# Package 'rexposome'

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# Contents

classification	3
clustering	4
correlation	5
expo	6
expos	7
ExposomeClust	
ExposomeCorr	8
ExposomePCA	10
ExposomeSet	12
exposureNames	17
expo_c	18
extract	
ExWAS	
exwas	
ex_imp	
familyNames	
get_robust_sd	
highAndLow	
ilod	
imExposomeSet	
imputation	
imputeLOD	
invExWAS	
loadExposome	
loadExposome_plain	
loadImputed	
me	
mExWAS	
mexwas	
ndim	
normalityTest	
pca	
phenotypeNames	
plot3PCA	
plotClassification	
plotCorrelation	
plotEffect	
plotEXP	46
plotExwas	47
plotFamily	48
plotHistogram	49
plotLOD	50
plotMissings	51
plotPCA	52
plotPHE	53
plotVolcano	
1	

#### classification

readExposome	54
rexposome	56
standardize	57
tableLOD	58
tableMissings	59
tef	60
toES	61
trans	61
volcano_plot	62
	64

# Index

classification Method to get the classification of the samples from an ExposomeClust.

# Description

Method to get the classification of the samples from an ExposomeClust.

#### Usage

```
classification(object)
```

# Arguments

object An ExposomeClust to get the samples' classification.

#### Value

A labelled vector with the classification of each exposure.

# See Also

clustering as a constructor for ExposomeClust, plotClassification to plot the groups

# Examples

```
data("eclust")
tt <- classification(expo_c)
table(tt)</pre>
```

#### clustering

#### Description

This method allows to create an ExposomeClust object from an ExposomeSet object by clustering samples through the exposure levels. The method is flexible to accept any clustering method (method) that allows to obtain a classification (cmethod) of the samples. The function assigned to argument method must have an argument called data, waiting for the matrix of exposures (samples as rows, exposures as columns). If the result object of the method has no accessor \$classification, then a cmethod is required and will be applied on the result of method to obtain a labelled vector with the samples' classification.

#### Usage

```
clustering(object, method, cmethod, ..., warnings = TRUE)
```

#### Arguments

object	ExposomeSet containing the exposures used for the clustering process
method	Function applied to the exposures of object. This function must has an argument named as data that will receive the matrix of exposures.
cmethod	(optional) Function to obtain the classification from the object generated with method.
	Passed to content of method.
warnings	(default TRUE) If set to FALSE warnings will not be displayed.

#### Value

ExposomeClust with the original exposures and the classification of each exposure.

#### Note

The function assigned to cmethod will be directly applied to the result of the method as: cmethod(model); being model the result of method.

#### See Also

classification to see how to obtain the classification of the samples from an ExposomeClust, plot-Classification to plot the groups

#### correlation

#### Examples

```
data("exposome")
# EXAMPLE 1: Clustering with mclust
library(mclust)
c <- clustering(expo[12:32, ], method = Mclust, G = 2)</pre>
table(classification(c)) # This works since the result of Mclust has an accessor
                   # $classification
# EXAMPLE 2: Cluseting with flexmix
library(flexmix)
# First we carete a function to apply flexmix to the ExposomeSet
flexmix_clust <- function(data, ...) {</pre>
  data <- as.matrix(data)</pre>
  flexmix(formula = data~1, ...)
}
# Then if we apply the method to the ExposomeSet it will crash:
# c <- clustering(expo[12:32, ], method = flexmix_clust, k = 2, model = FLXMCmvnorm())</pre>
# Because the method does not know how to obtain the classification for the result
# since flexmix has not an accessor called $classiciation
# We create a function to get the classification
flexmix_clas <- function(model, ...) {</pre>
  return(clusters(model))
}
# We put it to the ExposomeClust
c <- clustering(expo[12:32, ], method = flexmix_clust, cmethod = flexmix_clas,</pre>
    k = 2, model = FLXMCmvnorm())
classification(c) # This works because the ExposomeClust has a way to get
                  # the classification
```

```
correlation
```

Creation of an ExposomeCorr from an ExposomeSet.

#### Description

Method to calculate the correlation between the exposures of an ExposomeSet. The correlation method takes into account the nature of each pair of exposures: continuous vs. continuous uses cor function from R base, categorical vs. categorical uses cramerV function from lsr R package and categorical vs. continuous exposures correlation is calculated as the square root of the adjusted r-square obtained from fitting a lineal model with the categorical exposures as dependent variable and the continuous exposure as independent variable. The function creates and returns an ExposomeCorr object.

#### Usage

```
correlation(object, ..., warnings = TRUE)
```

#### Arguments

object	ExposomeSet which exposures will be used to calculate their correlation
	Other arguments passed to cor, cramersV or to lm.
warnings	(default TRUE) If set to FALSE warnings will not be displayed.

# Value

ExposomeCorr with the correlation between the selected exposures and their description

#### See Also

plotCorrelation to plot the correlations of an ExposomeCorr, clustering to see how the exposures can cluster samples, pca to compute PCA on exposures

# Examples

```
data("exposome")
expo.c <- correlation(expo)
expo.c
expo.c.table <- extract(expo.c)</pre>
```

expo

ExposomeSet for testing purpouses

#### Description

ExposomeSet with full set of 104 exposures, 1200 simulated samples and 4 phenotypes (asthma status, BMI measure, sex, and age).

# Usage

data("exposome")

#### Format

An object of class ExposomeSet of dimension 104 x 1200 x 4.

#### Value

An ExposomeSet object.

#### Examples

```
data("exposome")
dim(expo)
exposureNames(expo)
familyNames(expo)
sampleNames(expo)
phenotypeNames(expo)
```

expos

#### Description

Given an ExposomeSet it returns the inner matrix of exposures, having the exposures as columns and the samples as rows.

#### Usage

expos(object)

#### Arguments

object An ExposomeSet.

#### Value

A matrix of exposures

#### Examples

```
data("exposome")
expos(expo)[1:3, 1:3]
```

ExposomeClust Class ExposomeClust

#### Description

Class ExposomeClust obtained from clustering on an ExposomeSet object, represents the groups of samples created applying a clustering method on the ExposomeSet' exposures.

# Usage

```
## S4 method for signature 'ExposomeClust,ANY'
plot(x, y, ...)
## S4 method for signature 'ExposomeClust'
classification(object)
## S4 method for signature 'ExposomeClust'
plotClassification(object, type = "heatmap", ...)
## S4 method for signature 'ExposomeClust'
sampleNames(object)
```

#### Arguments

х	Object of class ExposomeClust
У	NOT USED
	Argument given to heatmap.2
object	An object of class ExposomeClust
type	(default "heatmap") Type of plot.

# Value

An object of class ExposomeClust

#### Methods (by generic)

- plot(x = ExposomeClust, y = ANY): Wrapper for plotClassification method.
- classification(ExposomeClust): Return classe asigned to each sample
- plotClassification(ExposomeClust): Draws a heatmap for the samples' classification.
- sampleNames(ExposomeClust): Method to obtain samples' names

#### Slots

model Result obtained on applying method on the exposures.

method Function used to perform the clustering of the exposures.

call Call used to create this object.

samples Name of the exposures after the clustering process.

#### See Also

clustering to apply a clustering on an ExposomeSet and create an ExposomeClust.

ExposomeCorr Class ExposomeCorr

# Description

Class ExposomeCorr contains a matrix of correlations between continuos exposures calculated using cor. It also contains the description of each exposures (fData of the original ExposomeSet) in order to maintain a coherence with the original source. It extends eSet-class.

# ExposomeCorr

#### Usage

```
## S4 method for signature 'ExposomeCorr,ANY'
plot(x, y, ...)
## S4 method for signature 'ExposomeCorr'
dim(x)
## S4 method for signature 'ExposomeCorr'
extract(object, sort, ...)
## S4 method for signature 'ExposomeCorr'
plotCorrelation(object, type = c("circos", "matrix"), ...)
```

#### Arguments

х	Object of class ExposomeCorr
У	NOT USED
	Arguments passed to corrplot when type="matrix".
object	An ExposomeCorr object.
sort	NOT USED
type	To choose between "circos" and "matrix".

#### Value

An object of class ExposomeCorr

#### Methods (by generic)

- plot(x = ExposomeCorr, y = ANY): Wrapper for plotClassification method.
- dim(ExposomeCorr): Return the dimension of the internat matrix of correlation.
- extract(ExposomeCorr): Return the raw correlation matrix
- plotCorrelation(ExposomeCorr): Draws both a matrix of circos plot of correlations

# Slots

assayData Contains the correlation matrix (see eSet, AssayData).

featureData Contains the description of the exposures including the family where they belong (see eSet, AnnotatedDataFrame).

#### See Also

pca to study the behavioud between samples and exposures in an ExposomeSet

ExposomePCA

#### Description

Class ExposomePCA contains a matrix of exposures used to compute the PCA, also a table of phenotypes and a set congtaing the multiple results of computing the PCA.

#### Usage

```
## S4 method for signature 'ExposomePCA,ANY'
plot(x, y, ...)
## S4 method for signature 'ExposomePCA'
exposureNames(object)
## S4 method for signature 'ExposomePCA'
extract(object, table = "exposures", ...)
## S4 method for signature 'ExposomePCA'
ndim(object)
## S4 method for signature 'ExposomePCA'
phenotypeNames(object)
## S4 method for signature 'ExposomePCA'
plot3PCA(
  object,
  cmpX,
  cmpY,
  cmpZ,
  phenotype,
 main,
  angle = 35,
  pch = 16,
  legend = TRUE,
  plines = TRUE
)
## S4 method for signature 'ExposomePCA'
plotEXP(object, exposure)
## S4 method for signature 'ExposomePCA'
plotPCA(
  object,
  set,
  cmpX = 1,
```

# ExposomePCA

```
cmpY = 2,
show.exposures = FALSE,
show.samples = FALSE,
phenotype
)
## S4 method for signature 'ExposomePCA'
plotPHE(object, phenotype, exp2fac = 5)
```

# Arguments

x	Object of class ExposomePCA
У	NOT USED
	NOT USED
object	An ExposomePCA object
table	Can takes values "exposures" or "individuals".
стрХ	PC to place in X-axis
cmpY	PC to place in Y-axis
cmpZ	PC to place in Z-axis
phenotype	(when set="samples") A phenotype can be selected so the samples are coloured by its value.
main	Title
angle	Point of view
pch	Size of the dots
legend	Boolean to show or hide the legend
plines	Boolean to show of hide the dotted lines that helps to place the dots in the X/Y axes
exposure	Vector of exposures to be shown in the plot
set	Can take values "exposures", "samples" or "all"
show.exposures	When set to TRUE, labels for exposures are shown
show.samples	When set to TRUE, labels for samples are shown
exp2fac	Number of different values to considere an exposures continuous

# Value

An object of class ExposomePCA

# Methods (by generic)

- plot(x = ExposomePCA, y = ANY): Wrapper for plotPCA method.
- exposureNames(ExposomePCA): Getter to obtain the exposures's names.
- extract(ExposomePCA): Method to extract the raw results of the PCA.
- ndim(ExposomePCA): Number of principal components in an ExposomePCA.

