# Package 'OmnipathR'

December 6, 2024

Type Package

Title OmniPath web service client and more

**Version** 3.14.0

**Description** A client for the OmniPath web service (https://www.omnipathdb.org) and many other resources.

It also includes functions to transform and pretty print some of the downloaded data, functions to access a number of other resources such as BioPlex, ConsensusPathDB, EVEX, Gene Ontology, Guide to Pharmacology (IUPHAR/BPS), Harmonizome, HTRIdb, Human Phenotype Ontology, InWeb InBioMap, KEGG Pathway, Pathway Commons, Ramilowski et al. 2015, RegNetwork, ReMap, TF census, TRRUST and Vinayagam et al. 2011. Furthermore, OmnipathR features a close integration with the NicheNet method for ligand activity prediction from transcriptomics data, and its R implementation `nichenetr` (available only on github).

License MIT + file LICENSE

URL https://r.omnipathdb.org/

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

biocViews GraphAndNetwork, Network, Pathways, Software, ThirdPartyClient, DataImport, DataRepresentation, GeneSignaling, GeneRegulation, SystemsBiology, Transcriptomics, SingleCell, Annotation, KEGG

**Encoding** UTF-8

VignetteBuilder knitr

**Depends** R(>=4.0)

Imports checkmate, crayon, curl, digest, dplyr(>= 1.1.0), httr, igraph, jsonlite, later, logger, lubridate, magrittr, progress, purrr, rappdirs, readr(>= 2.0.0), readxl, rlang, rmarkdown, RSQLite, R.utils, rvest, stats, stringi, stringr, tibble, tidyr, tidyselect, tools, utils, vctrs, withr, XML, xml2, yaml, zip

**Suggests** BiocStyle, bookdown, ggplot2, ggraph, gprofiler2, knitr, mlrMBO, parallelMap, ParamHelpers, Rgraphviz, R.matlab, sigmajs, smoof, supraHex, testthat

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```
.omnipathr_options_defaults
```

Default values for the package options

## **Description**

These options describe the default settings for OmnipathR so you do not need to pass these parameters at each function call. Currently the only option useful for the public web service at omnipathdb.org is "omnipathr.license". If you are a for-profit user set it to "commercial" to make sure all the data you download from OmniPath is legally allowed for commercial use. Otherwise just leave it as it is: "academic". If you don't use omnipathdb.org but within your organization you deployed your own pypath server and want to share data whith a limited availability to outside users, you may want to use a password. For this you can use the "omnipathr.password" option. Also if you want the R package to work from another pypath server instead of omnipathdb.org, you can change the option "omnipathr.url".

## Usage

.omnipathr\_options\_defaults

#### **Format**

An object of class list of length 25.

#### Value

Nothing, this is not a function but a list.

all\_uniprots

A table with all UniProt records

## **Description**

Retrieves a table from UniProt with all proteins for a certain organism.

## Usage

```
all_uniprots(fields = "accession", reviewed = TRUE, organism = 9606L)
```

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### **Arguments**

fields Character vector of fields as defined by UniProt. For possible values please refer

to https://www.uniprot.org/help/return\_fields

reviewed ('TRUE'), only unreviewed ('FALSE') or both ('NULL').

organism Character or integer: name or identifier of the organism.

## Value

Data frame (tibble) with the requested UniProt entries and fields.

## **Examples**

```
human_swissprot_entries <- all_uniprots(fields = 'id')
human_swissprot_entries
# # A tibble: 20,396 x 1
# `Entry name`
# <chr>
# 1 OR4K3_HUMAN
# 2 052A1_HUMAN
# 3 02AG1_HUMAN
# 4 010S1_HUMAN
# 5 011G2_HUMAN
# 5 011G2_HUMAN
# # . with 20,386 more rows
```

all\_uniprot\_acs

All UniProt ACs for one organism

## **Description**

All UniProt ACs for one organism

## Usage

```
all_uniprot_acs(organism = 9606, reviewed = TRUE)
```

## Arguments

organism Character or integer: name or identifier of the organism.

reviewed ('TRUE'), only unreviewed ('FALSE') or both ('NULL').

## Value

Character vector of UniProt accession numbers.

```
human_swissprot_acs <- all_uniprot_acs()
human_swissprot_acs[1:5]
# [1] "P51451" "A6H8Y1" "060885" "Q9Y3X0" "P22223"
length(human_swissprot_acs)
# [1] 20376
mouse_swissprot_acs <- all_uniprot_acs("mouse")</pre>
```

ancestors 9

ancestors

All ancestors in the ontology tree

## Description

Starting from the selected nodes, recursively walks the ontology tree until it reaches the root. Collects all visited nodes, which are the ancestors (parents) of the starting nodes.

## Usage

## Arguments

terms	Character vector of ontology term IDs or names. A mixture of IDs and names can be provided.
db_key	Character: key to identify the ontology database. For the available keys see omnipath_show_db.
ids	Logical: whether to return IDs or term names.
relations	Character vector of ontology relation types. Only these relations will be used.

### **Details**

Note: this function relies on the database manager, the first call might take long because of the database load process. Subsequent calls within a short period should be faster. See <a href="mailto:get\_ontology\_db">get\_ontology\_db</a>.

## Value

Character vector of ontology IDs. If the input terms are all root nodes, NULL is returned. The starting nodes won't be included in the result unless some of them are ancestors of other starting nodes.

```
ancestors('GO:0005035', ids = FALSE)
# [1] "molecular_function"
# [2] "transmembrane signaling receptor activity"
# [3] "signaling receptor activity"
# [4] "molecular transducer activity"
```

10 annotated\_network

annotated\_network

Network interactions with annotations

### **Description**

Annotations are often useful in a network context, e.g. one might want to label the interacting partners by their pathway membership. This function takes a network data frame and joins an annotation data frame from both the left and the right side, so both the source and target molecular entities will be labeled by their annotations. If one entity has many annotations these will yield many rows, hence the interacting pairs won't be unique across the data frame any more. Also if one entity has really many annotations the resulting data frame might be huge, we recommend to be careful with that. Finally, if you want to do the same but with intercell annotations, there is the import\_intercell\_network function.

## Usage

```
annotated_network(
 network = NULL,
  annot = NULL,
 network_args = list(),
  annot_args = list(),
)
```

## **Arguments**

network	Behaviour depends on type: if list, will be passed as arguments to omnipath_interactions to obtain a network data frame; if a data frame or tibble, it will be used as a network data frame; if a character vector, will be assumed to be a set of resource names and interactions will be queried from these resources.
annot	Either the name of an annotation resource (for a list of available resources call annotation_resources), or an annotation data frame. If the data frame contains more than one resources, only the first one will be used.
network_args	List: if 'network' is a resource name, pass these additional arguments to omnipath_interactions.
annot_args	List: if 'annot' is a resource name, pass these additional arguments to annotations.
• • •	Column names selected from the annotation data frame (passed to dplyr::select, if empty all columns will be selected.)

### Value

A data frame of interactions with annotations for both interacting entities.

```
signalink_with_pathways <-</pre>
    annotated_network("SignaLink3", "SignaLink_pathway")
```

annotations 11

annotations

Protein and gene annotations from OmniPath

#### **Description**

Protein and gene annotations about function, localization, expression, structure and other properties, from the <a href="https://omnipathdb.org/annotations">https://omnipathdb.org/annotations</a> endpoint of the OmniPath web service. Note: there might be also a few miRNAs annotated; a vast majority of protein complex annotations are inferred from the annotations of the members: if all members carry the same annotation the complex inherits.

#### Usage

```
annotations(proteins = NULL, wide = FALSE, ...)
```

#### **Arguments**

proteins

Vector containing the genes or proteins for whom annotations will be retrieved (UniProt IDs or HGNC Gene Symbols or miRBase IDs). It is also possible to donwload annotations for protein complexes. To do so, write 'COMPLEX:' right before the genesymbols of the genes integrating the complex. Check the vignette for examples.

wide

Convert the annotation table to wide format, which corresponds more or less to the original resource. If the data comes from more than one resource a list of wide tables will be returned. See examples at pivot\_annotations.

. . .

Arguments passed on to omnipath\_query

organism Character or integer: name or NCBI Taxonomy ID of the organism. OmniPath is built of human data, and the web service provides orthology translated interactions and enzyme-substrate relationships for mouse and rat. For other organisms and query types, orthology translation will be called automatically on the downloaded human data before returning the result.

resources Character vector: name of one or more resources. Restrict the data to these resources. For a complete list of available resources, call the '<query\_type>\_resources' functions for the query type of interst.

genesymbols Character or logical: TRUE or FALS or "yes" or "no". Include the 'genesymbols' column in the results. OmniPath uses UniProt IDs as the primary identifiers, gene symbols are optional.

fields Character vector: additional fields to include in the result. For a list of available fields, call 'query\_info("interactions")'.

default\_fields Logical: if TRUE, the default fields will be included.

silent Logical: if TRUE, no messages will be printed. By default a summary message is printed upon successful download.

logicals Character vector: fields to be cast to logical.

format Character: if "json", JSON will be retrieved and processed into a nested list; any other value will return data frame.

download\_args List: parameters to pass to the download function, which is 'readr::read\_tsv' by default, and 'jsonlite::safe\_load'.

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add\_counts Logical: if TRUE, the number of references and number of resources for each record will be added to the result.

- license Character: license restrictions. By default, data from resources allowing "academic" use is returned by OmniPath. If you use the data for work in a company, you can provide "commercial" or "for-profit", which will restrict the data to those records which are supported by resources that allow for-profit use.
- password Character: password for the OmniPath web service. You can provide a special password here which enables the use of 'license = "ignore" option, completely bypassing the license filter.
- exclude Character vector: resource or dataset names to be excluded. The data will be filtered after download to remove records of the excluded datasets and resources.
- strict\_evidences Logical: reconstruct the "sources" and "references" columns of interaction data frames based on the "evidences" column, strictly filtering them to the queried datasets and resources. Without this, the "sources" and "references" fields for each record might contain information for datasets and resources other than the queried ones, because the downloaded records are a result of a simple filtering of an already integrated data frame.
- genesymbol\_resource Character: "uniprot" (default) or "ensembl". The OmniPath web service uses the primary gene symbols as provided by UniProt. By passing "ensembl" here, the UniProt gene symbols will be replaced by the ones used in Ensembl. This translation results in a loss of a few records, and multiplication of another few records due to ambiguous translation.
- cache Logical: use caching, load data from and save to the. The cache directory by default belongs to the user, located in the user's default cache directory, and named "OmnipathR". Find out about it by getOption("omnipathr.cachedir"). Can be changed by omnipath\_set\_cachedir.

### **Details**

Downloading the full annotations dataset is disabled by default because the size of this data is around 1GB. We recommend to retrieve the annotations for a set of proteins or only from a few resources, depending on your interest. You can always download the full database from <a href="https://archive.omnipathdb.org/omnipath\_webservice\_annotations\_\_recent.tsv">https://archive.omnipathdb.org/omnipath\_webservice\_annotations\_\_recent.tsv</a> using any standard R or readr method.

#### Value

A data frame or list of data frames:

- If wide=FALSE (default), all the requested resources will be in a single long format data frame.
- If wide=TRUE: one or more data frames with columns specific to the requested resources. If more than one resources is requested a list of data frames is returned.

#### See Also

- annotation\_resources
- pivot\_annotations
- query\_info
- omnipath\_query
- annotated\_network

annotation\_categories 13

#### **Examples**

```
annotations <- annotations(
   proteins = c("TP53", "LMNA"),
   resources = c("HPA_subcellular")
)</pre>
```

annotation\_categories Annotation categories and resources

## **Description**

A full list of annotation resources, keys and values.

## Usage

