GSVA

March 24, 2012

computeGeneSetsOverlap

Compute gene-sets overlap

Description

Calculates the overlap among every pair of gene-sets given as input.

Usage

```
## S4 method for signature 'list, character'
computeGeneSetsOverlap(gSets, uniqGenes, min.sz=1, max.sz=Inf)
## S4 method for signature 'list, ExpressionSet'
computeGeneSetsOverlap(gSets, uniqGenes, min.sz=1, max.sz=Inf)
## S4 method for signature 'GeneSetCollection, character'
computeGeneSetsOverlap(gSets, uniqGenes, min.sz=1, max.sz=Inf)
## S4 method for signature 'GeneSetCollection, ExpressionSet'
computeGeneSetsOverlap(gSets, uniqGenes, min.sz=1, max.sz=Inf)
```

Arguments

gSets	Gene sets given either as a list or a GeneSetCollection object.
uniqGenes	Vector of unique genes to be considered when calculating the overlaps.
min.sz	Minimum size.
max.sz	Maximum size.

Details

This function calculates the overlap between every pair of gene sets of the input argument gSets. Before this calculation takes place, the gene sets in gSets are firstly filtered to discard genes that do not match to the identifiers in uniqGenes. Secondly, they are further filtered to meet the minimum and/or maximum size specified with the arguments min.sz and max.sz. The overlap between two gene sets is calculated as the number of common genes between the two gene sets divided by the smallest size of the two gene sets.

Value

A gene-set by gene-set matrix of the overlap among every pair of gene sets.

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Author(s)

J. Guinney

References

H\"anzelmann, S., Castelo, R. and Guinney, J. GSVA: Gene Set Variation Analysis, submitted

See Also

```
filterGeneSets
```

Examples

```
geneSets <- list(set1=as.character(1:4), set2=as.character(4:10))
computeGeneSetsOverlap(geneSets, unique(unlist(geneSets)))</pre>
```

filterGeneSets

Filter gene sets

Description

Filters gene sets through a given minimum and maximum set size.

Usage

```
## S4 method for signature 'list'
filterGeneSets(gSets, min.sz=1, max.sz=Inf)
## S4 method for signature 'GeneSetCollection'
filterGeneSets(gSets, min.sz=1, max.sz=Inf)
```

Arguments

gSets	Gene sets given either as a list or a GeneSetCollection object.
min.sz	Minimum size.
max.sz	Maximum size.

Details

This function filters the input gene sets according to a given minimum and maximum set size.

Value

A collection of gene sets that meet the given minimum and maximum set size.

Author(s)

J. Guinney

References

H\"anzelmann, S., Castelo, R. and Guinney, J. GSVA: Gene Set Variation Analysis, submitted

See Also

```
computeGeneSetsOverlap
```

Examples

```
geneSets <- list(set1=as.character(1:4), set2=as.character(4:10))
filterGeneSets(geneSets, min.sz=5)</pre>
```

gsva

Gene Set Variation Analysis

Description

Estimates GSVA enrichment scores.

Usage

```
## S4 method for signature 'ExpressionSet, list'
gsva(expr, gset.idx.list,
    abs.ranking=FALSE,
   min.sz=1,
   max.sz=Inf,
    no.bootstraps=0,
    bootstrap.percent = .632,
    parallel.sz=0,
    parallel.type="SOCK",
    verbose=TRUE,
    mx.diff=TRUE)
## S4 method for signature 'ExpressionSet, GeneSetCollection'
gsva(expr, gset.idx.list,
    abs.ranking=FALSE,
   min.sz=1,
    \max.sz=Inf
    no.bootstraps=0,
   bootstrap.percent = .632,
    parallel.sz=0,
    parallel.type="SOCK",
   verbose=TRUE,
   mx.diff=TRUE)
## S4 method for signature 'matrix, list'
gsva(expr, gset.idx.list,
    abs.ranking=FALSE,
    min.sz=1,
    \max.sz=Inf,
```

```
no.bootstraps=0,
bootstrap.percent = .632,
parallel.sz=0,
parallel.type="SOCK",
verbose=TRUE,
mx.diff=TRUE)
```