- phenotypeNames(ExposomePCA): Getter to obtain the phenotype's names.
- plot3PCA(ExposomePCA): Methdo to draw a 3D plot for PCA
- plotEXP(ExposomePCA): Plot correlation between exposures and PCA
- plotPCA(ExposomePCA): Methdo to draw a 2D plot for PCA
- plotPHE(ExposomePCA): Plot association score between phentoypes and PCA

#### Slots

pca list containing all elements of the PCA

phenoData Contains the phenotypes or variables experimenter-supplied (see eSet, AnnotatedDataFrame).

featureData Contains the description of the exposures including the family where they belong (see eSet, AnnotatedDataFrame).

#### See Also

correlation to study the correlation between exposures in a ExposomeSet

ExposomeSet

Class ExposomeSet

#### Description

Class ExposomeSet contains the exposure levels, the exposure's description and the samples phenotype. It is the starting object for rexposome package and extends eSet.

#### Usage

```
## S4 method for signature 'ExposomeSet,ANY'
plot(x, y, ...)
## S4 method for signature 'ExposomeSet'
clustering(object, method, cmethod, ..., warnings = TRUE)
## S4 method for signature 'ExposomeSet'
correlation(object, ..., warnings = TRUE)
## S4 method for signature 'ExposomeSet'
dim(x)
## S4 method for signature 'ExposomeSet'
expos(object)
## S4 method for signature 'ExposomeSet'
exposureNames(object)
```

# ExposomeSet

```
## S4 method for signature 'ExposomeSet'
exwas(
  object,
  formula,
  filter,
  family,
  . . . ,
  baselevels,
  tef = TRUE,
  verbose = FALSE,
 warnings = TRUE,
  robust = FALSE
)
## S4 method for signature 'ExposomeSet'
familyNames(object, by.exposure = FALSE)
## S4 method for signature 'ExposomeSet'
highAndLow(
  object,
  ngroups = 3,
  intervals = c("standard", "extreme"),
  select,
  drop = FALSE,
 warnings = TRUE
)
## S4 method for signature 'ExposomeSet'
ilod(
  object,
  seed = 1234,
  lod.col = "LOD",
  pNA = 0.2,
  tLog = FALSE,
 method = "QRILC",
 warnings = TRUE,
  . . .
)
## S4 method for signature 'ExposomeSet'
imputation(object, select, ..., messages = FALSE)
## S4 method for signature 'ExposomeSet'
invExWAS(object, formula, filter, tef = TRUE, verbose = FALSE, warnings = TRUE)
## S4 method for signature 'ExposomeSet'
mexwas(object, phenotype, family, warnings = TRUE)
```

```
## S4 method for signature 'ExposomeSet'
normalityTest(
  object,
  exposure,
  th = 0.05,
 min.val = 5,
 na.rm = TRUE,
 warnings = TRUE
)
## S4 method for signature 'ExposomeSet'
pca(object, npc = 10, pca = FALSE, ...)
## S4 method for signature 'ExposomeSet'
phenotypeNames(object)
## S4 method for signature 'ExposomeSet'
plotFamily(x, family, group, group2, scatter = TRUE, na.omit = TRUE)
## S4 method for signature 'ExposomeSet'
plotHistogram(x, select, density = TRUE, show.trans = FALSE)
## S4 method for signature 'ExposomeSet'
plotLOD(object, lod.col = "LOD", x.max = 100, sort = TRUE)
## S4 method for signature 'ExposomeSet'
plotMissings(
 object,
  set = c("exposures", "phenotypes"),
 x.max = 100,
  sort = TRUE
)
## S4 method for signature 'ExposomeSet'
standardize(object, select, method = "normal", na.rm = TRUE, warnings = TRUE)
## S4 method for signature 'ExposomeSet'
Summary(x, set = c("exposures", "phenotypes"), select, ..., na.rm = FALSE)
## S4 method for signature 'ExposomeSet'
tableLOD(object, output = "n", lod.col = "LOD", sort = TRUE)
## S4 method for signature 'ExposomeSet'
tableMissings(
  object,
  set = c("exposures", "phenotypes"),
  output = "n",
  sort = TRUE
```

# ExposomeSet

# )

```
## S4 method for signature 'ExposomeSet'
trans(object, fun, select, by.exposure = FALSE, ...)
```

# Arguments

9	
x	An ExposomeSet object.
У	NOT USED
	Arguments to be passed to imputeFAMD
object	An ExposomeSet object.
method	Method to be used.
cmethod	Function implementing a systsme to retrieve classification from clustering out- put
warnings	If set to TRUE it prints the warning messsages.
formula	Formula, not including exposures, to be tested. No need to provide response (left term)
filter	Expression to be used to filter ExposomeSet
family	Family descriving the nature of the health outcome
baselevels	Labeled vector with the default base level for categorical exposures.
tef	If TRUE it computed the threshold for effective tests.
verbose	If set to TRUE is shows messages on progression.
by.exposure	If set to TRUE ir returns the family which each exposure belongs
ngroups	Number of intervals to be used
intervals	If "standard" all levels are kept, if "extreme" intermetiate levels are set to NA.
select	Vector selecting thee xposures to be used.
drop	If set to TRUE exposures are replaced
seed	Numeric seed
lod.col	Indicator of the column where the LOD is located
pNA	Maximum percentage allowed of values under LOD
tLog	If set to TRUE it transforms all the exposures to lod before the imputation.
messages	If set to TRUE messages from mice's function will be displayed.
phenotype	Health outcome to be used as dependent variable.
exposure	Vecror of exposures to be used.
th	Threshold of P-Value used to considere normalit
min.val	Minimum number of observations to perform test
na.rm	If set to TRUE removes NA values
npc	Number of PC to be kept
рса	Perform PCA (only numerical variables) or FAMD (numerical and categorical)

group	Phenotype to group exposures
group2	Phenotype to group exposures
scatter	If set to true it shows the samples value in the plot
na.omit	If set to TRUE, NA values are discarded
density	If set to TRUE a desntiry plot is draw on the histogram
show.trans	If set to TRUE, three extra plots are drawn with usual transformations
x.max	Threshold for x axis (in %)
sort	If set to TRUE, results are ordered
set	Cantake values "exposures" or "phentoypes".
output	Can take values "n" (count) ot "p" (percentage)
fun	Function to bt used in the transformation process

#### Value

An object of class ExposomeSet

#### Methods (by generic)

- plot(x = ExposomeSet, y = ANY): Wrapper for plotFamily method.
- clustering(ExposomeSet): Performs clustering on samples based on exposure levels.
- correlation(ExposomeSet): Computes correlation on exposures.
- dim(ExposomeSet): Returns the number of exsures, samples and phenotypes.
- expos(ExposomeSet): Returns a data.frame with exposures.
- exposureNames(ExposomeSet): Getter to obtain the exposures's names.
- exwas(ExposomeSet): Performs an EXposome-Wide Association Study
- familyNames(ExposomeSet): Getter to obtain the families's names of the family of each exposure.
- highAndLow(ExposomeSet): Performs a discretization of continuous exposures.
- ilod(ExposomeSet): Imputation of under-LOD values of exposures.
- imputation(ExposomeSet): Imputation of missing values of exposures.
- invExWAS(ExposomeSet): Performs an EXposome-Wide Association Study (modelling the exposures as response)
- mexwas(ExposomeSet): Performs a Multiple-EXposure-Wide Association Study.
- normalityTest(ExposomeSet): Test the normality of each exposure.
- pca(ExposomeSet): Performs a PCA
- phenotypeNames(ExposomeSet): Getter to obtain the phenotypes's names.
- plotFamily(ExposomeSet): Draws a boxplot or accumulated-bar plot for each exposure in a given family.
- plotHistogram(ExposomeSet): Draws an histogram of a given continuous exposure or a pie chart if a given categorycal exposure.

#### exposureNames

- plotLOD(ExposomeSet): Draws a barchart with the amount of under-LOD values.
- plotMissings(ExposomeSet): Draws a bar-plot with the amount of missing values.
- standardize(ExposomeSet): Standardization of exposures.
- Summary(ExposomeSet): Summary of both continuous and categorical exposures
- tableLOD(ExposomeSet): Returns a vector with the number of under-LOD values per exposure.
- tableMissings(ExposomeSet): Returns a vector with the number of missing values per exposure.
- trans(ExposomeSet): Transformation of exposures.

#### Slots

- assayData Contains the exposures matrix with column number equal to nrow(phenoData) (see eSet, AssayData).
- featureData Contains the description of the exposures including the family where they belong (see eSet, AnnotatedDataFrame).
- phenoData Contains the phenotypes or variables experimenter-supplied (see eSet, AnnotatedDataFrame).

#### See Also

readExposome to create an ExposomeSet from files, loadExposome to create an ExposomeSet from
data.frames

exposureNames	Getter to obtain the exposures's names of an ExposomeSet or Expo-
	somePCA

#### Description

Getter to obtain the exposures's names of an ExposomeSet or ExposomePCA

#### Usage

```
exposureNames(object)
```

#### Arguments

object ExposomeSet that will be queried for the exposures's names.