```
annotation_categories()
```

#### Value

A data frame with resource names, annotation key labels and for each key all possible values.

#### **Examples**

```
annot_cat <- annotation_categories()</pre>
annot_cat
# # A tibble: 46,307 x 3
    source
                      label
                               value
#
     <chr>
                     <chr>
                               <chr>
# 1 connectomeDB2020 role
                               ligand
  2 connectomeDB2020 role
                               receptor
  3 connectomeDB2020 location ECM
  4 connectomeDB2020 location plasma membrane
  5 connectomeDB2020 location secreted
                     pathway Alanine, aspartate and glutamate metabolism
  6 KEGG-PC
# 7 KEGG-PC
# 8 KEGG-PC
# 9 KEGG-PC
                      pathway Amino sugar and nucleotide sugar metabolism
                     pathway Aminoacyl-tRNA biosynthesis
                      pathway Arachidonic acid metabolism
# 10 KEGG-PC
                      pathway Arginine and proline metabolism
```

annotation\_resources Retrieves a list of available resources in the annotations database of OmniPath

## **Description**

Get the names of the resources from <a href="https://omnipathdb.org/annotations">https://omnipathdb.org/annotations</a>.

#### Usage

```
annotation_resources(dataset = NULL, ...)
```

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## **Arguments**

```
dataset ignored for this query type
... optional additional arguments
```

## Value

character vector with the names of the annotation resources

#### See Also

- resources
- annotations

## **Examples**

```
annotation_resources()
```

biomart\_query

Query the Ensembl BioMart web service

## **Description**

Query the Ensembl BioMart web service

## Usage

```
biomart_query(
  attrs = NULL,
  filters = NULL,
  transcript = FALSE,
  peptide = FALSE,
  gene = FALSE,
  dataset = "hsapiens_gene_ensembl"
)
```

## **Arguments**

attrs Character vector: one or more Ensembl attribute names.

filters Character vector: one or more Ensembl filter names.

transcript Logical: include Ensembl transcript IDs in the result.

peptide Logical: include Ensembl peptide IDs in the result.

gene Logical: include Ensembl gene IDs in the result.

dataset Character: An Ensembl dataset name.

## Value

Data frame with the query result

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### **Examples**

```
cel_genes <- biomart_query(</pre>
    attrs = c("external_gene_name", "start_position", "end_position"),
    gene = TRUE,
    dataset = "celegans_gene_ensembl"
)
cel_genes
# # A tibble: 46,934 \times 4
   ensembl_gene_id external_gene_name start_position end_position
                    <chr>
                                                <dbl>
                                                             <dbl>
# 1 WBGene00000001 aap-1
                                              5107843
                                                            5110183
# 2 WBGene00000002 aat-1
                                              9599178
                                                           9601695
# 3 WBGene00000003 aat-2
                                              9244402
                                                            9246360
# 4 WBGene00000004 aat-3
                                              2552260
                                                            2557736
# 5 WBGene00000005 aat-4
                                              6272529
                                                            6275721
# # . with 46,924 more rows
```

bioplex1

Downloads the BioPlex version 1.0 interaction dataset

## Description

This dataset contains ~24,000 interactions detected in HEK293T cells using 2,594 baits. More details at https://bioplex.hms.harvard.edu/interactions.php.

## Usage

```
bioplex1()
```

## Value

Data frame (tibble) with interactions.

### See Also

- bioplex2
- bioplex3
- bioplex\_hct116\_1
- bioplex\_all

```
bioplex_interactions <- bioplex1()
nrow(bioplex_interactions)
# [1] 23744
colnames(bioplex_interactions)
# [1] "GeneA" "GeneB" "UniprotA" "UniprotB"
# [5] "SymbolA" "SymbolB" "p_wrong" "p_no_interaction"
# [9] "p_interaction"</pre>
```

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bioplex2

Downloads the BioPlex version 2.0 interaction dataset

## **Description**

This dataset contains ~56,000 interactions detected in HEK293T cells using 5,891 baits. More details at https://bioplex.hms.harvard.edu/interactions.php

## Usage

```
bioplex2()
```

#### Value

Data frame (tibble) with interactions.

#### See Also

- bioplex1
- bioplex3
- bioplex\_hct116\_1
- bioplex\_all

#### **Examples**

```
bioplex_interactions <- bioplex2()
nrow(bioplex_interactions)
# [1] 56553
colnames(bioplex_interactions)
# [1] "GeneA" "GeneB" "UniprotA" "UniprotB"
# [5] "SymbolA" "SymbolB" "p_wrong" "p_no_interaction"
# [9] "p_interaction"</pre>
```

bioplex3

Downloads the BioPlex version 3.0 interaction dataset

## **Description**

This dataset contains ~120,000 interactions detected in HEK293T cells using 10,128 baits. More details at https://bioplex.hms.harvard.edu/interactions.php.

## Usage

```
bioplex3()
```

#### Value

Data frame (tibble) with interactions.

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#### See Also

- bioplex1
- bioplex2
- bioplex\_hct116\_1
- bioplex\_all

## **Examples**

```
bioplex_interactions <- bioplex3()
nrow(bioplex_interactions)
# [1] 118162
colnames(bioplex_interactions)
# [1] "GeneA" "GeneB" "UniprotA" "UniprotB"
# [5] "SymbolA" "SymbolB" "p_wrong" "p_no_interaction"
# [9] "p_interaction"</pre>
```

bioplex\_all

Downloads all BioPlex interaction datasets

### **Description**

BioPlex provides four interaction datasets: version 1.0, 2.0, 3.0 and HCT116 version 1.0. This function downloads all of them, merges them to one data frame, removes the duplicates (based on unique pairs of UniProt IDs) and separates the isoform numbers from the UniProt IDs. More details at https://bioplex.hms.harvard.edu/interactions.php.

## Usage

```
bioplex_all(unique = TRUE)
```

## **Arguments**

unique

Logical. Collapse the duplicate interactions into single rows or keep them as they are. In case of merging duplicate records the maximum p value will be choosen for each record.

## Value

Data frame (tibble) with interactions.

#### See Also

- bioplex1
- bioplex2
- bioplex3
- bioplex\_hct116\_1

bioplex\_hct116\_1

#### **Examples**

```
bioplex_interactions <- bioplex_all()</pre>
bioplex_interactions
# # A tibble: 195,538 x 11
    UniprotA IsoformA UniprotB IsoformB GeneA GeneB SymbolA SymbolB
#
            NA 84561 11000 SLC12A8 SLC27A3
# 1 A0AV02
               2 Q5K4L6
                 2 Q8N5V2
# 2 A0AV02
                                 NA 84561 25791 SLC12A8 NGEF
                2 Q9H6S3
                               NA 84561 64787 SLC12A8 EPS8L2
# 3 A0AV02
                 2 000425
# 4 A0AV96
                                 2 54502 10643 RBM47 IGF2BP3
                               NA 54502 5286 RBM47 PIK3C2A
# 5 A0AV96
                 2 000443
                              NA 54502 8867 RBM47 SYNJ1
                2 043426
# 6 A0AV96
# 7 A0AV96
                2 075127
                               NA 54502 26024 RBM47 PTCD1
# 8 A0AV96
                2 095208
                                 2 54502 22905 RBM47 EPN2
# 9 A0AV96 2 095900 NA 54502 26995 RBM47 TRUB2
# 10 A0AV96 2 P07910 2 54502 3183 RBM47 HNRNPC
# # . with 195,528 more rows, and 3 more variables: p_wrong <dbl>,
# # p_no_interaction <dbl>, p_interaction <dbl>
```

bioplex\_hct116\_1

Downloads the BioPlex HCT116 version 1.0 interaction dataset

## Description

This dataset contains ~71,000 interactions detected in HCT116 cells using 5,522 baits. More details at https://bioplex.hms.harvard.edu/interactions.php.

### Usage

```
bioplex_hct116_1()
```

### Value

Data frame (tibble) with interactions.

## See Also

- bioplex1
- bioplex2
- bioplex3
- bioplex\_all

```
bioplex_interactions <- bioplex_hct116_1()
nrow(bioplex_interactions)
# [1] 70966
colnames(bioplex_interactions)
# [1] "GeneA" "GeneB" "UniprotA" "UniprotB"
# [5] "SymbolA" "SymbolB" "p_wrong" "p_no_interaction"
# [9] "p_interaction"</pre>
```

bma\_motif\_es 19

bma\_motif\_es

BMA motifs from a sequence of edges

## **Description**

These motifs can be added to a BMA canvas.

#### Usage

```
bma_motif_es(edge_seq, G, granularity = 2)
```

## **Arguments**

edge\_seq An igraph edge sequence.

G An igraph graph object.

granularity Numeric: granularity value.

### Value

Character: BMA motifs as a single string.

## **Examples**

```
interactions <- omnipath(resources = "ARN")
graph <- interaction_graph(interactions)
motifs <- bma_motif_es(igraph::E(graph)[1], graph)</pre>
```

bma\_motif\_vs

Prints a BMA motif to the screen from a sequence of nodes, which can be copy/pasted into the BMA canvas

## Description

Intended to parallel print\_path\_vs

## Usage

```
bma_motif_vs(node_seq, G)
```

## **Arguments**

node\_seq An igraph node sequence.

G An igraph graph object.

## Value

Character: BMA motifs as a single string.

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#### **Examples**

```
interactions <- omnipath(resources = "ARN")
graph <- interaction_graph(interactions)
bma_string <- bma_motif_vs(
    igraph::all_shortest_paths(
        graph,
        from = 'ULK1',
        to = 'ATG13'
    )$res,
    graph
)</pre>
```

chalmers\_gem

Genome scale metabolic model by Wang et al. 2021

## Description

Process the GEMs from Wang et al., 2021 (https://github.com/SysBioChalmers) into convenient tables.

#### Usage

```
chalmers_gem(organism = "Human", orphans = TRUE)
```

## **Arguments**

organism Character or integer: an organism (taxon) identifier. Supported taxons are 9606

(Homo sapiens), 10090 (Mus musculus), 10116 (Rattus norvegicus), 7955 (Danio rerio), 7227 (Drosophila melanogaster) and 6239 (Caenorhabditis elegans).

orphans Logical: include orphan reactions (reactions without known enzyme).

## Value

List containing the following elements:

- reactions: tibble of reaction data;
- metabolites: tibble of metabolite data;
- reaction\_ids: translation table of reaction identifiers;
- metabolite\_ids: translation table of metabolite identifiers;
- S: Stoichiometric matrix (sparse).

#### References

Wang H, Robinson JL, Kocabas P, Gustafsson J, Anton M, Cholley PE, Huang S, Gobom J, Svensson T, Uhlen M, Zetterberg H, Nielsen J. Genome-scale metabolic network reconstruction of model animals as a platform for translational research. Proc Natl Acad Sci U S A. 2021 Jul 27;118(30):e2102344118. doi: doi:10.1073/pnas.2102344118.

## See Also

- chalmers\_gem\_network
- chalmers\_gem\_metabolites
- chalmers\_gem\_reactions
- chalmers\_gem\_raw
- chalmers\_gem\_id\_mapping\_table
- cosmos\_pkn

### **Examples**

```
gem <- chalmers_gem()</pre>
```

```
chalmers_gem_id_mapping_table
```

Metabolite ID translation tables from Chalmers Sysbio

## **Description**

Metabolite ID translation tables from Chalmers Sysbio

## Usage

```
chalmers_gem_id_mapping_table(to, from = "metabolicatlas", organism = "Human")
```

## **Arguments**

to Character: type of ID to translate to, either label used internally in this package,

or a column name from "metabolites.tsv" distributed by Chalmers Sysbio. NSE

is supported.

from Character: type of ID to translate from, same format as "to".

organism Character or integer: name or identifier of the organism. Supported taxons are

9606 (Homo sapiens), 10090 (Mus musculus), 10116 (Rattus norvegicu), 7955 (Danio rerio), 7227 (Drosophila melanogaster) and 6239 (Caenorhabditis ele-

gans).

#### Value

Tibble with two columns, "From" and "To", with the corresponding ID types.

