

Package ‘Cascade’

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Suggests R.rsp, CascadeData, knitr

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Cascade-package	<i>The Cascade Package: Selection, Reverse-Engineering and Prediction in Cascade Networks</i>
-----------------	---

Description

A modeling tool allowing gene selection, reverse engineering, and prediction in cascade networks.

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014) <doi:10.1093/bioinformatics/btt705>.

Author(s)

This package has been written by Frédéric Bertrand, Myriam Maumy-Bertrand and Nicolas Jung with biological insights from Laurent Vallat. Maintainer: Frédéric Bertrand <frederic.bertrand@math.unistra.fr>

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

analyze_network, network-method
Analysing the network

Description

Calculates some indicators for each node in the network.

Usage

```
## S4 method for signature 'network'
analyze_network(Omega, nv, label_v = NULL)
```

Arguments

Omega	a network object
nv	the level of cutoff at which the analysis should be done
label_v	(optional) the name of the genes

Value

A matrix containing, for each node, its betweenness, its degree, its output, its closeness.

Author(s)

Nicolas Jung, Frédéric Bertrand, Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```
data(network)
analyze_network(network, nv=0)
```

as.micro_array	<i>Coerce a matrix into a micro_array object.</i>
----------------	---

Description

Coerce a matrix into a micro_array object.

Usage

```
as.micro_array(M, time, subject)
```

Arguments

M	A matrix. Contains the microarray measurements. Should of size $N * K$, with N the number of genes and $K=T*P$ with T the number of time points, and P the number of individuals. This matrix should be created using <code>cbind(M1,M2,...)</code> with $M1$ a $N*T$ matrix with the measurements for individual 1, $M2$ a $N*T$ matrix with the measurements for individual 2.
time	A vector. The time points measurements.
subject	The number of subjects.

Value

A micro_array object.

Author(s)

Nicolas Jung, Frédéric Bertrand , Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```

    if(require(CascadeData)){
    data(micro_US)
    micro_US<-as.micro_array(micro_US,time=c(60,90,210,390),subject=6)
    }

```

compare-methods	<i>Some basic criteria of comparison between actual and inferred network.</i>
-----------------	---

Description

Allows comparison between actual and inferred network.

Usage

```

## S4 method for signature 'network,network,numeric'
compare(Net, Net_inf, nv = 1)

```

Arguments

Net	A network object containing the actual network.
Net_inf	A network object containing the inferred network.
nv	A number that indicates at which level of cutoff the comparison should be done.

Value

A vector containing : sensibility, predictive positive value, and the F-score

Methods

```
list("signature(Net = \"network\", Net_inf = \"network\", nv = \"numeric\")")
```

Author(s)

Nicolas Jung, Frédéric Bertrand , Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```

data(Net)
data(Net_inf)

#Comparing true and inferred networks
F_score=NULL

#Here are the cutoff level tested
test.seq<-seq(0,max(abs(Net_inf@network*0.9)),length.out=200)
for(u in test.seq){
F_score<-rbind(F_score,Cascade::compare(Net,Net_inf,u))
}
matplot(test.seq,F_score,type="l",ylab="criterion value",xlab="cutoff level",lwd=2)

```

cutoff,network-method *Choose the best cutoff*

Description

Allows estimating the best cutoff, in function of the scale-freeness of the network. For a sequence of cutoff, the corresponding p-value is then calculated.

Usage

```

## S4 method for signature 'network'
cutoff(Omega, sequence = NULL, x_min = 0)

```

Arguments

Omega	a network object
sequence	(optional) a vector corresponding to the sequence of cutoffs that will be tested.
x_min	(optional) an integer ; only values over x_min are further retained for performing the test.

Value

A list containing two objects :

p.value	the p values corresponding to the sequence of cutoff
p.value.inter	the smoothed p value vector, using the loess function

Author(s)

Nicolas Jung, Frédéric Bertrand , Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```
data(network)
cutoff(network)
#See vignette for more details
```

dim	<i>Dimension of the data</i>
-----	------------------------------

Description

Dimension of the data

Usage

```
## S4 method for signature 'micro_array'
dim(x)
```

Arguments

x an object of class "micro-array"

Methods

list("signature(x = \"micro_array\")") Gives the dimension of the matrix of measurements.

Examples

```
if(require(CascadeData)){
  data(micro_US)
  micro_US<-as.micro_array(micro_US,time=c(60,90,210,390),subject=6)
  dim(micro_US)
}
```

 evolution, network-method

See the evolution of the network with change of cutoff

Description

See the evolution of the network with change of cutoff. This function may be usefull to see if the global topology is changed while increasing the cutoff.

