Package 'MOMA'

October 30, 2025

Title Multi Omic Master Regulator Analysis

Version 1.23.0

Description This package implements the inference of candidate master regulator proteins from multi-omics' data (MOMA) algorithm, as well as ancillary analysis and visualization functions.

Depends R (>= 4.0)

License GPL-3

Encoding UTF-8

LazyData true

BugReports https://github.com/califano-lab/MOMA/issues

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Transcriptomics, SystemsBiology

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Suggests BiocStyle, knitr, rmarkdown, testthat, viper

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Author Evan Paull [aut],

Sunny Jones [aut, cre],

Mariano Alvarez [aut]

Maintainer Sunny Jones <sunnyjjones@gmail.com>

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Description

aREA.enrich Compute aREA enrichment between all pairwise combinations of VIPER proteins and gene-level events

Usage

```
areaEnrich(events.mat, vipermat, event.type, verbose)
```

Arguments

events.mat	A Binary 0/1 matrix with columns as samples, and rows as proteins
vipermat	A VIPER network of inferred activity scores with columns as samples, and rows as proteins
event.type	Name of the event type for printing purposes
verbose	whether to print extra progress statements

Value

A matrix of enrichment scores with rows as event/gene names and columns as VIPER protein names

associateEvents	Use 'aREA' to calculate the enrichment between each genomic event - VIPER inferred protein pair.
	- VIFEK injerrea proiein pair.

Description

Requires pre-computed VIPER scores and a binary events matrix. Will use only samples in both event and VIPER matrices.

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Usage

```
associateEvents(
  vipermat,
  events.mat,
  min.events = NA,
  whitelist = NA,
  event.type = c("Amplifications", "Deletions", "Mutations", "Fusions", NA),
  verbose
)
```

Arguments

vipermat	Pre-computed VIPER scores with samples as columns and proteins as rows
events.mat	Binary 0/1 events matrix with samples as columns and genes or events as rows
min.events	Only compute enrichment if the number of samples with these events is GTE to this
whitelist	Only compute associations for events in this list
event.type	Name of the event type being analyzed
verbose	whether to print extra progress statements

Value

A matrix of aREA scores, dimensions are nrow(events.mat) x nrow(vipermat)

|--|--|

Description

Check Gene Map

Usage

```
checkGeneMap(gene.loc.mapping)
```

Arguments

```
gene.loc.mapping
```

dataframe with gene names, entrez ids and cytoband locations

Value

nothing

checkList 5

checkList

Check List of Assays

Description

Check List of Assays

Usage

```
checkList(assaylist)
```

Arguments

assaylist

list of assays (viper, cnv, mut and fusion)

Value

updated/filter assaylist obj

checkMAE

 ${\it Check\ MultiAssayExperiment}$

Description

Check MultiAssayExperiment

Usage

checkMAE(mae)

Arguments

mae

MultiAssayExperiment object

Value

updated/filtered MAE

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checkPathways

Check Pathways

Description

Check Pathways

Usage

```
checkPathways(pathways, x, type)
```

Arguments

pathways A named list of lists. Each named list represents interactions between proteins

(keys) and their associated partners

x the MAE or Assaylist

type whether x is MAE or Assaylist

Value

nothing

clusterRange

Cluster Range

Description

This function generate an cluster structure with 'k' groups and computes the cluster reliability score where 'k' is a range of values

```
clusterRange(
  dis,
  range = c(2, 100),
  step = 1,
  cores = 1,
  method = c("pam", "kmeans"),
  data = NULL
)
```

clusterReliability 7

Arguments

dis	Distance object
range	vector with start and end 'k'
step	Integer indicating the incremental number of clusters to add in each iteration
cores	Maximum number of CPU cores to use
method	Either 'pam' k-mediods or kmeans. Must supply the original data matrix if using kmeans