```
chalmers_gem_id_mapping_table('metabolicatlas', 'hmdb')
```

## Description

Metabolite identifier type label used in Chalmers Sysbio GEM

## Usage

```
chalmers_gem_id_type(label)
```

## **Arguments**

label

Character: an ID type label, as shown in the table at translate\_ids

#### Value

Character: the Chalmers GEM specific ID type label, or the input unchanged if it could not be translated (still might be a valid identifier name). These labels should be column names from the "metabolites.tsv" distributed with the GEMs.

#### See Also

- hmdb\_id\_type
- uniprot\_id\_type
- ensembl\_id\_type
- uploadlists\_id\_type

## **Examples**

```
chalmers_gem_id_type("metabolicatlas")
# [1] "metsNoComp"
```

```
chalmers_gem_metabolites
```

Metabolites from the Chalmers SysBio GEM (Wang et al., 2021)

## Description

Metabolites from the Chalmers SysBio GEM (Wang et al., 2021)

## Usage

```
chalmers_gem_metabolites(organism = "Human")
```

### **Arguments**

organism

Character or integer: an organism (taxon) identifier. Supported taxons are 9606 (Homo sapiens), 10090 (Mus musculus), 10116 (Rattus norvegicu), 7955 (Danio rerio), 7227 (Drosophila melanogaster) and 6239 (Caenorhabditis elegans).

#### Value

Data frame of metabolite identifiers.

#### References

Wang H, Robinson JL, Kocabas P, Gustafsson J, Anton M, Cholley PE, Huang S, Gobom J, Svensson T, Uhlen M, Zetterberg H, Nielsen J. Genome-scale metabolic network reconstruction of model animals as a platform for translational research. Proc Natl Acad Sci U S A. 2021 Jul 27;118(30):e2102344118. doi: doi:10.1073/pnas.2102344118.

## See Also

- chalmers\_gem\_network
- chalmers\_gem\_reactions
- chalmers\_gem
- chalmers\_gem\_raw
- chalmers\_gem\_id\_mapping\_table
- cosmos\_pkn

#### **Examples**

```
chalmers_gem_metabolites()
```

chalmers\_gem\_network Chalmers SysBio GEM in the form of gene-metabolite interactions

## **Description**

Processing GEMs from Wang et al., 2021 (https://github.com/SysBioChalmers) to generate PKN for COSMOS

### Usage

```
chalmers_gem_network(
  organism_or_gem = "Human",
  metab_max_degree = 400L,
  protein_ids = c("uniprot", "genesymbol"),
  metabolite_ids = c("hmdb", "kegg")
)
```

## **Arguments**

```
organism_or_gem
```

Character or integer or list or data frame: either an organism (taxon) identifier or a list containing the "reactions" data frame as it is provided by chalmers\_gem, or the reactions data frame itself. Supported taxons are 9606 (Homo sapiens), 10090 (Mus musculus), 10116 (Rattus norvegicus), 7955 (Danio rerio), 7227 (Drosophila melanogaster) and 6239 (Caenorhabditis elegans).

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metab\_max\_degree

Degree cutoff used to prune metabolites with high degree assuming they are cofactors (400 by default).

protein\_ids

Character: translate the protein identifiers to these ID types. Each ID type results two extra columns in the output, for the "a" and "b" sides of the interaction, respectively. The default ID type for proteins is Esembl Gene ID, and by default UniProt IDs and Gene Symbols are included.

metabolite\_ids Character: translate the protein identifiers to these ID types. Each ID type results two extra columns in the output, for the "a" and "b" sides of the interaction, respectively. The default ID type for metabolites is Metabolic Atlas ID, and HMDB IDs and KEGG IDs are included.

#### Value

Data frame (tibble) of gene-metabolite interactions.

#### References

Wang H, Robinson JL, Kocabas P, Gustafsson J, Anton M, Cholley PE, Huang S, Gobom J, Svensson T, Uhlen M, Zetterberg H, Nielsen J. Genome-scale metabolic network reconstruction of model animals as a platform for translational research. Proc Natl Acad Sci U S A. 2021 Jul 27;118(30):e2102344118. doi: doi:10.1073/pnas.2102344118.

#### See Also

- chalmers\_gem
- chalmers\_gem\_metabolites
- chalmers\_gem\_reactions
- chalmers\_gem\_raw
- chalmers\_gem\_id\_mapping\_table
- cosmos\_pkn

## **Examples**

```
gem <- chalmers_gem_network()</pre>
```

chalmers\_gem\_raw

GEM matlab file from Chalmers Sysbio (Wang et al., 2021)

## **Description**

Downloads and imports the matlab file containing the genome scale metabolic models created by Chalmers SysBio.

## Usage

```
chalmers_gem_raw(organism = "Human")
```

### **Arguments**

organism

Character or integer: name or identifier of the organism. Supported taxons are 9606 (Homo sapiens), 10090 (Mus musculus), 10116 (Rattus norvegicu), 7955 (Danio rerio), 7227 (Drosophila melanogaster) and 6239 (Caenorhabditis elegans).

#### **Details**

The Matlab object is parsed into a nested list containing a number of vectors and two sparse matrices. The top level contains a single element under the name "ihuman" for human; under this key there is an array of 31 elements. These elements are labeled by the row names of the array.

#### Value

Matlab object containing the GEM.

#### References

Wang H, Robinson JL, Kocabas P, Gustafsson J, Anton M, Cholley PE, Huang S, Gobom J, Svensson T, Uhlen M, Zetterberg H, Nielsen J. Genome-scale metabolic network reconstruction of model animals as a platform for translational research. Proc Natl Acad Sci U S A. 2021 Jul 27;118(30):e2102344118. doi: doi:10.1073/pnas.2102344118.

#### See Also

- chalmers\_gem\_network
- chalmers\_gem\_reactions
- chalmers\_gem
- chalmers\_gem\_reactions
- chalmers\_gem\_id\_mapping\_table
- cosmos\_pkn

## **Examples**

```
chalmers_gem_raw()
```

chalmers\_gem\_reactions

Reactions from the Chalmers SysBio GEM (Wang et al., 2021)

## Description

Reactions from the Chalmers SysBio GEM (Wang et al., 2021)

## Usage

```
chalmers_gem_reactions(organism = "Human")
```

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#### **Arguments**

organism

Character or integer: an organism (taxon) identifier. Supported taxons are 9606 (Homo sapiens), 10090 (Mus musculus), 10116 (Rattus norvegicu), 7955 (Danio rerio), 7227 (Drosophila melanogaster) and 6239 (Caenorhabditis elegans).

#### Value

Data frame of reaction identifiers.

#### References

Wang H, Robinson JL, Kocabas P, Gustafsson J, Anton M, Cholley PE, Huang S, Gobom J, Svensson T, Uhlen M, Zetterberg H, Nielsen J. Genome-scale metabolic network reconstruction of model animals as a platform for translational research. Proc Natl Acad Sci U S A. 2021 Jul 27;118(30):e2102344118. doi: doi:10.1073/pnas.2102344118.

#### See Also

- chalmers\_gem\_network
- chalmers\_gem\_metabolites
- chalmers\_gem
- chalmers\_gem\_raw
- chalmers\_gem\_id\_mapping\_table
- cosmos\_pkn

## **Examples**

```
{\tt chalmers\_gem\_reactions()}
```

common\_name

Common (English) names of organisms

### **Description**

Common (English) names of organisms

#### Usage

```
common_name(name)
```

## **Arguments**

name

Vector with any kind of organism name or identifier, can be also mixed type.

## Value

Character vector with common (English) taxon names, NA if a name in the input could not be found.

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#### See Also

- ncbi\_taxid
- latin\_name
- ensembl\_name

#### **Examples**

```
common_name(c(10090, "cjacchus", "Vicugna pacos"))
# [1] "Mouse" "White-tufted-ear marmoset" "Alpaca"
```

complexes

Protein complexes from OmniPath

#### **Description**

A comprehensive dataset of protein complexes from the <a href="https://omnipathdb.org/complexes">https://omnipathdb.org/complexes</a> endpoint of the OmniPath web service.

## Usage

```
complexes(...)
```

## **Arguments**

... Arguments passed on to omnipath\_query

organism Character or integer: name or NCBI Taxonomy ID of the organism. OmniPath is built of human data, and the web service provides orthology translated interactions and enzyme-substrate relationships for mouse and rat. For other organisms and query types, orthology translation will be called automatically on the downloaded human data before returning the result.

resources Character vector: name of one or more resources. Restrict the data to these resources. For a complete list of available resources, call the '<query\_type>\_resources' functions for the query type of interst.

genesymbols Character or logical: TRUE or FALS or "yes" or "no". Include the 'genesymbols' column in the results. OmniPath uses UniProt IDs as the primary identifiers, gene symbols are optional.

fields Character vector: additional fields to include in the result. For a list of available fields, call 'query\_info("interactions")'.

default\_fields Logical: if TRUE, the default fields will be included.

silent Logical: if TRUE, no messages will be printed. By default a summary message is printed upon successful download.

logicals Character vector: fields to be cast to logical.

format Character: if "json", JSON will be retrieved and processed into a nested list; any other value will return data frame.

download\_args List: parameters to pass to the download function, which is 'readr::read\_tsv' by default, and 'jsonlite::safe\_load'.

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add\_counts Logical: if TRUE, the number of references and number of resources for each record will be added to the result.

- license Character: license restrictions. By default, data from resources allowing "academic" use is returned by OmniPath. If you use the data for work in a company, you can provide "commercial" or "for-profit", which will restrict the data to those records which are supported by resources that allow for-profit use.
- password Character: password for the OmniPath web service. You can provide a special password here which enables the use of 'license = "ignore" option, completely bypassing the license filter.
- exclude Character vector: resource or dataset names to be excluded. The data will be filtered after download to remove records of the excluded datasets and resources.
- strict\_evidences Logical: reconstruct the "sources" and "references" columns of interaction data frames based on the "evidences" column, strictly filtering them to the queried datasets and resources. Without this, the "sources" and "references" fields for each record might contain information for datasets and resources other than the queried ones, because the downloaded records are a result of a simple filtering of an already integrated data frame.
- genesymbol\_resource Character: "uniprot" (default) or "ensembl". The OmniPath web service uses the primary gene symbols as provided by UniProt. By passing "ensembl" here, the UniProt gene symbols will be replaced by the ones used in Ensembl. This translation results in a loss of a few records, and multiplication of another few records due to ambiguous translation.
- cache Logical: use caching, load data from and save to the. The cache directory by default belongs to the user, located in the user's default cache directory, and named "OmnipathR". Find out about it by getOption("omnipathr.cachedir"). Can be changed by omnipath\_set\_cachedir.

#### Value

A data frame of protein complexes.

#### See Also

- complex\_resources
- query\_info
- omnipath\_query

#### **Examples**

```
cplx <- complexes(resources = c("CORUM", "hu.MAP"))</pre>
```

complex\_genes

Get all the molecular complexes for a given gene(s)

### **Description**

This function returns all the molecular complexes where an input set of genes participate. User can choose to retrieve every complex where any of the input genes participate or just retrieve these complexes where all the genes in input set participate together.

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### Usage