Usage

```
## S4 method for signature 'network'
evolution(
  net,
  list_nv,
  gr = NULL,
  color.vertex = NULL,
  fix = TRUE,
  gif = TRUE,
  taille = c(2000, 1000),
  label_v = 1:dim(net@network)[1],
  legend.position = "topleft",
  frame.color = "black",
  label.hub = FALSE
)
```

Arguments

net	a network object
list_nv	a vector of cutoff at which the network should be shown
gr	a vector giving the group of each gene
color.vertex	a vector giving the color of each node
fix	logical, should the position of the node in the network be calculated once at the beginning ? Defaults to TRUE.
gif	logical, TRUE
taille	vector giving the size of the plot. Default to c(2000,1000)
label_v	(optional) the name of the genes
legend.position	(optional) the position of the legend, defaults to "topleft"
frame.color	(optional) the color of the frame, defaults to "black"
label.hub	(optional) boolean, defaults to FALSE

Value

A HTML page with the evolution of the network.

Author(s)

Nicolas Jung, Frédéric Bertrand , Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```
data(network)
sequence<-seq(0,0.2,length.out=20)
#setwd("inst/animation")
#evolution(network,sequence)
```

geneNeighborhood, network-method

Find the neighborhood of a set of nodes.

Description

Find the neighborhood of a set of nodes.

Usage

```
## S4 method for signature 'network'
geneNeighborhood(
  net,
  targets,
  nv = 0,
  order = length(net@time_pt) - 1,
  label_v = NULL,
  ini = NULL,
  frame.color = "white",
  label.hub = FALSE,
  graph = TRUE,
  names = FALSE
)
```

Arguments

net	a network object
targets	a vector containing the set of nodes
nv	the level of cutoff. Default to 0.
order	of the neighborhood. Default to 'length(net@time_pt)-1'.
label_v	vector defining the vertex labels.
ini	using the "position" function, you can fix the position of the nodes.
frame.color	color of the frames.
label.hub	logical ; if TRUE only the hubs are labeled.
graph	plot graph of the network. Defaults to 'TRUE'.
names	return names of the neighbors. Defaults to 'FALSE'.

Value

The neighborhood of the targeted genes.

Author(s)

Nicolas Jung, Frédéric Bertrand , Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```
data(Selection)
data(network)
#A nv value can chosen using the cutoff function
nv=.11
EGR1<-which(match(Selection@name, "EGR1")==1)
P<-position(network, nv=nv)

geneNeighborhood(network, targets=EGR1, nv=nv, ini=P,
label_v=network@name)
```

geneSelection	<i>Methods for selecting genes</i>
---------------	------------------------------------

Description

Selection of differentially expressed genes.

Usage

```
## S4 method for signature 'micro_array,micro_array,numeric'
geneSelection(
  x,
  y,
  tot.number,
  data_log = TRUE,
  wanted.patterns = NULL,
  forbidden.patterns = NULL,
  peak = NULL,
  alpha = 0.05,
  Design = NULL,
  lfc = 0
)

## S4 method for signature 'list,list,numeric'
geneSelection(
  x,
  y,
  tot.number,
  data_log = TRUE,
  alpha = 0.05,
  cont = FALSE,
  lfc = 0,
  f.asso = NULL
)

## S4 method for signature 'micro_array,numeric'
genePeakSelection(
  x,
  peak,
  y = NULL,
  data_log = TRUE,
  durPeak = c(1, 1),
  abs_val = TRUE,
  alpha_diff = 0.05
)
```

