Package 'BioEnricher'

December 6, 2023

seamless integration for enrichment analysis, encompassing diverse functionalities

Title Integrate Analysis and Visualization for Bioinformatic Enrichment Analyzer

Description This package lies in addressing two issues: firstly, it facilitates the

```
such as GO, KEGG, WikiPathways, Reactome, MsigDB, Disease Ontology, Cancer Gene
     Network, DisGeNET, CellMarker, and CMAP (drugs); infers the activities of
      transcription factors and PROGENy cancer pathways; searches the gene information,
     PubMed records and GEO metadata based on the input terms. Secondly, it encapsulates
     advanced visualization functions, streamlining the process for faster and more
     convenient data presentation.
License MIT + file LICENSE
Encoding UTF-8
ByteCompile true
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RoxygenNote 7.2.3
Depends R (>= 4.3.0)
Imports clusterProfiler,
     dorothea,
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     europepmc,
     enrichplot,
     ggplot2,
     GSVA,
     HGNChelper,
     Hmisc,
     magrittr,
     msigdbr,
     openssl,
     pathview,
     progeny,
     ReactomePA,
     rlang,
      vroom,
     RColorBrewer,
```

Type Package

Version 0.1.0

RCurl, cowplot,

2 R topics documented:

fst, paletteer, showtext, stringi, xml2

Suggests knitr, rmarkdown

VignetteBuilder knitr

biocViews Software, GeneSetEnrichment, Visualization

 $\pmb{BugReports} \ \, \texttt{https://github.com/Zaoqu-Liu/BioEnricher/issues}$

 $\boldsymbol{URL} \text{ https://github.com/Zaoqu-Liu/BioEnricher}$

Additional_repositories https://bioconductor.org/packages/release/bioc

R topics documented:

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CMAPfromDSEATM

A dataframe including drugs and their related genes

Description

A dataframe including drugs and their related genes

Usage

CMAPfromDSEATM

Format

A dataframe with four columns from DSEATM

cols_brown_green

A vector of colors

Description

A vector of colors

Usage

```
cols_brown_green
```

Format

A vector with 11 types of colors.

gene.info

Get gene related information

Description

Get the basic information of genes. This function is from genekitr package.

```
gene.info(
  id = NULL,
  org = "hs",
  unique = FALSE,
  keepNA = TRUE,
  hgVersion = c("v38", "v19")
)
```

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Arguments

id Gene id (symbol, ensembl or entrez id) or uniprot id. If this argument is NULL,

return all gene info.

org Latin organism shortname from ensOrg_name. Default is human.

unique Logical, if one-to-many mapping occurs, only keep one record with fewest NA.

Default is FALSE.

keepNA If some id has no match at all, keep it or not. Default is TRUE.

hgVersion Select human genome build version from "v38" (default) and "v19".

Value

A data.frame.

Examples

```
# input list with fake id and one-to-many mapping id
x <- gene.info(id = c(
  "MCM10", "CDC20", "S100A9", "MMP1", "BCC7",
  "FAKEID", "TP53", "HBD", "NUDT10"
))
# use hg19 data
x <- gene.info(id = c("TP53", "BCC7"), hgVersion = "v19")</pre>
```

listEnrichMethod

List of enrichment methods

Description

List of enrichment methods, including GO, KEGG, MKEGG, WikiPathways, Reactome, MsigDB, DO, CGN, DisGeNET, CellMarker, and CMAP.

Usage

```
listEnrichMethod()
```

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

Examples

listEnrichMethod()

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lzq_getEF

Get enrichment factor from enrichResult

Description

Get enrichment factor from enrichResult (clusterProfiler).

Usage

```
lzq_getEF(res)
```

Arguments

res

enrichResult from clusterProfiler.

Value

A new result with enrichment factor.

Author(s)

```
Zaoqu Liu; E-mail: liuzaoqu@163.com
```

Examples

```
genes <- c("CANX", "HSPA1B", "KLRC2", "PSMC6", "RFXAP", "TAP1")

obj <- clusterProfiler::enrichGO(genes, org.Hs.eg.db::org.Hs.eg.db,
  keyType = "SYMBOL", ont = "BP"
)

obj2 <- BioEnricher::lzq_getEF(obj)
obj2@result$EnrichmentFactor</pre>
```

lzq_getGR_BR

Get numeric GeneRatio and BgRatio from enrichResult

Description

Get numeric GeneRatio and BgRatio from enrichResult (clusterProfiler).

Usage

```
lzq_getGR_BR(res)
```

Arguments

res

enrichResult from clusterProfiler.

Value

A new result with numeric GeneRatio and BgRatio.

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Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

Examples

```
genes <- c("CANX", "HSPA1B", "KLRC2", "PSMC6", "RFXAP", "TAP1")

obj <- clusterProfiler::enrichGO(genes, org.Hs.eg.db::org.Hs.eg.db,
   keyType = "SYMBOL", ont = "BP"
)

obj2 <- BioEnricher::lzq_getGR_BR(obj)

obj2@result$GeneRatio
obj2@result$BgRatio</pre>
```

1zq_GSEA

Gene set enrichment analysis

Description

Perform gene set enrichment analysis included GO, KEGG, WikiPathways, Reactome, MsigDB, Disease Ontoloty, Cancer Gene Network, DisGeNET, CellMarker, and CMAP.

Usage

```
lzq_GSEA(
   genes,
   gene.type = "SYMBOL",
   enrich.type,
   organism = "Human",
   GO.ont = "BP",
   GO.simplify = T,
   KEGG.use.internal.data = F,
   MsigDB.category = "H",
   CMAP.min.Geneset.Size = 3,
   pvalue.cutoff = 0.05,
   padjust.method = "BH",
   min.Geneset.Size = 10,
   max.Geneset.Size = 1000
)
```

Arguments

gene.type Keytype of input gene.

enrich.type Select an enrichment method. One of GO, KEGG, MKEGG, WikiPathways, Reactome, MsigDB, DO, CGN, DisGeNET, CellMarker, and CMAP. WikiPathways can be replaced by WP, Reactome can be replaced by RP, and CellMarker can be replaced by CM.

organism Specify species, currently support only Human and Mouse.

GO ont GO parameter. One of "BP", "MF", and "CC" subontologies, or "ALL" for all three.