```
complex_genes(complexes = complexes(), genes, all_genes = FALSE)
```

### **Arguments**

 ${\bf Complexes} \qquad \qquad {\bf Data\ frame\ of\ protein\ complexes\ (obtained\ using\ {\bf complexes})}.$ 

genes Character: search complexes where these genes present.

all\_genes Logical: select only complexes where all of the genes present together. By

default complexes where any of the genes can be found are returned.

#### Value

Data frame of complexes

#### See Also

complexes

## **Examples**

complex\_resources

Retrieve a list of complex resources available in Omnipath

## Description

Get the names of the resources from <a href="https://omnipathdb.org/complexes">https://omnipathdb.org/complexes</a>

## Usage

```
complex_resources(dataset = NULL)
```

## Arguments

dataset ignored for this query type

#### Value

character vector with the names of the databases

### See Also

- resources
- complexes

```
complex_resources()
```

consensuspathdb\_download

Retrieves the ConsensusPathDB network

#### **Description**

Compiles a table of binary interactions from ConsensusPathDB (http://cpdb.molgen.mpg.de/) and translates the UniProtKB ACs to Gene Symbols.

#### Usage

```
consensuspathdb_download(complex_max_size = 4, min_score = 0.9)
```

#### **Arguments**

```
complex_max_size
```

Numeric: do not expand complexes with a higher number of elements than this. ConsensusPathDB does not contain conventional interactions but lists of participants, which might be members of complexes. Some records include dozens of participants and expanding them to binary interactions result thousands, sometimes hundreds of thousands of interactions from one single record. At the end, this process consumes >10GB of memory and results rather unusable data, hence it is recommended to limit the complex sizes at some low number.

min\_score

Numeric: each record in ConsensusPathDB comes with a confidence score, expressing the amount of evidences. The default value, a minimum score of 0.9 retains approx. the top 30 percent of the interactions.

#### Value

Data frame (tibble) with interactions.

```
## Not run:
cpdb_data <- consensuspathdb_download(</pre>
    complex_max_size = 1,
    min_score = .99
)
nrow(cpdb_data)
# [1] 252302
colnames(cpdb_data)
                  "references" "uniprot_a"
# [1] "databases"
                                               "confidence"
                                                              "record_id"
# [6] "uniprot_b" "in_complex" "genesymbol_a" "genesymbol_b"
cpdb_data
# # A tibble: 252,302 x 9
     databases references uniprot_a confidence record_id uniprot_b in_com
                                        <dbl> <int> <chr>
     <chr>
            <chr> <chr>
                                                                  <lg1>
  1 Reactome NA
                         SUMF2_HU.
                                                      1 SUMF1_HU. TRUE
                                        1
# 2 Reactome NA
                         SUMF1_HU.
                                                      1 SUMF2_HU. TRUE
                                        1
# 3 DIP,Reac. 22210847,. STIM1_HU.
                                        0.998
                                                      2 TRPC1_HU. TRUE
# 4 DIP,Reac. 22210847,. TRPC1_HU.
                                        0.998
                                                      2 STIM1_HU. TRUE
# # . with 252,292 more rows, and 2 more variables: genesymbol_a <chr>,
      genesymbol_b <chr</pre>
```

```
## End(Not run)
```

consensuspathdb\_raw\_table

Downloads interaction data from ConsensusPathDB

## Description

Downloads interaction data from ConsensusPathDB

## Usage

```
consensuspathdb_raw_table()
```

## Value

Data frame (tibble) with interactions.

## **Examples**

```
cpdb_raw <- consensuspathdb_raw_table()</pre>
```

cookie

Acquire a cookie if necessary

## Description

Acquire a cookie if necessary

## Usage

```
cookie(
  url,
  init_url = NULL,
  post = NULL,
  payload = NULL,
  init_post = NULL,
  init_payload = NULL,
  curl_verbose = FALSE
)
```

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### **Arguments**

url	Character. URL to download to get the cookie.
init_url	Character. An initial URL to download to get the cookie, before downloading "url" with the cookie.
post	List: HTTP POST parameters.
payload	Data to send as payload.
init_post	List: HTTP POST parameters for "init_url".
init_payload	Data to send as payload with "init_url".
curl_verbose	Logical. Perform CURL requests in verbose mode for debugging purposes.

#### Value

A list with cache file path, cookies and response headers.

cosmos_pkn	Prior knowledge network (PKN) for COSMOS

## **Description**

The prior knowledge network (PKN) used by COSMOS is a network of heterogenous causal interactions: it contains protein-protein, reactant-enzyme and enzyme-product interactions. It is a combination of multiple resources:

- Genome-scale metabolic model (GEM) from Chalmers Sysbio (Wang et al., 2021.)
- Network of chemical-protein interactions from STITCH (https://stitch.embl.de/)
- Protein-protein interactions from Omnipath (Türei et al., 2021)

This function downloads, processes and combines the resources above. With all downloads and processing the build might take 30-40 minutes. Data is cached at various levels of processing, shortening processing times. With all data downloaded and HMDB ID translation data preprocessed, the build takes 3-4 minutes; the complete PKN is also saved in the cache, if this is available, loading it takes only a few seconds.

## Usage

```
cosmos_pkn(
  organism = "human",
  protein_ids = c("uniprot", "genesymbol"),
  metabolite_ids = c("hmdb", "kegg"),
  chalmers_gem_metab_max_degree = 400L,
  stitch_score = 700L,
  ...
)
```

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#### **Arguments**

organism

Character or integer: name or NCBI Taxonomy ID of an organism. Supported organisms vary by resource: the Chalmers GEM is available only for human, mouse, rat, fish, fly and worm. OmniPath can be translated by orthology, but for non-vertebrate or less researched taxa very few orthologues are available. STITCH is available for a large number of organisms, please refer to their web page: https://stitch.embl.de/.

protein\_ids

Character: translate the protein identifiers to these ID types. Each ID type results two extra columns in the output, for the "source" and "target" sides of the interaction, respectively. The default ID type for proteins depends on the resource, hence the "source" and "target" columns are heterogenous. By default UniProt IDs and Gene Symbols are included. The Gene Symbols used in the COSMOS PKN are provided by Ensembl, and do not completely agree with the ones provided by UniProt and used in OmniPath data by default.

metabolite\_ids Character: translate the metabolite identifiers to these ID types. Each ID type results two extra columns in the output, for the "source" and "target" sides of the interaction, respectively. The default ID type for metabolites depends on the resource, hence the "source" and "target" columns are heterogenous. By default HMDB IDs and KEGG IDs are included.

chalmers\_gem\_metab\_max\_degree

Numeric: remove metabolites from the Chalmers GEM network with defgrees larger than this. Useful to remove cofactors and over-promiscuous metabolites.

stitch\_score

Include interactions from STITCH with combined confidence score larger than

Further parameters to omnipath\_interactions.

## Value

A data frame of binary causal interations with effect signs, resource specific attributes and translated to the desired identifiers. The "record\_id" column identifies the original records within each resource. If one "record\_id" yields multiple records in the final data frame, it is the result of one-tomany ID translation or other processing steps. Before use, it is recommended to select one pair of ID type columns (by combining the preferred ones) and perform "distinct" by the identifier columns and sign.

#### References

Wang H, Robinson JL, Kocabas P, Gustafsson J, Anton M, Cholley PE, et al. Genome-scale metabolic network reconstruction of model animals as a platform for translational research. Proceedings of the National Academy of Sciences. 2021 Jul 27;118(30):e2102344118.

Türei D, Valdeolivas A, Gul L, Palacio-Escat N, Klein M, Ivanova O, et al. Integrated intra- and intercellular signaling knowledge for multicellular omics analysis. Molecular Systems Biology. 2021 Mar;17(3):e9923.

#### See Also

- chalmers\_gem\_network
- stitch\_network
- omnipath\_for\_cosmos
- omnipath-interactions

#### **Examples**

```
## Not run:
    human_cosmos <- cosmos_pkn(organism = "human")
## End(Not run)</pre>
```

curated\_ligand\_receptor\_interactions

Curated ligand-receptor interactions

### **Description**

The OmniPath *intercell* database annotates individual proteins and complexes, and we combine these annotations with network interactions on the client side, using <code>import\_intercell\_network</code>. The architecture of this database is complex, aiming to cover a broad range of knowledge on various levels of details and confidence. We can use the <code>intercell\_consensus\_filter</code> and <code>filter\_intercell\_network</code> functions for automated, data driven quality filtering, in order to enrich the cell-cell communication network in higher confidence interactions. However, for many users, a simple combination of the most established, expert curated ligand-receptor resources, provided by this function, fits better their purpose.

## Usage

```
curated_ligand_receptor_interactions(
  curated_resources = c("Guide2Pharma", "HPMR", "ICELLNET", "Kirouac2010", "CellTalkDB",
        "CellChatDB", "connectomeDB2020"),
  cellphonedb = TRUE,
  cellinker = TRUE,
  talklr = TRUE,
  signalink = TRUE,
  ...
)
```

## **Arguments**

curated\_resources

Character vector of the resource names which are considered to be expert curated. You can include any post-translational network resource here, but if you include non ligand-receptor or non curated resources, the result will not fulfill

the original intention of this function.

cellphonedb Logical: include the curated interactions from CellPhoneDB (not the whole

CellPhoneDB but a subset of it).

cellinker Logical: include the curated interactions from Cellinker (not the whole Cellinker

but a subset of it).

talklr Logical: include the curated interactions from talklr (not the whole talklr but a

subset of it).

signalink Logical: include the ligand-receptor interactions from SignaLink. These are all

expert curated.

... Passed to import\_post\_translational\_interactions: further parameters

for the interaction data. Should not contain 'resources' argument as that would

interfere with the downstream calls.

curated\_ligrec\_stats 35

#### **Details**

Some resources are a mixture of curated and bulk imported interactions, and sometimes it's not trivial to separate these, we take care of these here. This function does not use the *intercell* database of OmniPath, but retrieves and filters a handful of network resources. The returned data frame has the layout of *interactions* (network) data frames, and the *source* and *target* partners implicitly correspond to *ligand* and *receptor*. The data frame shows all resources and references for all interactions, but each interaction is supported by at least one ligand-receptor resource which is supposed to based on expert curation in a ligand-receptor context.

#### Value

A data frame similar to *interactions* (network) data frames, the *source* and *target* partners being ligand and receptor, respectively.

#### See Also

- import\_intercell\_network
- filter\_intercell\_network
- annotated\_network
- import\_post\_translational\_interactions
- import\_ligrecextra\_interactions
- curated\_ligrec\_stats

### **Examples**

```
lr <- curated_ligand_receptor_interactions()
lr</pre>
```

## **Description**

Statistics about literature curated ligand-receptor interactions

## Usage

```
curated_ligrec_stats(...)
```

## **Arguments**

... Passed to curated\_ligand\_receptor\_interactions, determines the set of all curated L-R interactions which will be compared against each of the individual resources.

36 database\_summary

#### **Details**

The data frame contains the total number of interactions, the number of interactions which overlap with the set of curated interactions (*curated\_overlap*), the number of interactions with literature references from the given resource (*literature*) and the number of interactions which are curated by the given resource (*curated\_self*). This latter we defined according to our best knowledge, in many cases it's not possible to distinguish curated interactions). All these numbers are also presented as a percent of the total. Importantly, here we consider interactions curated only if they've been curated in a cell-cell communication context.

#### Value

A data frame with estimated counts of curated ligand-receptor interactions for each L-R resource.

#### See Also

```
curated_ligand_receptor_interactions
```

#### **Examples**

```
clr <- curated_ligrec_stats()
clr</pre>
```

database\_summary

Summary of the annotations and intercell database contents

## Description

The 'annotations\_summary' and 'intercell\_summary' query types return detailed information on the contents of these databases. It includes all the available resources, fields and values in the database.

#### Usage

```
database_summary(query_type, return_df = FALSE)
```

### Arguments

```
query_type Character: either "annotations" or "intercell".
return_df Logical: return a data frame instead of list.
```

## Value

Summary of the database contents: the available resources, fields, and their possible values. As a nested list if format is "json", otherwise a data frame.