Value

data

list of cluster reliability scores by 'k', 'clustering' (the vector solution) and 'reliability' as well as 'medoids' labels

clusterReliability

Cluster membership reliability estimated by enrichment analysis

Description

This function estimates the cluster membership reliability using aREA

Original data matrix

Usage

```
clusterReliability(
  cluster,
  similarity,
  xlim = NULL,
  method = c("element", "cluster", "global")
)
```

Arguments

cluster Vector of cluster memberships or list of cluster memberships

similarity Similarity matrix

xlim Optional vector of 2 components indicating the limits for computing AUC

method Character string indicating the mthod to compute reliability, either by element,

by cluster or global

Value

Reliability score for each element

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cnvScoreStouffer

Integrate CNV scores

Description

Integrate CNV scores

Usage

```
cnvScoreStouffer(
  mapping,
  diggit.interactions,
  cytoband = TRUE,
  from.p = FALSE,
  pos.nes.only = TRUE
)
```

Arguments

mapping a named vector of genomic locations/cytoband IDs. names are the gene names

for each-i.e. a many to one mapping from HUGO or entrez IDs to cytoband

location

diggit.interactions

list indexed by MR/TF name in Entrez Space each points to a named vector of

NES / z-scores associated with entrez IDs for each interacting event.

cytoband Boolean to use cytoband locations for computing final integrated score

from.p Boolean, set TRUE if diggit.interaction values are p-values instead of z-scores

pos.nes.only Boolean, only consider positive DIGGIT association scores when ranking can-

didate MRs (default=TRUE)

Value

A vector of z-scores, named by the Master Regulators in 'diggit.interactions'

Description

Implements the conditional Bayes model to combine VIPER scores with diggit and pathway scores

```
conditionalModel(viper.scores, diggit.scores, pathway.scores)
```

conditionalP 9

Arguments

viper.scores numeric Vector

diggit.scores List indexed by type char, with numeric score vectors in [0,R+] for each

pathway.scores List, double indexed by each pathway dataset, then with type char. Each points

to a numeric score vectors in [0,R+] for each

Value

a named vector of empirical p-values for each protein/candidate Master Regulator

conditionalP

Get the conditional p-value of a gene

Description

Get the conditional p-value of a gene

Usage

```
conditionalP(gene.name, condition.on, x)
```

Arguments

gene.name Character

condition.on named Vector of scores for the distribution we are conditioning ON

x named Vector of scores for the dependent distribution

Value

a numeric p-value between 0 and 1

empiricalP

Get the empirical p-value from a distribution (vector)

Description

Get the empirical p-value from a distribution (vector)

```
empiricalP(gene.name, x)
```

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Arguments

gene.name Character

x named Vector of scores for the distribution

Value

a numeric p-value between 0 and 1

example.gbm.mae

Glioblastoma (GBM) Example Dataset

Description

MultiAssayExperiment Object containing all the genomic assays needed to run the example code for MOMA

Usage

```
example.gbm.mae
```

Format

An MultiAssayExperiment object with 4 different sets of GBM assays

viper matrix of viper scores with samples in columns and regulators across the rows

mut matrix of samples and genes with potential mutations. 0 for no mutation, 1 for presence of some non-silent mutation

cnv matrix of samples and genes with copy number variant scores

fitCurvePercent

Fit based on fractional overall coverage of genomic events

Description

Fit based on fractional overall coverage of genomic events

Usage

```
fitCurvePercent(sweep, frac = 0.85)
```

Arguments

sweep Numeric vector of genomic coverage values, named by -k- threshold frac Fraction of coverage to use as a threshold (default .85 = 85 percent)

Value

The -k- integer where coverage is acheived

gbm.pathways 11

gbm.pathways

Glioblastoma (GBM) Pathways

Description

Object containing information about the biological pathways that will be used in the analysis

Usage

gbm.pathways

Format

A list of lists named "cindy" and "preppi" respectively

cindy list of regulators, each with a set of modulators and p values representing their CINDY inferred association

preppi list of regulators, each with a set of potential binding partners and PREPPi inferred p values for probability of binding

gene.map

Gene Location Mapping

Description

Table used for converting between different forms of gene information. Downloaded from HGNC's custom download portal using the "Approved Symbol", "NCBI Gene ID", "Chromosome" and "Ensembl Gene ID" curated data options and only those with "Approved" status. Updated December 2019.