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```
GO.simplify
                 GO parameter. Whether to remove redundancy of enriched GO terms.
KEGG.use.internal.data
                 KEGG parameter. Logical, use KEGG.db or latest online KEGG data.
MsigDB.category
                 MsigDB parameter. MSigDB collection abbreviation, such as All, H, C1, C2,
                 C3, C4, C5, C6, C7.
CMAP.min.Geneset.Size
                 CMAP parameter. Minimal size of CMAP genes annotated for testing. Recom-
                 mended use 3.
pvalue.cutoff
                 pvalue cutoff on enrichment tests to report as significant.
padjust.method one of "holm", "hochberg", "hommel", "bonferroni", "BH", "BY", "fdr", "none".
min.Geneset.Size
                 Minimal size of genes annotated for testing. Not suitable for CMAP.
max.Geneset.Size
```

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Maximal size of genes annotated for testing.

Author(s)

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```
library(airway)
library(DESeq2)
library(tidyverse)
library(clusterProfiler)
library(org.Hs.eg.db)
data(airway)
se <- airway
se$dex <- relevel(se$dex, "untrt")</pre>
res <- DESeqDataSet(se, design = ~ cell + dex) %>%
  estimateSizeFactors() %>%
  DESeq() %>%
  results() %>%
  as.data.frame() %>%
ann <- bitr(rownames(res), "ENSEMBL", "SYMBOL", org.Hs.eg.db)</pre>
res <- merge(ann, res, by.x = 1, by.y = 0) %>% dplyr::distinct(SYMBOL, .keep_all = TRUE)
# Obtain an order ranked geneList.
grlist <- res$log2FoldChange</pre>
names(grlist) <- res$SYMBOL</pre>
grlist <- sort(grlist, decreasing = TRUE)</pre>
# Set enrich.type using an enrichment analysis method mentioned above.
fit <- lzq_GSEA(grlist, enrich.type = "GO")</pre>
```

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lzq_GSEA.barplot1

Enrichment barplot for positive or negative GSEA results

Description

Plot enrichment barplot for positive or negative GSEA results.

Usage

```
lzq_GSEA.barplot1(
  enrich.obj,
  type = "Positive",
  show.term.num = 15,
  Selct.P = "FDR",
  cutoff.P = 0.05,
 colors = c("#003c30", "#01665e", "#35978f", "#80cdc1", "#c7eae5", "#f5f5f5", "#f6e8c3",
    "#dfc27d", "#bf812d", "#8c510a", "#543005"),
  add.bar.border = T,
 bar.width = 0.6,
 y.label.position = "right",
  title = NULL,
  legend.position = "bottom",
  theme.plot = theme_bw(base_rect_size = 1.5),
 use.Chinese = F,
  appid = "20231122001888718"
  key = "5GpDqe8F3pmXfn0kEKGQ"
)
```

Arguments

key

enrich.obj A GSEA enrichment object from clusterProfiler. Specify whether you want to show positive or negative results. type show.term.num A number or a list of terms. If it is a number, the first n terms will be displayed. If it is a list of terms, the selected terms will be displayed. Selct.P Nominal P value (NP) or adjust P value (FDR) were selected to define significant terms. cutoff.P A cutoff value for Select P. A color vector for the bars. colors add.bar.border Logical. Whether to add the black border of bars. bar.width Width of bar in the plot. y.label.position Y label position. right or left. title Title of the plot. legend.position Position of legend. 'none', 'right', 'left' or two numeric variables. theme.plot ggtheme of plot. use.Chinese Logical. Whether to use Chinese annotation in the barplot. User app id from baidu translation api. https://fanyi-api.baidu.com/manage/developer. appid User Key from baidu translation api. https://fanyi-api.baidu.com/manage/developer.

lzq_GSEA.barplot2

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

Examples

```
library(airway)
library(DESeq2)
library(tidyverse)
library(clusterProfiler)
library(org.Hs.eg.db)
data(airway)
se <- airway
se$dex <- relevel(se$dex, "untrt")</pre>
res <- DESeqDataSet(se, design = ~ cell + dex) %>%
  estimateSizeFactors() %>%
 DESeq() %>%
 results() %>%
 as.data.frame() %>%
 na.omit()
ann <- bitr(rownames(res), "ENSEMBL", "SYMBOL", org.Hs.eg.db)</pre>
res <- merge(ann, res, by.x = 1, by.y = 0) %>% dplyr::distinct(SYMBOL, .keep_all = TRUE)
# Obtain an order ranked geneList.
grlist <- res$log2FoldChange</pre>
names(grlist) <- res$SYMBOL</pre>
grlist <- sort(grlist, decreasing = TRUE)</pre>
# Integrative enrichment analysis of the ranked gene list
fit <- lzq_GSEA.integrated(genes = grlist)</pre>
lzq_GSEA.barplot1(enrich.obj = fit$simplyGO, type = "pos")
```

lzq_GSEA.barplot2

Enrichment barplot for positive and negative GSEA results

Description

Plot enrichment barplot for positive and negative GSEA results.

```
lzq_GSEA.barplot2(
  enrich.obj,
  Selct.P = "FDR",
  cutoff.P = 0.05,
  types = c("Positive", "Negative"),
  type.colors = c("#ED6355", "#3E94B5"),
  pos.top.pathway.num = 10,
  neg.top.pathway.num = 10,
  bar.width = 0.6,
  add.bar.border = T,
  x.limit.fold = 1.05,
  label.size = 3.5,
```

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```
legend.position = "bottom",
use.Chinese = F,
appid = "20231122001888718",
key = "5GpDqe8F3pmXfnOkEKGQ")
```

Arguments

enrich.obj A GSEA enrichment object from clusterProfiler.

Selct.P Nominal P value (NP) or adjust P value (FDR) were selected to define significant

terms.

cutoff.P A cutoff value for Select_P.

types Two characters for defining the types of two objects.

type.colors Two colors for the types of two objects.

pos.top.pathway.num

The number of top pathways in positive terms. Based on the significant test.

neg.top.pathway.num

The number of top pathways in negative terms. Based on the significant test.

bar.width Width of bar in the plot.

add.bar.border Logical. Whether to add the black border of bars.

x.limit.fold Specify the fold of x limitation. Because some terms is too long.

label.size Fontsize of label.

legend.position

none, left, right, top, bottom; Or Two numeric variables indicated x and y posi-

tions, respectively.

use. Chinese Logical. Whether to use Chinese annotation in the barplot.

appid User app id from baidu translation api. https://fanyi-api.baidu.com/manage/developer.

key User Key from baidu translation api. https://fanyi-api.baidu.com/manage/developer.

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

