Package 'ADAM'

October 14, 2021

Type Package

Title ADAM: Activity and Diversity Analysis Module

Version 1.8.0 **Date** 2018-10-5

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Description ADAM is a GSEA R package created to group a set of genes from comparative samples (control versus experiment) belonging to different species according to their respective functions (Gene Ontology and KEGG pathways as default) and show their significance by calculating p-values referring togene diversity and activity. Each group of genes is called GFAG (Group of Functionally Associated Genes).

Depends R(>=3.5), stats, utils, methods

Imports Rcpp (>= 0.12.18), GO.db (>= 3.6.0), KEGGREST (>= 1.20.2), knitr, pbapply (>= 1.3-4), dplyr (>= 0.7.6), DT (>= 0.4), stringr (>= 1.3.1), SummarizedExperiment (>= 1.10.1)

Suggests testthat

LinkingTo Rcpp

SystemRequirements C++11

Encoding UTF-8

RoxygenNote 6.1.0

LazyData TRUE

License GPL (>= 2)

biocViews GeneSetEnrichment, Pathways, KEGG

VignetteBuilder knitr

LazyLoad yes

NeedsCompilation yes

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git_url https://git.bioconductor.org/packages/ADAM
git_branch RELEASE_3_13
git_last_commit e062ab5
git_last_commit_date 2021-05-19
Date/Publication 2021-10-14

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Description

A sample fragment of gene expression from an RNA-seq experiment of Aedes aegypti mosquito.

Format

A data frame with 34 rows and 5 variables

References

AKBARI, O. S. et al. The developmental transcriptome of the mosquito aedes aegypti, an invasive species and major arbovirus vector. G3: Genes— Genomes— Genetics, Genetics Society of America, v. 3, n. 9, p. 1493–1509, 2013.

Examples

data(ExpressionAedes)

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GFAGAnalysis	Group of Functionally Associated Genes (GFAG) analysis

Description

Analysis of functionally associated gene groups, based on gene diversity and activity, for different species according to existing annotation packages or user's personal annotations.

Usage

```
GFAGAnalysis(ComparisonID, ExpressionData, MinGene, MaxGene, SeedNumber, BootstrapNumber, PCorrection, DBSpecies, PCorrectionMethod, WilcoxonTest, FisherTest, AnalysisDomain, GeneIdentifier)

ADAnalysis(ComparisonID, ExpressionData, MinGene, MaxGene,
```

Arguments

ComparisonID	Sample comparisons	identification.	It must be a	vector in which each el	lement
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corresponds to 2 sample columns from the expression data. The data's sample columns in each element from the vector are comma separated. The first one refers to the control sample, while the second refers to the experiment. This argument must be informed by the user. There is no default value for it.

ExpressionData Gene expression data (microarray or RNA-seq, for example). It must be a Sum-

marizedExperiment object, a data frame or a path for a text file tab separated containing at least 3 columns. First column mandatory corresponds to the gene identification, according to GeneIdentifier argument. Second, third and the other columns correspond to the gene expression values realated to the genes in the first column and each of these columns correspond to a different sample (control versus experiment). This argument must be informed by the user. There is no

default value for it.

DBSpecies, AnalysisDomain, GeneIdentifier)

MinGene Minimum number of genes per GFAG. It must be a positive integer value differ-

ent from zero and lower than MaxGene argument. Default is 3.

Maximum number of genes per GFAG. It must be a positive integer value dif-

ferent from zero and greater than MinGene argument. Default is 2000.

SeedNumber Seed for bootstrap. A numeric positive value used as seed for random number

generating by bootstrap function. Default is 1049.

BootstrapNumber

Number of bootstraps. A numeric value greater than zero, used by bootstrap

function generates p-values for each GFAG. Default is 1000.

PCorrection Cutoff for p-value correction. A numeric value between 0 and 1. Default is 0.05.

DBSpecies A string corresponding to the name of an OrgDb species gene annotation pack-

age: org.Ag.eg.db (Anopheles gambiae), org.At.tair.db (Arabdopsis thaliana), org.Bt.eg.db (Bos taurus), org.Ce.eg.db (Caenorhabditis elegans), org.Cf.eg.db

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(Canis familiaris), org.Dm.eg.db (Drosophila melanogaster), org.Dr.eg.db (Danio rerio), org.EcK12.eg.db (Escherichia coli K12), org.EcSakai.eg.db (Escherichia coli Sakai), org.Gg.eg.db (Gallus gallus), org.Hs.eg.db (Homo sapiens), org.Mm.eg.db (Mus musculus), org.Mmu.eg.db (Macaca mulatta), org.Pf.plasmo.db (Plasmodium falciparum), org.Pt.eg.db (Pan troglodytes), org.Rn.eg.db (Rattus norvegicus), org.Sc.sgd.db (Saccharomyces cerevisiae), org.Ss.eg.db (Sus scrofa) and org.Xl.eg.db (Xenopus laevis). If there is no package, it's possible for the user to create a personal gene annotation file, tab separated, containing 3 columns: gene, term annotation code and description of the term annotation. So, istead of a string with an OrgDb name, inform a data frame or a path for the file. This argument must be informed by the user. There is no default value for it.

