression data
######
library(geNetClassifier)
# Load an expressionSet:
library(leukemiasEset)
data(leukemiasEset)
######
# Generic expression profile plot
######
# Default settings: Receiving only an expressionSet.
# Attempts to plot all available genes
# (If fileName is not provided, it shows up to 20 genes on screen)
plotExpressionProfiles(leukemiasEset)
# Plot expression of specific genes:
myGenes <- c("ENSG00000169575", "ENSG00000078399", "ENSG00000005381", "ENSG00000154511")
plotExpressionProfiles(leukemiasEset, genes=myGenes, sampleLabels="LeukemiaType")
######
# Expression profiles related to a classifier
######
# Train a classifier or load a trained one:
trainSamples<- c(1:10, 13:22, 25:34, 37:46, 49:58)
```

plotNetwork 37

plotNetwork

Plot GenesNetwork

#### **Description**

Plots the coexpression and/or mutual information network for the given genes.

#### Usage

```
plotNetwork(genesNetwork, classificationGenes=NULL, genesRanking=NULL,
genesInfo=NULL, geneLabels=NULL, returniGraphs=FALSE,
plotType="dynamic", fileName=NULL,
plotAllNodesNetwork=TRUE, plotOnlyConnectedNodesNetwork=FALSE,
plotClassificationGenesNetwork=FALSE,
labelSize=0.5, vertexSize=NULL, width=NULL, height=NULL, verbose=TRUE)
```

## Arguments

plotAllNodesNetwork

```
genesNetwork List of GenesNetwork returned by geNetClassifier. (@genesNetwork)
classificationGenes
                 Matrix or classificationGenes returned by geNetClassifier. (@classificationGenes)
genesRanking Matrix or genesRanking returned by geNetClassifier. (@genesRanking)
                 List or data frame with the properties of the genes to plot: genesDetails (_@genesRanking)
genesInfo
                 Vector or Matrix. Gene name, ID or label which should be shown in the returned
geneLabels
                 results and plots.
returniGraphs
                 Logical. If TRUE, returns a list with the plotted networks as igraph objects.
                 Character. "dynamic": Interactive plot. "static": One canvas split for the differ-
plotType
                 ent networks. "pdf": All the networks are saved into a pdf file.
                 Character. File name to save the plot with. If not provided, the plots will be
fileName
                 shown through the standard output device.
```

Logical. If TRUE, plots a network only with all the available genes

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```
plotOnlyConnectedNodesNetwork
```

Logical. If TRUE, plots a network only with the connected nodes/genes

plotClassifcationGenesNetwork

Logical. If TRUE, plots a network only with the classification genes

labelSize Integer. Gene/node label size for static and pdf plots.

vertexSize Integer. Vertex minimum size.

width Numeric. Dinamic or pdf plot width. height Numeric. Dinamic or pdf plot height.

verbose Logical. If TRUE, messages indicating the execution progress will be shown.

#### Value

```
Graph list List with the plotted igraph objects. Network plots
```

Shown throught the standard output devide or saved in the working directory as 'fileName.pdf' if fileName was provided.

#### Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

#### References

Main package function and classifier training: geNetClassifier

```
Package igraph
```

## See Also

plot. GenesNetwork () is an alias to this function. It can all so be called as i.e. plotNetwork (clGenesSubNet Note: The slot @genesNetwork returned by geNetClassifier is a List of GenesNetworks!

queryGeNetClassifier 39

queryGeNetClassifier

Queries the classifier trained with geNetClassifier.

#### **Description**

Queries the classifier trained by geNetClassifier in order to find out the class of new samples.

#### Usage

```
queryGeNetClassifier(classifier, eset, minProbAssignCoeff = 1,
    minDiffAssignCoeff = 0.8, verbose = TRUE)
```

#### **Arguments**

classifier Classifier returned by geNetClassifier. (@classifier)
eset ExpressionSet or Matrix. Gene expression matrix of the new samples.
minProbAssignCoeff

Numeric. Coefficient to modify the minimum probability requird to assign a sample to a class. Reduce to improve call rate. Increase to reduce error. 0: Removes this restriction. The sample will always be assigned to the class with the highest probability. between 0 and 1: Reduces the required probability to assign a sample to a class. >1: Increases the required probability. Warning: if minProbCoef is equal to 2\*number of classes, all the samples will be left as 'NotAssigned'.

 ${\tt minDiffAssignCoeff}$ 

Numeric. Coefficient to modify the required difference between the two most likelly classes. Reduce to improve call rate. Increase to reduce error. 0: Removes this restriction. The probability of the second most-likely class will not be taken into account. between 1 and 1: Reduces the required difference to assign the sample. >1: Increases the required difference. Warning: if minDiffAssignCoeff is equal to the number of classes, all the samples will be left as 'NotAssigned'.

verbose

Logical. If TRUE, messages indicating the execution progress will be printed on screen.