Usage

gene.map

Format

A Data frame with 4 columns

Gene.Symbol Approved Symbol gene name

Entrez.IDs NCBI Gene ID

Cytoband Chromosome location

Ensembl gene ID

@source https://www.genenames.org/download/custom/

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genomicPlotSmall

Make small genomic plot

Description

Make small genomic plot

Usage

```
genomicPlotSmall(input.df, fraction = 0.85, tissue.cluster = NULL)
```

Arguments

input.df : tissue.coverage.df with mean, k, fraction and unique events. raction : what fraction coverage to use for genomic curve threshold

tissue.cluster: which cluster subsample to look at

Value

output .png

getCoverage

Get coverage of interactions

Description

Get coverage of interactions

Usage

```
getCoverage(
  MomaObject,
  cMR.ranking,
  viper.samples,
  topN = 100,
  mutation.filter = NULL,
  verbose = FALSE
)
```

Arguments

MomaObject A numeric vector with cluster membership, names are samples

cMR.ranking A vector entrez IDs, in order

viper.samples Calculate the genomic coverage only for these sample topN Compute coverage for only the top -N- Master Regulators

mutation.filter

Retain only mutation events in this (positive) list

getDataFrame 13

Value

A list of lists, indexed by sample name, with coverage statistics for each sample

getDataFrame

Helper function to get data frame for bar plot plot.events function

Description

Helper function to get data frame for bar plot plot.events function

Usage

```
getDataFrame(
  data,
  highlight.genes,
  genomeBand_2_gene,
  max.muts = 10,
  max.cnv = 5
)
```

Arguments

data data.frame with \$type, \$id, \$Freq per event

highlight.genes

genes to look for in mutations/cnv lists (if looking for specific genes because of

prior knowledge)

genomeBand_2_gene

mapping of genomic location IDs to gene name: vector of HUGO gene ids,

named by genomic loci

max.muts maximum number of mutations to get per sample, default is 10

max.cnv maximum number of cnvs to per sample, default is 5

Value

ordered data frame with each genomic event and it's frequency

14 getEmpiricalQvals

```
getDiggitEmpiricalQvalues
```

Compute the empirical q-values of each genomic-event/VIPER gene pair

Description

Use against the background distribution of associations with a given set of 'null' VIPER genes (i.e. low activity TFs)

Usage

```
getDiggitEmpiricalQvalues(vipermat, nes, null.TFs, alternative = "both")
```

Arguments

viper inferences matrix, samples are columns, rows are TF entrez gene IDs

nes scores for each mutation (rows) against each TF (columns)
null.TFs low-importance TFs used to calculate null distributions

alternative Alternative defaults to 'both': significant p-values can come from both sides of

the null distribution

Value

A named list of qvalues for each TF/cMR protein. Each entry contains a vector of q-values for all associated events; names are gene ids

getEmpiricalQvals

Get empirical qvals

Description

Get empirical qvals

Usage

```
getEmpiricalQvals(test.statistics, null.statistics, alternative = "both")
```

Arguments

test.statistics

P-values generated from the test comparisons

null.statistics

P-values generated under the null (permutation) model

alternative Optional: 1 or 2 tails used to generate the p-value

getPvalsMatrix 15

Value

A list with both the qvalues and empirical p-values from the supplied test and null stats

getPvalsMatrix

Utility function

Description

Utility function

Usage

```
getPvalsMatrix(corrected.scores)
```

Arguments

```
corrected.scores
```

- corrected p-values processed by 'qvals' package

Value

A matrix of p-values for scores between genes/events (rows) and TFs (columns)

```
getSubtypeEventTables Helper function to get subtype specific events
```