```
library(airway)
library(DESeq2)
library(tidyverse)
library(clusterProfiler)
library(org.Hs.eg.db)
data(airway)
se <- airway
se$dex <- relevel(se$dex, "untrt")</pre>
res <- DESeqDataSet(se, design = ~ cell + dex) %>%
  estimateSizeFactors() %>%
 DESeq() %>%
  results() %>%
  as.data.frame() %>%
  na.omit()
ann <- bitr(rownames(res), "ENSEMBL", "SYMBOL", org.Hs.eg.db)</pre>
res <- merge(ann, res, by.x = 1, by.y = 0) %>% dplyr::distinct(SYMBOL, .keep_all = TRUE)
```

lzq_GSEA.dotplot1

```
# Obtain an order ranked geneList.
grlist <- res$log2FoldChange
names(grlist) <- res$SYMBOL
grlist <- sort(grlist, decreasing = TRUE)

# Integrative enrichment analysis of the ranked gene list
fit <- lzq_GSEA.integrated(genes = grlist)

lzq_GSEA.barplot2(enrich.obj = fit$simplyGO)</pre>
```

lzq_GSEA.dotplot1

Enrichment dotplot for positive or negative GSEA results

Description

Plot enrichment dotplot for positive or negative GSEA results.

Usage

```
lzq_GSEA.dotplot1(
  enrich.obj,
  type = "neg",
  show.term.num = 15,
  Selct.P = "FDR",
  cutoff.P = 0.05,
 colors = c("#003c30", "#01665e", "#35978f", "#80cdc1", "#c7eae5", "#f5f5f5", "#f6e8c3",
    "#dfc27d", "#bf812d", "#8c510a", "#543005"),
  size.range = c(3, 8),
  y.label.position = "right",
  title = NULL,
  legend.position = "bottom",
  theme.plot = theme_bw(base_rect_size = 1.5),
  use.Chinese = F,
  appid = "20231122001888718",
  key = "5GpDqe8F3pmXfnOkEKGQ"
)
```

Arguments

enrich.obj	A GSEA enrichment object from clusterProfiler.
type	Specify whether you want to show positive or negative results.
show.term.num	A number or a list of terms. If it is a number, the first n terms will be displayed. If it is a list of terms, the selected terms will be displayed.
Selct.P	Nominal P value (NP) or adjust P value (FDR) were selected to define significant terms.
cutoff.P	A cutoff value for Select_P.
colors	A color vector for the bars.
size.range	Two numeric variables, the first is minimal value and the first is maximal value.

```
y.label.position
Y label position. right or left.

title Title of the plot.

legend.position
Position of legend. 'none', 'right', 'left' or two numeric variables.

theme.plot ggtheme of plot.

use.Chinese Logical. Whether to use Chinese annotation in the barplot.

appid User app id from baidu translation api. https://fanyi-api.baidu.com/manage/developer.

key User Key from baidu translation api. https://fanyi-api.baidu.com/manage/developer.
```

Author(s)

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Examples

```
library(airway)
library(DESeq2)
library(tidyverse)
library(clusterProfiler)
library(org.Hs.eg.db)
data(airway)
se <- airway
se$dex <- relevel(se$dex, "untrt")</pre>
res <- DESeqDataSet(se, design = ~ cell + dex) %>%
  estimateSizeFactors() %>%
  DESeq() %>%
  results() %>%
  as.data.frame() %>%
 na.omit()
ann <- bitr(rownames(res), "ENSEMBL", "SYMBOL", org.Hs.eg.db)</pre>
res <- merge(ann, res, by.x = 1, by.y = 0) \% dplyr::distinct(SYMBOL, .keep_all = TRUE)
# Obtain an order ranked geneList.
grlist <- res$log2FoldChange</pre>
names(grlist) <- res$SYMBOL</pre>
grlist <- sort(grlist, decreasing = TRUE)</pre>
# Integrative enrichment analysis of the ranked gene list
fit <- lzq_GSEA.integrated(genes = grlist)</pre>
lzq_GSEA.dotplot1(enrich.obj = fit$simplyGO, type = "pos")
```

Description

Perform integrated gene set enrichment analysis included GO, KEGG, WikiPathways, Reactome, MsigDB, Disease Ontoloty, Cancer Gene Network, DisGeNET, CellMarker, and CMAP.

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Usage

```
lzq_GSEA.integrated(
  genes,
  gene.type = "SYMBOL",
  organism = "Human",
  GO.ont = "BP",
  KEGG.use.internal.data = F,
  perform.WikiPathways = F,
  perform.Reactome = F,
  perform.MsigDB = F,
  MsigDB.category = "H",
  perform.disease.ontoloty = F,
  perform.Cancer.Gene.Network = F,
  perform.DisGeNET = F,
  perform.CellMarker = F,
  perform.CMAP = T,
  pvalue.cutoff = 0.05,
  padjust.method = "BH",
  min.Geneset.Size = 10,
  max.Geneset.Size = 1000,
  CMAP.min.Geneset.Size = 3
)
```

Arguments

perform.CMAP

pvalue.cutoff

```
A vector of gene id.
genes
                 Keytype of input gene.
gene.type
organism
                 Specify species, currently support only Human and Mouse.
GO.ont
                 One of "BP", "MF", and "CC" subontologies, or "ALL" for all three.
KEGG.use.internal.data
                 Logical, use KEGG.db or latest online KEGG data.
perform.WikiPathways
                 Whether to perform WikiPathways enrichment.
perform.Reactome
                 Whether to perform Reactome enrichment.
perform. MsigDB Whether to perform MsigDB enrichment.
MsigDB.category
                 MSigDB collection abbreviation, such as All, H, C1, C2, C3, C4, C5, C6, C7.
perform.disease.ontoloty
                 Whether to perform DO enrichment.
perform.Cancer.Gene.Network
                 Whether to perform CGN enrichment.
perform.DisGeNET
                 Whether to perform DisGeNET enrichment.
perform.CellMarker
```

pvalue cutoff on enrichment tests to report as significant.

Whether to perform CellMarker enrichment. Marker from cellmarker database. Whether to perform CMAP enrichment. Marker from CMAP database (in DSEATM

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```
padjust.method one of "holm", "hochberg", "hommel", "bonferroni", "BH", "BY", "fdr", "none".

min.Geneset.Size

Minimal size of genes annotated for testing. Not suitable for CMAP.

max.Geneset.Size

Maximal size of genes annotated for testing.

CMAP.min.Geneset.Size

Minimal size of CMAP genes annotated for testing. Recommended use 3.
```