#### Value

The name of the exposures as a character vector.

#### Warning

exposureNames collides with featureNames of eSet. Although in rexposome 1.0.0 both function can be used as synonyms, this usage is discouraged and it is not assured.

# See Also

phenotypeNames to get the phenotypes, familyNames to get the families of exposures

# Examples

```
data("exposome")
exposureNames(expo)
```

expo\_c

#### ExposomeClust for testing purpouses

# Description

ExposomeClust created from an ExposomeSet with full set of 104 exposures, 1200 simulated samples and 4 phenotypes (asthma status, BMI measure, sex, and age). The clustering was done using hclust and cutree with k = 3.

#### Usage

data("eclust")

### Format

An object of class ExposomeClust of dimension 99 x 1200 x 5.

# Value

An ExposomeSet object.

# Examples

```
data("eclust")
dim(expo_c)
table(classification(expo_c))
```

extract

#### Description

Returns internal table of results of objects of class ExWAS, ExposomeClust and ExposomeCorr.

#### Usage

```
extract(object, ...)
```

#### Arguments

object	Object of class ExWAS, ExposomeClust or ExposomeCorr
	NO USED

# Value

A data.frame containing the raw result from PsyGeNET or a data.frame with the result Jaccard Index for each disease.

ExWAS

Class ExWAS

# Description

Class ExWAS obtained from exwas method of an ExposomeSet object, contains the result of testing the association of exposures of an ExposomeSet to its phenotypes. "ExWAS" is the acronym of "Exposome-Wide Association Study". The function can be applied to one of to many phenotypes in the ExposomeSet object.

#### Usage

```
## S4 method for signature 'ExWAS,ANY'
plot(x, y, ...)
## S4 method for signature 'ExWAS'
extract(object, sort = TRUE, ...)
## S4 method for signature 'ExWAS'
get_robust_sd(object, sort = TRUE, ...)
## S4 method for signature 'ExWAS'
names(x)
```

```
## S4 method for signature 'ExWAS'
plotEffect(x, y, select, labels, xlab, ylab)
## S4 method for signature 'ExWAS'
plotExwas(
 object,
  ...,
 subtitles,
 color,
 exp.order,
 labels,
 show.effective = TRUE
)
## S4 method for signature 'ExWAS'
plotVolcano(x, p.value = -log10(0.001), labels, show.effect = FALSE)
## S4 method for signature 'ExWAS'
tef(object)
```

# Arguments

Х	An ExWAS object
У	An ExWAS object
	NOT USED
object	An object of class ExWAS, mExWAS or nlExWAS.
sort	If TRUE, the results are ordered by pvalue.
select	(optional) Vector with the selected exposures
labels	(optional) Character vector with the labels for each exposure. It must be labeled vector.
xlab	(optional) Label for x-axis
ylab	(optional) Label for y-axis
subtitles	Character vector with the "subtitle" for each plot when given more than one ExWAS.
color	(optional) A vector of colors. The vector must have length equal to the number of families. The vector must be names with the name of the families.
exp.order	(optional) Order of the exposures.
show.effective	(default TRUE) draws a brown line on the threshold given by the effective number of tests.
p.value	(default $-\log 10(0.001)$ ) Is the threshold from where the exposures can be taken as significants.
show.effect	(default false) Apply exp to obtained beta.

#### Value

An object of class ExWAS

#### exwas

#### Methods (by generic)

- plot(x = ExWAS, y = ANY): Wrapper for plotExwas method.
- extract(ExWAS): Method to obtain the matrix of association scores
- get\_robust\_sd(ExWAS): Method to obtain the matrix of association scores
- names(ExWAS): Method to obtain the Threshold for effective tests (TEF)
- plotEffect(ExWAS): Draws a plot with the confidence interval of each exposure. Alows to compare two ExWAS instances.
- plotExwas(ExWAS): Method to plot a manhatan plot for association between exposures and phenitypes
- plotVolcano(ExWAS): Method to plot a volcano plot for association between exposures and phenotypes
- tef(ExWAS): Method to obtain the Threshold for effective tests (TEF)

#### Slots

effective Number containing the effective number of tests.

formula Tested formula.

comparison Result of performing the test to find association between levels of exposures and phenotype.

description Description of the exposures used in the ExWAS. (in description file).

#### See Also

exwas to perform an Exposome-Wide Association Study and to create an ExWAS, mexwas to perform a Multivariate Exposome-Wide Association Study and to create a mExWAS

Testing the association between an exposure and a phenotype of an *ExposomeSet* 

#### Description

The exwas method performs an "Exposome-Wide Association Study" (ExWAS) using the exposures in ExposomeSet and one of its phenotype.

#### Usage

exwas( object, formula, filter, family, ..., baselevels,

```
tef = TRUE,
verbose = FALSE,
warnings = TRUE,
robust = FALSE
)
```

# Arguments

object	ExposomeSet that will be used for the ExWAS.
formula	formula indicating the test to be done. If any exposure is included it will be used as covariate. exwas metho will perform the test for each exposure.
filter	expression to be used to filter the individuals included into the test.
family	Family of the distribution followed by the health outcome to be tested (gaussian, bionomal, check glm).
	NOT USED
baselevels	(optional) If set, must be a labeled vector with the default base level for categor- ical exposures.
tef	(default TRUE) If TRUE it computed the effective number of tests and the threhold for the effective number of tests. Usually it needs imputed data.
verbose	(default FALSE) If set o true messages along the tests are shown.
warnings	(default TRUE) If set to FALSE warnings will not be displayed.

#### Value

An codeExWAS object with the result of the association study

#### References

An Environment-Wide Association Study (ExWAS) on Type 2 Diabetes Mellitus. Chirag J. Patel, Jayanta Bhattacharya, Atul J. Butte. May 20, 2010 Plos One

Evaluating the effective numbers of independent tests and significant p-value thresholds in commercial genotyping arrays and public imputation reference datasets. Miao-Xin Li, Juilian M. Y. Yeung, Stacey S. Cherny and Pak C. Sham. May 2012 Hum Genet.

# See Also

extract to obtain a table with the result of the ExWAS, plotExwas to plot the results of the ExWAS

# Examples

```
data(exposome)
w1 <- exwas(expo[1:5, ], asthma~1, family = "binomial")
w2 <- exwas(expo[1:5, ], asthma~sex+age, family = "binomial")
w3 <- exwas(expo[1:5, ], asthma~age, family = "binomial", filter = sex=="boy")</pre>
```

ex\_imp

# Description

imExposomeSet created from the raw data included in the package. The creation process is ilustrated in the vignette.

# Usage

```
data("ex_imp")
```

#### Format

An object of class imExposomeSet of dimension 47 x 109 x 10.

#### Value

An imExposomeSet object.

#### Examples

data("ex\_imp")
ex\_imp

familyNames

Getter to obtain the exposures's names of an ExposomeSet.

#### Description

This method returns the name of the families in an ExposomeSet, but it can return a vector, labeled with the exposures in the ExposomeSet, containing the family belonging to each exposure.

#### Usage

```
familyNames(object, by.exposure = FALSE)
```

# Arguments

object	ExposomeSet that will be queried for the exposures's family-names.
by.exposure	(default FALSE) If TRUE a vector labeled with each exposure name will be re- turned with the family of each exposures. If FALSE a vector with the (unique)
	name of the families of exposures will be returned.

# Value

The families of the exposures into the ExposomeSet, or the family of each exposure into the ExposomeSet.

# See Also

exposureNames to get the name of the exposures, phenotypeNames to get the phenotypes

# Examples

```
data("exposome")
# Get families
familyNames(expo)
# Get the family of each exposure
familyNames(expo, by.exposure = TRUE)
```

get\_robust\_sd Raw data from ExWAS

# Description

Returns internal table of robust SD of objects of class ExWAS

# Usage

```
get_robust_sd(object, ...)
```

#### Arguments

object	Object of class ExWAS
	NO USED

# Value

A data.frame containing the raw result from robust SD

highAndLow

# Description

This method allows to convert continuous exposures of an ExposomeSet to categorical exposures using the n-percentile groups, defined by ngroups argument. By default, all levels are kept but if intervals is set to "extrem", the levels between the extrems (aka. lowes and highest) are discarted and their values set to NA.

#### Usage

```
highAndLow(
   object,
   ngroups = 3,
   intervals = "standard",
   select,
   drop = FALSE,
   warnings = TRUE
)
```

#### Arguments

object	An object of class ExposomeSet.
ngroups	(default 3) Number of intervals to be created.
intervals	(default "standard") If set to "sctandard" all levels are set. If set to "extreme" obly lowes and highest levels are kept (others are set to NA).
select	(optional) Subset of exposures where the discretization is applied. If missing, all exposures are used.
drop	(default FALSE) If set to FALSE original exposures are kept and discretized exposures are add to ExposomeSet. If set to TRUE, original exposures are replaced by categorical exposures.
warnings	(defaulr TRUE) If set to FALSE warnings are not shown.

#### Value

A new ExposomeSet with categorical exposures.

# See Also

trans to transform exposures, standardize to standardize exposures.

# Examples

```
# No drop
data("exposome")
exp.hl <- highAndLow(expo, intervals = "standard", select = "ldde_lip")</pre>
dim(exp.hl)
# exposures samples phenotyes
#
      105
              1200
                            4
dim(expo)
             samples phenotyes
# exposures
#
      104
               1200
                             4
# exps.hl has 107 exposures: the original 104 plus the new
         3 factored exposures
#
# Drop
exp.hl <- highAndLow(expo, intervals = "standard",</pre>
   select = "ldde_lip", drop = TRUE)
dim(exp.hl)
# exposures
             samples phenotyes
       104
               1200
#
                             4
dim(expo)
             samples phenotyes
# exposures
       104
                1200
#
                             4
```

ilod

Function to impute under-LOD values from an ExposomeSet

# Description

This function is a wrapper of the functions impute.MinProb from the package imputeLCMD.