Description

Helper function to get subtype specific events

Usage

```
getSubtypeEventTables(saturation.data, sample.clustering, checkpoints)
```

Arguments

saturation.data

: genomic saturation object from MOMA. List indexed by cluster then sample then regulator with the number of events associated with each additional regula-

nina

sample.clustering

: clustering vector with sample names and cluster designations

checkpoints : from momaObj

Value

a table that has counts of how many times a particular event happens in a cluster

integrateTZ

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Numerical integration of functions

Description

Integrates numerically a function over a range using the trapezoid method

Usage

```
integrateFunction(f, xmin, xmax, steps = 100, ...)
```

Arguments

f	Function of 1 variable (first argument)
xmin	Number indicating the min x value
xmax	Number indicating the max x value

steps Integer indicating the number of steps to evaluate

... Additional arguments for f

Value

Number

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Integration with trapezoid method

Description

This function integrate over a numerical range using the trapezoid method

Usage

```
integrateTZ(x, y)
```

Arguments

X	Numeric vector of x values
У	Numeric vector of y values

Value

Number

makeCoverageDf 17

makeCoverageDf

Helper function for making the coverage dataframe

Description

Helper function for making the coverage dataframe

Usage

```
makeCoverageDf(coverage.list, cutoff)
```

Arguments

coverage.list : List indexed by sample name, contains mut/fus/amp/del interactions

cutoff : number of regulators to include

Value

dataframe with each sample and which events are captured by the checkpoint mrs

makeSaturationPlots

Main function to generate the summary plots of the analysis

Description

Main function to generate the summary plots of the analysis

Usage

```
makeSaturationPlots(
  momaObj,
  clustering.solution = NULL,
  important.genes = NULL,
  fCNV = NULL,
  max.events = 30
)
```

Arguments

momaObj : momaObj that has already run the saturationCalculation function

clustering.solution

: clustering vector with sample names and cluster designations

important.genes

: vector of gene names to prioritize when plotting. Can be general genes of

interest, oncogenes, tumor supressors etc

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fCNV : vector of confirmed functional CNVs if calculated. Will filter for only those

CNVs

max.events : maximum number of events to plot for the oncoplots

Value

object with both types of summary plot for each subtype

Examples

```
## Not run:
makeSaturationPlots(momaObj, max.events = 20)
## End(Not run)
```

mapEntrez

Convert from entrez ids to hugo gene names

Description

Convert from entrez ids to hugo gene names

Usage

```
mapEntrez(entrez.ids)
```

Arguments

entrez.ids : vector of entrez ids requires hugo2entrez to be loaded

Value

: vector of hugo gene names

See Also

mapHugo

Examples

```
mapEntrez(c("29974", "5728"))
```

mapHugo 19

mapHugo

Convert from hugo gene names to entrez ids

Description

Convert from hugo gene names to entrez ids

Usage

```
mapHugo(hugo.ids)
```

Arguments

hugo.ids

: vector of hugo gene names, requires hugo2entrez to be loaded

Value

: vector of entrez ids

See Also

```
{\tt mapEntrez}
```

Examples

```
mapHugo(c("A1CF","PTEN"))
```

 ${\tt mapScoresCnvBand}$

Map scores to cytoband location

Description

Map scores to cytoband location

```
mapScoresCnvBand(
  mapping,
  diggit.interactions,
  from.p = FALSE,
  pos.nes.only = TRUE
)
```

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Arguments

mapping a named vector of genomic locations/cytoband IDs. names are the gene names

for each-i.e. a many to one mapping from HUGO or entrez IDs to cytoband

location

diggit.interactions

list indexed by MR/TF name in Entrez Space

from.p DIGGIT interactions are in p-value format instead of z-score (default=FALSE)

pos.nes.only Only consider positive associations with NES scores (default=TRUE) each points

to a named vector of NES / z-scores associated with entrez IDs for each inter-

acting event.

