Package 'PubScore'

October 17, 2020

Type Package

Title Automatic calculation of literature relevance of genes

Version 1.0.0

Description
Calculates the importance score for a given gene. The importance score is the total counts of articles in the pubmed database that are a result for that gene AND each term of a list.

Imports ggplot2, igraph, ggrepel, rentrez, progress, graphics, dplyr, utils, methods, intergraph, network, sna

Suggests FCBF, plotly, SummarizedExperiment, SingleCellExperiment, knitr, rmarkdown, testthat (>= 2.1.0), BiocManager, biomaRt

biocViews GeneSetEnrichment, GeneExpression, SystemsBiology, Genetics, Epigenetics, BiomedicalInformatics, Visualization, SingleCell

VignetteBuilder knitr

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Author Tiago Lubiana [aut, cre],
Helder Nakaya [aut, ths]

Maintainer Tiago Lubiana <tiago.lubiana.alves@usp.br>

R topics documented:

.getSimulation_test   *Auxiliary function for the test method*

Description

Auxiliary function for the test method

Usage

```
.getSimulation_test(pub, ambiguous = c(), n_simulations)
```

Arguments

pub	An object of class `PubScore`
ambiguous	A character vector with possible ambiguous gene names
n_simulations	The number of simulations to run.

Value

A data-frame with a simulation of literature scores for random samplings

---

.query_pubmed   *#' .query_pubmed*

Description

Auxiliary function for getting the list score

Usage

```
.query_pubmed(search_topic, wait_time = 0, ret_max = 1)
```

Arguments

search_topic	Item to search on PubMed via rentrez
wait_time	Time between searches
ret_max	Number of IDs to be returned. Defaults to 1.

Value

The rentrez search result (a list)

---

all_counts	*all_counts*

Description

A dataframe with all pubmed counts for the genes in the Dengue dataset in relation to the term "Dengue".

Usage

```
data(all_counts)
```

Format

An object of class `data.frame`

Details

Outcome of the test_score method of the pubscore class. As this function may take a long time, this dataset speeds up the vignette.

Contains: 3 columns: #tax_id: The reference ID for the taxon. All are 9606 (humans). GeneID: The Entrez ID code for a given gene. PubMedID: A PubMed ID for a paper that mentions the gene in the "Gene ID" column.

1335548 rows: gene-paper associations in the gene2pubmed database.

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gene2pubmed_db	*human genes on gene2pubmed_db*

Description

A subset of the gene2pubmed database downloaded via FTP from ftp://ftp.ncbi.nlm.nih.gov/gene/DATA/gene2pubmed.gz #' The subset contains only the rows corresponding to humans (#tax_id = 906) The table was downloaded in October 2019.

Usage

```
data(gene2pubmed_db)
```

Format

An object of class `data.frame`

Details

Contains: 3 columns: #tax_id: The reference ID for the taxon. All are 9606 (humans). GeneID: The Entrez ID code for a given gene. PubMedID: A PubMed ID for a paper that mentions the gene in the "Gene ID" column.

1335548 rows: gene-paper associations in the gene2pubmed database.

References

Maglott, Donna, et al. 'Entrez Gene: gene-centered information at NCBI.' Nucleic acids research 33.suppl_1 (2005): D54-D58.

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getScore   *Retrieve the literature_score attribute*

Description

Retrieve the literature_score attribute

Usage

```
getScore(pub)

## S4 method for signature 'PubScore'
getScore(pub)
```

Arguments

pub   Object of class `PubScore`

Value

A "numeric" with the literature score for this gene x term combination

Examples

```
# Create a new pubscore object
pub <- pubscore(genes = c('cd4','cd8'),terms_of_interest = c('blabla','immunity'))
plot(networkViz(pub))
```

---

get_all_counts   *Retrieve the all_counts attribute*

Description

Retrieve the all_counts attribute

Usage

```
get_all_counts(pub)

## S4 method for signature 'PubScore'
get_all_counts(pub)
```

Arguments

pub   Object of class PubScore

Value

A dataframe containing the counts table for all genes.

Examples

```
# Create a new pubscore object
pub <- pubscore(genes = c('cd4','cd8'),terms_of_interest = c('blabla','immunity'))
plot(networkViz(pub))
```

---

get_literature_score   *get_literature_score*

Description

Calculates the importance score for a given gene. The importance score is the total counts of articles
in the pubmed database that are a result for that gene AND each term of a list

Usage

```
get_literature_score(genes, terms_of_interest, gene2pubmed = FALSE,
  return_all = FALSE, wait_time = 0, show_progress = TRUE,
  verbose = FALSE)
```

Arguments

genes   A vector with multiple genes.

terms_of_interest
                       A list of terms of interest related to the topic you want to find the relevance for

gene2pubmed   logical defining if gene2pubmed db is going to be used. If used, the vector of
                       genes has to be of HUMAN genes in the hgcn_symbol format.

return_all	Only to be used with gene2pubmed! logical defining if the all_counts table is going to be returned here. Usually it is generated by the test_score function.
wait_time	How long should be the interval between two requests to the ENTREZ database when it fails. Defaults to 0. In seconds.
show_progress	If TRUE, a progress bar is displayed. Defaults to TRUE.
verbose	If TRUE, will display the index of the search occuring. Defaults to FALSE.