Arguments

x	either a <code>micro_array</code> object or a list of <code>micro_array</code> objects. In the first case, the <code>micro_array</code> object represents the stimulated measurements. In the second case, the control unstimulated data (if present) should be the first element of the list.
y	either a <code>micro_array</code> object or a list of strings. In the first case, the <code>micro_array</code> object represents the stimulated measurements. In the second case, the list is the way to specify the contrast: First element: condition, condition&time or pattern. The condition specification is used when the overall is to compare two conditions. The condition&time specification is used when comparing two conditions at two precise time points. The pattern specification allows to decide which time point should be differentially expressed. Second element: a vector of length 2. The two conditions which should be compared. If a condition is used as control, it should be the first element of the vector. However, if this control is not measured through time, the option <code>cont=TRUE</code> should be used. Third element: depends on the first element. It is no needed if condition has been specified. If condition&time has been specified, then this is a vector containing the time point at which the comparison should be done. If pattern has been specified, then this is a vector of 0 and 1 of length T, where T is the number of time points. The time points with desired differential expression are provided with 1.
tot.number	an integer. The number of selected genes. If <code>tot.number < 0</code> all differentially genes are selected. If <code>tot.number > 1</code> , <code>tot.number</code> is the maximum of differentially genes that will be selected. If <code>0 < tot.number < 1</code> , <code>tot.number</code> represents the proportion of differentially genes that are selected.
data_log	logical (default to TRUE); should data be logged ?
wanted.patterns	a matrix with wanted patterns [only for <code>geneSelection</code>].
forbidden.patterns	a matrix with forbidden patterns [only for <code>geneSelection</code>].
peak	integer. At which time points measurements should the genes be selected [optional for <code>geneSelection</code>].
alpha	float; the risk level. Default to 'alpha=0.05'
Design	the design matrix of the experiment. Defaults to 'NULL'.
lfc	log fold change value used in limma's 'topTable'. Defaults to 0.
cont	use contrasts. Defaults to 'FALSE'.
f.asso	function used to assess the association between the genes. The default value 'NULL' implies the use of the usual 'mean' function.
durPeak	vector of size 2 (default to <code>c(1,1)</code>); the first elements gives the length of the peak at the left, the second at the right. [only for <code>genePeakSelection</code>]
abs_val	logical (default to TRUE); should genes be selected on the basis of their absolute value expression ? [only for <code>genePeakSelection</code>]
alpha_diff	float; the risk level

Value

A micro_array object.

Author(s)

Nicolas Jung, Frédéric Bertrand , Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```

if(require(CascadeData)){
data(micro_US)
micro_US<-as.micro_array(micro_US,time=c(60,90,210,390),subject=6)
data(micro_S)
micro_S<-as.micro_array(micro_S,time=c(60,90,210,390),subject=6)

#Basically, to find the 50 more significant expressed genes you will use:
Selection_1<-geneSelection(x=micro_S,y=micro_US,
tot.number=50,data_log=TRUE)
summary(Selection_1)

#If we want to select genes that are differentially
#at time t60 or t90 :
Selection_2<-geneSelection(x=micro_S,y=micro_US,tot.number=30,
wanted.patterns=
rbind(c(0,1,0,0),c(1,0,0,0),c(1,1,0,0)))
summary(Selection_2)

#To select genes that have a differential maximum of expression at a specific time point.

Selection_3<-genePeakSelection(x=micro_S,y=micro_US,peak=1,
abs_val=FALSE,alpha_diff=0.01)
summary(Selection_3)
}

if(require(CascadeData)){
data(micro_US)
micro_US<-as.micro_array(micro_US,time=c(60,90,210,390),subject=6)
data(micro_S)
micro_S<-as.micro_array(micro_S,time=c(60,90,210,390),subject=6)
#Genes with differential expression at t1

```

```

Selection1<-geneSelection(x=micro_S,y=micro_US,20,wanted.patterns= rbind(c(1,0,0,0)))
#Genes with differential expression at t2
Selection2<-geneSelection(x=micro_S,y=micro_US,20,wanted.patterns= rbind(c(0,1,0,0)))
#Genes with differential expression at t3
Selection3<-geneSelection(x=micro_S,y=micro_US,20,wanted.patterns= rbind(c(0,0,1,0)))
#Genes with differential expression at t4
Selection4<-geneSelection(x=micro_S,y=micro_US,20,wanted.patterns= rbind(c(0,0,0,1)))
#Genes with global differential expression
Selection5<-geneSelection(x=micro_S,y=micro_US,20)

#We then merge these selections:
Selection<-unionMicro(list(Selection1,Selection2,Selection3,Selection4,Selection5))
print(Selection)

#Prints the correlation graphics Figure 4:
summary(Selection,3)

##Uncomment this code to retrieve geneids.
#library(org.Hs.eg.db)
#
#ff<-function(x){substr(x, 1, nchar(x)-3)}
#ff<-Vectorize(ff)
#
##Here is the function to transform the probeset names to gene ID.
#
#library("hgu133plus2.db")
#
#probe_to_id<-function(n){
#x <- hgu133plus2SYMBOL
#mp<-mappedkeys(x)
#xx <- unlist(as.list(x[mp]))
#genes_all = xx[(n)]
#genes_all[is.na(genes_all)]<-"unknown"
#return(genes_all)
#}
#Selection@name<-probe_to_id(Selection@name)
}

```

gene_expr_simulation, network-method

Simulates microarray data based on a given network.

Description

Simulates microarray data based on a given network.

Usage

