# Package 'decoupleR'

June 29, 2025

Type Package

**Title** decoupleR: Ensemble of computational methods to infer biological activities from omics data

**Version** 2.14.0

Description Many methods allow us to extract biological activities from omics data using information from prior knowledge resources, reducing the dimensionality for increased statistical power and better interpretability. Here, we present decoupleR, a Bioconductor package containing different statistical methods to extract these signatures within a unified framework. decoupleR allows the user to flexibly test any method with any resource. It incorporates methods that take into account the sign and weight of network interactions. decoupleR can be used with any omic, as long as its features can be linked to a biological process based on prior knowledge. For example, in transcriptomics gene sets regulated by a transcription factor, or in phospho-proteomics phosphosites that are targeted by a kinase.

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URL https://saezlab.github.io/decoupleR/

BugReports https://github.com/saezlab/decoupleR/issues

**Depends** R (>= 4.0)

**Imports** BiocParallel, broom, dplyr, magrittr, Matrix, parallelly, purrr, rlang, stats, stringr, tibble, tidyr, tidyselect, withr

**Suggests** glmnet (>= 4.1-7), GSVA, viper, fgsea (>= 1.15.4), AUCell, SummarizedExperiment, rpart, ranger, BiocStyle, covr, knitr, pkgdown, RefManageR, rmarkdown, roxygen2, sessioninfo, pheatmap, testthat, OmnipathR, Seurat, ggplot2, ggrepel, patchwork, magick

VignetteBuilder knitr

**biocViews** DifferentialExpression, FunctionalGenomics, GeneExpression, GeneRegulation, Network, Software, StatisticalMethod, Transcription,

Config/testthat/edition 3

**Encoding UTF-8** 

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```
LazyData false
Roxygen list(markdown = TRUE)
RoxygenNote 7.3.0
git_url https://git.bioconductor.org/packages/decoupleR
git_branch RELEASE_3_21
git_last_commit 5ed37a4
git_last_commit_date 2025-04-15
Repository Bioconductor 3.21
Date/Publication 2025-06-29
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## **Description**

Many methods allow us to extract biological activities from omics data using information from prior knowledge resources, reducing the dimensionality for increased statistical power and better interpretability. Here, we present decoupleR, a Bioconductor package containing different statistical methods to extract these signatures within a unified framework. decoupleR allows the user to flexibly test any method with any resource. It incorporates methods that take into account the sign and weight of network interactions. decoupleR can be used with any omic, as long as its features can be linked to a biological process based on prior knowledge. For example, in transcriptomics gene sets regulated by a transcription factor, or in phospho-proteomics phosphosites that are targeted by a kinase.

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#### See Also

Useful links:

- https://saezlab.github.io/decoupleR/
- Report bugs at https://github.com/saezlab/decoupleR/issues

 $. \verb|decoupler_mat_format| \\ \textit{DecoupleR mat format}$ 

## **Description**

DecoupleR mat format

## **Arguments**

mat

Matrix to evaluate (e.g. expression matrix). Target nodes in rows and conditions in columns. rownames(mat) must have at least one intersection with the elements in network .target column.

#### See Also

 $Other\ decouple R\ formats:\ .decoupler\_network\_format$ 

.decoupler\_network\_format

DecoupleR network format

#### **Description**

A network passed to any run\_ method in the package must contain at least two attributes: .source and .target. In addition, the methods must map their corresponding metadata associated with their edges.

## **Arguments**

network Tibble or dataframe with edges and it's associated metadata.

. source Column with source nodes.. target Column with target nodes.

.mor Column with edge mode of regulation (i.e. mor)..likelihood Deprecated argument. Now it will always be set to 1.

#### **Details**

- All the attributes to be mapped are prefixed by .
- The idea of using this type of mapping is to provide flexibility to different types of networks, be they regulatory, metabolic, or of any other type. This way, you should only consider having your network or networks in a long format and these can easily be manipulated by functions within the tidyverse ecosystem.

#### See Also

Other decoupleR formats: .decoupler\_mat\_format

 $.fit\_preprocessing$ 

Pre-processing for methods that fit networks.

## **Description**

• If center is true, then the expression values are centered by the mean of expression across the samples.

```
.fit_preprocessing(network, mat, center, na.rm, sparse)
```

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## **Arguments**

network	Tibble or dataframe with edges and it's associated metadata.
mat	Matrix to evaluate (e.g. expression matrix). Target nodes in rows and conditions in columns. rownames(mat) must have at least one intersection with the elements in network .target column.
center	Logical value indicating if mat must be centered by base::rowMeans().
na.rm	Should missing values (including NaN) be omitted from the calculations of base::rowMeans()?
sparse	Deprecated parameter.

## Value

A named list of matrices to evaluate in methods that fit models, like .mlm\_analysis().

- mat: Features as rows and samples as columns.
- mor\_mat: Features as rows and columns as source.

## **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
net <- rename_net(net, source, target, mor)
.fit_preprocessing(net, mat, center = FALSE, na.rm = FALSE, sparse = FALSE)</pre>
```

check\_corr

Check correlation (colinearity)

## Description

Checks the correlation across the regulators in a network.

```
check_corr(
  network,
  .source = "source",
  .target = "target",
  .mor = "mor",
  .likelihood = NULL
)
```

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#### Arguments

network Tibble or dataframe with edges and it's associated metadata.

. source Column with source nodes.. target Column with target nodes.

.mor Column with edge mode of regulation (i.e. mor)..likelihood Deprecated argument. Now it will always be set to 1.

#### Value

Correlation pairs tibble.

#### **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
net <- readRDS(file.path(inputs_dir, "net.rds"))
check_corr(net, .source='source')</pre>
```

convert\_f\_defaults

Rename columns and add defaults values if column not present

#### **Description**

convert\_f\_defaults() combine the dplyr::rename() way of working and with the tibble::add\_column() to add columns with default values in case they don't exist after renaming data.

#### Usage

```
convert_f_defaults(.data, ..., .def_col_val = c(), .use_dots = TRUE)
```

## **Arguments**

.data	A data frame, data frame extension (e.g. a tibble), or a lazy data frame (e.g. from dbplyr or dtplyr). See <i>Methods</i> , below, for more details.
•••	For rename(): <tidy-select> Use new_name = old_name to rename selected variables.</tidy-select>
	For rename_with(): additional arguments passed onto .fn.
.def_col_val	Named vector with columns with default values if none exist after rename.