PCorrectionMethod

Method p-value correction: holm, hochberg, hommel, bonferroni, bh, by or fdr.

Default is "fdr".

WilcoxonTest A logical value indicating whether or not to perform Wilcoxon test (TRUE for

performing and FALSE for not performing). Default is FALSE.

FisherTest A logical value indicating whether or not to perform Fisher's exact test (TRUE

for performing and FALSE for not performing). Default is FALSE.

AnalysisDomain Analysis domain to be considered for building GFAGs, according: gobp (Gene

Ontology - Biological Processes), gocc (Gene Ontology - Celular Components), gomf (Gene Ontology - Molecular Functions), kegg (KEGG Pathways) or own (if there is no annotation package - the annotations were defined in a file by the user). This argument must be informed by the user. There is no default value for

it.

GeneIdentifier Gene nomenclature to be used: entrez or tairID for Arabdopsis thaliana, entrez

or orfID for Saccharomyces cerevisiae, symbol or orfID for Plasmodium falciparum and symbol or entrez for the other species. If there is no annotation package, just put the gene nomenclature present in the user's personal annotations. This argument must be informed by the user. There is no default value for

it.

Details

The genes present in the expression data are grouped by their respective functions according to the domains described by AnalysisDomain argument. The relationship between genes and functions are made based on the species annotation package. If there is no annotation package, a three column file (gene, function and function description) must be provided. For each GFAG, gene diversity and activity in each sample are calculated. As the package always compare two samples (control versus experiment), relative gene diversity and activity for each GFAG are calculated. Using bootstrap method, for each GFAG, according to relative gene diversity and activity, two p-values are calculated. The p-values are then corrected, according to the correction method defined by PCorrectionMethod argument, generating a q-value. The significative GFAGs will be those whoose q-value stay under the cutoff set by PCorrection argument. Optionally, it's possible to run Wilcoxon test and/or Fisher's exact test. These tests also provide a corrected p-value, and significative groups can be seen through them. GFAGAnalysis function allows to run a complete analysis, using all available arguments. ADAnalysis function allows to run a partial analysis, where is calculated just gene diversity and activity of each GFAG with no signicance by bootrstrap, Wilcoxon or Fisher.

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Value

Return a list with two elements. The first one refers to a data frame with the GFAGs and their respective genes. The second one is a a list where each position is a data frame presenting the result of the analysis, according to ComparisonID argument.

Author(s)

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References

CASTRO, M. A., RYBARCZYK-FILHO, J. L., et al. Viacomplex: software for landscape analysis of gene expression networks in genomic context. Bioinformatics, Oxford Univ Press, v. 25, n. 11, p. 1468–1469, 2009.

Examples

```
#Complete Analysis with Aedes aetypti through GFAGAnalysis function
data(ExpressionAedes)
data(KeggPathwaysAedes)
ResultAnalysis <- GFAGAnalysis(ComparisonID = c("control1, experiment1",</pre>
"control2, experiment2"), ExpressionData = ExpressionAedes, MinGene = 3L,
MaxGene = 20L, SeedNumber = 1049, BootstrapNumber = 1000L,
PCorrection = 0.05, DBSpecies = KeggPathwaysAedes,
PCorrectionMethod = "fdr", WilcoxonTest = TRUE, FisherTest = TRUE,
AnalysisDomain = "own", GeneIdentifier = "gene")
## Not run:
head(ResultAnalysis[[1]]) #Relation between genes and functions
head(ResultAnalysis[[2]][1]) #Result comparison 1
head(ResultAnalysis[[2]][2]) #Result comparison 2
## End(Not run)
#Partial Analysis with Aedes aegypti through ADAnalysis function
data(ExpressionHs)
ResultAnalysis <- ADAnalysis(ComparisonID = c("control1,experiment1"),</pre>
ExpressionData = ExpressionAedes, MinGene = 3L, MaxGene = 20L,
DBSpecies = KeggPathwaysAedes, AnalysisDomain = "own",
GeneIdentifier = "geneStableID")
head(ResultAnalysis[[1]]) #Relation between genes and functions
head(ResultAnalysis[[2]][1]) #Result comparison 1
## End(Not run)
```

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KeggPathwaysAedes	Relation between Aedes aegypti genes and KEGG pathways as ADAM input

Description

A relation between the genes in the ExpressionAedes data and their respective KEGG pathways (GFAGs).

Format

A data frame with 200 rows and 2 variables

References

Molan, A. L. 2018. "Construction of a Tool for Multispecies Genic Functional Enrichment Analysis Among Comparative Samples." Master's thesis Institute of Biosciences of Botucatu – Univ. Estadual Paulista. http://hdl.handle.net/11449/157105.

Examples

data(KeggPathwaysAedes)

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