```
## S4 method for signature 'network'  
gene_expr_simulation(network, time_label = 1:4, subject = 5, level_peak = 100)
```

Arguments

network	A network object.
time_label	a vector containing the time labels.
subject	the number of subjects
level_peak	the mean level of peaks.

Value

A micro_array object.

Author(s)

Nicolas Jung, Frédéric Bertrand , Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```
data(Net)  
set.seed(1)  
  
#We simulate gene expression according to the network Net  
Msim<-gene_expr_simulation(  
  network=Net,  
  time_label=rep(1:4,each=25),  
  subject=5,  
  level_peak=200)  
head(Msim)
```

head,micro_array-method

Overview of a micro_array object

Description

Overview of a micro_array object.

Usage

```
## S4 method for signature 'micro_array'  
head(x, ...)
```

Arguments

x an object of class 'micro_array'.
... additional parameters

Methods

list("signature(x = \"ANY\")") Gives an overview.

list("signature(x = \"micro_array\")") Gives an overview.

Examples

```
if(require(CascadeData)){  
  data(micro_US)  
  micro_US<-as.micro_array(micro_US,time=c(60,90,210,390),subject=6)  
  head(micro_US)  
}
```

inference,micro_array-method

Reverse-engineer the network

Description

Reverse-engineer the network.

Usage

```
## S4 method for signature 'micro_array'
inference(
  M,
  tour.max = 30,
  g = function(x) {
    1/x
  },
  conv = 0.001,
  cv.subjects = TRUE,
  nb.folds = NULL,
  eps = 10^-5,
  type.inf = "iterative"
)
```

Arguments

M	a micro_array object.
tour.max	maximal number of steps. Defaults to 'tour.max=30'
g	the new solution is chosen as (the old solution + g(x) * the new solution)/(1+g(x)) where x is the number of steps. Defaults to 'g=function(x) 1/x'
conv	convergence criterion. Defaults to 'conv=10e-3'
cv.subjects	should the cross validation be done removing the subject one by one ? Defaults to 'cv.subjects=TRUE'.
nb.folds	Relevant only if cv.subjects is FALSE. The number of folds in cross validation. Defaults to 'NULL'.
eps	machine zero. Defaults to '10e-5'.
type.inf	"iterative" or "noniterative" : should the algorithm be computed iteratively. Defaults to "'iterative"'

Value

A network object.

Author(s)

Nicolas Jung, Frédéric Bertrand , Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```
#With simulated data
data(M)
infM <- inference(M)
str(infM)

#With selection of genes from GSE39411
data(Selection)
infSel <- inference(Selection)
str(infSel)
```

M	<i>Simulated M data for examples.</i>
---	---------------------------------------

Description

Simulated M microarray.

Examples

```
data(M)
head(M)
```

micropredict-class	<i>Class "micropredict"</i>
--------------------	-----------------------------

Description

The "micropredict" class

Objects from the Class

Objects can be created by calls of the form `new("micropredict", ...)`.

Examples

```
showClass("micropredict")
```

micro_array-class	<i>Class "micro_array"</i>
-------------------	----------------------------

Description

The "micro_array" class

Objects from the Class

Objects can be created by calls of the form `new("micro_array", ...)`.

Examples

```
showClass("micro_array")
```

Net	<i>Simulated network data for examples.</i>
-----	---

Description

Simulated network.

Examples

```
data(Net)
str(Net)
```

network	<i>A network object data.</i>
---------	-------------------------------

Description

A network object. It is the same as the result in the vignette for the inference of the network.

Examples

```
data(network)
plot(network)
print(network)
```

network-class	<i>Class "network"</i>
---------------	------------------------

Description

The "network" class

Objects from the Class

Objects can be created by calls of the form `new("network", ...)`.

Examples

```
showClass("network")
```

network_random	<i>Generates a network.</i>
----------------	-----------------------------

Description

Generates a network.

Usage