Value

A list of input scores, now named by cytoband location

mergeData

Helper function for mergeDataBySubtype

Description

Helper function for mergeDataBySubtype

Usage

```
mergeData(coverage.range, topN)
```

Arguments

coverage.range : genomic saturation for a particular subtype

topN : max number of top regulators to search through

Value

dataframe with coverage data for genomic events

mergeDataBySubtype 21

mergeDataBySubtype	Create data frame from coverage data, including number of total
	events 'covered' and unique events

Description

Create data frame from coverage data, including number of total events 'covered' and unique events

Usage

```
mergeDataBySubtype(genomic.saturation, sample.clustering, topN = 100)
```

Arguments

genomic.saturation

: data from genomic saturation function

sample.clustering

: clustering vector with sample names and cluster designations

topN : number of regulators to look through. default is 100

Value

dataframe with coverage data for genomic events

```
mergeGenomicSaturation
```

mergeGenomicSaturation Create data frame from coverage data, including number of total events 'covered' and unique events

Description

mergeGenomicSaturation Create data frame from coverage data, including number of total events 'covered' and unique events

Usage

```
mergeGenomicSaturation(coverage.range, topN)
```

Arguments

coverage.range List indexed by sample, then sub-indexed by # of master regulators, then by event type (mut/amp/del/fus). Holds all events by sample
topN Maximum number of master regulators to compute coverage

Value

A data frame with summary statistics for genomic saturation at each k

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mergeLists

Helper function

Description

Helper function

Usage

```
mergeLists(11, 12)
```

Arguments

11 list 112 list 2

Value

single merged list

Moma-class

MOMA Object

Description

Main class encapsulating the input data and logic of the MOMA algorithm

Fields

viper matrix of inferred activity score inferred by viper
mut binary mutation matrix 1 for presence of mutation, 0 for not, NA if not determined
cnv matrix of cnv values. Can be binary or a range.
fusions binary matrix of fusion events if appliable
pathways list of pathways/connections to consider as extra evidence in the analysis
gene.blacklist character vector of genes to not include because of high mutation frequency
output.folder character vector of location to save files if desired
gene.loc.mapping data frame of gene names, entrez ids and cytoband locations
nes field for saving Normalized Enrichment Matrices from the associate events step
interactions field for saving the MR-interactions list
clustering.results results from clustering are saved here
ranks results field for ranking of MRs based on event association analysis
hypotheses results field for saving events that have enough occurences to be considered

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```
genomic.saturation results field for genomic saturation analysis

coverage.summaryStats results field for genomic saturation analysis

checkpoints results field with the MRs determined to be the checkpoint for each cluster

sample.clustering field to save sample clustering vector. Numbers are cluster assignments,

names are sample ids
```

Methods

```
Cluster (clus.eval = c("reliability", "silhouette"), use.parallel = FALSE, cores = 1)
        Cluster the samples after applying the MOMA weights to the VIPER scores

makeInteractions(genomic.event.types = c("amp", "del", "mut", "fus"), cindy.only = FALSE)
        Make interaction web for significant MRs based on their associated events

Rank(use.cindy = TRUE, genomic.event.types = c("amp", "del", "mut", "fus"), use.parallel = FALSE, cores =
        Rank MRs based on DIGGIT scores and number of associated events

runDIGGIT(fCNV = NULL, cnvthr = 0.5, min.events = 4, verbose = FALSE) Run DIGGIT association function to get associations for driver genomic events

saturationCalculation(clustering.solution = NULL, cov.fraction = 0.85, topN = 100, verbose = FALSE)
        Calculate the number of MRs it takes to represent the desired coverage fraction of events
```

MomaConstructor

MOMA Constructor Function

Description

Create MOMA Object from either a MultiAssayExperiment object or a list of assays. See vignette for more information on how to set up and run the MOMA object