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

Examples

```
library(airway)
library(DESeq2)
library(tidyverse)
library(clusterProfiler)
library(org.Hs.eg.db)
data(airway)
se <- airway
se$dex <- relevel(se$dex, "untrt")</pre>
res <- DESeqDataSet(se, design = ~ cell + dex) %>%
  estimateSizeFactors() %>%
 DESeq() %>%
  results() %>%
  as.data.frame() %>%
ann <- bitr(rownames(res), "ENSEMBL", "SYMBOL", org.Hs.eg.db)</pre>
res <- merge(ann, res, by.x = 1, by.y = 0) %>% dplyr::distinct(SYMBOL, .keep_all = TRUE)
# Obtain an order ranked geneList.
grlist <- res$log2FoldChange</pre>
names(grlist) <- res$SYMBOL</pre>
grlist <- sort(grlist, decreasing = TRUE)</pre>
# Integrative enrichment analysis of the ranked gene list
fit <- lzq_GSEA.integrated(genes = grlist)</pre>
```

lzq_gseaplot

Visualize analyzing result of GSEA.

Description

Visualize analyzing result of GSEA.

```
lzq_gseaplot(
  GSEA.result,
  Pathway.ID,
  heatbar = T,
  rank = T,
```

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```
line.color = "#41A98E",
  rank.colors = viridis::viridis(10),
  heatbar.colors = c(rev(RColorBrewer::brewer.pal(5, "Blues")),
    RColorBrewer::brewer.pal(5, "Reds")),
  add.x.ann = T,
  x.lab = "Gene ranks",
  line.y.lab = "Enrichment score",
  rank.y.lab = "logFC",
  statistic.position = c(0.5, 0.2),
  statistic.face = "italic",
  statistic.size = 3.5,
  rel.heights = c(1.5, 0.2, 1),
  theme.plot = theme_bw(base_rect_size = 1.5)
)
```

Arguments

GSEA.result GSEA results from clusterProfiler::GSEA() function. Corresponding pathway term of the output plot. Pathway.ID Whether to add heatbar. Default True. heatbar rank Whether to add Rank map. Default True. line.color Line color for running score. rank.colors Color scheme of rank lines. A vector. heatbar.colors Color scheme of heatbar. A vector. add.x.ann Whether to add the title, text, and ticks of X axis. x.lab X label. line.y.lab Y label of running score plot. rank.y.lab Y label of rank plot. statistic.position Position of statistics in the running score plot. statistic.face Font face of statistics. statistic.size Font size of statistics. Relative heights of subplots. rel.heights

A theme object from ggplot2.

Author(s)

theme.plot

Zaoqu Liu; E-mail: liuzaoqu@163.com

```
library(airway)
library(DESeq2)
library(tidyverse)
library(clusterProfiler)
library(org.Hs.eg.db)
data(airway)
se <- airway
se$dex <- relevel(se$dex, "untrt")</pre>
```

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```
res <- DESegDataSet(se, design = ~ cell + dex) %>%
  estimateSizeFactors() %>%
 DESeq() %>%
  results() %>%
  as.data.frame() %>%
 na.omit()
ann <- bitr(rownames(res), "ENSEMBL", "SYMBOL", org.Hs.eg.db)</pre>
res <- merge(ann, res, by.x = 1, by.y = 0) %>% dplyr::distinct(SYMBOL, .keep_all = TRUE)
# Obtain an order ranked geneList.
grlist <- res$log2FoldChange</pre>
names(grlist) <- res$SYMBOL</pre>
grlist <- sort(grlist, decreasing = TRUE)</pre>
# Integrative enrichment analysis of the ranked gene list
fit <- lzq_GSEA.integrated(genes = grlist)</pre>
lzq_gseaplot(
  fit$simplyGO,
  Pathway.ID = "G0:0030016",
  rank = F,
  statistic.position = c(0.71, 0.85),
  rel.heights = c(1, 0.4)
```

lzq_inferTF

Perform VIPER analysis

Description

This function performs Virtual Inference of Protein-activity by Enriched Regulon analysis

Usage

```
lzq_inferTF(
  exp,
  organism = "Human",
  use.cancer.regulons = F,
  confidence = c("A", "B", "C")
)
```

Arguments

exp Numeric matrix containing the expression data or gene expression signatures,

with samples in columns and genes in rows.

organism Specify species, currently support only Human and Mouse.

use.cancer.regulons

Use TF-target interactions for cancer application.

confidence The score comprises five categories, ranging from A (highest confidence) to E

(lowest confidence). The scoring criteria are described in PMID: 31340985.

Value

A matrix of inferred activity for each regulator gene in the network across all samples.

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Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

Examples

```
gene_expression <- as.matrix(read.csv(system.file("extdata", "human_input.csv", package = "progeny"),
   row.names = 1
))
s <- lzq_inferTF(gene_expression)</pre>
```

lzq_KEGGview

KEGG pathway visualization

Description

Simple visualization of KEGG pathway based on pathview package.

Usage

```
lzq_KEGGview(
  gene.data = NULL,
  gene.type = "SYMBOL",
  pathway.id,
  species = "hsa",
  figure.suffix = ""
)
```

Arguments

gene.data

either vector (single sample) or a matrix-like data (multiple sample). Vector should be numeric with gene IDs as names or it may also be character of gene IDs. Character vector is treated as discrete or count data. Matrix-like data structure has genes as rows and samples as columns. Row names should be gene IDs. Here gene ID is a generic concepts, including multiple types of gene, transcript and protein uniquely mappable to KEGG gene IDs. KEGG ortholog IDs are also treated as gene IDs as to handle metagenomic data. Check details for mappable ID types. Default gene.data=NULL.

gene.type

character, ID type used for the gene.data, case insensitive. Default gene.idtype="entrez", i.e. Entrez Gene, which are the primary KEGG gene ID for many common model organisms. For other species, gene.idtype should be set to "KEGG" as KEGG use other types of gene IDs. For the common model organisms (to check the list, do: data(bods); bods), you may also specify other types of valid IDs. To check the ID list, do: data(gene.idtype.list); gene.idtype.list.

pathway.id

character vector, the KEGG pathway ID(s), usually 5 digit, may also include the 3 letter KEGG species code.

species

character, either the kegg code, scientific name or the common name of the target species. This applies to both pathway and gene.data or cpd.data. When KEGG ortholog pathway is considered, species="ko". Default species="hsa", it is equivalent to use either "Homo sapiens" (scientific name) or "human" (common name).