# Usage

```
ilod(
   object,
   seed = 1234,
   lod.col = "LOD",
   pNA = 0.2,
   tLog = FALSE,
   method = "QRILC",
   warnings = TRUE,
   ...
)
```

# Arguments

object	ExposomeSet which exposures will be imputed.
seed	(default 1234) Seed to make the imputation reproducible.
lod.col	(default "LOD") Name of the column in fData with the threshold of the LOD.

# imExposomeSet

pNA	(default 0.2) Maximum percentage allowed of values under LOD
tLog	(default FALSE) If set to TRUE it transforms all the exposures to lod before the imputation.
method	(default "QRILC") Method to be used to impute the under-LOD values. Two al- lowed: QRILC method (value "QRILC") and stochastic minimal value approach (value "MinProb").
warnings	(default TRUE) If set to FALSE warnings will not be displayed.
	Arguments passed to impute.QRILC or impute.MinProb from imputeLCMD.

# Value

A new ExposomeSet with the imputed exposures.

# See Also

plotMissings to plot the missing data of an ExposomeSet, tableMissings to get a table with the missing data of an ExposomeSet

#### Examples

```
## Not run:
#Being x an ExposomeSet
x <- ilod(x)</pre>
```

## End(Not run)

imExposomeSet Class imExposomeSet

#### Description

Class imExposomeSet was designed to store the exposures obtained after a multiple imputation process done using mice. The data.frame obtained from mice has the particulative to contain the columns .imp and .id joint to phenotypes and exposures. The slots assayData, featureData, and phenoData are coordinated.

#### Usage

```
## S4 method for signature 'imExposomeSet,ANY'
plot(x, y, ...)
## S4 method for signature 'imExposomeSet'
dim(x)
## S4 method for signature 'imExposomeSet'
expos(object)
```

```
## S4 method for signature 'imExposomeSet'
exposureNames(object)
## S4 method for signature 'imExposomeSet'
extract(object, rid = -1, ...)
## S4 method for signature 'imExposomeSet'
exwas(
  object,
  formula,
  filter,
  family,
  ...,
  baselevels,
  tef = TRUE,
  verbose = FALSE,
 warnings = TRUE,
  robust = FALSE
)
## S4 method for signature 'imExposomeSet'
familyNames(object, by.exposure = FALSE)
## S4 method for signature 'imExposomeSet'
phenotypeNames(object)
## S4 method for signature 'imExposomeSet'
plotFamily(x, family, group, group2, scatter = FALSE, na.omit = TRUE)
## S4 method for signature 'imExposomeSet'
sampleNames(object)
## S4 method for signature 'imExposomeSet'
x[[i]]
## S4 method for signature 'imExposomeSet,ANY,ANY,ANY'
x[i, j, k, ..., drop = FALSE]
## S4 method for signature 'imExposomeSet'
toES(object, rid = 1)
```

#### Arguments

х	An imExposomeSet object.
у	NOT USED
	Other used arguments
object	An object of class imExposomeSet
rid	Number of the imputation to be extracted

#### imExposomeSet

formula	Formula, not including exposures, to be tested.
filter	Expression to be used to filter ExposomeSet
family	Family descriving the nature of the health outcome
baselevels	Labeled vector with the base-level of the categorical exposures
tef	If set to TRUE the threshold for effective test is computed.
verbose	If set to TRUE it shows messages on progression.
warnings	If set to TRUE it shows warnings on progession.
by.exposure	If set to TRUE ir returns the family which each exposure belongs
group	Phenotype to group exposures
group2	NOT USED
scatter	If set to true it shows the samples value in the plot
na.omit	NOT USED
i	Character coresponding to selected exposures.
j	Character corresponding to selected sample names.
k	Character corresponding to selected phenotypes.
drop	NOT USED

# Value

An object of class imExposomeSet

#### Methods (by generic)

- plot(x = imExposomeSet, y = ANY): Wrapper for plotFamily method.
- dim(imExposomeSet): Returns the number of exsures, samples and phenotypes.
- expos(imExposomeSet): Returns a data.frame with exposures.
- exposureNames(imExposomeSet): Method to obtain samples' names
- extract(imExposomeSet): Method to extract exposures for a single imputation
- exwas(imExposomeSet): Performs an EXposome-Wide Association Study
- familyNames(imExposomeSet): Getter to obtain the families's names of the family of each exposure.
- phenotypeNames(imExposomeSet): Getter to obtain the families's names of the family of each exposure.
- plotFamily(imExposomeSet): Draws a boxplot or accumulated-bar plot for each exposure in the all imputed sets.
- sampleNames(imExposomeSet): Method to obtain samples' names
- [[: Get an ExposomeSet with the selected imputation
- x[i: Subset an imExposomeSet
- toES(imExposomeSet): Returns an ExposomeSet with ethe given imputation.

nimputation Number of imputations done with mice. assayData data.frame containing .imp, .id and the exposures. featureData data.frame containing the description of the exposures. phenoData data.frame containing .imp, .id and the phenotypes.

#### Note

Sample order is not guarantee

#### See Also

loadImputed to create an imExposomeSet from data.frames

imputation

Function to impute missing values from an ExposomeSet

# Description

This function is a wrapper of the functions mice and complete from the package mice. Also to the impute from the package Hmisc. The function is designed to use those functions to impute missing values on exposures (not in phenotypes).

#### Usage

imputation(object, select, ..., messages = FALSE)

#### Arguments

object	ExposomeSet which exposures will be imputed.
select	Exposures to be imputed. If missing, all exposes will be imputed.
	Argument given to function mice of pakcage mice (printFlag is set from messages).
messages	(default FALSE) If set to TRUE messages from mice's function will be displayed.

# Value

A new ExposomeSet with the imputed exposures.

#### See Also

plotMissings to plot the missing data of an ExposomeSet, tableMissings to get a table with the missing data of an ExposomeSet

# imputeLOD

# Examples

```
## Not run:
#Being x an ExposomeSet
x <- imputation(x)</pre>
```

## End(Not run)

```
imputeLOD
```

Function to impute values under limit of detection

# Description

Homologous function to the ilod method of ExposomeSet for a given data.frame of exposures and a vector of threshold. The use of this function is related to HELIX Project.

# Usage

```
imputeLOD(
    x,
    lod,
    description = NULL,
    pNA = 0.2,
    pLOD = 0,
    log = TRUE,
    seed = NULL
)
```

#### Arguments

data.frame containing the exposures as columns and the samples as rows.
vectro containing the
1 means values under LOD while 2 means quantifiable value, 3
(default: 0.2) maximum percentage of allowed missing data
(default: 0) minimum percentave of values under LOD
(default: NA) log transformation to normalize data
(default: NULL)

#### Value

A new data.frame with the imputed exposures.

# Examples

## End(Not run)

invExWAS

# Description

The invExWAS method performs an "Exposome-Wide Association Study" (ExWAS) using the exposures in ExposomeSet and one of its phenotype. (modelling the exposures as response)

#### Usage

```
invExWAS(object, formula, filter, tef = TRUE, verbose = FALSE, warnings = TRUE)
```

#### Arguments

object	ExposomeSet that will be used for the ExWAS.
formula	formula indicating the test to be done. If any exposure is included it will be used as covariate. exwas metho will perform the test for each exposure.
filter	expression to be used to filter the individuals included into the test.
tef	(default TRUE) If TRUE it computed the effective number of tests and the threhold for the effective number of tests. Usually it needs imputed data.
verbose	(default FALSE) If set o true messages along the tests are shown.
warnings	(default TRUE) If set to FALSE warnings will not be displayed.

#### Value

An codeExWAS object with the result of the association study

# See Also

extract to obtain a table with the result of the ExWAS, plotExwas to plot the results of the association

# Examples

```
data(exposome)
w1 <- invExWAS(expo, ~BMI)
w2 <- invExWAS(expo, ~BMI + sex)
plotExwas(w1, w2)</pre>
```

loadExposome

#### Description

Given three data.frames that defines the exposome (measures of exposome, exposome description and individuals phentype) it loads them and creates an object of type ExposomeSet.

#### Usage

```
loadExposome(
    exposures,
    description,
    phenotype,
    description.famCol = "family",
    exposures.asFactor = 5,
    warnings = TRUE
)
```

# Arguments

exposures	data.frame of exposures.	
description	data.frame with the description of the exposures (relation between exposures and exposure-family).	
phenotype	data.frame with the phenotypes of interest.	
description.famCol		
	(default "family") Index where the family's name (per exposures) if found in file "description". It can be both numeric or character.	
exposures.asFactor		
	(default 5) The exposures with more than this number of unique items will be considered as "continuous" while the exposures with less or equal number of items will be considered as "factor".	
warnings	(default TRUE) If TRUE shows useful information/warnings from the process of loading the exposome.	

#### Details

The rows of the exposure's data.frames, that corresponds to samples' names, must be the same than the phenotype's data.frames. In the same way, the columns in exposure's data.frames must be the same in description data.frame.

#### Value

An object of class ExposomeSet.

ExposomeSet's fData will contain some inner columns called .std, .trn, .fct and .type in order to trace the transformations an exposure suffers and to know, at eny moment, if an exposure is categorical or continuous. The "description" file can contains a column called type with values "factor" and "numeric" to speficy how an exposure needs to be understood. If given, this column will be renamed to .type. If not given, it will be created using exposures.asFactor value.

# See Also

ExposomeSet for class description, readExposome for constructor from txt/csv files.