```
MomaConstructor(
    x,
    pathways,
    gene.blacklist = NA_character_,
    output.folder = NA_character_,
    gene.loc.mapping = gene.map,
    viperAssay = "viper",
    mutMat = "mut",
    cnvMat = "cnv",
    fusionMat = "fusion"
)
```

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Arguments

Χ

A MultiAssayExerperiment object or list object with the following assays: (note: by default assays must have these exact names. Otherwise they can be changed using the viperAssay, mutMat, cnvMat and fusionMat parameters.)

viper VIPER protein activity matrix with samples as columns and rows as protein IDs

mut An indicator matrix (0/1) of mutation events with samples as columns and genes as rows

cnv A matrix of CNV scores (typically SNP6 array scores from TCGA) with samples as columns and genes as rows

fusion An indicator matrix (0/1) of fusion events with samples as columns and genes as rows

pathways A named list of lists. Each named list represents interactions between proteins

(keys) and their associated partners

gene.blacklist A vector of genes to exclude from the analysis output.folder Location to store output and intermediate results

gene.loc.mapping

A data frame of band locations and Entrez IDs

name associated with the viper assay in the assay object viperAssay name associated with the mutation matrix in the assay object mutMat cnvMat name associated with the cnv matrix in the assay object

fusionMat name associated with the fusion matrix in the assay object

Value

an instance of class Moma

Examples

momaObj <- MomaConstructor(example.gbm.mae, gbm.pathways)</pre>

mutSig MutSig Blacklisted genes

Description

List of genes to not include in the DIGGIT mutation inference because they have been found to be mutated more often than expected by chance given background mutation processes.

Usage

mutSig

oncoprintPlot 25

Format

A character vector of Entrez Gene IDs

Source

https://software.broadinstitute.org/cancer/cga/mutsig

oncoprintPlot

Function to plot genomic events in the style of oncoPrint/cBioPortal

Description

Function to plot genomic events in the style of oncoPrint/cBioPortal

Usage

```
oncoprintPlot(
   summary.vec,
   snpmat.thisClus,
   amps.thisClus,
   dels.thisClus,
   fusions.thisClus,
   important.genes,
   band2gene,
   max.events,
   k
)
```

Arguments

summary.vec : named vector of the counts, named 'Event name':'Type' where type is 'mut',

'amp', 'del', 'fus'. Mutations are in Entrez ID Amp/Deletion CNV events are in

genomic band location

snpmat.thisClus

: SNP matrix subset to samples in current cluster

amps.thisClus : CNV matrix subset to samples in current cluster (just amplifications)

dels.thisClus : CNV matrix subset to samples in current cluster (just deletions)

fusions.thisClus

: Fusion matrix subset to samples in current cluster

important.genes

: well known genes to highlight in the analysis

band2gene : mapping of genomic location IDs to gene name: vector of HUGO gene ids,

named by genomic location

max.events : maximum number of events to plot for the oncoplots

k : current cluster number

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Value

oncoprint event plot

```
pathwayDiggitIntersect
```

Combine DIGGIT inferences with pathway knowledge

Description

Combine DIGGIT inferences with pathway knowledge

Usage

```
pathwayDiggitIntersect(diggit.int, pathway, pos.nes.only = TRUE, cores = 1)
```

Arguments

diggit.int List of interactions between MRs - Genomic events, inferred by DIGGIT

pathway - a list indexed by TF/MR entrez ID, contains the named vector of p-values for

interactions

pos.nes.only Only use positive associations between MR activity and presence of events (de-

fault = True)

cores Number of cores to use if parallel is selected

Value

numeric vector, zscores for each TF/MR

plotEvents

Plot barchart of genomic events

Description

Plot barchart of genomic events