Value

A dataframe with the literature scores.

Examples

```
genes <- c('CD8A', 'CD4')
terms_of_interest <-
  c(
    "CD4 T cell",
    "CD14+ Monocyte",
    "B cell",
    "CD8 T cell",
    "FCGR3A+ Monocyte",
    "NK cell",
    "Dendritic cell",
    "Megakaryocyte",
    'immunity'
  )
get_literature_score(genes, terms_of_interest)
```

---

heatmapViz   *Retrieve the heatmap attribute*

Description

Retrieve the heatmap attribute

Usage

```
heatmapViz(pub)

## S4 method for signature 'PubScore'
heatmapViz(pub)
```

Arguments

pub	Object of class PubScore

Value

A "gg" object, from ggplot2, containing a heatmap from the counts table.

Examples

```
#Create a new pubscore object
pub <- pubscore(genes = c('cd4','cd8'),terms_of_interest = c('blabla','immunity'))
plot(heatmapViz(pub))
```

---

hgcn_entrez_reference   *hgcn_entrez_reference*

Description

Contains the result of a query to the biomaRt service done in October, 2019.

Usage

```
data(hgcn_entrez_reference)
```

Format

An object of class `data.frame`

Details

2 columns: entrezgene_id (containing the Entrez ids) and hgnc_symbol (containing gene symbols from the HUGO gene nomenclature consortium)

20491 rows, for the mapping between the two nomenclatures for human genes.

References

Mapping identifiers for the integration of genomic datasets with the R/Bioconductor package biomaRt. Steffen Durinck,Paul T. Spellman, Ewan Birney and Wolfgang Huber, Nature Protocols 4, 1184-1191 (2009).

---

initialize,PubScore-method
                                initialize

Description

initialize

Usage

```
## S4 method for signature 'PubScore'
initialize(.Object, genes, terms_of_interest,
  gene2pubmed = FALSE)
```

Arguments

`.Object`	The object of signature PubScore that is foinf to be created
`genes`	The genes to which you want to calculate and visualize the literature score.
`terms_of_interest`	
	A list of terms of interest related to the topic you want to find the relevance.
`gene2pubmed`	Logical (TRUE / FALSE) defining if gene2pubmed db is going to be used.

Value

A object of the PubScore class

`networkViz`	*Retrieve the network attribute*

Description

Retrieve the network attribute

Usage

```
networkViz(pub)

## S4 method for signature 'PubScore'
networkViz(pub)
```

Arguments

`pub`	Object of class `PubScore`

Value

A "gg" object, from ggplot2, containing a network from the counts table.

Examples

```
# Create a new pubscore object
pub <- pubscore(genes = c('cd4','cd8'),terms_of_interest = c('blabla','immunity'))
plot(networkViz(pub))
```

---

plot_literature_graph *#'plot_literature_graph*

Description

Plot a graph inspired in CEMiTool's graphs

Usage

```
plot_literature_graph(plot_counts, name, color = "#B30000FF",
  max_number_of_labels = 10)
```

Arguments

`plot_counts`	The dataframe returned from the get_literature_score function
`name`	The name of the plot.
`color`	The color of the plot. Defaults to a shade of red ("#B30000FF").
`max_number_of_labels`	
	The max number of gene labels to show. Defaults to 10.

Value

A ploty/ggplot2 object is either returned or directly plotted

Examples

```
gene <- c('CD4','CD14', "AIF1", "ACVR1", "CDY2A")
terms_of_interest <- c("CD4 T cell", "CD14+ Monocyte")
literature_counts <- get_literature_score(gene, terms_of_interest)
 pl <- plot_literature_graph(literature_counts, name = 'test')
 pl
```

---

plot_literature_score *plot_literature_score*

Description

Plots a non-clusterized heatmap of the article counts for the combination of gene list and list of terms NOTE: the object has to be exactly the one returned by get_literature_score.R . Otherwise ggplot2 will not be able to identify the correct columns.

Usage

```
plot_literature_score(plot_counts, return_ggplot = FALSE,
  is_plotly = FALSE)
```

Arguments

`plot_counts`	The dataframe returned from the get_literature_score function
`return_ggplot`	If TRUE, returns a ggplot2 object instead of plotting. Defaults to FALSE.
`is_plotly`	If TRUE, a interactive plot is plotted in the place o static ggplot. Defaults to FALSE.

Value

A ggplot2 object is either returned or directly plotted

Examples

```
gene <- c('CD4','CD14', "AIF1", "ACVR1", "CDY2A")
terms_of_interest <- c("CD4 T cell", "CD14+ Monocyte", "B cell",
"CD8 T cell","FCGR3A+ Monocyte", "NK cell", "Dendritic cell",
"Megakaryocyte", 'immunity')
literature_counts <- get_literature_score(gene, terms_of_interest)
P <-plot_literature_score(literature_counts, return_ggplot = TRUE)
plot(P)
```

---

pubscore	*PubScore fundamental analysis*

Description

Runs the initalization and the basic functions for querying pubmed and getting the literature scores.