# Examples

```
path <- file.path(path.package("rexposome"), "extdata")</pre>
description <- file.path(path, "description.csv")</pre>
phenotype <- file.path(path, "phenotypes.csv")</pre>
exposures <- file.path(path, "exposures.csv")</pre>
dd <- read.csv(description, header=TRUE)</pre>
ee <- read.csv(exposures, header=TRUE)</pre>
pp <- read.csv(phenotype, header=TRUE)</pre>
rownames(dd) <- dd[, 2]</pre>
dd <- dd[ , -2]
rownames(ee) <- ee[ , 1]</pre>
ee <- ee[ , -1]
rownames(pp) <- pp[ , 1]</pre>
pp <- pp[ , -1]
exp <- loadExposome(</pre>
exposures = ee,
description = dd,
phenotype = pp,
description.famCol = "Family"
)
```

loadExposome\_plain Creation of an ExposomeSet from single data.frame

#### Description

Creation of an ExposomeSet from single data.frame

### Usage

```
loadExposome_plain(
   data,
   data_id,
   sep = ",",
   pheno_cols,
   na.strings = c("NA", "-", "?", " ", ""),
```

34

# Note

```
families = NULL,
exposures.asFactor = 5,
warnings = TRUE
)
```

#### Arguments

	data	data.frame With the exposures and phenotypes (in no particular order!) or string with the path to a file (.csv, .tsv, .txt) with the table of exposures and phenotypes
	data_id	character Name of the column on the selected table that contains the ID
	sep	character (default ", ") Separator used by read.table to load the files "expo- sures", "description" and "phenotype". Only applies when providing a path on the data argument
	pheno_cols	character Character vector of the phenotype columns (all the other columns are considered exposures)
	na.strings	character (default c("NA", "-", "?", " ", "")) Character defining the NA values in expsome's files.
	families	<pre>list (default NULL) List to specify the families of the exposures, construct it as: list(Family1 = c("exposure_1_1", "exposure_1_2", "exposure_1_n"), Family2 = c("exposure_2_1", "exposure_2_2", "exposure_2_n"), FamilyM = c("exposure_M_1", "exposure_M_2", "exposure_M_n")). All the exposures on the data table have to be on this provided list with their respective families. The family classifica- tion is optional, input NULL to bypass the family classifier</pre>
exposures.asFactor		
		numeric (default 5) The exposures with more than this number of unique items will be considered as "continuous" while the exposures with less or equal number of items will be considered as "factor".
	warnings	(default TRUE) If TRUE shows useful information/warnings from the process of loading the exposome.

# Value

An object of class ExposomeSet.

#### Examples

```
path <- file.path(path.package("rexposome"), "extdata")
phenotype <- file.path(path, "phenotypes.csv")
exposures <- file.path(path, "exposures.csv")
ee <- read.csv(exposures, header=TRUE)
pp <- read.csv(phenotype, header=TRUE)
# Create fake dataset with exposures and phenotypes combined
data <- cbind(ee, pp)</pre>
```

```
loadExposome_plain <- function(data, data_id = "idnum",
pheno_cols = c("rhinitis", "wheezing", "sex", "age", "cbmi", "blood_pre", "whistling_chest", "flu"))</pre>
```

loadImputed

#### Description

Given a data.frame from code with the multiple imputations of both exposures and phenotypes, join with a data.frame with exposures' description, and object of class imExposomeSet is created.

# Usage

```
loadImputed(
   data,
   description,
   description.famCol = "family",
   description.expCol = "exposure",
   exposures.asFactor = 5,
   warnings = TRUE
)
```

# Arguments

data	The data.frame of both exposures and phentoypes obtained from mice.	
description	data.frame with the description of the exposures (relation between exposures and exposure-family).	
description.famCol		
	(default "family") Index where the family's name (per exposures) if found in	
	file "description". It can be both numeric or character.	
description.expCol		
	(default "exposures") Index where the exposure's name if found in file "de- scription". It can be both numeric or character.	
exposures.asFactor		
	(default 5) The exposures with more than this number of unique items will be considered as "continuous" while the exposures with less or equal number of items will be considered as "factor".	
warnings	(default TRUE) If TRUE shows useful information/warnings from the process of loading the exposome.	

# Details

The coordination of the information is based in the columns .imp and .id from the data.frame obtained from mice. The division of exposures and phenotypes is based in description data.frame, that are the exposures. Hence, the elements in the main data.frame that are not in the description, are the phentoypes.

# Value

An object of class imExposomeSet.
me

# See Also

imExposomeSet for class description

# Examples

```
data("me") # me is an imputed matrix of exposure and phenotyes
path <- file.path(path.package("rexposome"), "extdata")
description <- file.path(path, "description.csv")
dd <- read.csv(description, header=TRUE, stringsAsFactors=FALSE)
dd <- dd[dd$Exposure %in% colnames(me), ]
ex_imp <- loadImputed(data = me, description = dd,
description.famCol = "Family",
description.expCol = "Exposure")
```

me

# data.frame for testing purpouses

# Description

data.frame created with mice used to test and ilustrate the creation of imExposomeSet.

#### Usage

data("me")

# Format

An object of class data. frame with 654 rows and 57 columns.

## Value

A data.frame.

#### Examples

data("me")
dim(me)
colnames(me)

mExWAS

# Description

Class mExWAS obtained from mexwas method of an ExposomeSet object, contains the result of testing the multiple models of exposures of an ExposomeSet to a set of given phenotypes. "mExWAS" is the #' acronym of "Multivariate Exposome-Wide Association Study".

# Usage

```
## S4 method for signature 'mExWAS,ANY'
plot(x, y, ...)
## S4 method for signature 'mExWAS'
extract(object, type = "test", sort = TRUE)
## S4 method for signature 'mExWAS'
plotExwas(
   object,
    ...,
   subtitles,
   color,
   exp.order,
   labels,
   show.effective = TRUE
)
```

x	Object of class mExWAS
У	NOT USED
	Other used arguments.
object	object of class mExWAS.
type	(default "test"). Can take "test" or "raw" to obtain a data.frame of glmnet result.
sort	NOT USED
subtitles	NOT USED
color	NOT USED
exp.order	NOT USED
labels	NOT USED
show.effective	NOT USED

#### mexwas

#### Value

An object of class mExWAS

#### Methods (by generic)

- plot(x = mExWAS, y = ANY): Wrapper for plotExwas method.
- extract(mExWAS): Method to obtain the generated model
- plotExwas(mExWAS): Method to plot a heap-map with the coeficient of each exposure

# Slots

result klist with the fitted model and result.

phenotype Name of the phenotype used in the analysys.

description feature data from original ExposomeSet.

#### See Also

mexwas to perform a Multivariate Exposome-Wide Association Study and to create a mExWAS, exwas to perform an Exposome-Wide Association Study and to create an ExWAS

mexwas	Testing the association between an exposure and a phenotype of an
	ExposomeSet using a multivariate aproach.

#### Description

The mexwas method performs an "Multi Exposome-Wide Association Study" (m-ExWAS) using the exposures in ExposomeSet and one of its phenotype. It uses the packages glmnet and partDSA.

## Usage

mexwas(object, phenotype, family, warnings = TRUE)

#### Arguments

object	ExposomeSet that will be used for the ExWAS.
phenotype	Target phenotype used for the study. If missing all the phenotypes in the Expo- someSet will be used.
family	It must decrive the nature of the outcome. Can take values "gaussian", "binomial", "poisson", "multinomial", "cox" or "mgaussian".
warnings	(default TRUE) If set to FALSE warnings will not be displayed.

# Value

Returns an object of class mExWAS

#### See Also

extract to obtain a table with the result of the ExWAS, plotExwas to plot the results of the ExWAS

#### Examples

```
data("exposome")
wt <- mexwas(expo[3:7, 1:100], phenotype = "asthma", family = "binomial")</pre>
```

ndim

Number of principal components in an ExposomePCA.

#### Description

Number of principal components in an ExposomePCA.

#### Usage

ndim(object)

#### Arguments

object ExposomePCA to obtain the number of components.

# Value

The number of components in the ExposomePCA.

# See Also

plotPCA to plot the PCA values of an ExposomePCA, clustering to see how the exposures can cluster samples, correlation to compute the correlation between exposures

#### Examples

```
data("exposome")
epca <- pca(expo[3:7, 1:100])
ndim(epca)</pre>
```

40

normalityTest

It creates a data.frame of boolean indicating if the exposures follows a normal distribution or not.

## Description

This functions uses shapiro.test to test the normality of the exposures and returns a data.frame with a boolean value and a p-value for each exposure.

#### Usage

```
normalityTest(
   object,
   exposure,
   th = 0.05,
   min.val = 5,
   na.rm = TRUE,
   warnings = TRUE
)
```

#### Arguments

object	ExposomeSet with the exposome to be tested.
exposure	Name of the exposure to be tested, if missing all the exposures will be tested.
th	(by default $0.05$ ) Threshold to considere an exposure to follow a normal distribution.
min.val	(by default 5) Minimum number of values not missings to test the exposures.
na.rm	(by default TRUE) Removes the NA values to test the normality on the exposure.
warnings	(by default: TRUE) Show warnings if required.

#### Value

A data.frame with three columns: exposure, normality and p.value. "exposure" column contains the name of each exposure. "normalty" column contains a logical value indicating if the obtained p-value is under the given threshold. "p.value" column contains the obtained p-value from shapiro.test.

#### See Also

plotHistogram to draw the shape of an exposure, plotMissings to draw a plot with the missing data an ExposomeSet, imputation to impute missing data on the exposures of an ExposomeSet

```
data("exposome")
normalityTest(expo)
```

# Description

Method to calculate a PCA based on the exposures of an ExposomeSet. Only numerical-exposures (non categorical) will be computed. The function resurns an ExposomePCA object. This PCA is performed by using FactoMineR package.

#### Usage

pca(object, npc = 10, pca = FALSE, ...)

#### Arguments

object	ExposomeSet which exposures will be used for the PCA
npc	(by default 10) number of dimensions kept in the results
рса	(default FALSE) Set to TRUE to Perform PCA (only numerical variables) or FALSE to perform FAMD (numerical and categorical)
	Arguments to be passed to imputeFAMD

# Value

An ExposomePCA with the values of the PCA.

#### See Also

plotPCA to plot the PCA values of an ExposomePCA, clustering to see how the exposures can cluster samples, correlation to compute the correlation between exposures

# Examples

```
data("exposome")
epca <- pca(expo[12:20, ])</pre>
```

phenotypeNames	Getter to obtain the phenotype's names of an ExposomeSet or Expo-
	somePCA.

# Description

Getter to obtain the phenotype's names of an ExposomeSet or ExposomePCA.

#### Usage

phenotypeNames(object)

# рса

# plot3PCA

#### Arguments

object ExposomeSet that will be queried for the phenotype's names.