```
plotEvents(
   summary.vec,
   highlight.genes = NULL,
   genomeBand_2_gene = NULL,
   samples.total,
   max.muts = 10,
   max.cnv = 5
)
```

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Arguments

summary.vec : named vector of the counts, named 'Event name': 'Type' where type is 'mut',

'amp', 'del', 'fus'. Mutations are in Entrez ID Amp/Deletion CNV events are in

genomic band location

highlight.genes

: well known genes to highlight in the analysis in

genomeBand_2_gene

: mapping of genomic location IDs to gene name: vector of HUGO gene ids,

named by genomic loci

samples.total : number of samples in the subtype, used to calculate percentages max.muts : maximum number of mutations to get per sample, default is 10

max.cnv : maximum number of cnvs to per sample, default is 5

Value

plot object

rea	This function calculates an Enrichment Score of Association based on	
	how the features rank on the samples sorted by a specific gene	

Description

This function calculates an Enrichment Score of Association based on how the features rank on the samples sorted by a specific gene

Usage

```
rea(eset, regulon, minsize = 1, maxsize = Inf, event.type = NA, verbose)
```

Arguments

eset	Numerical matrix
regulon	A list with genomic features as its names and samples as its entries, indicating presence of event
minsize	The minimum number of events to use when calculating enrichment
maxsize	The maximum number of events to use when calculating enrichment
event.type	Type of event being analyzed
verbose	whether to print extra progress statements

Value

A list containing two elements:

groups Regulon-specific NULL model containing the enrichment scores ss Direction of the regulon-specific NULL model

28 sampleNameFilter

reaNULL	This function generates the NULL model function, which computes the normalized enrichment score and associated p-value
	r

Description

This function generates the NULL model function, which computes the normalized enrichment score and associated p-value

Usage

```
reaNULL(regulon, minsize = 1, maxsize = Inf)
```

Arguments

regulon	A list with genomic features as its names and samples as its entries

minsize Minimum number of event (or size of regulon)
maxsize Maximum number of event (or size of regulon)

Value

A list of functions to compute NES and p-value

sampleNameFilter Retain TCGA sample ids without the final letter designation ('A/B	'/C')
--	-------

Description

Retain TCGA sample ids without the final letter designation ('A/B/C')

Usage

```
sampleNameFilter(input, desired.len = 15)
```

Arguments

input Matrix of expression or protein activity scores. Columns are sample names,

rows are genes. Input can also just be an input vector of sample names.

desired.len length to reduce strings to. Default is 15 because of TCGA naming conventions

Value

An identical matrix with new (shorter) column names, or a vector with the shortened names.

Examples

```
sample.names <- c("TCGA-14-1825-01A", "TCGA-76-4931-01B", "TCGA-06-5418-01A") sampleNameFilter(sample.names)
```

sampleOverlap 29

sampleOverlap	The core function to compute which sample-specific alterations over- lap with genomic events that are explained via DIGGIT.
	tup with genomic events that are explained via D10011.

Description

The core function to compute which sample-specific alterations overlap with genomic events that are explained via DIGGIT.

Usage

```
sampleOverlap(
  MomaObject,
  viper.samples,
  selected.tfs,
  interaction.map,
  cnv.threshold = 0.5,
  mutation.filter = NULL,
  idx.range = NULL,
  verbose = FALSE
)
```

Arguments

MomaObject	Object reference of momaRunner class	
viper.samples	Sample vector to restrict sample-specific analysis to	
selected.tfs	Transcription factors being analyzed	
interaction.map		
	List object of events 'covered' by the supplied interactions of type mut/amp/del/fus	
cnv.threshold	Numeric absolute value to threshold SNP6 and/or GISTIC or other CNV scores. Above that absolute value is considered a positive event.	
mutation.filter		
	A vector of whitelisted mutation events, in entrez gene IDs	
idx.range	Number of tfs to check for genomic saturation calculation, default is 1253	
verbose	Output status during the run (default=FALSE)	

Value

A list of lists, indexed by sample name, with coverage statistics/data for each sample

30 sREA

sigInteractorsDIGGIT Filter interactions from NES (DIGGIT) scores and corresponding background-corrected scores.

Description

Use this version in the Bayes model to rank TFs

Usage