```
network_random(
  nb,
  time_label,
  exp,
  init,
  regul,
  min_expr,
  max_expr,
  casc.level
)
```

Arguments

nb	Integer. The number of genes.
time_label	Vector. The time points measurements.
exp	The exponential parameter, as in the <code>barabasi.game</code> function in <code>igraph</code> package.
init	The attractiveness of the vertices with no adjacent edges. See <code>barabasi.game</code> function.
regul	A vector mapping each gene with its number of regulators.

min_expr	Minimum of strength of a non-zero link
max_expr	Maximum of strength of a non-zero link
casc.level	...

Value

A network object.

Author(s)

Nicolas Jung, Frédéric Bertrand , Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```
set.seed(1)
Net<-network_random(
  nb=100,
  time_label=rep(1:4,each=25),
  exp=1,
  init=1,
  regul=round(rexp(100,1))+1,
  min_expr=0.1,
  max_expr=2,
  casc.level=0.4
)
plot(Net)
```

Net_inf

Reverse-engineered network of the simulated data.

Description

The reverse-engineered network of the simulated data (M and Net).

Examples

```
data(Net_inf)
str(Net_inf)
```

plot-methods

Plot

Description

Considering the class of the argument which is passed to plot, the graphical output differs.

Usage

```
## S4 method for signature 'micro_array,ANY'
plot(x, y, ...)
```

```
## S4 method for signature 'network,ANY'
```

```
plot(
  x,
  y,
  choice = "network",
  nv = 0,
  gr = NULL,
  ini = NULL,
  color.vertex = NULL,
  video = TRUE,
  weight.node = NULL,
  ani = FALSE,
  taille = c(2000, 1000),
  label_v = 1:dim(x@network)[1],
  horiz = TRUE,
  legend.position = "topleft",
  frame.color = "black",
  label.hub = FALSE,
  ...
)
```

```
## S4 method for signature 'micropredict,ANY'
```

```
plot(
  x,
  time = NULL,
  label_v = NULL,
  frame.color = "white",
  ini = NULL,
  label.hub = FALSE,
```

```

    edge.arrow.size = 0.7,
    edge.thickness = 1
)

```

Arguments

<code>x</code>	a <code>micro_array</code> object, a network object or a <code>micropredict</code> object
<code>y</code>	optional and not used if <code>x</code> is an appropriate structure
<code>...</code>	additional parameters
<code>choice</code>	what graphic should be plotted: either "F" (for a representation of the matrices F) or "network".
<code>nv</code>	the level of cutoff. Default to '0'.
<code>gr</code>	a vector giving the group of each gene
<code>ini</code>	using the "position" function, you can fix the position of the nodes.
<code>color.vertex</code>	a vector defining the color of the vertex.
<code>video</code>	if <code>ani</code> is TRUE and <code>video</code> is TRUE, the result of the animation is saved as an animated GIF.
<code>weight.node</code>	nodes weighting. Defaults to 'NULL'.
<code>ani</code>	animated plot?
<code>taille</code>	vector giving the size of the plot. Default to 'c(2000,1000)'.
<code>label_v</code>	vector defining the vertex labels.
<code>horiz</code>	landscape? Defaults to 'TRUE'.
<code>legend.position</code>	position of the legend.
<code>frame.color</code>	color of the frames.
<code>label.hub</code>	logical ; if TRUE only the hubs are labeled.
<code>time</code>	sets the time for plot of the prediction. Defaults to 'NULL'
<code>edge.arrow.size</code>	size of the arrows ; default to 0.7.
<code>edge.thickness</code>	edge thickness ; default to 1.

Methods

list("signature(x = \"micro_array\", y = \"ANY\",...)) `x` a `micro_array` object

list_nv a vector of cutoff at which the network should be shown

list("signature(x = \"network\", y = \"ANY\",...)) `x` a network object

list() Optionnal arguments:

gr a vector giving the group of each gene

choice what graphic should be plotted: either "F" (for a representation of the matrices F) or "network".

nv the level of cutoff. Defaut to 0.

ini using the "position" function, you can fix the position of the nodes

color.vertex a vector defining the color of the vertex
ani animated plot?
size vector giving the size of the plot. Default to c(2000,1000)
video if ani is TRUE and video is TRUE, the animation result is a GIF video
label_v vector defining the vertex labels
legend.position position of the legend
frame.color color of the frames
label.hub logical ; if TRUE only the hubs are labeled
edge.arrow.size size of the arrows ; default to 0.7
edge.thickness edge thickness ; default to 1.

list("signature(x = \"micropredict\", y = \"ANY\",...)\") x a micropredict object

list() Optionnal arguments: see plot for network

Examples

```
data(Net)
plot(Net)
```

```
data(M)
plot(M)
```

```
data(Selection)
data(network)
nv<-0.11
plot(network,choice="network",gr=Selection@group,nv=nv,label_v=Selection@name,
edge.arrow.size=0.9,edge.thickness=1.5)
```

position-methods

Returns the position of edges in the network

Description

Returns the position of edges in the network

Usage

```
## S4 method for signature 'network'
position(net, nv = 0)
```

Arguments

net a network object
nv the level of cutoff at which the analysis should be done

Methods

list("signature(net = \"network\")") Returns a matrix with the position of the node. This matrix can then be used as an argument in the plot function.