# Value

The name of the phenotypes as a a character vector.

# See Also

exposureNames to get the name of the exposures, familyNames to get the families of exposures

# Examples

data("exposome")
phenotypeNames(expo)

plot3PCA

Ploting PCA in a 3D space

# Description

Methdo to draw a plot for samples using three PC contained in an ExposomePCA

#### Usage

```
plot3PCA(
  object,
  cmpX,
  cmpY,
  cmpZ,
  phenotype,
  main,
  angle = 35,
  pch = 16,
  legend = TRUE,
  plines = TRUE
)
```

object	An onbject of class ExposomePCA
стрХ	Component to be placed at X axis
cmpY	Component to be placed at Y axis
cmpZ	Component to be placed at Z axis
phenotype	Used to color samples by phentoype
main	Title for the plot

angle	(default 35) angle between x and y axis.
pch	(default 16) plotting "character", i.e. symbol to use.
legend	(default TRUE) If TRUE shows the legend.
plines	(default TRUE) If TRUE it draws the lines from each dot to the base plane.

# Value

A list with different graphics option from scatterplot3d.

# See Also

pca to compite PCA on an ExposomeSet, plotPCA to plot the PCA, ExposomePCA as main class data("exposome") epca <- pca(expo[3:7, 1:100]) plot3PCA(epca, cmpX = 1, cmpY = 2, cmpZ = 3, phenotype = "sex")

plotClassification	Draw the profile of the levels exposures after a classification with a
	clustering method

# Description

Draw the profile of the levels exposures after a classification with a clustering method

## Usage

```
plotClassification(object, type = "heatmap", ...)
```

# Arguments

object	Object of class Exposomeclust
type	Two types are available: "heatmap" or "valuemap".
	NOT USED

## Value

A list with different graphics parameters.

# See Also

clustering as a constructor for ExposomeClust, classification to see how to obtain the classification of the samples from an ExposomeClust

#### Examples

```
## Not run:
data("eclust")
plotClassification(expo_c)
```

## End(Not run)

plotCorrelation

# Description

While the circos plot can be used to see the general behaviours intra and extra families of exposures, the matrix plot allows for a detailed view of the correlations within an ExposomeCorr object.

#### Usage

```
plotCorrelation(object, type = "circos", ...)
```

# Arguments

object	ExposomeCorr which correlations will be plotted.
type	(default "circos") Can take both "circos" or "matrix".
	Arguments given to corrplot of package corrplot if a matrix is draw. Moreover extra arguments are can be passed to inner functions to draw both the matrix and the circos of correlations.

# Value

A list with different graphics parameters.

# See Also

correlation as a constructor for ExposomeCorr objects, pca to compute PCA on exposures

#### Examples

```
data("exposome")
expo.c <- correlation(expo)
plotCorrelation(expo.c, type="circos")
plotCorrelation(expo.c, type="matrix")</pre>
```

plotEffect

```
Function to draw a plot of the pvalues stored in an ExWAS object
```

#### Description

This function draws a sort of manhattan plots using the p-value of the association of the exposures with phenotypes of an ExWAS object.

#### Usage

plotEffect(x, y, select, labels, xlab, ylab)

#### Arguments

х	An ExWAS object which effect will be ploted.
У	(optional) Another ExWAS object. If provded its effects will be ploted in Y-axis.
select	(optional) Character with exposures to be shown.
labels	(optional) Character vector with the labels for each exposure. It must be labeled vector.
xlab	(optional) Label for X-axis.
ylab	(optional) Label for Y-axis.

## Value

An object of class ggplot.

# See Also

exwas as a constructor for ExWAS objects, extract to obtain a table with the result of the ExWAS

# Examples

```
data(exposome)
w1 <- exwas(expo[1:5, ], asthma~1, family = "binomial")
w2 <- exwas(expo[1:5, ], asthma~sex+age, family = "binomial")
plotEffect(w1, w2)</pre>
```

plotEXP

Plot correlation between exposures and PCA

#### Description

Function used to plot the correlation between the exposures in an ExposomePCA and the values for each component of the PCA in the same ExposomePCA

#### Usage

```
plotEXP(object, exposure)
```

# Arguments

object	An object of class ExposomePCA
exposure	(optional) to select a set of exposures to be ploted. If not given all are used.

## Value

An object of class ggplot.

# plotExwas

#### See Also

pca to compute PCA on an ExposomeSet, plotPHE to plot the P-Value of association between phenotypes ans PCA, ExposomePCA as main class

#### Examples

```
data("exposome")
epca <- pca(expo[3:7, 1:100])
plotEXP(epca)</pre>
```

plotExwas

Function to draw a plot of the pvalues stored in an ExWAS object

#### Description

This function draws a sort of manhattan plots using the p-value of the association of the exposures with phenotypes of an ExWAS object.

# Usage

```
plotExwas(
   object,
   ...,
   subtitles,
   color,
   exp.order,
   labels,
   show.effective = TRUE
)
```

#### Arguments

object	An ExWAS object which p-values will be plotted.
	Other objects of class ExWAS.
subtitles	(optional) Characters used as "substitle" when more than one ExWAS is given.
color	(optional) Character vector of HTML colors, labeled with family's names. Used to colore the exposures.
exp.order	(optional) Character vector of exposures used to order and subset the plot.
labels	(optional) Character vector with the labels for each exposure. It must be labeled vector.
show.effective	(default TRUE) If set to FALSE, line showing effective test threshold is not shown.

#### Value

An object of class ggplot.

#### See Also

exwas as a constructor for ExWAS objects, extract to obtain a table with the result of the ExWAS

#### Examples

```
data(exposome)
w1 <- exwas(expo[1:5, ], asthma~1, family = "binomial")
plotExwas(w1)</pre>
```

plotFamily

```
It draws the profile of the exposome in an ExposomeSet
```

# Description

This function draw a profile of the full exposome into an ExposomeSet or the profile of the exposures in a specific family. For continuous families, box-plots are drawn; while for categorical families accumulative bar-charts.

## Usage

```
plotFamily(x, family, group, group2, scatter = TRUE, na.omit = TRUE)
```

## Arguments

х	ExposomeSet which exposome will be plotted.
family	Name of the familty that will be drawn. 'all' is allowed to draw a grid with all the families.
group	If set it displays the family grouped by the given phenotype.
group2	If set it displays the family grouped by the given phenotype.
scatter	(default TRUE) If the family to be plotted is continuous, the samples will be shown.
na.omit	(default TRUE) Do not show NA values.

## Value

A ggplot object if a family was selected. invisible() if argument family was set to "all".

## See Also

plotHistogram to draw the shape of an exposure, plotMissings to plot the missing data from an ExposomeSet

48

# plotHistogram

# Examples

```
data("exposome")
plt <- plotFamily(expo, family = "Metals")
plt <- plt + ggplot2::ggtitle("Metals")
plt
plt <- plotFamily(expo, family = "Indoor air")
plt <- plt + ggplot2::ggtitle("Indoor air")
plt</pre>
```

plotHistogram It draws a histogram for each exposure in an ExposomeSet

#### Description

It draws a grid with an histogram per exposure in an ExposomeSet.

# Usage

```
plotHistogram(x, select, density = TRUE, show.trans = FALSE)
```

# Arguments

х	ExposomeSet which exposome will be plotted.
select	Name fo the exposure to be plotted. If missing, all exposures will be used.
density	(default TRUE) If TRUE a density plot is drawn overlapping the histogram.
show.trans	(default FALSE) If set to TRUE it will draw a panel of auxiliar plots with the continuous exposure transformed with log and sqrt.

#### Value

A ggplot object.

## See Also

plotFamily to draw the profile of a family of exposures, plotMissings to plot the missing data from an ExposomeSet

```
data("exposome")
plotHistogram(expo[1:3, ], select = "ldde_lip")
```

plotLOD

# Description

This function can be used to draw the amount of under-LOD values in the exposures of an ExposomeSet.

## Usage

plotLOD(object, lod.col = "LOD", x.max = 100, sort = TRUE)

# Arguments

object	ExposomeSet which exposome will be plotted.
lod.col	(default "LOD") Name of the column in fData containing the LOD thresholds.
x.max	(default 100) Fix the maxium value of the X-axis.
sort	(default TRUE) If TRUE the chart will be ordered from the features with less miss- ing data to the ones with more under-LOD values.

# Value

A ggplot object.

# See Also

plotFamily to draw the profile of a family of exposures, plotHistogram to draw the shape of an exposure, tableMissings to get a table with the missing data of an ExposomeSet, imputation to impute missing data on the exposures of an ExposomeSet

```
data("exposome")
# The included has no missing data
plotLOD(expo)
```

plotMissings

#### Description

This function can be used to draw the number of missing data in exposures and in phenotypes of an ExposomeSet.

#### Usage

plotMissings(object, set, x.max = 100, sort = TRUE)

## Arguments

object	ExposomeSet which exposome will be plotted.
set	Can be set to "exposures" or to "phenotypes".
x.max	(default 100) Fix the maxium value of the X-axis.
sort	(default TRUE) If TRUE the chart will be ordered from the features with less miss- ing data to the ones with more missing data.

## Value

A ggplot object.

# See Also

plotFamily to draw the profile of a family of exposures, plotHistogram to draw the shape of an exposure, tableMissings to get a table with the missing data of an ExposomeSet, imputation to impute missing data on the exposures of an ExposomeSet

```
data("exposome")
# The included has no missing data
plotMissings(expo, set = "exposures")
plotMissings(expo, set = "phenotypes")
```

plotPCA

# Description

Methdo to draw a plot for PCA contained in an ExposomePCA

## Usage

```
plotPCA(
   object,
   set,
   cmpX = 1,
   cmpY = 2,
   show.exposures = FALSE,
   show.samples = FALSE,
   phenotype
)
```

# Arguments

object	An onbject of class ExposomePCA
set	Group ("all", "samples" or "exposures") taht will be ploted.
cmpX	(default: 1) component to be placed at X axis
cmpY	(default: 2) component to be placed at Y axis
show.exposures	(default: FALSE) If set to true, labels indicating the exposures are shown.
show.samples	(default: FALSE) If set to true, labels indicating the samples are shown.
phenotype	If set is set to "samples" can be used to color samples by phenotype

#### Value

An object of class ggplot or an object of class gtable if argument set was set to "all".