. use\_dots Should a dot prefix be added to renamed variables? This will allow swapping of columns.

#### **Details**

The objective of using .use\_dots is to be able to swap columns which, by default, is not allowed by the dplyr::rename() function. The same behavior can be replicated by simply using the dplyr::select(), however, the select evaluation allows much more flexibility so that unexpected results could be obtained. Despite this, a future implementation will consider this form of execution to allow renaming the same column to multiple ones (i.e. extend dataframe extension).

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## Value

An object of the same type as .data. The output has the following properties:

- Rows are not affected.
- · Column names are changed.
- Column order is the same as that of the function call.

## **Examples**

```
df <- tibble::tibble(x = 1, y = 2, z = 3)
# Rename columns
df <- tibble::tibble(x = 1, y = 2)
convert_f_defaults(
    .data = df,
    new_x = x,
    new_y = y,
    new_z = NULL,
    .def_col_val = c(new_z = 3)
)</pre>
```

decouple

Evaluate multiple statistics with same input data

## Description

Calculate the source activity per sample out of a gene expression matrix by coupling a regulatory network with a variety of statistics.

```
decouple(
  mat,
  network,
  .source = source,
  .target = target,
  statistics = NULL,
  args = list(NULL),
  consensus_score = TRUE,
  consensus_stats = NULL,
  include_time = FALSE,
  show_toy_call = FALSE,
  minsize = 5
)
```

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#### **Arguments**

mat Matrix to evaluate (e.g. expression matrix). Target nodes in rows and condi-

tions in columns. rownames(mat) must have at least one intersection with the

elements in network .target column.

network Tibble or dataframe with edges and it's associated metadata.

. source Column with source nodes.. target Column with target nodes.

statistics Statistical methods to be run sequentially. If none are provided, only top per-

former methods are run (mlm, ulm and wsum).

args A list of argument-lists the same length as statistics (or length 1). The default

argument, list(NULL), will be recycled to the same length as statistics, and will call each function with no arguments (apart from mat, network, .source

and, .target).

consensus\_score

Boolean whether to run a consensus score between methods.

consensus\_stats

List of estimate names to use for the calculation of the consensus score. This is used to filter out extra estimations from some methods, for example wsum returns wsum, corr\_wsum and norm\_wsum. If none are provided, and also no statistics where provided, only top performer methods are used (mlm, ulm and norm\_wsum). Else, it will use all available estimates after running all methods

in the statistics argument.

include\_time Should the time per statistic evaluated be informed?

show\_toy\_call The call of each statistic must be informed?

minsize Integer indicating the minimum number of targets per source.

#### Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. run\_id: Indicates the order in which the methods have been executed.
- 2. statistic: Indicates which method is associated with which score.
- 3. source: Source nodes of network.
- 4. condition: Condition representing each column of mat.
- 5. score: Regulatory activity (enrichment score).
- 6. statistic\_time: If requested, internal execution time indicator.
- 7. p\_value: p-value (if available) of the obtained score.

#### See Also

Other decoupleR statistics: run\_aucell(), run\_fgsea(), run\_gsva(), run\_mdt(), run\_mlm(), run\_ora(), run\_udt(), run\_ulm(), run\_viper(), run\_wmean(), run\_wsum()

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#### **Examples**

```
if (FALSE) {
    inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")</pre>
   mat <- readRDS(file.path(inputs_dir, "mat.rds"))</pre>
   net <- readRDS(file.path(inputs_dir, "net.rds"))</pre>
    decouple(
        mat = mat,
        network = net,
        .source = "source",
        .target = "target",
        statistics = c("gsva", "wmean", "wsum", "ulm", "aucell"),
        args = list(
            gsva = list(verbose = FALSE),
            wmean = list(.mor = "mor", .likelihood = "likelihood"),
            wsum = list(.mor = "mor"),
            ulm = list(.mor = "mor")
        ),
        minsize = 0
}
```

extract\_sets

Extract sets

## **Description**

Extracts feature sets from a renamed network (see rename\_net).

## Usage

```
extract_sets(network)
```

#### **Arguments**

network

Tibble or dataframe with edges and it's associated metadata.

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
net <- rename_net(net, source, target, mor)
extract_sets(net)</pre>
```

filt\_minsize 11

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Filter sources with minsize targets

#### **Description**

Filter sources of a net with less than minsize targets

## Usage

```
filt_minsize(mat_f_names, network, minsize = 5)
```

### **Arguments**

mat\_f\_names Feature names of mat.

network Tibble or dataframe with edges and it's associated metadata.

minsize Integer indicating the minimum number of targets per source.

#### Value

Filtered network.

#### **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
net <- rename_net(net, source, target, mor)
filt_minsize(rownames(mat), net, minsize = 4)</pre>
```

get\_collectri

CollecTRI gene regulatory network. Wrapper to access CollecTRI gene regulatory network. CollecTRI is a comprehensive resource containing a curated collection of transcription factors (TFs) and their target genes. It is an expansion of DoRothEA. Each interaction is weighted by its mode of regulation (either positive or negative).

## **Description**

CollecTRI gene regulatory network. Wrapper to access CollecTRI gene regulatory network. CollecTRI is a comprehensive resource containing a curated collection of transcription factors (TFs) and their target genes. It is an expansion of DoRothEA. Each interaction is weighted by its mode of regulation (either positive or negative).

```
get_collectri(organism = "human", split_complexes = FALSE, ...)
```

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## **Arguments**

```
organism Which organism to use. Only human, mouse and rat are available.

split_complexes

Whether to split complexes into subunits. By default complexes are kept as they are.

... Ignored.
```

## Examples

```
collectri <- get_collectri(organism='human', split_complexes=FALSE)</pre>
```

get\_dorothea

DoRothEA gene regulatory network.

### **Description**

Wrapper to access DoRothEA gene regulatory network. DoRothEA is a comprehensive resource containing a curated collection of transcription factors (TFs) and their target genes. Each interaction is weighted by its mode of regulation (either positive or negative) and by its confidence level

## Usage

```
get_dorothea(
  organism = "human",
  levels = c("A", "B", "C"),
  weight_dict = list(A = 1, B = 2, C = 3, D = 4)
)
```

#### **Arguments**

organism Which organism to use. Only human, mouse and rat are available.

levels List of confidence levels to return. Goes from A to D, A being the most confident

and D being the less.

weight\_dict Dictionary of values to divide the mode of regulation (-1 or 1), one for each

confidence level. Bigger values will generate weights close to zero.

```
dorothea <- get_dorothea(organism='human', levels=c('A', 'B'))</pre>
```

get\_ksn\_omnipath 13

get_ksn_omnipath	OmniPath kinase-substrate network

#### **Description**

Retrieve a ready to use, curated kinase-substrate Network from the OmniPath database.