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figure.suffix character, the suffix to be added after the pathway name as part of the output graph file. Sample names or column names of the gene.data or cpd.data are also added when there are multiple samples. Default out.suffix="pathview".

Author(s)

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Examples

```
library(airway)
library(DESeq2)
library(tidyverse)
library(clusterProfiler)
library(org.Hs.eg.db)
data(airway)
se <- airway
se$dex <- relevel(se$dex, "untrt")</pre>
res <- DESeqDataSet(se, design = ~ cell + dex) %>%
  estimateSizeFactors() %>%
  DESeq() %>%
 results() %>%
 as.data.frame() %>%
 na.omit()
ann <- bitr(rownames(res), "ENSEMBL", "SYMBOL", org.Hs.eg.db)</pre>
res <- merge(ann, res, by.x = 1, by.y = 0) %>% dplyr::distinct(SYMBOL, .keep_all = TRUE)
# Set enrich.type using an enrichment analysis method mentioned above.
kegg <- lzq_ORA(
  genes = res$SYMBOL[res$log2FoldChange > 0 & res$padj < 0.05],</pre>
  enrich.type = "KEGG"
)
res2 <- res[res$log2FoldChange > 0 \& res$padj < 0.05, c(2, 4)]
res2 <- data.frame(row.names = res2$SYMBOL, R = res2$log2FoldChange)</pre>
lzq_KEGGview(gene.data = res2, pathway.id = "hsa04218")
```

lzq_ORA

Over-representative analysis

Description

Perform over-representative analysis included GO, KEGG, WikiPathways, Reactome, MsigDB, Disease Ontoloty, Cancer Gene Network, DisGeNET, CellMarker, and CMAP.

```
lzq_ORA(
  genes,
  background.genes = NULL,
  gene.type = "SYMBOL",
  enrich.type,
  organism = "Human",
```

Izq_ORA

```
GO.ont = "BP",
GO.simplify = T,
KEGG.use.internal.data = F,
MsigDB.category = "H",
CMAP.min.Geneset.Size = 3,
pvalue.cutoff = 0.05,
qvalue.cutoff = 0.05,
padjust.method = "BH",
min.Geneset.Size = 10,
max.Geneset.Size = 1000
)
```

Arguments

genes A vector of gene id.

background.genes

Background genes. If missing, the all genes listed in the database (eg TERM2GENE

table) will be used as background.

gene. type Keytype of input gene.

enrich.type Select an enrichment method. One of GO, KEGG, MKEGG, WikiPathways,

Reactome, MsigDB, DO, CGN, DisGeNET, CellMarker, and CMAP. WikiPathways can be replaced by WP, Reactome can be replaced by RP, and CellMarker

can be replaced by CM.

organism Specify species, currently support only Human and Mouse.

GO.ont GO parameter. One of "BP", "MF", and "CC" subontologies, or "ALL" for all

three.

GO.simplify GO parameter. Whether to remove redundancy of enriched GO terms.

KEGG.use.internal.data

KEGG parameter. Logical, use KEGG.db or latest online KEGG data.

MsigDB.category

MsigDB parameter. MSigDB collection abbreviation, such as All, H, C1, C2,

C3, C4, C5, C6, C7.

CMAP.min.Geneset.Size

CMAP parameter. Minimal size of CMAP genes annotated for testing. Recom-

mended use 3.

pvalue.cutoff pvalue cutoff on enrichment tests to report as significant.

qvalue.cutoff qvalue cutoff on enrichment tests to report as significant.

 $\verb|padjust.method| one of "holm", "hochberg", "hommel", "bonferroni", "BH", "BY", "fdr", "none". \\$

min.Geneset.Size

Minimal size of genes annotated for testing. Not suitable for CMAP.

max.Geneset.Size

Maximal size of genes annotated for testing.

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

```
genes <- c("CANX", "HSPA1B", "KLRC2", "PSMC6", "RFXAP", "TAP1")
res <- lzq_ORA(genes, enrich.type = "GO")</pre>
```

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lzq_ORA.barplot1

Enrichment barplot for one ORA enrichment object

Description

Plot enrichment barplot for one ORA enrichment object.

Usage

```
lzq_ORA.barplot1(
  enrich.obj,
  x = "GeneRatio",
  show.term.num = 15,
  color.by = "p.adjust",
 colors = c("#003c30", "#01665e", "#35978f", "#80cdc1", "#c7eae5", "#f5f5f5", "#f6e8c3",
    "#dfc27d", "#bf812d", "#8c510a", "#543005"),
  color.title = color.by,
  bar.width = 0.6,
  add.bar.border = F,
  y.label.position = "right",
  title = NULL,
  legend.position = "right",
  theme.plot = theme_bw(base_rect_size = 1.5),
  use.Chinese = F,
  appid = "20231122001888718",
  key = "5GpDqe8F3pmXfnOkEKGQ"
```

Arguments

enrich.obj	An object from clusterProfiler.
Х	variable for x-axis, one of 'GeneRatio', 'pvalue', 'p.adjust', 'Count', EnrichmentFactor.
show.term.num	A number or a list of terms. If it is a number, the first n terms will be displayed. If it is a list of terms, the selected terms will be displayed.
color.by	variable that used to color enriched terms, one of 'GeneRatio', 'pvalue', 'p.adjust', 'Count', EnrichmentFactor.
colors	A color vector for the bars.
color.title	Title of color annotation legend.
bar.width	Width of bars.
add.bar.border	Logical. Whether to add the black border of bars.
y.label.position	
	Y label position. right or left.
title	Title of the plot.
legend.position	
	Position of legend. 'none', 'right', 'left' or two numeric variables.
theme.plot	ggtheme of plot.