Arguments

expr Gene expression data which can be given either as an ExpressionSet object

or as a matrix of expression values where rows correspond to genes and columns

correspond to samples.

gset.idx.list

Gene sets provided either as a list object or as a GeneSetCollection

object.

abs.ranking Flag to determine whether genes should be ranked according to their sign (flag=FALSE)

or by absolute value (flag=TRUE). In the latter, pathways with genes enriched

on either extreme (high or low) will be regarded as 'highly' activated.

min.sz Minimum size of the resulting gene sets.

max.sz Maximum size of the resulting gene sets.

no.bootstraps

Number of bootstrap iterations to perform.

bootstrap.percent

.632 is the ideal percent samples bootstrapped.

parallel.sz Number of processors to use when doing the calculations in parallel. This

requires to previously load either the multicore or the snow library. If

multicore is loaded and this argument is left with its default value (parallel.sz=0) then it will use all available core processors unless we set this argument with a

smaller number. If snow is loaded then we must set this argument to a positive integer number that specifies the number of processors to employ in the parallel

calculation.

parallel.type

Type of cluster architecture when using snow.

verbose Gives information about each calculation step. Default: FALSE.

mx.diff Offers two approaches to calculate the enrichment statistic (ES) from the KS

random walk statistic. mx.diff=FALSE: ES is calculated as the maximum distance of the random walk from 0. mx.diff=TRUE (default): ES is calculated as the magnitude difference between the largest positive and negative

random walk deviations.

Details

GSVA assesses the relative enrichment of gene sets across samples using a non-parametric approach. Conceptually, GSVA transforms a p-gene by n-sample gene expression matrix into a geneset by n-sample pathway enrichment matrix. This facilitates many forms of statistical analysis in the 'space' of pathways rather than genes, providing a higher level of interpretability.

The gsva() function first maps the identifiers in the gene sets to the identifiers in the input expression data leading to a filtered collection of gene sets. This collection can be further filtered to require a minimum and/or maximum size of the gene sets for which we want to calculate GSVA enrichment scores, by using the arguments min.sz and max.sz.

Value

A gene-set by sample matrix of GSVA enrichment scores.

Author(s)

J. Guinney

References

H\"anzelmann, S., Castelo, R. and Guinney, J. GSVA: Gene Set Variation Analysis, submitted

See Also

filterGeneSets computeGeneSetsOverlap

Examples

```
library(limma)
p <- 10 ## number of genes
n <- 30 \ \# \ number of samples
nGrp1 <- 15 ## number of samples in group 1
nGrp2 <- n - nGrp1 ## number of samples in group 2
## consider three disjoint gene sets
geneSets <- list(set1=paste("g", 1:3, sep=""),</pre>
                 set2=paste("g", 4:6, sep=""),
                 set3=paste("g", 7:10, sep=""))
\#\# sample data from a normal distribution with mean 0 and st.dev. 1
y <- matrix(rnorm(n*p), nrow=p, ncol=n,
            dimnames=list(paste("g", 1:p, sep="") , paste("s", 1:n, sep="")))
## genes in set1 are expressed at higher levels in the last 10 samples
y[geneSets\$set1, (nGrp1+1):n] <- y[geneSets\$set1, (nGrp1+1):n] + 2
## build design matrix
design <- cbind(sampleGroup1=1, sampleGroup2vs1=c(rep(0, nGrp1), rep(1, nGrp2)))</pre>
## fit linear model
fit <- lmFit(y, design)</pre>
## estimate moderated t-statistics
fit <- eBayes(fit)</pre>
## genes in set1 are differentially expressed
topTable(fit, coef="sampleGroup2vs1")
## estimate GSVA enrichment scores for the three sets
gsva_es <- gsva(y, geneSets, mx.diff=1)$es.obs</pre>
## fit the same linear model now to the GSVA enrichment scores
fit <- lmFit(gsva_es, design)</pre>
## estimate moderated t-statistics
```

```
fit <- eBayes(fit)
## set1 is differentially expressed
topTable(fit, coef="sampleGroup2vs1")</pre>
```

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