## Usage

```
get_ksn_omnipath(...)
```

## **Arguments**

.. Passed to OmnipathR::import\_omnipath\_enzsub.

#### **Details**

Import enzyme-PTM network from OmniPath, then filter out anything that is not phospho or dephosphorilation. Then format the columns for use with decoupleR functions.

get\_profile\_of Complete a data frame with missing combinations of data

## Description

Turns implicit missing values into explicit missing values. This is a wrapper around expand(), dplyr::full\_join() and replace\_na() that's useful for completing missing combinations of data.

#### Usage

```
get_profile_of(data, sources, values_fill = NA)
```

#### **Arguments**

data A data frame.

sources A named vector or list with the values to expand and get profile.

values\_fill Optionally, a (scalar) value that specifies what each value should be filled in

with when missing.

This can be a named list if you want to apply different fill values to different

value columns.

### Value

A data frame with the expanded grid of the values passed in sources and filled as specified in the fill argument.

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#### See Also

complete expand

#### **Examples**

```
## Not run:
library(dplyr, warn.conflicts = FALSE)
df <- tibble(</pre>
   group = c(1:2, 1),
    item_id = c(1:2, 2),
   item_name = c("a", "b", "b"),
   value1 = 1:3,
   value2 = 4:6
)
to_get_profile \leftarrow list(group = c(1, 2, 3), item_id = c(1, 2))
# This will add the combinations of group 3 with the id of the items
df %>% get_profile_of(sources = to_get_profile)
# You can also choose to fill in missing values
# This only fill with "Unknown" the NA values of the column item_name
df %>% get_profile_of(
    sources = to_get_profile,
    values_fill = list(item_name = "Unknown")
)
# Replace all NAs with "Unkwnon"
df %>% get_profile_of(sources = to_get_profile, values_fill = "Unknown")
## End(Not run)
```

get\_progeny

Pathway RespOnsive GENes for activity inference (PROGENy).

## Description

Wrapper to access PROGENy model gene weights. Each pathway is defined with a collection of target genes, each interaction has an associated p-value and weight. The top significant interactions per pathway are returned.

## Usage

```
get_progeny(organism = "human", top = 500)
```

#### **Arguments**

organism Which organism to use. Only human and mouse are available.

top Number of genes per pathway to return.

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#### **Examples**

```
progeny <- get_progeny(organism='human', top=500)</pre>
```

get\_resource

Wrapper to access resources inside Omnipath. This wrapper allows to easily query different prior knowledge resources. To check available resources run decoupleR::show\_resources(). For more information visit the official website for Rhrefhttps://omnipathdb.org/Omnipath.

## **Description**

Wrapper to access resources inside Omnipath. This wrapper allows to easily query different prior knowledge resources. To check available resources run decoupleR::show\_resources(). For more information visit the official website for Omnipath.

#### Usage

```
get_resource(name, organism = "human", ...)
```

### **Arguments**

name Name of the resource to query.

organism name or NCBI Taxonomy ID.

... Passed to OmnipathR::import\_omnipath\_annotations.

## Examples

```
df <- decoupleR::get_resource('SIGNOR')</pre>
```

get\_toy\_data

Generate a toy mat and network.

#### **Description**

Generate a toy mat and network.

## Usage

```
get_toy_data(n_samples = 24, seed = 42)
```

#### **Arguments**

n\_samples Number of samples to simulate.

seed A single value, interpreted as an integer, or NULL for random number genera-

tion.

intersect\_regulons

## Value

List containing mat and network.

## **Examples**

```
data <- get_toy_data()
mat <- data$mat
network <- data$network</pre>
```

 $intersect\_regulons$ 

Intersect network target features with input matrix.

## Description

Keep only edges which its target features belong to the input matrix.

## Usage

```
intersect_regulons(mat, network, .source, .target, minsize)
```

## Arguments

mat	Matrix to evaluate (e.g. expression matrix). Target nodes in rows and conditions in columns. rownames(mat) must have at least one intersection with the elements in network .target column.
network	Tibble or dataframe with edges and it's associated metadata.
.source	Column with source nodes.
.target	Column with target nodes.
minsize	Minimum number of targets per source allowed.

### Value

Filtered tibble.

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
intersect_regulons(mat, net, source, target, minsize=4)</pre>
```

pivot\_wider\_profile 17

pivot\_wider\_profile

Pivot a data frame to wider and convert it to matrix

#### **Description**

Generates a kind of table where the rows come from id\_cols, the columns from names\_from and the values from values\_from.

### Usage

```
pivot_wider_profile(
  data,
  id_cols,
  names_from,
  values_from,
  values_fill = NA,
  to_matrix = FALSE,
  to_sparse = FALSE,
  ...
)
```

#### Arguments

data

A data frame to pivot.

id\_cols

<tidy-select> A set of columns that uniquely identify each observation. Typically used when you have redundant variables, i.e. variables whose values are perfectly correlated with existing variables.

Defaults to all columns in data except for the columns specified through names\_from and values\_from. If a tidyselect expression is supplied, it will be evaluated on data after removing the columns specified through names\_from and values\_from.

names\_from, values\_from

<tidy-select> A pair of arguments describing which column (or columns) to get the name of the output column (names\_from), and which column (or columns) to get the cell values from (values\_from).

If values\_from contains multiple values, the value will be added to the front of the output column.

values\_fill

Optionally, a (scalar) value that specifies what each value should be filled in

with when missing.

This can be a named list if you want to apply different fill values to different value columns.

to\_matrix

Logical value indicating if the result should be a matrix. Parameter is ignored in case sparse is TRUE.

case spai se is in

to\_sparse

Logical value indicating whether the resulting matrix should be sparse or not.

. . .

Additional arguments passed on to methods.

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#### **Details**

In the current state of the function, to ensure its operation, the id\_cols parameter is a single selector.

#### Value

"widened" data; it is increasing the number of columns and decreasing the number of rows.

## **Examples**