```
annotations_summary <- database_summary('annotations')</pre>
```

datasets\_one\_column 37

datasets_one_column	Create a column with dataset names listed
---------------------	---

### **Description**

From logical columns for each dataset, here we create a column that is a list of character vectors, containing dataset labels.

# Usage

```
datasets_one_column(data, remove_logicals = TRUE)
```

### **Arguments**

data Interactions data frame with dataset columns (i.e. queried with the option 'fields = "datasets"').
remove\_logicals

Logical: remove the per dataset logical columns.

### Value

The input data frame with the new column "datasets" added.

descendants All descendants in the ontology tree

# Description

Starting from the selected nodes, recursively walks the ontology tree until it reaches the leaf nodes. Collects all visited nodes, which are the descendants (children) of the starting nodes.

# Usage

### **Arguments**

terms	Character vector of ontology term IDs or names. A mixture of IDs and names can be provided.
db_key	Character: key to identify the ontology database. For the available keys see omnipath_show_db.
ids	Logical: whether to return IDs or term names.
relations	Character vector of ontology relation types. Only these relations will be used.

38 ensembl\_dataset

#### **Details**

Note: this function relies on the database manager, the first call might take long because of the database load process. Subsequent calls within a short period should be faster. See <a href="mailto:get\_ontology\_db">get\_ontology\_db</a>.

#### Value

Character vector of ontology IDs. If the input terms are all leaves NULL is returned. The starting nodes won't be included in the result unless some of them are descendants of other starting nodes.

# **Examples**

```
descendants('GO:0005035', ids = FALSE)
# [1] "tumor necrosis factor-activated receptor activity"
# [2] "TRAIL receptor activity"
# [3] "TNFSF11 receptor activity"
```

ensembl\_dataset

Ensembl dataset name from organism

# **Description**

Ensembl dataset name from organism

# Usage

```
ensembl_dataset(organism)
```

### **Arguments**

organism

Character or integer: an organism (taxon) name or identifier. If an Ensembl dataset name is provided

## Value

Character: name of an ensembl dataset.

```
ensembl_dataset(10090)
# [1] "mmusculus_gene_ensembl"
```

```
ensembl_id_mapping_table
```

Identifier translation table from Ensembl

### **Description**

Identifier translation table from Ensembl

### Usage

```
ensembl_id_mapping_table(to, from = "uniprot", organism = 9606)
```

### **Arguments**

to Character or symbol: target ID type. See Details for possible values.

from Character or symbol: source ID type. See Details for possible values.

organism Character or integer: NCBI Taxonomy ID or name of the organism (by default 9606 for human).

# Details

The arguments to and from can be provided either as character or as symbol (NSE). Their possible values are either Ensembl attribute names or synonyms listed at translate\_ids.

#### Value

A data frame (tibble) with columns 'From' and 'To'.

## See Also

- translate\_ids
- uniprot\_full\_id\_mapping\_table
- uniprot\_id\_mapping\_table
- hmdb\_id\_mapping\_table
- chalmers\_gem\_id\_mapping\_table

```
ensp_up <- ensembl_id_mapping_table("ensp")
ensp_up
# # A tibble: 119,129 × 2
# From To
# <chr> <chr> # 1 P03886 ENSP00000354687
# 2 P03891 ENSP00000355046
# 3 P00395 ENSP00000354499
# 4 P00403 ENSP00000354876
# 5 P03928 ENSP00000355265
# # . with 119,124 more rows
```

40 ensembl\_name

ensembl\_id\_type

Ensembl identifier type label

### **Description**

Ensembl identifier type label

#### Usage

```
ensembl_id_type(label)
```

### **Arguments**

label

Character: an ID type label, as shown in the table at translate\_ids

#### Value

Character: the Ensembl specific ID type label, or the input unchanged if it could not be translated (still might be a valid identifier name). These labels should be valid Ensembl attribute names, directly usable in Ensembl queries.

#### See Also

- uniprot\_id\_type
- uploadlists\_id\_type
- chalmers\_gem\_id\_type
- hmdb\_id\_type

# **Examples**

```
ensembl_id_type("uniprot")
# [1] "uniprotswissprot"
```

ensembl\_name

Ensembl identifiers of organisms

# Description

Ensembl identifiers of organisms

# Usage

```
ensembl_name(name)
```

### **Arguments**

name

Vector with any kind of organism name or identifier, can be also mixed type.

ensembl\_organisms 41

### Value

Character vector with Ensembl taxon names, NA if a name in the input could not be found.

### See Also

- ncbi\_taxid
- common\_name
- latin\_name

### **Examples**

```
ensembl_name(c(9606, "cat", "dog"))
# [1] "hsapiens" "fcatus" "clfamiliaris"
ensembl_name(c("human", "kitten", "cow"))
# [1] "hsapiens" NA "btaurus"
```

ensembl\_organisms

Organism names and identifiers from Ensembl

# Description

A table with various taxon names and identifiers: English common names, latin (scientific) names, Ensembl organism IDs and NCBI taxonomy IDs.

### Usage

```
ensembl_organisms()
```

#### Value

A data frame with the above mentioned columns.

```
ens_org <- ensembl_organisms()
ens_org</pre>
```

42 ensembl\_orthology

```
ensembl_organisms_raw Table of Ensembl organisms
```

# Description

A table with various taxon IDs and metadata about related Ensembl database contents, as shown at https://www.ensembl.org/info/about/species.html. The "Taxon ID" column contains the NCBI Taxonomy identifiers.

# Usage

```
ensembl_organisms_raw()
```

### Value

The table described above as a data frame.

### **Examples**

```
ens_org <- ensembl_organisms_raw()
ens_org</pre>
```

ensembl\_orthology

Orthologous gene pairs from Ensembl

### **Description**

Orthologous gene pairs from Ensembl

# Usage

```
ensembl_orthology(
  organism_a = 9606,
  organism_b = 10090,
  attrs_a = NULL,
  attrs_b = NULL,
  colrename = TRUE
)
```

# Arguments

organism_a	Character or integer: organism name or identifier for the left side organism. We query the Ensembl dataset of this organism and add the orthologues of the other organism to it. Ideally this is the organism you translate from.
organism_b	Character or integer: organism name or identifier for the right side organism. We add orthology information of this organism to the gene records of the left side organism.
attrs_a	Further attributes about organism_a genes. Will be simply added to the attributes list.

ensure\_igraph 43

attrs\_b Further attributes about organism\_b genes (orthologues). The available attributes

are: "associated\_gene\_name", "chromosome", "chrom\_start", "chrom\_end", "wga\_coverage",

 $"goc\_score", "perc\_id\_r1", "perc\_id", "subtype". Attributes included by default: "ensembl\_gene", "ensembl\_peptide", "canonical\_transcript\_protein", "or-perconduction", "or-perconductio$ 

thology\_confidence" and "orthology\_type".

colrename Logical: replace prefixes from organism\_b attribute column names, so the re-

turned table always have the same column names, no matter the organism. E.g. for mouse these columns all have the prefix "mmusculus\_homolog\_", which this

option changes to "b\_".

#### **Details**

Only the records with orthology information are returned. The order of columns is the following: defaults of organism\_a, extra attributes of organism\_b, defaults of organism\_b, extra attributes of organism\_b.

#### Value

A data frame of orthologous gene pairs with gene, transcript and peptide identifiers and confidence values.

#### **Examples**

```
## Not run:
sffish <- ensembl_orthology(</pre>
   organism_b = 'Siamese fighting fish',
   attrs_a = 'external_gene_name',
   attrs_b = 'associated_gene_name'
)
sffish
# # A tibble: 175,608 × 10
     \verb|ensembl_gene_id| ensembl_transcript_id| ensembl_peptide. | external_gene_n.
#
    <chr>
                    <chr>
                                           <chr>
                                                            <chr>
  1 ENSG00000277196 ENST00000621424
                                           ENSP00000481127 NA
# 2 ENSG00000277196 ENST00000615165
                                         ENSP00000482462 NA
# 3 ENSG00000278817 ENST00000613204
                                          ENSP00000482514 NA
# 4 ENSG00000274847 ENST00000400754
                                           ENSP00000478910 MAFIP
# 5 ENSG00000273748 ENST00000612919
                                           ENSP00000479921 NA
# # . with 175,603 more rows, and 6 more variables:
# # b_ensembl_peptide <chr>, b_ensembl_gene <chr>,
# #
     b_orthology_type <chr>, b_orthology_confidence <dbl>,
##
     b_canonical_transcript_protein <chr>, b_associated_gene_name <chr>
## End(Not run)
```

ensure\_igraph

Converts a network to igraph object unless it is already one

#### **Description**

Converts a network to igraph object unless it is already one

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### Usage

```
ensure_igraph(network)
```

### **Arguments**

network

Either an OmniPath interaction data frame, or an igraph graph object.

#### Value

An igraph graph object.

enzsub\_graph

Enzyme-substrate graph

# Description

Transforms the a data frame with enzyme-substrate relationships (obtained by enzyme\_substrate) to an igraph object.

# Usage

```
enzsub_graph(enzsub)
```

# Arguments

enzsub

Data frame created by enzyme\_substrate

## Value

An igraph directed graph object.

# See Also

- enzyme\_substrate
- giant\_component
- find\_all\_paths

```
\label{lem:control_enzy} $$\operatorname{enzyme\_substrate(resources = c('PhosphoSite', 'SIGNOR'))}$$ enzsub\_g <- enzsub\_graph(enzsub = enzsub)
```

enzsub\_resources 45

enzsub\_resources

Retrieves a list of enzyme-substrate resources available in OmniPath

# Description

Get the names of the enzyme-substrate relationship resources available in <a href="https://omnipathdb.org/enzsub">https://omnipathdb.org/enzsub</a>

# Usage

```
enzsub_resources(dataset = NULL)
```

# **Arguments**

dataset

ignored for this query type

### Value

character vector with the names of the enzyme-substrate resources

#### See Also

- resources
- enzyme\_substrate

### **Examples**

```
enzsub_resources()
```

enzyme\_substrate

Enzyme-substrate (PTM) relationships from OmniPath

# Description

Imports the enzyme-substrate (more exactly, enzyme-PTM) relationship database from <a href="https://omnipathdb.org/enzsub">https://omnipathdb.org/enzsub</a>. These are mostly kinase-substrate relationships, with some acetylation and other types of PTMs.

# Usage