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use.Chinese	Logical. Whether to use Chinese annotation in the barplot.
appid	User app id from baidu translation api. https://fanyi-api.baidu.com/manage/developer.
key	User Key from baidu translation api. https://fanyi-api.baidu.com/manage/developer.

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

Examples

```
genes <- c("CANX", "HSPA1B", "KLRC2", "PSMC6", "RFXAP", "TAP1")
res <- lzq_ORA(genes, enrich.type = "GO")
lzq_ORA.barplot1(res$simplyG0)</pre>
```

lzq_ORA.barplot2

Enrichment barplot for two ORA enrichment objects

Description

Plot enrichment barplot for two ORA enrichment objects.

Usage

```
lzq_ORA.barplot2(
  enrich.obj1,
  enrich.obj2,
  Selct.P = "FDR",
  cutoff.P = 0.05,
  obj.types = c("Up", "Down"),
  obj.type.colors = c("#ED6355", "#3E94B5"),
  obj1.top.pathway.num = 10,
  obj2.top.pathway.num = 10,
  bar.width = 0.6,
  add.bar.border = T,
  x.limit.fold = 1.05,
  label.size = 3.5,
  legend.position = "bottom",
  use.Chinese = F,
  appid = "20231122001888718",
  key = "5GpDqe8F3pmXfnOkEKGQ"
)
```

Arguments

enrich.obj1	An object from clusterProfiler.
enrich.obj2	An object from clusterProfiler.
Selct.P	Nominal P value (NP) or adjust P value (FDR) were selected to define significant terms.
cutoff.P	A cutoff value for Select_P.
obj.types	Two characters for defining the types of two objects.

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```
obj.type.colors
                  Two colors for the types of two objects.
obj1.top.pathway.num
                  The number of top pathways in object 1. Based on the significant test.
obj2.top.pathway.num
                  The number of top pathways in object 2. Based on the significant test.
bar.width
                  Width of bar in the plot.
add.bar.border Logical. Whether to add the black border of bars.
x.limit.fold
                  Specify the fold of x limitation. Because some terms is too long.
label.size
                  Fontsize of label.
legend.position
                  none, left, right, top, bottom; Or Two numeric variables indicated x and y posi-
                  tions, respectively.
                  Logical. Whether to use Chinese annotation in the barplot.
use.Chinese
                  User app id from baidu translation api. https://fanyi-api.baidu.com/manage/developer.
appid
key
                  User Key from baidu translation api. https://fanyi-api.baidu.com/manage/developer.
```

Author(s)

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```
library(airway)
library(DESeq2)
library(tidyverse)
library(clusterProfiler)
library(org.Hs.eg.db)
data(airway)
se <- airway
se$dex <- relevel(se$dex, "untrt")</pre>
res <- DESeqDataSet(se, design = ~ cell + dex) %>%
  estimateSizeFactors() %>%
 DESeq() %>%
  results() %>%
  as.data.frame() %>%
 na.omit()
ann <- bitr(rownames(res), "ENSEMBL", "SYMBOL", org.Hs.eg.db)</pre>
res <- merge(ann, res, by.x = 1, by.y = 0) %>% distinct(SYMBOL, .keep_all = T) # Very crude, just as an example
# Define an up-regulated gene list
up.genes <- res$SYMBOL[res$log2FoldChange > 2 & res$padj < 0.05]</pre>
# Define a down-regulated gene list
down.genes <- resSYMBOL[res$log2FoldChange < -2 & res$padj < 0.05]
# Integrative enrichment analysis of the up-regulated gene list
up.enrich <- lzq_ORA.integrated(genes = up.genes)</pre>
# Integrative enrichment analysis of the down-regulated gene list
down.enrich <- lzq_ORA.integrated(genes = down.genes)</pre>
lzq_ORA.barplot2(
  enrich.obj1 = up.enrich$simplyGO,
```

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```
enrich.obj2 = down.enrich$simplyGO,
  obj.types = c("Up", "Down")
)
```

lzq_ORA.dotplot1

Enrichment dotplot for one ORA enrichment object

Description

Plot enrichment dotplot for one ORA enrichment object.

Usage

```
lzq_ORA.dotplot1(
  enrich.obj,
  x = "GeneRatio",
  show.term.num = 15,
  color.by = "p.adjust",
 colors = c("#003c30", "#01665e", "#35978f", "#80cdc1", "#c7eae5", "#f5f5f5", "#f6e8c3",
    "#dfc27d", "#bf812d", "#8c510a", "#543005"),
  color.title = color.by,
  size.by = "Count",
  size.range = c(3, 8),
  size.title = size.by,
  y.label.position = "right",
  title = NULL,
  legend.position = "bottom",
  theme.plot = theme_bw(base_rect_size = 1.5),
  use.Chinese = F,
  appid = "20231122001888718",
  key = "5GpDqe8F3pmXfnOkEKGQ"
)
```

Arguments

enrich.obj	An object from clusterProfiler.
X	variable for x-axis, one of 'GeneRatio', 'pvalue', 'p.adjust', 'Count', EnrichmentFactor.
show.term.num	A number or a list of terms. If it is a number, the first n terms will be displayed. If it is a list of terms, the selected terms will be displayed.
color.by	variable that used to color enriched terms, one of 'GeneRatio', 'pvalue', 'p.adjust', 'Count', EnrichmentFactor.
colors	A color vector for the bars.
color.title	Title of color annotation legend.
size.by	variable that used to size enriched terms, one of 'GeneRatio', 'pvalue', 'p.adjust', 'Count', EnrichmentFactor.
size.range	Two numeric variables, the first is minimal value and the first is maximal value.
size.title	Title of size annotation legend.

```
y.label.position
Y label position. right or left.

title Title of the plot.
legend.position
Postion of legend. 'none', 'right', 'left' or two numeric variables.

theme.plot ggtheme of plot.

use.Chinese Logical. Whether to use Chinese annotation in the barplot.

appid User app id from baidu translation api. https://fanyi-api.baidu.com/manage/developer.

key User Key from baidu translation api. https://fanyi-api.baidu.com/manage/developer.
```

Author(s)

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Examples

```
genes <- c("CANX", "HSPA1B", "KLRC2", "PSMC6", "RFXAP", "TAP1")
res <- lzq_ORA(genes, enrich.type = "GO")
lzq_ORA.dotplot1(res$simplyGO)</pre>
```

lzq_ORA.integrated

Integrate over-representative analysis

Description

Perform integrated over-representative analysis included GO, KEGG, WikiPathways, Reactome, MsigDB, Disease Ontoloty, Cancer Gene Network, DisGeNET, CellMarker, and CMAP.