Examples

```
data(Net)
position(Net)
```

predict,micro_array-method

Prediction of the gene expressions after a knock-out experience
predict

Description

Prediction of the gene expressions after a knock-out experience

Usage

```
## S4 method for signature 'micro_array'
predict(object, Omega, nv = 0, targets = NULL, adapt = TRUE)
```

Arguments

object	a micro_array object
Omega	a network object.
nv	[=0] numeric; the level of the cutoff
targets	[NULL] vector; which genes are knocked out?
adapt	[TRUE] boolean; do not raise an error if used with vectors instead of one column matrices.

Author(s)

Nicolas Jung, Frédéric Bertrand , Myriam Maumy-Bertrand.

References

Jung, N., Bertrand, F., Bahram, S., Vallat, L., and Maumy-Bertrand, M. (2014). Cascade: a R-package to study, predict and simulate the diffusion of a signal through a temporal gene network. *Bioinformatics*, btt705.

Vallat, L., Kemper, C. A., Jung, N., Maumy-Bertrand, M., Bertrand, F., Meyer, N., ... & Bahram, S. (2013). Reverse-engineering the genetic circuitry of a cancer cell with predicted intervention in chronic lymphocytic leukemia. *Proceedings of the National Academy of Sciences*, 110(2), 459-464.

Examples

```
data(Selection)
data(network)
#A nv value can chosen using the cutoff function
nv=.11
EGR1<-which(match(Selection@name,"EGR1")==1)
P<-position(network,nv=nv)

#We predict gene expression modulations within the network if EGR1 is experimentaly knocked-out.
prediction_ko5<-predict(Selection,network,nv=nv,targets=EGR1)

#Then we plot the results. Here for example we see changes at time point t2:
plot(prediction_ko5,time=2,ini=P,label_v=Selection@name)
```

print-methods

Methods for Function print

Description

Methods for function print

Usage

```
## S4 method for signature 'micro_array'
print(x, ...)

## S4 method for signature 'network'
print(x, ...)
```

Arguments

x an object of class micro-array or network
... additional parameters

Examples

```
data(Net)
print(Net)

data(M)
print(M)
```

Selection	<i>Selection of genes.</i>
-----------	----------------------------

Description

20 (at most) genes with differential expression at t1, 20 (at most) genes with differential expression at t2, 20 (at most) genes with differential expression at t3, 20 (at most) genes with differential expression at t4 et 20 (at most) genes with global differential expression were selected.

Examples

```
data(Selection)
head(Selection)
summary(Selection,3)
```

summary-methods	<i>Methods for Function summary</i>
-----------------	-------------------------------------

Description

Methods for function summary

Usage

```
## S4 method for signature 'micro_array'
summary(object, nb.graph = NULL, ...)
```

Arguments

object	an object of class micro-array
nb.graph	(optionnal) choose the graph to plot. Displays all graphs by default.
...	additional parameters.

Examples

```
data(M)
summary(M)
```

unionMicro-methods *Makes the union between two micro_array objects.*

Description

Makes the union between two micro_array objects.

Usage

```
## S4 method for signature 'micro_array,micro_array'  
unionMicro(M1, M2)
```

Arguments

M1 a micro-array or a list of micro-arrays
M2 a micro-array or nothing if M1 is a list of micro-arrays

Methods

list("signature(M1 = \"micro_array\", M2 = \"micro_array\")") Returns a micro_array object which is the union of M1 and M2.

list("signature(M1 = \"list\", M2 = \"ANY\")") Returns a micro_array object which is the union of the elements of M1.

Examples

```
data(M)  
#Create another microarray object with 100 genes  
Mbis<-M  
#Rename the 100 genes  
Mbis@name<-paste(M@name,"bis")  
rownames(Mbis@microarray) <- Mbis@name  
#Union (merge without duplicated names) of the two microarrays.  
str(unionMicro(M,Mbis))
```

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