```
enzyme_substrate(...)
```

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### **Arguments**

.. Arguments passed on to omnipath\_query

- organism Character or integer: name or NCBI Taxonomy ID of the organism. OmniPath is built of human data, and the web service provides orthology translated interactions and enzyme-substrate relationships for mouse and rat. For other organisms and query types, orthology translation will be called automatically on the downloaded human data before returning the result.
- resources Character vector: name of one or more resources. Restrict the data to these resources. For a complete list of available resources, call the '<query\_type>\_resources' functions for the query type of interst.
- genesymbols Character or logical: TRUE or FALS or "yes" or "no". Include the 'genesymbols' column in the results. OmniPath uses UniProt IDs as the primary identifiers, gene symbols are optional.
- fields Character vector: additional fields to include in the result. For a list of available fields, call 'query\_info("interactions")'.
- default\_fields Logical: if TRUE, the default fields will be included.
- silent Logical: if TRUE, no messages will be printed. By default a summary message is printed upon successful download.
- logicals Character vector: fields to be cast to logical.
- format Character: if "json", JSON will be retrieved and processed into a nested list; any other value will return data frame.
- download\_args List: parameters to pass to the download function, which is 'readr::read\_tsv' by default, and 'jsonlite::safe\_load'.
- add\_counts Logical: if TRUE, the number of references and number of resources for each record will be added to the result.
- license Character: license restrictions. By default, data from resources allowing "academic" use is returned by OmniPath. If you use the data for work in a company, you can provide "commercial" or "for-profit", which will restrict the data to those records which are supported by resources that allow for-profit use.
- password Character: password for the OmniPath web service. You can provide a special password here which enables the use of 'license = "ignore" option, completely bypassing the license filter.
- exclude Character vector: resource or dataset names to be excluded. The data will be filtered after download to remove records of the excluded datasets and resources.
- strict\_evidences Logical: reconstruct the "sources" and "references" columns of interaction data frames based on the "evidences" column, strictly filtering them to the queried datasets and resources. Without this, the "sources" and "references" fields for each record might contain information for datasets and resources other than the queried ones, because the downloaded records are a result of a simple filtering of an already integrated data frame.
- genesymbol\_resource Character: "uniprot" (default) or "ensembl". The OmniPath web service uses the primary gene symbols as provided by UniProt. By passing "ensembl" here, the UniProt gene symbols will be replaced by the ones used in Ensembl. This translation results in a loss of a few records, and multiplication of another few records due to ambiguous translation.
- cache Logical: use caching, load data from and save to the. The cache directory by default belongs to the user, located in the user's default cache directory,

evex\_download 47

and named "OmnipathR". Find out about it by getOption("omnipathr.cachedir"). Can be changed by omnipath\_set\_cachedir.

#### Value

A data frame of enzymes and their PTM substrates.

#### See Also

- enzsub\_resources
- omnipath\_interactions
- enzsub\_graph
- print\_interactions
- query\_info
- omnipath\_query

### **Examples**

```
enzsub <- enzyme_substrate(
    resources = c("PhosphoSite", "SIGNOR"),
    organism = 9606
)</pre>
```

evex\_download

Interactions from the EVEX database

# Description

Downloads interactions from EVEX, a versatile text mining resource (http://evexdb.org). Translates the Entrez Gene IDs to Gene Symbols and combines the interactions and references into a single data frame.

### Usage

```
evex_download(
    min_confidence = NULL,
    remove_negatives = TRUE,
    top_confidence = NULL
)
```

### **Arguments**

min\_confidence Numeric: a threshold for confidence scores. EVEX confidence scores span roughly from -3 to 3. By providing a numeric value in this range the lower confidence interactions can be removed. If NULL no filtering performed.

remove\_negatives

Logical: remove the records with the "negation" attribute set.

top\_confidence Confidence cutoff as quantile (a number between 0 and 1). If NULL no filtering performed.

48 evidences

#### Value

Data frame (tibble) with interactions.

### **Examples**

```
evex_interactions <- evex_download()</pre>
evex_interactions
# # A tibble: 368,297 x 13
  general_event_id source_entrezge. target_entrezge. confidence negation
              <dbl> <chr>
#
                                   <chr>
                                                        <dbl> <dbl>
# 1
                98 8651
                                  6774
                                                      -1.45
                                                                   0
# 2
                                  6774
                                                      -1.45
               100 8431
                                                                    0
# 3
               205 6261
                                  6263
                                                       0.370
                                                                    0
                                                      -1.09
# 4
               435 1044
                                  1045
                                                                    0
\# . with 368,287 more rows, and 8 more variables: speculation <dbl>,
  coarse_type <chr>, coarse_polarity <chr>, refined_type <chr>,
   refined_polarity <chr>, source_genesymbol <chr>,
   target_genesymbol <chr>, references <chr>
```

evidences

Show evidences for an interaction

### **Description**

Show evidences for an interaction

# Usage

```
evidences(
  partner_a,
  partner_b,
  interactions = NULL,
  directed = FALSE,
  open = TRUE,
  browser = NULL,
  max_pages = 25L
```

### **Arguments**

partner_a	Identifier or name of one interacting partner. The order of the partners matter only if 'directed' is 'TRUE'. For both partners, vectors of more than one identifiers can be passed.
partner_b	Identifier or name of the other interacting partner.
interactions	An interaction data frame. If not provided, all interactions will be loaded within this function, but that takes noticeable time. If a 'list' is provided, it will be used as parameters for omnipath_interactions. This way you can define the organism, datasets or the interaction type.
directed	Logical: does the direction matter? If 'TRUE', only a $\rightarrow$ b interactions will be shown.
open	Logical: open online articles in a web browser.

extra\_attrs 49

browser Character: override the web browser executable used to open online articles.

max\_pages Numeric: largest number of pages to open. This is to prevent opening hundreds

or thousands of pages at once.

#### **Details**

If the number of references is larger than 'max\_pages', the most recent ones will be opened. URLs are passed to the browser in order of decreasing publication date, though browsers do not seem to respect the order at all. In addition Firefox, if it's not open already, tends to randomly open empty tab for the first or last URL, have no idea what to do about it.

#### Value

Nothing.

### **Examples**

```
## Not run:
evidences('CALM1', 'TRPC1', list(datasets = 'omnipath'))
## End(Not run)
```

extra\_attrs

Extra attribute names in an interaction data frame

#### **Description**

Interaction data frames might have an 'extra\_attrs' column if this field has been requested in the query by passing the 'fields = 'extra\_attrs' argument. This column contains resource specific attributes for the interactions. The names of the attributes consist of the name of the resource and the name of the attribute, separated by an underscore. This function returns the names of the extra attributes available in the provided data frame.

#### Usage

```
extra_attrs(data)
```

# Arguments

data

An interaction data frame, as provided by any of the omnipath-interactions functions.

#### Value

Character: the names of the extra attributes in the data frame.

### See Also

- extra\_attrs\_to\_cols
- has\_extra\_attrs
- with\_extra\_attrs
- filter\_extra\_attrs
- extra\_attr\_values

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#### **Examples**

```
i <- omnipath(fields = "extra_attrs")</pre>
extra_attrs(i)
```

New columns from extra attributes extra\_attrs\_to\_cols

### **Description**

New columns from extra attributes

# Usage

```
extra_attrs_to_cols(data, ..., flatten = FALSE, keep_empty = TRUE)
```

### **Arguments**

data An interaction data frame. The names of the extra attributes; NSE is supported. Custom column names can be provided as argument names. flatten Logical: unnest the list column even if some records have multiple values for the attributes; these will yield multiple records in the resulted data frame. Logical: if 'flatten' is 'TRUE', shall we keep the records which do not have the keep\_empty

attribute?

#### Value

Data frame with the new column created; the new column is list type if one interaction might have multiple values of the attribute, or character type if

### See Also

- extra\_attrs
- has\_extra\_attrs
- with\_extra\_attrs
- filter\_extra\_attrs
- extra\_attr\_values

```
i <- omnipath(fields = "extra_attrs")</pre>
extra_attrs_to_cols(i, Cellinker_type, Macrophage_type)
extra_attrs_to_cols(
    i,
    Cellinker_type,
    Macrophage_type,
    flatten = TRUE,
    keep_empty = FALSE
)
```

extra\_attr\_values 51

extra\_attr\_values

Possible values of an extra attribute

### **Description**

Extracts all unique values of an extra attribute occuring in this data frame.

# Usage

```
extra_attr_values(data, key)
```

# Arguments

data An interaction data frame with *extra\_attrs* column.

key The name of an extra attribute.

#### **Details**

Note, at the end we unlist the result, which means it works well for attributes which are atomic vectors but gives not so useful result if the attribute values are more complex objects. As the time of writing this, no such complex extra attribute exist in OmniPath.

# Value

A vector, most likely character, with the unique values of the extra attribute occuring in the data frame.

### See Also

- extra\_attrs\_to\_cols
- has\_extra\_attrs
- with\_extra\_attrs
- filter\_extra\_attrs
- extra\_attrs

```
op <- omnipath(fields = "extra_attrs")
extra_attr_values(op, SIGNOR_mechanism)</pre>
```

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filter\_by\_resource

Filters OmniPath data by resources

#### **Description**

Keeps only those records which are supported by any of the resources of interest.

### Usage

```
filter_by_resource(data, resources = NULL)
```

### Arguments

data A data frame downloaded from the OmniPath web service (interactions, enzyme-

substrate or complexes).

resources Character vector with resource names to keep.

#### Value

The data frame filtered.

### **Examples**

```
interactions <- omnipath()
signor <- filter_by_resource(interactions, resources = "SIGNOR")</pre>
```

filter\_evidences

Filter evidences by dataset, resource and license

### **Description**

Filter evidences by dataset, resource and license

### Usage

```
filter_evidences(data, ..., datasets = NULL, resources = NULL, exclude = NULL)
```

# Arguments

data	An interaction data frame with some columns containing evidences as nested
------	--

lists.

... The evidences columns to filter: tidyselect syntax is supported. By default the

columns "evidences", "positive", "negative", "directed" and "undirected" are fil-

tered, if present.

datasets A character vector of dataset names.

resources A character vector of resource names.

exclude Character vector of resource names to be excluded.

filter\_extra\_attrs 53

#### Value

The input data frame with the evidences in the selected columns filtered.

#### See Also

- only\_from
- unnest\_evidences
- from\_evidences

filter\_extra\_attrs

Filter interactions by extra attribute values

### **Description**

Filter interactions by extra attribute values

#### Usage

```
filter_extra_attrs(data, ..., na_ok = TRUE)
```

#### **Arguments**

data

An interaction data frame with extra\_attrs column.

. . .

Extra attribute names and values. The contents of the extra attribute *name* for each record will be checked against the values provided. The check by default is a set intersection: if any element is common between the user provided values and the values of the extra attribute for the record, the record will be matched. Alternatively, any value can be a custom function which accepts the value of the extra attribute and returns a single logical value. Finally, if the extra attribute name starts with a dot, the result of the check will be negated.

na\_ok

Logical: keep the records which do not have the extra attribute. Typically these are the records which are not from the resource providing the extra attribute.

#### Value

The input data frame with records removed according to the filtering criteria.

### See Also

- extra\_attrs
- has\_extra\_attrs
- extra\_attrs\_to\_cols
- with\_extra\_attrs
- extra\_attr\_values

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#### **Examples**

```
cl <- post_translational(
    resources = "Cellinker",
    fields = "extra_attrs"
)
# Only cell adhesion interactions from Cellinker
filter_extra_attrs(cl, Cellinker_type = "Cell adhesion")

op <- omnipath(fields = "extra_attrs")
# Any mechanism except phosphorylation
filter_extra_attrs(op, .SIGNOR_mechanism = "phosphorylation")</pre>
```

filter\_intercell

Filter intercell annotations

### **Description**

Filters a data frame retrieved by intercell.

### Usage