```
lzq_ORA.integrated(
  genes,
 background.genes = NULL,
  gene.type = "SYMBOL",
 organism = "Human",
 GO.ont = "BP",
 KEGG.use.internal.data = F,
 perform.WikiPathways = F,
 perform.Reactome = F,
 perform.MsigDB = F,
 MsigDB.category = "H",
 perform.disease.ontoloty = F,
 perform.Cancer.Gene.Network = F,
 perform.DisGeNET = F,
 perform.CellMarker = F,
 perform.CMAP = T,
 pvalue.cutoff = 0.05,
 qvalue.cutoff = 0.05,
 padjust.method = "BH",
 min.Geneset.Size = 10,
```

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```
max.Geneset.Size = 1000,
  CMAP.min.Geneset.Size = 3
)
```

Arguments

genes A vector of gene id.

background.genes

Background genes. If missing, the all genes listed in the database (eg TERM2GENE

table) will be used as background.

gene.type Keytype of input gene.

organism Specify species, currently support only Human and Mouse.

GO. ont One of "BP", "MF", and "CC" subontologies, or "ALL" for all three.

KEGG.use.internal.data

Logical, use KEGG.db or latest online KEGG data.

perform.WikiPathways

Whether to perform WikiPathways enrichment.

perform.Reactome

Whether to perform Reactome enrichment.

perform. MsigDB Whether to perform MsigDB enrichment.

MsigDB.category

MSigDB collection abbreviation, such as All, H, C1, C2, C3, C4, C5, C6, C7.

perform.disease.ontoloty

Whether to perform DO enrichment.

perform.Cancer.Gene.Network

Whether to perform CGN enrichment.

perform.DisGeNET

Whether to perform DisGeNET enrichment.

perform.CellMarker

Whether to perform CellMarker enrichment. Marker from cellmarker database.

perform. CMAP Whether to perform CMAP enrichment. Marker from CMAP database (in DSEATM

tool).

pvalue.cutoff pvalue cutoff on enrichment tests to report as significant.

qvalue.cutoff qvalue cutoff on enrichment tests to report as significant.

padjust.method one of "holm", "hochberg", "hommel", "bonferroni", "BH", "BY", "fdr", "none".

min.Geneset.Size

Minimal size of genes annotated for testing. Not suitable for CMAP.

max.Geneset.Size

Maximal size of genes annotated for testing.

CMAP.min.Geneset.Size

Minimal size of CMAP genes annotated for testing. Recommended use 3.

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

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Examples

```
library(airway)
library(DESeq2)
library(tidyverse)
library(clusterProfiler)
library(org.Hs.eg.db)
data(airway)
se <- airway
se$dex <- relevel(se$dex, "untrt")</pre>
res <- DESeqDataSet(se, design = ~ cell + dex) %>%
  estimateSizeFactors() %>%
  DESeq() %>%
  results() %>%
  as.data.frame() %>%
  na.omit()
ann <- bitr(rownames(res), "ENSEMBL", "SYMBOL", org.Hs.eg.db)</pre>
res <- merge(ann, res, by.x = 1, by.y = 0) %>% distinct(SYMBOL, .keep_all = T) # Very crude, just as an example
# Define an up-regulated gene list
up.genes <- res$SYMBOL[res$log2FoldChange > 2 & res$padj < 0.05]</pre>
# Define a down-regulated gene list
down.genes <- res$SYMBOL[res$log2FoldChange < -2 & res$padj < 0.05]</pre>
# Integrative enrichment analysis of the up-regulated gene list
# up.enrich <- lzq_ORA.integrated(genes = up.genes)</pre>
# Integrative enrichment analysis of the down-regulated gene list
# down.enrich <- lzq_ORA.integrated(genes = down.genes)</pre>
```

lzq_progeny

Perform PROGENy analysis

Description

Perform PROGENy analysis.

```
lzq_progeny(
  exp,
  scale = T,
  organism = "Human",
  top = 100,
  perm = 1,
  z_scores = F,
  get_nulldist = F,
  assay_name = "RNA",
  return_assay = F
)
```

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Arguments

A gene expression object with HGNC/MGI symbols in rows and samples in columns. In order to run PROGENy in single-cell RNAseq data, it also accepts Seurat and SingleCellExperiment object, taking the normalized counts for the computation.
A logical value indicating whether to scale the scores of each pathway to have a mean of zero and a standard deviation of one. It does not apply if we use permutations.
The model organism - "Human" or "Mouse".
The top n genes for generating the model matrix according to significance (p-value).
An interger detailing the number of permutations. No permutations by default (1). When Permutations larger than 1, we compute progeny pathway scores and assesses their significance using a gene sampling-based permutation strategy, for a series of experimental samples/contrasts.
Only applies if the number of permutations is greater than 1. A logical value. TRUE: the z-scores will be returned for the pathway activity estimations. FALSE: the function returns a normalized z-score value between -1 and 1.
Only applies if the number of permutations is greater than 1. A logical value. TRUE: the null distributions generated to assess the signifiance of the pathways scores is also returned.
Only applies if the input is a Seurat object. It selects the name of the assay on which Progeny will be run. Default to: RNA, i.e. normalized expression values.
Only applies if the input is a Seurat object. A logical value indicating whether to return progeny results as a new assay called Progeny in the Seurat object used as input. Default to FALSE.

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

Examples

```
gene_expression <- as.matrix(read.csv(system.file("extdata", "human_input.csv", package = "progeny"), row.nam
s <- lzq_progeny(gene_expression)</pre>
```

lzq_progeny.dea

Perform differential analysis for the PROGENy results.

Description

Perform differential analysis for the PROGENy results.

```
lzq_progeny.dea(
  progeny.res,
  groups,
  control.group,
  theme.plot = theme_classic(base_line_size = 0.8)
)
```

Arguments

```
progeny.res A PROGENy results with row samples and column pathways.
groups Group information that matches the counts column sample names.
control.group Specify the control group.
theme.plot ggtheme of plot.
```

Examples

```
gene_expression <- as.matrix(read.csv(system.file("extdata", "human_input.csv", package = "progeny"),
   row.names = 1
))
s <- lzq_progeny(gene_expression)
l <- lzq_progeny.dea(s, groups = rep(c("A", "B"), each = 4), "A")</pre>
```

lzq_progeny.gene.details

Generate the differential expression and PROGENy weight of genes.