```
## Not run:
df <- tibble::tibble(</pre>
    tf = c("tf_1", "tf_1", "tf_2", "tf_2"),
   gene = c("gene_1", "gene_2", "gene_1", "gene_2"),
   mor = c(1, -1, 1, -1)
)
# Return a tibble
pivot_wider_profile(
   data = df,
   id_cols = tf,
   names_from = gene,
   values_from = mor
)
# Return a matrix
pivot_wider_profile(
   data = df,
   id_cols = tf,
   names_from = gene,
   values_from = mor,
    to_matrix = TRUE
# Return a sparse Matrix of class "dgCMatrix"
pivot_wider_profile(
   data = df,
    id_cols = tf,
   names_from = gene,
    values_from = mor,
    to_sparse = TRUE
)
## End(Not run)
```

randomize\_matrix

Randomize matrix

## **Description**

Utility function used in functions that require permutations of the expression matrix

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#### Usage

```
randomize_matrix(mat, randomize_type = c("rows", "cols_independently"))
```

## **Arguments**

```
mat Matrix to randomize. randomize_type How to randomize.
```

#### Value

Randomized matrix

## **Examples**

```
## Not run:
mat <- matrix(seq_len(9), ncol = 3)
mat

set.seed(42)
randomize_matrix(mat, randomize_type = "rows")

set.seed(42)
randomize_matrix(mat, randomize_type = "cols_independently")
## End(Not run)</pre>
```

rename\_net

Rename network

## Description

Renames a given network to these column names: .source, .target, .mor, If .mor is not provided, then the function sets them to default values.

```
rename_net(
  network,
  .source,
  .target,
  .mor = NULL,
  .likelihood = NULL,
  def_mor = 1
)
```

20 run\_aucell

## Arguments

network Tibble or dataframe with edges and it's associated metadata.

. source Column with source nodes.. target Column with target nodes.

.mor Column with edge mode of regulation (i.e. mor)..likelihood Deprecated argument. Now it will always be set to 1.

def\_mor Default value for .mor when not provided.

## Examples

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
rename_net(net, source, target, mor)</pre>
```

run\_aucell

**AUCell** 

## **Description**

Calculates regulatory activities using AUCell.

## Usage

```
run_aucell(
  mat,
  network,
  .source = source,
  .target = target,
  aucMaxRank = ceiling(0.05 * nrow(rankings)),
  nproc = availableCores(),
  seed = 42,
  minsize = 5
)
```

#### **Arguments**

mat Matrix to evaluate (e.g. expression matrix). Target nodes in rows and condi-

tions in columns. rownames(mat) must have at least one intersection with the

elements in network .target column.

network Tibble or dataframe with edges and it's associated metadata.

. source Column with source nodes.. target Column with target nodes.aucMaxRank Threshold to calculate the AUC.

run\_consensus 21

nproc Number of cores to use for computation.

seed A single value, interpreted as an integer, or NULL for random number genera-

tion.

minsize Integer indicating the minimum number of targets per source.

#### **Details**

AUCell (Aibar et al., 2017) uses the Area Under the Curve (AUC) to calculate whether a set of targets is enriched within the molecular readouts of each sample. To do so, AUCell first ranks the molecular features of each sample from highest to lowest value, resolving ties randomly. Then, an AUC can be calculated using by default the top 5% molecular features in the ranking. Therefore, this metric, aucell, represents the proportion of abundant molecular features in the target set, and their relative abundance value compared to the other features within the sample.

Aibar S. et al. (2017) Scenic: single-cell regulatory network inference and clustering. Nat. Methods, 14, 1083–1086.

#### See Also

```
Other decoupleR statistics: decouple(), run_fgsea(), run_gsva(), run_mdt(), run_mlm(), run_ora(), run_udt(), run_ulm(), run_viper(), run_wmean(), run_wsum()
```

## **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_aucell(mat, net, minsize=0, nproc=1, aucMaxRank=3)</pre>
```

run\_consensus

Consensus score between methods

## **Description**

Function to generate a consensus score between methods from the result of the decouple function.

## Usage

```
run_consensus(df, include_time = FALSE, seed = NULL)
```

#### **Arguments**

df decouple data frame result

include\_time Should the time per statistic evaluated be informed?

seed Deprecated parameter.

run\_fgsea

#### Value

Updated tibble with the computed consensus score between methods

## **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")</pre>
mat <- readRDS(file.path(inputs_dir, "mat.rds"))</pre>
net <- readRDS(file.path(inputs_dir, "net.rds"))</pre>
results <- decouple(</pre>
   mat = mat,
   network = net,
   .source = "source",
   .target = "target",
   statistics = c("wmean", "ulm"),
   args = list(
            wmean = list(.mor = "mor", .likelihood = "likelihood"),
            ulm = list(.mor = "mor", .likelihood = "likelihood")
        ),
   consensus_score = FALSE,
   minsize = 0
run_consensus(results)
```

run\_fgsea

Fast Gene Set Enrichment Analysis (FGSEA)

## **Description**

Calculates regulatory activities using FGSEA.

```
run_fgsea(
  mat,
  network,
  .source = source,
  .target = target,
  times = 100,
  nproc = availableCores(),
  seed = 42,
  minsize = 5,
  ...
)
```

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#### **Arguments**

mat Matrix to evaluate (e.g. expression matrix). Target nodes in rows and condi-

tions in columns. rownames (mat) must have at least one intersection with the

elements in network .target column.

network Tibble or dataframe with edges and it's associated metadata.

. source Column with source nodes.. target Column with target nodes.

times How many permutations to do?

nproc Number of cores to use for computation.

seed A single value, interpreted as an integer, or NULL.

minsize Integer indicating the minimum number of targets per source.

... Arguments passed on to fgsea::fgseaMultilevel

sampleSize The size of a random set of genes which in turn has size = pathwaySize

minSize Minimal size of a gene set to test. All pathways below the threshold are excluded.

maxSize Maximal size of a gene set to test. All pathways above the threshold are excluded.

eps This parameter sets the boundary for calculating the p value.

scoreType This parameter defines the GSEA score type. Possible options are ("std", "pos", "neg"). By default ("std") the enrichment score is computed as in the original GSEA. The "pos" and "neg" score types are intended to be used for one-tailed tests (i.e. when one is interested only in positive ("pos") or negateive ("neg") enrichment).

gseaParam GSEA parameter value, all gene-level statis are raised to the power of 'gseaParam' before calculation of GSEA enrichment scores.