# See Also

pca to compite PCA on an ExposomeSet, plotPCA to plot the PCA, ExposomePCA as main class

```
data("exposome")
epca <- pca(expo[3:7, 1:100])
# A grid with exposures space, samples space and explained variance
plotPCA(epca, set = "all")
# Only exposures space
plotPCA(epca, set = "exposures") + ggplot2::theme(legend.position = "bottom")
# Only samples space</pre>
```

# plotPHE

```
plotPCA(epca, set = "samples")
# Only samples space but coloured by phenotype
plotPCA(epca, set = "samples", phenotype = "sex") +
ggplot2::theme(legend.position = "bottom")
```

plotPHE

#### Plot association score between phentoypes and PCA

# Description

Function used to plot the association between the phentoypes in an ExposomePCA and the values for each component of the PCA in the same ExposomePCA

# Usage

```
plotPHE(object, phenotype, exp2fac = 5)
```

# Arguments

object	An object oc class ExposomePCA
phenotype	(optional) to select a set of phenotypes to be ploted. If not given all are used.
exp2fac	(default, 5) Threshold to considere a phentoype categorical (less or equal to) or continuous (more than).

#### Value

An object of class ggplot.

#### See Also

pca to compute PCA on an ExposomeSet, plotEXP to plot the correlation between exposures ans PCA, ExposomePCA as main class

```
data("exposome")
epca <- pca(expo[3:7, 1:100])
plotPHE(epca)</pre>
```

plotVolcano

# Description

This function draws a sort of manhattan plots using the p-value of the association of the exposures with phenotypes of an ExWAS object.

#### Usage

```
plotVolcano(x, p.value = -log10(0.001), labels, show.effect = FALSE)
```

#### Arguments

х	An ExWAS object which effect will be ploted.
p.value	(default "-log10(0.001)") Threshold for P-Value.
labels	(optional) Character vector with the labels for each exposure. It must be labeled vector.
show.effect	(default FALSE) Applyes an exponential transformation on the effects of the exposures.

#### Value

An object of class ggplot.

# See Also

exwas as a constructor for ExWAS objects, extract to obtain a table with the result of the ExWAS, plotEffect to see or compare effects of one or two models.

 ${\tt readExposome}$ 

Creation of an ExposomeSet from files

## Description

Given the files that defines the exposome (measures of exposome, exposome description and individuals phentype) it loads them and creates an object of type ExposomeSet.

# readExposome

# Usage

```
readExposome(
    exposures,
    description,
    phenotype,
    sep = ",",
    na.strings = c("NA", "-", "?", " ", ""),
    exposures.samCol = "sample",
    description.expCol = "exposure",
    description.famCol = "family",
    phenotype.samCol = "sample",
    exposures.asFactor = 5,
    warnings = TRUE
)
```

exposures	String with the path to the file with the matrix of exposures.	
description	String with the path to the file with the description of the exposures (relation between exposures and exposure-family).	
phenotype	String with the path to the file with the phenotypes of interest.	
sep	(default ",") Separator used by read.table to load the files "exposures", "de- scription" and "phenotype".	
na.strings	(default c("NA", "-", "?", "", "")) Character defining the NA values in exp- some's files.	
exposures.samCc	1	
	(default "sample") Index where the samples' name are found in file "expo- sures". It can be both character or numeric.	
description.expCol		
	(default "exposure") Index where the exposures' name are found in file "de- scription". It can be both numeric or character.	
description.famCol		
	(default "family") Index where the family's name (per exposures) if found in file "description". It can be both numeric or character.	
phenotype.samCc	51	
	(default "sample") Index where the sample's name are found in file "pheno- type". It can be both numeric or character.	
exposures.asFactor		
	(default 5) The exposures with more than this number of unique items will be considered as "continuous" while the exposures with less or equal number of items will be considered as "factor".	
warnings	(default TRUE) If TRUE shows useful information/warnings from the process of loading the exposome.	

#### Details

The rows of the exposure's file, that corresponds to samples' names, must be the same than the phenotype's file. In the same way, the columns in exposure's file must be the same found as rows in description file.

#### Value

An object of class ExposomeSet.

#### Note

ExposomeSet's fData will contain some inner columns called .std, .trn, .fct and .type in order to trace the transformations an exposure suffers and to know, at eny moment, if an exposure is categorical or continuous. The "description" file can contains a column called type with values "factor" and "numeric" to speficy how an exposure needs to be understood. If given, this column will be renamed to .type. If not given, it will be created using exposures.asFactor value.

#### See Also

ExposomeSet for class description, loadExposome for constructor from loaded data.frames

#### Examples

```
## Locate the data-files
path <- file.path(path.package("rexposome"), "extdata")
description <- file.path(path, "description.csv")
phenotype <- file.path(path, "phenotypes.csv")
exposures <- file.path(path, "exposures.csv")
## Create ExposomeSet from files
exp <- readExposome(
    exposures = exposures,
    description = description,
    phenotype = phenotype,
    exposures.samCol = 1,
    description.famCol = 1,
    phenotype.samCol = 1
)</pre>
```

rexposome

rexposome: Package for exposome exploration and outcome data analysis

#### Description

#' @section exposures loading and exposures managment: rexposome offers two methods to bring exposome data to R and Bioconductor. readExposome allows to read three txt-like files (.csv, .tsv, ...) while loadExposome is sued with matrix and data.frames. The class obtained is an ExposomeSet, a class based in eSet for exposome data managment.

#### standardize

#### exposures processing

The packages offers a wide set of functions to preprocess exposome data. Method trans allow to transforms the exposures, method normalityTest allows to check for normality in exposome, standardize allows to standardize the exposures, among others. Finally, impute and ilod allow tu use mice, Hmisc and imputeLCMD for exposure missing data and exposure under-lod data imputation.

#### exposures analyses

the two methods exwas and mexwas allows to test the association between exposures and health outcomes (phenotpe data).

#### exposures plotting

The methods plotFamily allows to see how the exposures behaves within families. plotCorrelation helps to understand how exposures are related between themselves. plotClassification allos to visually detect cuslters of samples that share the same pattern of levels of exposures.

standardize	<ul> <li>#' Summary of an ExposomeSet. #' #' Given an ExposomeSet is shows a summary for its exposures or #' its phenotypes. #' #' @name Summary #' @rdname Summary-methods #' @aliases Summary #' @param object codeExposomeSet with 'set' will be summarized. #' @param set Set to be sumarized ("exposures" or "phenotypes"). #' @param select Subseting of exposures of phenotypes. #' @return A basic description of the exposures in the ExposomeSet #' @examples #' data("exposome") #' Summary(expo, set = "exposures") #' @export Summary setGeneric("Summary", function(object, set=c("exposures", "phenotypes"), select) standardGeneric("Summary") ) Standardize of an EmoscomeSet</li> </ul>
	an ExposomeSet.

# Description

Given an ExposomeSet it standardizes the exposures by using mean/sd if method is "normal" or by using median/mad if method is "robust".

# Usage

```
standardize(object, select, method = "normal", na.rm = TRUE, warnings = TRUE)
```

object	ExposomeSet with exposures to be standardized.
select	Subseting of exposures of phenotypes.
method	(default "normal") Character selecting the method to be applied ("normal" "iqr" or "robust").
na.rm	(by default TRUE) Removes NA data to perform standardization.
warnings	(defaulr TRUE) If set to FALSE warnings are not shown.

#### Value

An ExposomeSet with the selected exposures standardized and keeping the others exposures as the original input object.

#### See Also

highAndLow to transform the continuous exposures to levelled factors, trans to transform the exposures

## Examples

```
data("exposome")
exp.sn <- standardize(expo, method = "normal", select = "lbde100_lip")
exp.rs <- standardize(expo, method = "iqr", select = "lbde100_lip")
exp.rs <- standardize(expo, method = "robust", select = "lbde100_lip")</pre>
```

tableLOD
----------

It creates a vector with the amount of under-LOD exposures in an ExposomeSet

#### Description

This function can be used to obtain a table with the under-LOD data in exposures of an Exposome-Set.

#### Usage

```
tableLOD(object, output = "n", lod.col = "LOD", sort = TRUE)
```

#### Arguments

object	ExposomeSet which exposome will be plotted.
output	(default "n") Can be "n" for number of values, and "p" for percentage.
lod.col	(default "LOD") Name of the column in fData containing the LOD thresholds.
sort	(default TRUE) If TRUE the chart will be ordered from the features with less miss- ing data to the ones with more under-LOD values.

# Value

A numeric vector with number of udner-LOD values per exposure. The vector is labeled with the exposure's names.

#### See Also

plotFamily to draw the profile of a family of exposures, plotHistogram to draw the shape of an exposure, plotMissings to draw a plot with the missing data an ExposomeSet, imputation to impute missing data on the exposures of an ExposomeSet

58

# tableMissings

#### Examples

```
data("exposome")
# The included has no missing data
tableLOD(expo, output = "n")
```

tableMissings It creates a vector with the amount of missing in an ExposomeSet

# Description

This function can be used to obtain a table with the missing data in exposures and in phenotypes of an ExposomeSet.

#### Usage

```
tableMissings(object, set, output = "n", sort = TRUE)
```

## Arguments

object	ExposomeSet which exposome will be plotted.
set	Can be set to "exposures" or to "phenotypes".
output	(default "n") Can be "n" for number of values, and "p" for percentage.
sort	(default TRUE) If TRUE the chart will be ordered from the features with less miss- ing data to the ones with more missing data.

# Value

A numeric vector with number of missing values per exposure. The vector is labeled with the exposure's names.