#### **Details**

By default, in order to assign a sample two contitions must be met:

- if minProbAssignCoeff = 1The probability of belonging to the class should be at least double of the random probability.
- if minDiffAssignCoeff = 0.8The difference of probabilities between the most likely class and the second most likely class should be more than 80

This means, that in a 4-class classifier, in order to assing a sample, the highest probability should be at least 0.5 (2x0.25), and the next most-likely-class should have a probability at least 0.2 (80 If these conditions are not met, the sample will be left as notAssigned.

Modify the arguments values in order to modify these assignment conditions. Setting minProbAssignCoeff = and minDiffAssignCoeff = 0 all samples will be assigned to the most likely class without any further restrictions.

#### Value

List:

- call Command used to execute the function.
- classes Classes to wich each of the samples were asigned to.
- probabilities Probabilities to the 2 classes each sample is most likely to belong to.

#### Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

#### See Also

```
Main package function and classifier training: geNetClassifier

Query summary: querySummary

External validation stats: externalValidation.stats and externalValidation.probMatrix
```

querySummary 41

```
############################
## Classifier Query
#############################
# Select the samples to query the classifier
# - Real use: samples whose class we want to known
querySamples <- "GSM330154.CEL"
  - External validation: samples not used for training
querySamples <- c(1:60) [-trainSamples]</pre>
#### Make a query to the classifier ("ask" about what class the new samples are):
queryResult <- queryGeNetClassifier(leukemiasClassifier, leukemiasEset[,querySamples])</pre>
# See the class it assigned to each sample:
queryResult$class[1:5]
# Or the samples which it wasn't sure about:
t(queryResult$probabilities[,queryResult$class=="NotAssigned"])
# Obtain an overview of the results
querySummary(queryResult)
#### Optional: Modify assignment conditions
# (minDiffCoef=0, minProbCoef=0: All samples will be assigned to the most likely class)
queryResult_AssignAll <- queryGeNetClassifier(leukemiasClassifier,</pre>
    leukemiasEset[,querySamples], minDiffAssignCoeff=0, minProbAssignCoeff=0)
# No samples are left as "NotAssigned":
queryResult$probabilities[,queryResult_AssignAll$class=="NotAssigned"]
#### External validation:
# Confusion matrix:
confMatrix <- table(leukemiasEset[,querySamples]$LeukemiaType,</pre>
    queryResult_AssignAll$class)
# New accuracy, call rate, sensitivity and specificity:
externalValidation.stats(confMatrix)
# Probability matrix for the assigned samples
externalValidation.probMatrix(queryResult, leukemiasEset[,querySamples]$LeukemiaType)
```

querySummary

Summary of the query.

# Description

Counts the number of samples assigned to each class and calculates basic statistics regarding the assignment probabilities.

## Usage

```
querySummary(queryResult, showNotAssignedSamples = TRUE, numDecimals = 2,
    verbose = TRUE)
```

#### **Arguments**

```
queryResult Object returned by queryGeNetClassifier
```

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```
showNotAssignedSamples
```

Logical. Shows the two most likely classes for the NotAssigned samples and

the probabilities of belonging to each of them.

 ${\tt numDecimals} \quad Integer. \ Number of \ decimals \ to \ show \ on \ the \ statistics.$ 

verbose Logical. If TRUE, messages indicating the execution progress will be printed on screen.

#### Value

Returns a list with the following fields:

- callRate Count and percentage of assigned samples.
- assigned Number of samples assigned to each class and mean and SD of the assignment probabilities.
- notAssignedSamples Optional. Most likely classes for the Not Assigned samples.

#### Author(s)

Bioinformatics and Functional Genomics Group. Centro de Investigacion del Cancer (CIC-IBMCC, USAL-CSIC). Salamanca. Spain

#### See Also

```
Main package function and classifier training: geNetClassifier Query the classifier: queryGeNetClassifier
```

```
###########################
## Classifier training
##########################
library(geNetClassifier)
# Load an expressionSet:
library(leukemiasEset)
data(leukemiasEset)
# Select the train samples:
trainSamples <- c(1:10, 13:22, 25:34, 37:46, 49:58)
# Train a classifier or load a trained one:
# leukemiasClassifier <- geNetClassifier(leukemiasEset[,trainSamples],</pre>
     sampleLabels="LeukemiaType", plotsName="leukemiasClassifier")
data(leukemiasClassifier) # Sample trained classifier
############################
## Classifier query
#############################
# Select the samples to query the classifier
  - Real use: samples whose class we want to known
querySamples <- "GSM330154.CEL"
  - External validation: samples not used for training
querySamples <- c(1:60) [-trainSamples]</pre>
# Make a query to the classifier:
queryResult <- queryGeNetClassifier(leukemiasClassifier, leukemiasEset[,querySamples])
```

setProperties-methods 43

```
{\it Set Properties} \\
```

# Description

Allows setting or modifiying the GenesRanking properties.

## Methods

```
setProperties(object, geneLabels=NULL, discriminantPower=NULL,
meanDif=NULL, isRedundant=NULL, gERankMean=NULL)
```

#### See Also

Main package function and classifier training: geNetClassifier This method's class (GenesRanking) help page.

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