```
filter_intercell(
 data,
  categories = NULL,
 resources = NULL,
 parent = NULL,
  scope = NULL,
 aspect = NULL,
 source = NULL,
  transmitter = NULL,
 receiver = NULL,
  secreted = NULL,
 plasma_membrane_peripheral = NULL,
 plasma_membrane_transmembrane = NULL,
 proteins = NULL,
 causality = NULL,
  topology = NULL,
)
```

### **Arguments**

An intercell annotation data frame as provided by intercell.

Character: allow only these values in the category column.

Character: allow records only from these resources.

Character: filter for records with these parent categories.

Character: filter for records with these annotation scopes. Possible values are

generic and specific.

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aspect Character: filter for records with these annotation aspects. Possible values are functional and locational. source Character: filter for records with these annotation sources. Possible values are composite and resource\_specific. Logical: if TRUE only transmitters, if FALSE only non-transmitters will be setransmitter lected, if NULL it has no effect. receiver Logical: works the same way as transmitters. secreted Logical: works the same way as transmitters. plasma\_membrane\_peripheral Logical: works the same way as transmitters. plasma\_membrane\_transmembrane Logical: works the same way as transmitters. proteins Character: filter for annotations of these proteins. Gene symbols or UniProt IDs can be used. Character: filter for records with these causal roles. Possible values are transmitter causality and receiver. The filter applied simultaneously to the transmitter and receiver arguments, it's just a different notation for the same thing. Character: filter for records with these localization topologies. Possible values topology are secreced, plasma\_membrane\_peripheral and plasma\_membrane\_transmembrane; the shorter notations sec, pmp and pmtm can be used. Has the same effect as the logical type arguments, just uses a different notation. Ignored.

#### Value

The intercell annotation data frame filtered according to the specified conditions.

#### See Also

- intercell
- intercell\_categories
- intercell\_generic\_categories
- intercell\_summary
- intercell\_network

```
ic <- intercell()
ic <- filter_intercell(
    ic,
    transmitter = TRUE,
    secreted = TRUE,
    scope = "specific"
)</pre>
```

```
filter_intercell_network
```

Quality filter an intercell network

### **Description**

The intercell database of OmniPath covers a very broad range of possible ways of cell to cell communication, and the pieces of information, such as localization, topology, function and interaction, are combined from many, often independent sources. This unavoidably result some weird and unexpected combinations which are false positives in the context of intercellular communication. <a href="intercell\_network">intercell\_network</a> provides a shortcut (high\_confidence) to do basic quality filtering. For custom filtering or experimentation with the parameters we offer this function.

#### Usage

```
filter_intercell_network(
 network,
  transmitter_topology = c("secreted", "plasma_membrane_transmembrane",
    "plasma_membrane_peripheral"),
  receiver_topology = "plasma_membrane_transmembrane",
 min_curation_effort = 2,
 min_resources = 1,
 min_references = 0,
 min_provenances = 1,
  consensus_percentile = 50,
  loc_consensus_percentile = 30,
  ligand_receptor = FALSE,
  simplify = FALSE,
  unique_pairs = FALSE,
  omnipath = TRUE,
  ligrecextra = TRUE,
 kinaseextra = FALSE,
 pathwayextra = FALSE,
)
```

### **Arguments**

network An intercell network data frame, as provided by intercell\_network, without simplify.

transmitter\_topology

Character vector: topologies allowed for the entities in transmitter role. Abbreviations allowed: "sec", "pmtm" and "pmp".

receiver\_topology

Same as transmitter\_topology for the entities in the receiver role.

min\_curation\_effort

Numeric: a minimum value of curation effort (resource-reference pairs) for network interactions. Use zero to disable filtering.

min\_resources Numeric: minimum number of resources for interactions. The value 1 means no filtering.

min\_references Numeric: minimum number of references for interactions. Use zero to disable filtering.

min\_provenances

Numeric: minimum number of provenances (either resources or references) for interactions. Use zero or one to disable filtering.

# consensus\_percentile

Numeric: percentile threshold for the consensus score of generic categories in intercell annotations. The consensus score is the number of resources supporting the classification of an entity into a category based on combined information of many resources. Here you can apply a cut-off, keeping only the annotations supported by a higher number of resources than a certain percentile of each category. If NULL no filtering will be performed. The value is either in the 0-1 range, or will be divided by 100 if greater than 1. The percentiles will be calculated against the generic composite categories and then will be applied to their resource specific annotations and specific child categories.

#### loc\_consensus\_percentile

Numeric: similar to consensus\_percentile for major localizations. For example, with a value of 50, the secreted, plasma membrane transmembrane or peripheral attributes will be TRUE only where at least 50 percent of the resources support these.

ligand\_receptor

Logical. If TRUE, only *ligand* and *receptor* annotations will be used instead of the more generic *transmitter* and *receiver* categories.

simplify

Logical: keep only the most often used columns. This function combines a network data frame with two copies of the intercell annotation data frames, all of them already having quite some columns. With this option we keep only the names of the interacting pair, their intercellular communication roles, and the minimal information of the origin of both the interaction and the annotations.

unique\_pairs

Logical: instead of having separate rows for each pair of annotations, drop the annotations and reduce the data frame to unique interacting pairs. See unique\_intercell\_network for details.

omnipath ligrecextra kinaseextra pathwayextra Logical: shortcut to include the *omnipath* dataset in the interactions query.

Logical: shortcut to include the *ligrecextra* dataset in the interactions query.

Logical: shortcut to include the *kinaseextra* dataset in the interactions query.

Logical: shortcut to include the *pathwayextra* dataset in the interactions query.

If simplify or unique\_pairs is TRUE, additional column names can be passed

here to dplyr::select on the final data frame. Otherwise ignored.

#### Value

. . .

An intercell network data frame filtered.

#### See Also

- intercell\_network
- unique\_intercell\_network
- simplify\_intercell\_network
- intercell
- intercell\_categories
- intercell\_generic\_categories
- intercell\_summary

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### **Examples**

```
icn <- intercell_network()
icn_f <- filter_intercell_network(
    icn,
    consensus_percentile = 75,
    min_provenances = 3,
    simplify = TRUE
)</pre>
```

find\_all\_paths

All paths between two groups of vertices

### **Description**

Finds all paths up to length 'maxlen' between specified groups of vertices. This function is needed only becaues igraph's 'all\_shortest\_paths' finds only the shortest, not any path up to a defined length.

# Usage

```
find_all_paths(
    graph,
    start,
    end,
    attr = NULL,
    mode = 'OUT',
    maxlen = 2,
    progress = TRUE
)
```

# **Arguments**

graph	An igraph graph object.
start	Integer or character vector with the indices or names of one or more start vertices.
end	Integer or character vector with the indices or names of one or more end vertices.
attr	Character: name of the vertex attribute to identify the vertices by. Necessary if 'start' and 'end' are not igraph vertex ids but for example vertex names or labels.
mode	Character: IN, OUT or ALL. Default is OUT.
maxlen	Integer: maximum length of paths in steps, i.e. if maxlen = 3, then the longest path may consist of 3 edges and 4 nodes.
progress	Logical: show a progress bar.

# Value

List of vertex paths, each path is a character or integer vector.

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#### See Also

- interaction\_graph
- enzsub\_graph
- giant\_component

## **Examples**

```
interactions <- import_omnipath_interactions()
graph <- interaction_graph(interactions)
paths <- find_all_paths(
   graph = graph,
   start = c('EGFR', 'STAT3'),
   end = c('AKT1', 'ULK1'),
   attr = 'name'
)</pre>
```

from\_evidences

Recreate interaction records from evidences columns

#### **Description**

Recreate interaction records from evidences columns

#### Usage

```
from_evidences(data, .keep = FALSE)
```

### **Arguments**

data An interaction data frame from the OmniPath web service with evidences col-

umn.

.keep Logical: keep the original "evidences" column when unnesting to separate columns

by direction.

#### Details

The OmniPath interaction data frames specify interactions primarily by three columns: "is\_directed", "is\_stimulation" and "is\_inhibition". Besides these, there are the "sources" and "references" columns that are always included in data frames created by OmnipathR and list the resources and literature references for each interaction, respectively. The optional "evidences" column is required to find out which of the resources and references support the direction or effect sign of the interaction. To properly recover information for arbitrary subsets of resources or datasets, the evidences can be filtered first, and then the standard data frame columns can be reconstructed from the selected evidences. This function is able to do the latter. It expects either an "evidences" column or evidences in their wide format 4 columns layout. It overwrites the standard columns of interaction records based on data extracted from the evidences, including the "curation\_effort" and "consensus..." columns.

**Note:** The "curation\_effort" might be calculated slightly differently from the version included in the OmniPath web service. Here we count the resources and the also add the number of references for each resource. E.g. a resource without any literatur reference counts as 1, while a resource with 3 references adds 4 to the value of the curation effort.

 $get\_db$ 

**Note:** If the "evidences" column has been already unnested to multiple columns ("positive", "negative", etc.) by unnest\_evidences, then these will be used; otherwise, the column will be unnested within this function.

**Note:** This function (or rather its wrapper, only\_from) is automatically applied if the 'strict\_evidences' argument is passed to any function querying interactions (see omnipath-interactions).

#### Value

A copy of the input data frame with all the standard columns describing the direction, effect, resources and references of the interactions recreated based on the contents of the nested list evidences column(s).

#### See Also

- filter\_evidences
- unnest\_evidences
- only\_from

### **Examples**

```
## Not run:
ci <- collectri(evidences = TRUE)
ci <- unnest_evidences(ci)
ci <- filter_evidences(datasets = 'collectri')
ci <- from_evidences(ci)
# the three lines above are equivalent to only_from(ci)
# and all the four lines above is equivalent to:
# collectri(strict_evidences = TRUE)
## End(Not run)</pre>
```

get\_db

Access a built in database

### **Description**

Databases are resources which might be costly to load but can be used many times by functions which usually automatically load and retrieve them from the database manager. Each database has a lifetime and will be unloaded automatically upon expiry.

# Usage

```
get_db(key, param = NULL, reload = FALSE, ...)
```

# **Arguments**

key

Character: the key of the database to load. For a list of available keys see omnipath\_show\_db.

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param List: override the defaults or pass further parameters to the database loader func-

tion. See the loader functions and their default parameters in omnipath\_show\_db. If the database is already loaded with different parameters it will be reloaded

with the new parameters only if the reload option is TRUE.

reload Reload the database if param passed here is different from the parameters used

the last time the database was loaded. If different functions with different parameters access the database repeatedly and request reload the frequent reloads

might cost substantial time and resource use.

... Arguments for the loader function of the database. These override the default

arguments.

#### Value

An object with the database contents. The exact format depends on the database, most often it is a data frame or a list.

#### See Also

```
omnipath_show_db.
```

### **Examples**

```
organisms <- get_db('organisms')</pre>
```

get\_ontology\_db

Access an ontology database

## Description

Retrieves an ontology database with relations in the desired data structure. The database is automatically loaded and the requested data structure is constructed if necessary. The databases stay loaded up to a certain time period (see the option omnipathr.db\_lifetime). Hence the first one of repeated calls to this function might take long and the subsequent ones should be really quick.

### Usage

```
get_ontology_db(key, rel_fmt = "tbl", child_parents = TRUE)
```

# Arguments

key Character: key of the ontology database. For the available keys see omnipath\_show\_db.

rel\_fmt Character: the data structure of the ontology relations. Posible values are 1)

"tbl" a data frame, 2) "lst" a list or 3) "gra" a graph.

or from parent to children (FALSE).

#### Value

A list with the following elements: 1) "names" a table with term IDs and names; 2) "namespaces" a table to connect term IDs and namespaces they belong to; 3) "relations" a table with relations between terms and their parent terms; 4) "subsets" a table with terms and the subsets they are part of; 5) "obsolete" character vector with all the terms labeled as obsolete.

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#### See Also

- omnipath\_show\_db
- get\_db

### **Examples**