```
sigInteractorsDIGGIT(
  corrected.scores,
  nes.scores,
  cindy,
  p.thresh = 0.05,
  cindy.only = TRUE
)
```

Arguments

corrected.scores

A list indexed by the genomic event/gene with corresponding pvals and qvals

for each TF

nes. scores Matrix with tfs as columns, rows are genomic events

cindy CINDy algorithm output matrix p.thresh P-value threshold (default=0.05)

cindy.only Consider only CINDy validated interactions (default=TRUE)

Value

a list (indexed by VIPER protein) of significant genomic interactions and associated pvals over the background (null TF) model, and NES scores

SREA Simple one-tail rank based enrichment analysis sREA (for cluster analysis)

Description

This function performs simple 1-tail rank based enrichment analysis

```
sREA(signatures, groups)
```

stoufferIntegrate 31

Arguments

signatures Numeric matrix of signatures

groups List containing the groups as vectors of sample names

Value

Matrix of Normalized Enrichment Zcores

stoufferIntegrate

dispatch method for either CNV location corrected or SNV

Description

dispatch method for either CNV location corrected or SNV

Usage

```
stoufferIntegrate(interactions, cytoband.map = NULL)
```

Arguments

interactions List of MR - Genomic Event interactions, inferred by DIGGIT

cytoband.map Data.frame mapping Entrez.IDs to cytoband locations

Value

Z-scores for each MR

stoufferIntegrateDiggit

Use Stouffer's method to combine z-scores of DIGGIT interactions for each cMR protein.

Description

This function combines only positively associated DIGGIT scores by default to create a culmulative DIGGIT score for each cMR.

```
stoufferIntegrateDiggit(interactions, from.p = FALSE, pos.nes.only = TRUE)
```

32 validDiggitInteractions

Arguments

interactions A list indexed by TF, includes z-scores or p-values for each interacting event

from.p Integrate p-values or z-scores (default z-scores; from.p = FALSE)
pos.nes.only Use only positive NES scores to rank proteins (default TRUE)

Value

A list indexed by TF, a stouffer integrated z-score

subsetListInteractions

Helper function: subset a list to the set of keys supplied return the names of interactions with positive values, in a list structure

Description

Helper function: subset a list to the set of keys supplied return the names of interactions with positive values, in a list structure

Usage

```
subsetListInteractions(int.1, keys)
```

Arguments

int.1 List of interactions, at each index this is a numeric named vector

keys Keys used to reduce interactions

Value

Returns a filtered list of interactions in the same format as the input

validDiggitInteractions

Return a set of events 'covered' by specified cMR-event interactions

Description

Return a set of events 'covered' by specified cMR-event interactions

Usage

validDiggitInteractions(interactions, gene.loc.mapping, selected.tfs)

viperGetSigTFS 33

Arguments

interactions List indexed by amp/mut/del/fus from cMRs to interacting events gene.loc.mapping

Data.frame mapping entrezIDs to cytoband locations

selected.tfs For each event type list, search within only these cMRS

Value

a list of events 'covered' by the supplied interactions of type mut/amp/del/fus

viperGetSigTFS Calculate p-values from pseudo zscores / VIPER aREA scores, threshold

Description

Calculate p-values from pseudo zscores / VIPER aREA scores, threshold

Usage

```
viperGetSigTFS(zscores, fdr.thresh = 0.05)
```

Arguments

zscores Vector of normally distributed z-scores representing protein activities.

fdr. thresh Threshold for false discovery rate, default is 0.05

Value

Get the names of proteins with significant z-scores, after multi-hypothesis correction

viperGetTFScores Function to normalize TF scores

Description

Function to normalize TF scores

Usage

```
viperGetTFScores(vipermat, fdr.thresh = 0.05)
```

Arguments

vipermat - matrix of VIPER scores with columns as samples, rows as protein names

fdr. thresh - BH-FDR threshold (default 0.05 FDR rate)

34 viperGetTFScores

Value

A vector of normalized z-scores, named by TF id

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