Usage

```
pubscore(terms_of_interest, genes, gene2pubmed = FALSE)
```

Arguments

`terms_of_interest`	
	A list of terms of interest related to the topic you want to find the relevance for
`genes`	A vector with multiple genes.
`gene2pubmed`	Logical (TRUE / FALSE) defining if gene2pubmed db is going to be used. Defaults to FALSE.

Value

Object of class `PubScore`

PubScore-class   *An S4 class to represent PubScore results*

Description

The S4 class to PubScore and its basic initializate and show methods.

Slots

terms_of_interest A list of terms of interest related to the topic you want to find the relevance.

genes The genes to which you want to calculate and visualize the literature score.

date The date when the object was initialized. PubScore counts will likely increase with time.

gene2pubmed Logical (TRUE / FALSE) noting if gene to pubmed was used or not.

counts A data frame with the counts retrieved on PubMed

network A visualization of the results found in a network

heatmap A visualization of the results found in a heatmap

---

set_all_counts<-   *Set the all_counts attribute*

Description

Set the all_counts attribute

Usage

```
set_all_counts(pub) <- value

## S4 replacement method for signature 'PubScore'
set_all_counts(pub) <- value
```

Arguments

pub	Object of class PubScore
value	The table with all gene x term article counts from the "test_score" method.

Value

A dataframe containing the counts table for all genes.

Examples

```
terms_of_interest <- c('Dengue')
pub <- pubscore(terms_of_interest = terms_of_interest, genes = c("CD4", "CD8", "CD14") )
print(getScore(pub))
data("all_counts")
set_all_counts(pub) <- all_counts
```

---

test_score   *Test the literature enrichment score*

Description

Test the literature enrichment score

Usage

```
test_score(pub, total_genes, show_progress = TRUE,
  remove_ambiguous = TRUE, verbose = FALSE, nsim = 1e+05,
  ambiguous_terms = c("PC", "JUN", "IMPACT", "ACHE", "SRI", "SET", "CS",
  "PROC", "MET", "SHE", "CAD", "DDT", "PIGS", "SARS", "REST", "GC", "CP",
  "STAR", "SI", "GAN", "MARS", "SDS", "AGA", "NHS", "CPE", "POR", "MAX",
  "CAT", "LUM", "ANG", "POLE", "CLOCK", "TANK", "ITCH", "SDS", "AES",
  "CIC", "FST", "CAPS", "COPE", "F2", "AFM", "SPR", "PALM", "C2", "BAD",
  "GPI", "CA2", "SMS", "INVS", "WARS", "HP", "GAL", "SON", "AFM", "BORA",
  "MBP", "MAK", "MALL", "COIL", "CAST "))

## S4 method for signature 'PubScore'
test_score(pub, total_genes, show_progress = TRUE,
  remove_ambiguous = TRUE, verbose = FALSE, nsim = 1e+05,
  ambiguous_terms = c("PC", "JUN", "IMPACT", "ACHE", "SRI", "SET", "CS",
  "PROC", "MET", "SHE", "CAD", "DDT", "PIGS", "SARS", "REST", "GC", "CP",
  "STAR", "SI", "GAN", "MARS", "SDS", "AGA", "NHS", "CPE", "POR", "MAX",
  "CAT", "LUM", "ANG", "POLE", "CLOCK", "TANK", "ITCH", "SDS", "AES",
  "CIC", "FST", "CAPS", "COPE", "F2", "AFM", "SPR", "PALM", "C2", "BAD",
  "GPI", "CA2", "SMS", "INVS", "WARS", "HP", "GAL", "SON", "AFM", "BORA",
  "MBP", "MAK", "MALL", "COIL", "CAST "))
```

Arguments

pub	Object of class PubScore
total_genes	A list of all the possible genes in your study. Usually all the names in the rows of an "exprs" object.
show_progress	If TRUE, a progress bar is displayed. Defaults to True.
remove_ambiguous	
	If TRUE, ambiguously named genes (such as "MARCH") will be removed. Defaults to TRUE.
verbose	If TRUE, will display the index of the search occuring. Defaults to false.
nsim	The number of simulations to run. Defaults to 100000.
ambiguous_terms	
	A character vector of the ambiguous terms to use instead of the default. The default includes 60 genes pre-selected as ambiguous (as IMPACT, MARCH and ACHE).

Value

A "gg" object, from ggplot2, containing a network from the counts table.

Examples

```
# Create a new pubscore object
pub <- pubscore(genes = c('cd4','cd8'),
terms_of_interest = c('blabla','immunity'))
pub <- test_score(pub,
total_genes = c('notagene1', 'notagene2', 'cd4', 'cd8'),
remove_ambiguous = TRUE)
```

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