BPPARAM Parallelization parameter used in bplapply. Can be used to specify cluster to run. If not initialized explicitly or by setting 'nproc' default value 'bpparam()' is used.

absEps deprecated, use 'eps' parameter instead

#### **Details**

GSEA (Aravind et al., 2005) starts by transforming the input molecular readouts in mat to ranks for each sample. Then, an enrichment score fgsea is calculated by walking down the list of features, increasing a running-sum statistic when a feature in the target feature set is encountered and decreasing it when it is not. The final score is the maximum deviation from zero encountered in the random walk. Finally, a normalized score norm\_fgsea, can be obtained by computing the z-score of the estimate compared to a null distribution obtained from N random permutations. The used implementation is taken from the package fgsea (Korotkevich et al., 2021).

Aravind S. et al. (2005) Gene set enrichment analysis: A knowledge-based approach for interpreting genome-wide expression profiles. PNAS. 102, 43.

Korotkevich G. et al. (2021) Fast gene set enrichment analysis. bioRxiv. DOI: https://doi.org/10.1101/060012.

zun\_gsva

#### Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. statistic: Indicates which method is associated with which score.
- 2. source: Source nodes of network.
- 3. condition: Condition representing each column of mat.
- 4. score: Regulatory activity (enrichment score).

#### See Also

```
Other decoupleR statistics: decouple(), run_aucell(), run_gsva(), run_mdt(), run_mlm(), run_ora(), run_udt(), run_ulm(), run_viper(), run_wmean(), run_wsum()
```

## **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_fgsea(mat, net, minsize=0, nproc=1)</pre>
```

run\_gsva

Gene Set Variation Analysis (GSVA)

## **Description**

Calculates regulatory activities using GSVA.

```
run_gsva(
  mat,
  network,
  .source = source,
  .target = target,
  verbose = FALSE,
  method = c("gsva", "plage", "ssgsea", "zscore"),
  minsize = 5L,
  maxsize = Inf,
  ...
)
```

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#### **Arguments**

mat Matrix to evaluate (e.g. expression matrix). Target nodes in rows and condi-

tions in columns. rownames(mat) must have at least one intersection with the

elements in network . target column.

Tibble or dataframe with edges and it's associated metadata. network

.source Column with source nodes.

.target Column with target nodes.

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

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). Further avail-

able methods are "plage", "ssgsea" and "zscore". Read more in the manual of

GSVA::gsva.

Integer indicating the minimum number of targets per source. Must be greater

than 0.

Integer indicating the maximum number of targets per source. maxsize

Arguments passed on to GSVA::gsvaParam, GSVA::ssgseaParam

assay The name of the assay to use in case exprData is a multi-assay container, otherwise ignored. By default, the first assay is used.

annotation The name of a Bioconductor annotation package for the gene identifiers occurring in the row names of the expression data matrix. This can be used to map gene identifiers occurring in the gene sets if those are provided in a GeneSetCollection. By default gene identifiers used in expression data matrix and gene sets are matched directly.

kcdf Character vector of length 1 denoting the kernel to use during the nonparametric estimation of the cumulative distribution function of expression levels across samples. By default, kcdf="Gaussian" which is suitable when input expression values are continuous, such as microarray fluorescent units in logarithmic scale, RNA-seq log-CPMs, log-RPKMs or log-TPMs. When input expression values are integer counts, such as those derived from RNA-seq experiments, then this argument should be set to kcdf="Poisson".

tau Numeric vector of length 1. The exponent defining the weight of the tail in the random walk performed by the GSVA (Hänzelmann et al., 2013) method. The default value is 1 as described in the paper.

maxDiff Logical vector of length 1 which offers two approaches to calculate the enrichment statistic (ES) from the KS random walk statistic.

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

absRanking Logical vector of length 1 used only when maxDiff=TRUE. When absRanking=FALSE (default) a modified Kuiper statistic is used to calculate enrichment scores, taking the magnitude difference between the largest positive and negative random walk deviations. When absRanking=TRUE the

minsize

. . .

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original Kuiper statistic that sums the largest positive and negative random walk deviations, is used. In this latter case, gene sets with genes enriched on either extreme (high or low) will be regarded as 'highly' activated.

alpha Numeric vector of length 1. The exponent defining the weight of the tail in the random walk performed by the ssGSEA (Barbie et al., 2009) method. The default value is 0.25 as described in the paper.

normalize Logical vector of length 1; if TRUE runs the ssGSEA method from Barbie et al. (2009) normalizing the scores by the absolute difference between the minimum and the maximum, as described in their paper. Otherwise this last normalization step is skipped.

#### **Details**

GSVA (Hänzelmann et al., 2013) starts by transforming the input molecular readouts in mat to a readout-level statistic using Gaussian kernel estimation of the cumulative density function. Then, readout-level statistics are ranked per sample and normalized to up-weight the two tails of the rank distribution. Afterwards, an enrichment score gsva is calculated using a running sum statistic that is normalized by subtracting the largest negative estimate from the largest positive one.

Hänzelmann S. et al. (2013) GSVA: gene set variation analysis for microarray and RNA-seq data. BMC Bioinformatics, 14, 7.

#### Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. statistic: Indicates which method is associated with which score.
- 2. source: Source nodes of network.
- 3. condition: Condition representing each column of mat.
- 4. score: Regulatory activity (enrichment score).

#### See Also

```
Other decoupleR statistics: decouple(), run_aucell(), run_fgsea(), run_mdt(), run_mlm(), run_ora(), run_udt(), run_ulm(), run_viper(), run_wmean(), run_wsum()
```

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_gsva(mat, net, minsize=1, verbose = FALSE)</pre>
```

run\_mdt 27

 $\verb"run_mdt"$ 

Multivariate Decision Trees (MDT)

## Description

Calculates regulatory activities using MDT.

## Usage

```
run_mdt(
 mat,
 network,
  .source = source,
  .target = target,
  .mor = mor,
  .likelihood = likelihood,
  sparse = FALSE,
  center = FALSE,
 na.rm = FALSE,
  trees = 10,
 min_n = 20,
 nproc = availableCores(),
  seed = 42,
 minsize = 5
)
```

## Arguments

mat	Matrix to evaluate (e.g. expression matrix). Target nodes in rows and conditions in columns. rownames(mat) must have at least one intersection with the elements in network .target column.
network	Tibble or dataframe with edges and it's associated metadata.
.source	Column with source nodes.
.target	Column with target nodes.
.mor	Column with edge mode of regulation (i.e. mor).
.likelihood	Deprecated argument. Now it will always be set to 1.
sparse	Deprecated parameter.
center	Logical value indicating if mat must be centered by base::rowMeans().
na.rm	Should missing values (including NaN) be omitted from the calculations of base::rowMeans()?
trees	An integer for the number of trees contained in the ensemble.
min_n	An integer for the minimum number of data points in a node that are required for the node to be split further.
nproc	Number of cores to use for computation.

zun\_mlm

seed A single value, interpreted as an integer, or NULL for random number generation.

minsize Integer indicating the minimum number of targets per source.