#### See Also

plotFamily to draw the profile of a family of exposures, plotHistogram to draw the shape of an exposure, plotMissings to draw a plot with the missing data an ExposomeSet, imputation to impute missing data on the exposures of an ExposomeSet

```
data("exposome")
# The included has no missing data
tableMissings(expo, set = "exposures")
tableMissings(expo, set = "phenotypes")
```

tef

# Description

Function to get the Threshold for effective tests (TEF)

#### Usage

tef(object)

#### Arguments

object An ExWAS object

# Value

A number indicating the efective threshold.

# References

Evaluating the effective numbers of independent tests and significant p-value thresholds in commercial genotyping arrays and public imputation reference datasets. Miao-Xin Li, Juilian M. Y. Yeung, Stacey S. Cherny and Pak C. Sham. May 2012 Hum Genet.

# See Also

exwas as a constructor for ExWAS objects

```
data(exposome)
w1 <- exwas(expo[1:5, ], asthma~1, family = "binomial")
w2 <- exwas(expo[1:5, ], asthma~sex+age, family = "binomial")
tef(w1)
tef(w2)</pre>
```

toES

# Description

This methods allows to select an imputed-set and use it to create an ExposomeSet from an imExposomeSet.

## Usage

toES(object, rid = 1)

# Arguments

object	An object of class imExposomeSet
rid	(default 1) Number of the imputation to be extracted

# Value

An object of class ExposomeSet-

# Examples

data("ex\_imp")
toES(ex\_imp, rid = 1)

trans	Function to apply a transformation to the exposures of an Exposome-
	Set

# Description

The exposures in an ExposomeSet can be transformed using this function. transform apples a function fun to the selected exposures.

# Usage

```
trans(object, fun, select, by.exposure = FALSE, ...)
```

object	ExposomeSet which exposures will be transformed.
fun	Function to be applied on the exposures.
select	If not set, receive the name of all exposures. It can takes a character vector to select specific exposures.
by.exposure	(default FALSE) If TRUE applies fun to each exposure (given as a vector ). If FALSE the matrix of exposures is fiven to fun.
	Argument given to fun.

A new ExposomeSet with selected exposures transformed with fun.

## See Also

highAndLow to transform the continuous exposures to levelled factors, standardize to standardize by normal or robust methods the exposures

# Examples

```
data("exposome")
exp.t <- trans(expo, fun = log, select = "ldde_lip")</pre>
```

volcano\_plot Function to draw a Volcano Plot

# Description

Function that takes two numeric vectors (P-Value and fold change) and draws a volcano plot using ggplot2

# Usage

```
volcano_plot(
   pval,
   fc,
   names,
   size = 2,
   tFC = 2,
   tPV = -log10(0.001),
   show.effect = FALSE
)
```

pval	numeric vector of P.Values
fc	numeric vector of fold change
names	character vector with the feature's names.
size	(default 2) Sice of the labels in case they are placed.
tFC	(default 2) fold change threshold. It can be set to NULL to do not filter.
tPV	(default $-\log 10(0.001)$ ) P-Value threshold. It can be set to NULL to not filter.
show.effect	(default FALSE) If set to TRUE, the X-axis will should 2 <sup>logFC</sup> instead to teh default logFC.

# volcano\_plot

# Value

A ggplot object

```
data(exposome)
w1 <- extract(exwas(expo[1:20, ], asthma~1, family = "binomial"))
volcano_plot(w1$pvalue, w1$effect, rownames(w1))</pre>
```

# Index

```
AnnotatedDataFrame, 9, 12, 17
AssayData, 9, 17
```

classification, 3, 4, 44 classification, ExposomeClust-method (ExposomeClust), 7 clustering, 3, 4, 6-8, 40, 42, 44 clustering,ExposomeSet-method (ExposomeSet), 12 cor, 6, 8 correlation, 5, 12, 40, 42, 45 correlation, ExposomeSet-method (ExposomeSet), 12 corrplot, 45 cramersV, 6 dim, ExposomeCorr-method (ExposomeCorr), 8 dim, ExposomeSet-method (ExposomeSet), 12 dim, imExposomeSet-method (imExposomeSet), 27 eSet, 9, 12, 17 eSet-class, 8 ex\_imp, 23 expo, 6 expo\_c, 18 expos, 7 expos,ExposomeSet-method (ExposomeSet),

12

expos, imExposomeSet-method (imExposomeSet), 27 ExposomeClust, 3, 4, 7, 8, 44 ExposomeClust-class (ExposomeClust), 7 ExposomeCorr, 5, 6, 8, 9, 45 ExposomeCorr-class (ExposomeCorr), 8 ExposomePCA, 10, 11, 40, 42-44, 46, 47, 52, 53 ExposomePCA-class (ExposomePCA), 10 ExposomeSet, 4, 5, 7-9, 12, 12, 15, 17, 19, 21, 23, 24, 29, 31–35, 38, 39, 41, 42, 44, 47-54, 56-59, 61 ExposomeSet-class(ExposomeSet), 12 exposureNames, 17, 17, 24, 43 exposureNames, ExposomePCA-method (ExposomePCA), 10 exposureNames, ExposomeSet-method (ExposomeSet), 12 exposureNames, imExposomeSet-method (imExposomeSet), 27 extract, 19, 22, 32, 40, 46, 48, 54 extract (get\_robust\_sd), 24 extract, ExposomeCorr-method (ExposomeCorr), 8 extract,ExposomePCA-method (ExposomePCA), 10 extract, ExWAS-method (ExWAS), 19 extract, imExposomeSet-method (imExposomeSet), 27 extract, mExWAS-method (mExWAS), 38 ExWAS, 19, 20, 21, 39, 46, 48, 54, 60 exwas, 19, 21, 21, 39, 46, 48, 54, 57, 60 exwas, ExposomeSet-method (ExposomeSet), 12 exwas, imExposomeSet-method (imExposomeSet), 27 ExWAS-class (ExWAS), 19

FactoMineR, *42* familyNames, *18*, *23*, *43* 

# INDEX

familyNames,ExposomeSet-method
 (ExposomeSet), 12
familyNames,imExposomeSet-method
 (imExposomeSet), 27
featureNames, 17

get\_robust\_sd, 24
get\_robust\_sd, ExWAS-method (ExWAS), 19
ggplot2, 62
glm, 22

highAndLow, 25, 58, 62 highAndLow, ExposomeSet-method (ExposomeSet), 12

ilod, 26, 31, 57 ilod,ExposomeSet-method(ExposomeSet), 12 imExposomeSet, 27, 28, 36, 37, 61 imExposomeSet-class(imExposomeSet), 27 imExposomeSet-methods (imExposomeSet), 27 imputation, 30, 41, 50, 51, 58, 59 imputation,ExposomeSet-method (ExposomeSet), 12 impute, 57 impute.MinProb, 27 imputeLOD, 31 invExWAS, 32 invExWAS, ExposomeSet-method (ExposomeSet), 12

# lm, <mark>6</mark>

loadExposome, 17, 33, 56
loadExposome\_plain, 34
loadImputed, 30, 36

normalityTest,ExposomeSet-method (ExposomeSet), 12 pca, 6, 9, 42, 44, 45, 47, 52, 53 pca,ExposomeSet-method(ExposomeSet), 12 phenotypeNames, 18, 24, 42 phenotypeNames, ExposomePCA-method (ExposomePCA), 10 phenotypeNames, ExposomeSet-method (ExposomeSet), 12 phenotypeNames, imExposomeSet-method (imExposomeSet), 27 plot,ExposomeClust,ANY-method (ExposomeClust), 7 plot, ExposomeCorr, ANY-method (ExposomeCorr), 8 plot, ExposomePCA, ANY-method (ExposomePCA), 10 plot, ExposomeSet, ANY-method (ExposomeSet), 12 plot, ExWAS, ANY-method (ExWAS), 19 plot,imExposomeSet,ANY-method (imExposomeSet), 27 plot, mExWAS, ANY-method (mExWAS), 38 plot3PCA, 43 plot3PCA, ExposomePCA-method (ExposomePCA), 10 plotClassification, 3, 4, 44, 57 plotClassification, ExposomeClust-method (ExposomeClust), 7 plotCorrelation, 6, 45, 57 plotCorrelation, ExposomeCorr-method (ExposomeCorr), 8 plotEffect, 45, 54 plotEffect, ExWAS-method (ExWAS), 19 plotEXP, 46, 53 plotEXP,ExposomePCA-method (ExposomePCA), 10 plotExwas, 22, 32, 40, 47 plotExwas, ExWAS-method (ExWAS), 19 plotExwas, mExWAS-method (mExWAS), 38 plotFamily, 48, 49-51, 57-59 plotFamily,ExposomeSet-method (ExposomeSet), 12 plotFamily,imExposomeSet-method (imExposomeSet), 27 plotHistogram, 41, 48, 49, 50, 51, 58, 59 plotHistogram,ExposomeSet-method (ExposomeSet), 12

INDEX

plotLOD, 50 plotLOD,ExposomeSet-method (ExposomeSet), 12 plotMissings, 27, 30, 41, 48, 49, 51, 58, 59 plotMissings,ExposomeSet-method (ExposomeSet), 12 plotPCA, 40, 42, 44, 52, 52 plotPCA,ExposomePCA-method (ExposomePCA), 10 plotPHE, 47, 53 plotPHE,ExposomePCA-method (ExposomePCA), 10 plotVolcano, 54 plotVolcano, ExWAS-method (ExWAS), 19 read.table, 35, 55 readExposome, 17, 34, 54, 56 rexposome, 56 sampleNames,ExposomeClust-method (ExposomeClust), 7 sampleNames,imExposomeSet-method (imExposomeSet), 27 shapiro.test, 41 standardize, 25, 57, 57, 62 standardize,ExposomeSet-method (ExposomeSet), 12 Summary, ExposomeSet-method (ExposomeSet), 12 tableLOD, 58 tableLOD,ExposomeSet-method (ExposomeSet), 12 tableMissings, 27, 30, 50, 51, 59 tableMissings,ExposomeSet-method (ExposomeSet), 12 tef, 60 tef, ExWAS-method (ExWAS), 19 toES, 61 toES, imExposomeSet-method (imExposomeSet), 27 trans, 25, 57, 58, 61 trans,ExposomeSet-method(ExposomeSet), 12

volcano\_plot, 62

66