Description

Generate the differential expression and PROGENy weight of genes.

Usage

```
lzq_progeny.gene.details(
  dea.table,
  pathway,
  organism = "Human",
  top = 100,
  y.lab = bquote(~Log[2] ~ "(Fold change)"),
  colors = c("#3E94B5", "grey70", "#ED6355"),
  point.size = 2,
  label.size = 4,
  theme.plot = theme_classic(base_line_size = 0.8)
)
```

Arguments

dea.table	A dataframe with two columns, the first is gene id and the second is logFC/-logP/Stat.
pathway	Specific a pathway, such as Androgen, EGFR, Estrogen, Hypoxia, JAK-STAT, MAPK, NFkB, p53, PI3K, TGFb, TNFa, Trail, VEGF, and WNT.
organism	The model organism - "Human" or "Mouse".
top	The top n genes for generating the model matrix according to significance (p-value).
y.lab	Y label for scatter.
colors	Colors for different types of points.
point.size	Size of point.
label.size	Size of gene label.
theme.plot	ggtheme of plot.

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Examples

```
gene_expression <- as.matrix(read.csv(system.file("extdata", "human_input.csv", package = "progeny"), row.nam
s <- lzq_progeny(gene_expression)</pre>
```

1zq_ssGSES

Generate single-sample gene-set enrichment score

Description

Estimate gene-set enrichment score across all samples.

Usage

```
lzq_ssGSES(exp, gene.list, method = "ssgsea")
```

Arguments

exp

Numeric matrix containing the expression data or gene expression signatures, with samples in columns and genes in rows.

gene.list

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

method

Method to employ in the estimation of gene-set enrichment scores per sample. By default this is set to gsva (Hänzelmann et al, 2013) and other options are ssgsea (Barbie et al, 2009), zscore (Lee et al, 2008) or plage (Tomfohr et al, 2005). The latter two standardize first expression profiles into z-scores over the samples and, in the case of zscore, it combines them together as their sum divided by the square-root of the size of the gene set, while in the case of plage they are used to calculate the singular value decomposition (SVD) over the genes in the gene set and use the coefficients of the first right-singular vector as pathway activity profile.

Value

A gene-set by sample matrix (of matrix or dgCMatrix type, depending on the input) of gene-set enrichment scores.

Author(s)

```
Zaoqu Liu; E-mail: liuzaoqu@163.com
```

```
gene_expression <- as.matrix(read.csv(system.file("extdata", "human_input.csv", package = "progeny"),
    row.names = 1
))
gl <- list(A = rownames(gene_expression)[1:10], B = rownames(gene_expression)[11:20])
s <- lzq_ssGSES(gene_expression, gl, method = "gsva")</pre>
```

30 lzq_updateSymbol

lzq_translate

Baidu translation

Description

Perform Baidu translation.

Usage

```
lzq_translate(
    sentence,
    from = "en",
    to = "zh",
    appid = "20231122001888718",
    key = "5GpDqe8F3pmXfnOkEKGQ"
)
```

Arguments

sentence A sentence or word need to be translated.

from Input language type. to Output language type.

appid User app id from baidu translation api. https://fanyi-api.baidu.com/manage/developer. key User Key from baidu translation api. https://fanyi-api.baidu.com/manage/developer.

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

Examples

lzq_translate("BioEnricher is a simple and useful package!")

 $\begin{tabular}{ll} ${\it Liq}$ update Symbol & \it Title Identify outdated or Excel-mogrified gene symbols for a gene vector \\ \hline \end{tabular}$

Description

Title Identify outdated or Excel-mogrified gene symbols for a gene vector

Usage

```
lzq_updateSymbol(genes, unmapGene_keep = F)
```

Arguments

```
genes A gene vector.
unmapGene_keep whether to keep unmapped genes.
```

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

Examples

```
lzq_updateSymbol("PD-L1")
```

Description

Title Identify outdated or Excel-mogrified gene symbols for a dataframe

Usage

```
lzq_updateSymbolforDL(data, unmapGene_keep = F)
```

Arguments

data A expression dataframe with genename rows and sample columns. unmapGene_keep whether to keep unmapped genes.

Author(s)

Zaoqu Liu; E-mail: liuzaoqu@163.com

searchGEO Searching GEO metadata

Description

Searching GEO metadata based on the input term. This function is from genekitr package.

Usage

```
searchGEO(searchterm, minnum = 0, maxnum = 1000)
```

Arguments

searchterm input searching terms as GEO database keywords, multiple terms are seperated

by blanks

minnum The minimum return records, default is 0 maxnum The maximum return records, default is 1000

Value

A data.frame.

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Examples

```
meta <- searchGEO("ezh2 knockout", maxnum = 5)</pre>
```

searchPubmed

Get 'PubMed' paper records by searching abstract

Description

Get 'PubMed' paper records by searching abstract. This function is from europepmc package.

Usage

```
searchPubmed(term, add_term = NULL, num = 100)
```

Arguments

term query terms e.g. gene id, GO/KEGG pathway add_term other searching terms Default is NULL num limit the number of records . Default is 100.

Value

A list of dataframe for PubMed records

Examples

```
term <- c("Tp53", "Brca1", "Tet2")
add_term <- c("stem cell", "mouse")
1 <- searchPubmed(term, add_term, num = 30)</pre>
```

searchPubmedTrend

Get the yearly number of hits for a query and the total yearly number of hits for a given period

Description

Get the yearly number of hits for a query and the total yearly number of hits for a given period. This function is from europepmc package.

Usage

```
searchPubmedTrend(term, add_term = NULL, period)
```

Arguments

term query terms e.g. gene id, GO/KEGG pathway add_term other searching terms Default is NULL

period a vector of years (numeric) over which to perform the search.

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Value

a data.frame (dplyr tbl_df) with year, total number of hits (all_hits) and number of hits for the query (query_hits).

```
term <- c("Tp53", "Brca1", "Tet2")
add_term <- c("stem cell", "mouse")
1 <- searchPubmedTrend(term, add_term, period = 2020:2023)</pre>
```

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