#### **Details**

MDT fits a multivariate regression random forest for each sample, where the observed molecular readouts in mat are the response variable and the regulator weights in net are the covariates. Target features with no associated weight are set to zero. The obtained feature importances from the fitted model are the activities mdt of the regulators in net.

#### Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. statistic: Indicates which method is associated with which score.
- 2. source: Source nodes of network.
- 3. condition: Condition representing each column of mat.
- 4. score: Regulatory activity (enrichment score).

### See Also

```
Other decoupleR statistics: decouple(), run_aucell(), run_fgsea(), run_gsva(), run_mlm(), run_ora(), run_udt(), run_ulm(), run_viper(), run_wmean(), run_wsum()
```

## **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_mdt(mat, net, minsize=0)</pre>
```

run\_mlm

Multivariate Linear Model (MLM)

#### **Description**

Calculates regulatory activities using MLM.

run\_mlm 29

#### Usage

```
run_mlm(
   mat,
   network,
   .source = source,
   .target = target,
   .mor = mor,
   .likelihood = likelihood,
   sparse = FALSE,
   center = FALSE,
   na.rm = FALSE,
   minsize = 5
)
```

#### **Arguments**

mat	Matrix to evaluate (e.g. expression matrix). Target nodes in rows and conditions in columns. rownames(mat) must have at least one intersection with the elements in network .target column.
network	Tibble or dataframe with edges and it's associated metadata.
.source	Column with source nodes.
.target	Column with target nodes.
.mor	Column with edge mode of regulation (i.e. mor).
.likelihood	Deprecated argument. Now it will always be set to 1.
sparse	Deprecated parameter.
center	Logical value indicating if mat must be centered by base::rowMeans().
na.rm	Should missing values (including NaN) be omitted from the calculations of base::rowMeans()?
minsize	Integer indicating the minimum number of targets per source.

## **Details**

MLM fits a multivariate linear model for each sample, where the observed molecular readouts in mat are the response variable and the regulator weights in net are the covariates. Target features with no associated weight are set to zero. The obtained t-values from the fitted model are the activities (mlm) of the regulators in net.

#### Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. statistic: Indicates which method is associated with which score.
- 2. source: Source nodes of network.
- 3. condition: Condition representing each column of mat.
- 4. score: Regulatory activity (enrichment score).

run\_ora

#### See Also

```
Other decoupleR statistics: decouple(), run_aucell(), run_fgsea(), run_gsva(), run_mdt(), run_ora(), run_udt(), run_viper(), run_wmean(), run_wsum()
```

## **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_mlm(mat, net, minsize=0)</pre>
```

run\_ora

Over Representation Analysis (ORA)

## **Description**

Calculates regulatory activities using ORA.

## Usage

```
run_ora(
  mat,
  network,
  .source = source,
  .target = target,
  n_up = ceiling(0.05 * nrow(mat)),
  n_bottom = 0,
  n_background = 20000,
  with_ties = TRUE,
  seed = 42,
  minsize = 5,
  ...
)
```

## Arguments

mat	Matrix to evaluate (e.g. expression matrix). Target nodes in rows and conditions in columns. rownames(mat) must have at least one intersection with the elements in network .target column.
network	Tibble or dataframe with edges and it's associated metadata.
.source	Column with source nodes.
.target	Column with target nodes.
n_up	Integer indicating the number of top targets to slice from mat.
n_bottom	Integer indicating the number of bottom targets to slice from mat.

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n\_background Integer indicating the background size of the sliced targets. If not specified the number of background targets is determined by the total number of unique targets in the union of mat and network.

with\_ties Should ties be kept together? The default, TRUE, may return more rows than you request. Use FALSE to ignore ties, and return the first n rows.

seed A single value, interpreted as an integer, or NULL for random number genera-

minsize Integer indicating the minimum number of targets per source.

.. Arguments passed on to stats::fisher.test

workspace an integer specifying the size of the workspace used in the network algorithm. In units of 4 bytes. Only used for non-simulated p-values larger than  $2 \times 2$  tables. Since R version 3.5.0, this also increases the internal stack size which allows larger problems to be solved, however sometimes needing hours. In such cases, simulate.p.values=TRUE may be more reasonable.

hybrid a logical. Only used for larger than  $2 \times 2$  tables, in which cases it indicates whether the exact probabilities (default) or a hybrid approximation thereof should be computed.

hybridPars a numeric vector of length 3, by default describing "Cochran's conditions" for the validity of the chisquare approximation, see 'Details'.

control a list with named components for low level algorithm control. At present the only one used is "mult", a positive integer  $\geq 2$  with default 30 used only for larger than  $2 \times 2$  tables. This says how many times as much space should be allocated to paths as to keys: see file 'fexact.c' in the sources of this package.

or the hypothesized odds ratio. Only used in the  $2 \times 2$  case.

alternative indicates the alternative hypothesis and must be one of "two.sided", "greater" or "less". You can specify just the initial letter. Only used in the  $2\times 2$  case.

conf.int logical indicating if a confidence interval for the odds ratio in a  $2 \times 2$  table should be computed (and returned).

conf.level confidence level for the returned confidence interval. Only used in the  $2\times 2$  case and if conf.int = TRUE.

simulate.p.value a logical indicating whether to compute p-values by Monte Carlo simulation, in larger than  $2\times 2$  tables.

B an integer specifying the number of replicates used in the Monte Carlo test.

#### **Details**

ORA measures the overlap between the target feature set and a list of most altered molecular features in mat. The most altered molecular features can be selected from the top and or bottom of the molecular readout distribution, by default it is the top 5% positive values. With these, a contingency table is build and a one-tailed Fisher's exact test is computed to determine if a regulator's set of features are over-represented in the selected features from the data. The resulting score, ora, is the minus log10 of the obtained p-value.

run\_udt

## Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. statistic: Indicates which method is associated with which score.
- 2. source: Source nodes of network.
- 3. condition: Condition representing each column of mat.
- 4. score: Regulatory activity (enrichment score).

#### See Also

```
Other decoupleR statistics: decouple(), run_aucell(), run_fgsea(), run_gsva(), run_mdt(), run_mlm(), run_udt(), run_viper(), run_wmean(), run_wsum()
```

## Examples

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_ora(mat, net, minsize=0)</pre>
```

run\_udt

*Univariate Decision Tree (UDT)* 

## **Description**

Calculates regulatory activities by using UDT.

```
run_udt(
  mat,
  network,
  .source = source,
  .target = target,
  .mor = mor,
  .likelihood = likelihood,
  sparse = FALSE,
  center = FALSE,
  ma.rm = FALSE,
  min_n = 20,
  seed = 42,
  minsize = 5
)
```

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## Arguments

mat	Matrix to evaluate (e.g. expression matrix). Target nodes in rows and conditions in columns. rownames(mat) must have at least one intersection with the elements in network .target column.
network	Tibble or dataframe with edges and it's associated metadata.
.source	Column with source nodes.
.target	Column with target nodes.
.mor	Column with edge mode of regulation (i.e. mor).
.likelihood	Deprecated argument. Now it will always be set to 1.
sparse	Deprecated parameter.
center	Logical value indicating if mat must be centered by base::rowMeans().
na.rm	Should missing values (including NaN) be omitted from the calculations of $base::rowMeans()$ ?
min_n	An integer for the minimum number of data points in a node that are required for the node to be split further.
seed	A single value, interpreted as an integer, or NULL for random number generation.
minsize	Integer indicating the minimum number of targets per source.

#### **Details**

UDT fits a single regression decision tree for each sample and regulator, where the observed molecular readouts in mat are the response variable and the regulator weights in net are the explanatory one. Target features with no associated weight are set to zero. The obtained feature importance from the fitted model is the activity udt of a given regulator.

#### Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. statistic: Indicates which method is associated with which score.
- 2. source: Source nodes of network.
- $3.\ \ condition:$  Condition representing each column of mat.
- 4. score: Regulatory activity (enrichment score).

#### See Also

```
Other decoupleR statistics: decouple(), run_aucell(), run_fgsea(), run_gsva(), run_mdt(), run_mlm(), run_ora(), run_ulm(), run_viper(), run_wmean(), run_wsum()
```

run\_ulm

## **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_udt(mat, net, minsize=0)</pre>
```

run\_ulm

Univariate Linear Model (ULM)

## Description

Calculates regulatory activities using ULM.

## Usage

```
run_ulm(
  mat,
  network,
  .source = source,
  .target = target,
  .mor = mor,
  .likelihood = likelihood,
  sparse = FALSE,
  center = FALSE,
  na.rm = FALSE,
  minsize = 5L
)
```

## Arguments

mat	Matrix to evaluate (e.g. expression matrix). Target nodes in rows and conditions in columns. rownames(mat) must have at least one intersection with the elements in network . target column.
network	Tibble or dataframe with edges and it's associated metadata.
.source	Column with source nodes.
.target	Column with target nodes.
.mor	Column with edge mode of regulation (i.e. mor).
.likelihood	Deprecated argument. Now it will always be set to 1.
sparse	Deprecated parameter.
center	Logical value indicating if mat must be centered by base::rowMeans().
na.rm	Should missing values (including NaN) be omitted from the calculations of base::rowMeans()?
minsize	Integer indicating the minimum number of targets per source.

run\_viper 35

#### **Details**

ULM fits a linear model for each sample and regulator, where the observed molecular readouts in mat are the response variable and the regulator weights in net are the explanatory one. Target features with no associated weight are set to zero. The obtained t-value from the fitted model is the activity ulm of a given regulator.

#### Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. statistic: Indicates which method is associated with which score.
- 2. source: Source nodes of network.
- 3. condition: Condition representing each column of mat.
- 4. score: Regulatory activity (enrichment score).

#### See Also

```
Other decoupleR statistics: decouple(), run_aucell(), run_fgsea(), run_gsva(), run_mdt(), run_mlm(), run_ora(), run_udt(), run_viper(), run_wmean(), run_wsum()
```

## **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_ulm(mat, net, minsize=0)</pre>
```

run\_viper

Virtual Inference of Protein-activity by Enriched Regulon analysis (VIPER)

#### **Description**

Calculates regulatory activities using VIPER.

```
run_viper(
  mat,
  network,
  .source = source,
  .target = target,
  .mor = mor,
  .likelihood = likelihood,
```

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```
verbose = FALSE,
minsize = 5,
pleiotropy = TRUE,
eset.filter = FALSE,
...
)
```

#### **Arguments**

Matrix to evaluate (e.g. expression matrix). Target nodes in rows and conditions in columns. rownames(mat) must have at least one intersection with the

elements in network .target column.

network Tibble or dataframe with edges and it's associated metadata.

. source Column with source nodes.. target Column with target nodes.

.mor Column with edge mode of regulation (i.e. mor)..likelihood Deprecated argument. Now it will always be set to 1.

verbose Logical, whether progression messages should be printed in the terminal.

minsize Integer indicating the minimum number of targets per source.

pleiotropy Logical, whether correction for pleiotropic regulation should be performed.

eset.filter Logical, whether the dataset should be limited only to the genes represented in

the interactome.

... Arguments passed on to viper::viper

dnull Numeric matrix for the null model, usually generated by nullTtest nes Logical, whether the enrichment score reported should be normalized method Character string indicating the method for computing the single samples signature, either scale, rank, mad, ttest or none

bootstraps Integer indicating the number of bootstraps iterations to perform. Only the scale method is implemented with bootstraps.

adaptive.size Logical, whether the weighting scores should be taken into account for computing the regulon size

pleiotropyArgs list of 5 numbers for the pleotropy correction indicating: regulators p-value threshold, pleiotropic interaction p-value threshold, minimum number of targets in the overlap between pleiotropic regulators, penalty for the pleiotropic interactions and the method for computing the pleiotropy, either absolute or adaptive

cores Integer indicating the number of cores to use (only 1 in Windows-based systems)

#### **Details**

VIPER (Alvarez et al., 2016) estimates biological activities by performing a three-tailed enrichment score calculation. For further information check the supplementary information of the decoupler manuscript or the original publication.

Alvarez M.J.et al. (2016) Functional characterization of somatic mutations in cancer using network-based inference of protein activity. Nat. Genet., 48, 838–847.

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#### Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. statistic: Indicates which method is associated with which score.
- 2. source: Source nodes of network.
- 3. condition: Condition representing each column of mat.
- 4. score: Regulatory activity (enrichment score).

#### See Also

```
Other decoupleR statistics: decouple(), run_aucell(), run_fgsea(), run_gsva(), run_mdt(), run_mlm(), run_ora(), run_udt(), run_ulm(), run_wmean(), run_wsum()
```

#### **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_viper(mat, net, minsize=0, verbose = FALSE)</pre>
```

run\_wmean

Weighted Mean (WMEAN)

## **Description**

Calculates regulatory activities using WMEAN.

```
run_wmean(
  mat,
  network,
  .source = source,
  .target = target,
  .mor = mor,
  .likelihood = likelihood,
  times = 100,
  seed = 42,
  sparse = TRUE,
  randomize_type = "rows",
  minsize = 5
)
```

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#### **Arguments**

mat Matrix to evaluate (e.g. expression matrix). Target nodes in rows and condi-

tions in columns. rownames (mat) must have at least one intersection with the

elements in network .target column.

network Tibble or dataframe with edges and it's associated metadata.

. source Column with source nodes.. target Column with target nodes.

.mor Column with edge mode of regulation (i.e. mor).

.likelihood Deprecated argument. Now it will always be set to 1.

times How many permutations to do?

seed A single value, interpreted as an integer, or NULL for random number genera-

tion.

sparse Should the matrices used for the calculation be sparse?

randomize\_type How to randomize the expression matrix.

minsize Integer indicating the minimum number of targets per source.

#### **Details**

WMEAN infers regulator activities by first multiplying each target feature by its associated weight which then are summed to an enrichment score wmean. Furthermore, permutations of random target features can be performed to obtain a null distribution that can be used to compute a z-score norm\_wmean, or a corrected estimate corr\_wmean by multiplying wmean by the minus log10 of the obtained empirical p-value.

#### Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. statistic: Indicates which method is associated with which score.
- 2. source: Source nodes of network.
- 3. condition: Condition representing each column of mat.
- 4. score: Regulatory activity (enrichment score).
- 5. p\_value: p-value for the score of the method.

#### See Also

Other decoupleR statistics: decouple(), run\_aucell(), run\_fgsea(), run\_gsva(), run\_mdt(), run\_mlm(), run\_ora(), run\_udt(), run\_viper(), run\_wsum()

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#### **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_wmean(mat, net, minsize=0)</pre>
```

run\_wsum

Weighted Sum (WSUM)

## **Description**

Calculates regulatory activities using WSUM.

### Usage

```
run_wsum(
  mat,
  network,
  .source = source,
  .target = target,
  .mor = mor,
  .likelihood = likelihood,
  times = 100,
  seed = 42,
  sparse = TRUE,
  randomize_type = "rows",
  minsize = 5
)
```

#### Arguments

mat Matrix to evaluate (e.g. expression matrix). Target nodes in rows and condi-

tions in columns. rownames (mat) must have at least one intersection with the

elements in network .target column.

network Tibble or dataframe with edges and it's associated metadata.

. source Column with source nodes.. target Column with target nodes.

.mor Column with edge mode of regulation (i.e. mor)..likelihood Deprecated argument. Now it will always be set to 1.

times How many permutations to do?

seed A single value, interpreted as an integer, or NULL for random number genera-

tion.

sparse Should the matrices used for the calculation be sparse?

randomize\_type How to randomize the expression matrix.

minsize Integer indicating the minimum number of targets per source.

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#### **Details**

WSUM infers regulator activities by first multiplying each target feature by its associated weight which then are summed to an enrichment score wsum. Furthermore, permutations of random target features can be performed to obtain a null distribution that can be used to compute a z-score norm\_wsum, or a corrected estimate corr\_wsum by multiplying wsum by the minus log10 of the obtained empirical p-value.

#### Value

A long format tibble of the enrichment scores for each source across the samples. Resulting tibble contains the following columns:

- 1. statistic: Indicates which method is associated with which score.
- 2. source: Source nodes of network.
- 3. condition: Condition representing each column of mat.
- 4. score: Regulatory activity (enrichment score).
- 5. p\_value: p-value for the score of the method.

#### See Also

```
Other decoupleR statistics: decouple(), run_aucell(), run_fgsea(), run_gsva(), run_mdt(), run_mlm(), run_ora(), run_udt(), run_viper(), run_wmean()
```

#### **Examples**

```
inputs_dir <- system.file("testdata", "inputs", package = "decoupleR")
mat <- readRDS(file.path(inputs_dir, "mat.rds"))
net <- readRDS(file.path(inputs_dir, "net.rds"))
run_wsum(mat, net, minsize=0)</pre>
```

show\_methods

Show methods

#### **Description**

Prints the methods available in decoupleR. The first column correspond to the function name in decoupleR and the second to the method's full name.

#### Usage

```
show_methods()
```

```
show_methods()
```

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show\_resources

Shows available resources in Omnipath. For more information visit the official website for Rhrefhttps://omnipathdb.org/Omnipath.

#### **Description**

Shows available resources in Omnipath. For more information visit the official website for Omnipath.

### Usage

```
show_resources()
```

#### **Examples**

```
decoupleR::show_resources()
```

tidyeval

Tidy eval helpers

#### **Description**

- rlang::sym() creates a symbol from a string and syms() creates a list of symbols from a character vector.
- enquo() and enquos() delay the execution of one or several function arguments. enquo() returns a single quoted expression, which is like a blueprint for the delayed computation. enquos() returns a list of such quoted expressions.
- expr() quotes a new expression *locally*. It is mostly useful to build new expressions around arguments captured with enquo() or enquos(): expr(mean(!!enquo(arg), na.rm = TRUE)).
- rlang::as\_name() transforms a quoted variable name into a string. Supplying something else than a quoted variable name is an error.

That's unlike rlang::as\_label() which also returns a single string but supports any kind of R object as input, including quoted function calls and vectors. Its purpose is to summarise that object into a single label. That label is often suitable as a default name.

If you don't know what a quoted expression contains (for instance expressions captured with enquo() could be a variable name, a call to a function, or an unquoted constant), then use as\_label(). If you know you have quoted a simple variable name, or would like to enforce this, use as\_name().

To learn more about tidy eval and how to use these tools, visit https://tidyeval.tidyverse.org and the Metaprogramming section of Advanced R.

```
if (FALSE) {
   help("nse-defuse", package = "rlang")
}
```

42 %>%

%>%

Pipe operator

## Description

```
See magrittr::%>% for details.
```

## Usage

lhs %>% rhs

## Value

Pipe an object forward into a function or call expression.

```
c(1, 2, 3) %>% sum()
```

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