```
go <- get_ontology_db('go_basic', child_parents = FALSE)</pre>
```

giant\_component

Giant component of a graph

### **Description**

For an igraph graph object returns its giant component.

#### Usage

```
giant_component(graph)
```

# Arguments

graph

An igraph graph object.

#### Value

An igraph graph object containing only the giant component.

## **Examples**

```
interactions <- import_post_translational_interactions()
graph <- interaction_graph(interactions)
graph_gc <- giant_component(graph)</pre>
```

go\_annot\_download

Gene annotations from Gene Ontology

# Description

Gene Ontology is an ontology of gene subcellular localizations, molecular functions and involvement in biological processes. Gene products across many organisms are annotated with the ontology terms. This function downloads the gene-ontology term associations for certain model organisms or all organisms. For a description of the columns see <a href="http://geneontology.org/docs/go-annotation-file-gaf-format-2.2/">http://geneontology.org/docs/go-annotation-file-gaf-format-2.2/</a>.

#### Usage

```
go_annot_download(organism = "human", aspects = c("C", "F", "P"), slim = NULL)
```

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#### **Arguments**

organism	Character: either "chicken", "cow", "dog", "human", "pig" or "uniprot_all".
aspects	Character vector with some of the following elements: "C" (cellular component), "F" (molecular function) and "P" (biological process). Gene Ontology is three separate ontologies called as three aspects. By this parameter you can control which aspects to include in the output.
slim	Character: if not NULL, the name of a GOsubset (slim). instead of the full GO annotation, the slim annotation will be returned. See details at go_annot_slim. If TRUE, the "generic" slim will be used.

#### Value

A tibble (data frame) of annotations as it is provided by the database

### **Examples**

```
goa_data <- go_annot_download()</pre>
goa_data
# # A tibble: 606,840 x 17
             db_object_id db_object_symbol qualifier go_id
#
    db
                                                          db_ref
#
    <fct>
             <chr> <chr> <chr>
                                                           <chr>
# 1 UniProt. A0A024RBG1 NUDT4B
                                         NA
                                                   GO:000. GO_REF:00.
# 2 UniProt. A0A024RBG1 NUDT4B
                                         NA
                                                   GO:000. GO_REF:00.
# 3 UniProt. A0A024RBG1 NUDT4B
                                         NA
                                                   GO:004. GO_REF:00.
# 4 UniProt. A0A024RBG1
                        NUDT4B
                                         NA
                                                   GO:005. GO_REF:00.
# 5 UniProt. A0A024RBG1
                        NUDT4B
                                                   GO:005. GO_REF:00.
# # . with 606,830 more rows, and 11 more variables:
     evidence_code <fct>, with_or_from <chr>, aspect <fct>,
# #
     db_object_name <chr>, db_object_synonym <chr>,
     db_object_type <fct>, taxon <fct>, date <date>,
# #
     assigned_by <fct>, annotation_extension <chr>,
# #
     gene_product_from_id <chr>
```

go\_annot\_slim

GO slim gene annotations

#### **Description**

GO slims are subsets of the full GO which "give a broad overview of the ontology content without the detail of the specific fine grained terms". In order to annotate genes with GO slim terms, we take the annotations and search all ancestors of the terms up to the root of the ontology tree. From the ancestors we select the terms which are part of the slim subset.

#### Usage

```
go_annot_slim(
  organism = "human",
  slim = "generic",
  aspects = c("C", "F", "P"),
  cache = TRUE
)
```

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### **Arguments**

organism	Character: either "chicken", "cow", "dog", "human", "pig" or "uniprot_all".
slim	Character: the GO subset (GO slim) name. Available GO slims are: "agr" (Alliance for Genomics Resources), "generic", "aspergillus", "candida", "drosophila". "chembl", "metagenomic", "mouse", "plant", "pir" (Protein Information Resource), "pombe" and "yeast".
aspects	Character vector with some of the following elements: "C" (cellular component), "F" (molecular function) and "P" (biological process). Gene Ontology is three separate ontologies called as three aspects. By this parameter you can control which aspects to include in the output.
cache	Logical: Load the result from cache if available.

#### **Details**

Building the GO slim is resource intensive in its current implementation. For human annotation and generic GO slim it might take around 20 minutes. The result is saved into the cache so next time loading the data from there is really quick. If the cache option is FALSE the data will be built fresh (the annotation and ontology files still might come from cache), and the newly build GO slim will overwrite the cache instance.

#### Value

A tibble (data frame) of genes annotated with ontology terms in in the GO slim (subset).

### See Also

- go\_annot\_download
- go\_ontology\_download
- get\_db

```
## Not run:
goslim <- go_annot_slim(organism = 'human', slim = 'generic')</pre>
goslim
# # A tibble: 276,371 x 8
         db_object_id db_object_symbol go_id aspect db_object_name
    db
    <fct> <chr>
                      <chr>
                                        <chr> <fct>
                                                     <chr>
  1 UniPr. A0A024RBG1 NUDT4B
                                        GO:0. F
                                                     Diphosphoinosito.
  2 UniPr. A0A024RBG1 NUDT4B
                                        GO:0. F
                                                     Diphosphoinosito.
  3 UniPr. A0A024RBG1 NUDT4B
                                        GO:0. C
                                                     Diphosphoinosito.
  4 UniPr. A0A024RBG1 NUDT4B
                                        GO:0. C
                                                     Diphosphoinosito.
  5 UniPr. A0A024RBG1 NUDT4B
                                                     Diphosphoinosito.
                                        GO:0. C
\# # . with 276,366 more rows, and 2 more variables:
     db_object_synonym <chr>, db_object_type <fct>
## End(Not run)
```

go\_ontology\_download The Gene Ontology tree

### **Description**

The Gene Ontology tree

### Usage

```
go_ontology_download(
  basic = TRUE,
  tables = TRUE,
  subset = NULL,
 relations = c("is_a", "part_of", "occurs_in", "regulates", "positively_regulates",
    "negatively_regulates")
)
```

#### **Arguments**

basic Logical: use the basic or the full version of GO. As written on the GO home

> page: "the basic version of the GO is filtered such that the graph is guaranteed to be acyclic and annotations can be propagated up the graph. The relations included are is a, part of, regulates, negatively regulates and positively regulates. This version excludes relationships that cross the 3 GO hierarchies. This version

should be used with most GO-based annotation tools."

tables In the result return data frames or nested lists. These later can be converted to

each other if necessary. However converting from table to list is faster.

Character: the GO subset (GO slim) name. GO slims are subsets of the full subset

GO which "give a broad overview of the ontology content without the detail of the specific fine grained terms". This option, if not NULL, overrides the basic parameter. Available GO slims are: "agr" (Alliance for Genomics Resources), "generic", "aspergillus", "candida", "drosophila", "chembl", "metagenomic", "mouse", "plant", "pir" (Protein Information Resource), "pombe" and

"yeast".

relations Character vector: the relations to include in the processed data.

### Value

A list with the following elements: 1) "names" a list with terms as names and names as values; 2) "namespaces" a list with terms as names and namespaces as values; 3) "relations" a list with relations between terms: terms are keys, values are lists with relations as names and character vectors of related terms as values; 4) "subsets" a list with terms as keys and character vectors of subset names as values (or NULL if the term does not belong to any subset); 5) "obsolete" character vector with all the terms labeled as obsolete. If the tables parameter is TRUE, "names", "namespaces", "relations" and "subsets" will be data frames (tibbles).

```
# retrieve the generic GO slim, a small subset of the full ontology
go <- go_ontology_download(subset = 'generic')</pre>
```

graph\_interaction

Interaction data frame from igraph graph object

### **Description**

Convert an igraph graph object to interaction data frame. This is the reverse of the operation done by thje interaction\_graph function. Networks can be easily converted to igraph objects, then you can make use of all igaph methods, and at the end, get back the interactions in a data frame, along with all new edge and node attributes.

## Usage

```
graph_interaction(graph, implode = FALSE)
```

### **Arguments**

graph An igraph graph object created formerly from an OmniPath interactions data

frame.

implode Logical: restore the original state of the list type columns by imploding them to

character vectors, subitems separated by semicolons.

#### Value

An interaction data frame.

### See Also

interaction\_graph

 $\verb"guide2pharma_download" \textit{Downloads interactions from the Guide to Pharmacology database}$ 

#### **Description**

Downloads ligand-receptor interactions from the Guide to Pharmacology (IUPHAR/BPS) database (https://www.guidetopharmacology.org/).

### Usage

guide2pharma\_download()

#### Value

A tibble (data frame) of interactions as it is provided by the database

#### **Examples**

```
g2p_data <- guide2pharma_download()</pre>
g2p_data
# # A tibble: 21,586 x 38
    target target_id target_gene_sym. target_uniprot target_ensembl_.
#
           <dbl> <chr> <chr> <chr>
    <chr>
# 1 12S-L.
               1387 ALOX12
                                    P18054
                                                   ENSG00000108839
# 2 15-L0.
              1388 ALOX15
                                   P16050
                                                  ENSG00000161905
  3 15-L0.
               1388 ALOX15
                                    P16050
                                                   ENSG00000161905
# 4 15-L0.
               1388 ALOX15
                                    P16050
                                                   ENSG00000161905
# # . with 21,576 more rows, and 33 more variables: target_ligand <chr>,
# # target_ligand_id <chr>, target_ligand_gene_symbol <chr>,
# ... (truncated)
```

harmonizome\_download Downloads a Harmonizome network dataset

### **Description**

Downloads a single network dataset from Harmonizome https://maayanlab.cloud/Harmonizome.

### Usage

harmonizome\_download(dataset)

## Arguments

dataset

The dataset part of the URL. Please refer to the download section of the Harmonizome webpage.

### Value

Data frame (tibble) with interactions.

```
harmonizome_data <- harmonizome_download('phosphositeplus')</pre>
harmonizome_data
# # A tibble: 6,013 x 7
#
    source source_id target target_desc target_id weight
            <chr> <dbl> <chr> <dr>
                                                   <dbl> <dbl>
    <chr>
# 1 TP53
                           7157 STK17A na
                                                      9263
            na
                                                               1
# 2 TP53
                           7157 TP53RK na
                                                    112858
                                                               1
# 3 TP53
            na
                           7157 SMG1 na
                                                     23049
                                                               1
# 4 UPF1
            na
                            5976 SMG1 na
                                                     23049
                                                               1
\# # . with 6,003 more rows
```

has\_extra\_attrs

Tells if an interaction data frame has an extra\_attrs column

# Description

Tells if an interaction data frame has an extra\_attrs column

### Usage

```
has_extra_attrs(data)
```

### **Arguments**

data

An interaction data frame.

#### Value

Logical: TRUE if the data frame has the "extra\_attrs" column.

#### See Also

- extra\_attrs
- extra\_attrs\_to\_cols
- with\_extra\_attrs
- filter\_extra\_attrs
- extra\_attr\_values

### **Examples**

```
i <- omnipath(fields = "extra_attrs")
has_extra_attrs(i)</pre>
```

### **Description**

Identifier translation table from HMDB

# Usage

```
hmdb_id_mapping_table(to, from, entity_type = "metabolite")
```

#### **Arguments**

to Character or symbol: target ID type. See Details for possible values. from Character or symbol: source ID type. See Details for possible values.

entity\_type Character: "gene" and "smol" are short symbols for proteins, genes and small

molecules respectively. Several other synonyms are also accepted.

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#### **Details**

The arguments to and from can be provided either as character or as symbol (NSE). Their possible values are either HMDB XML tag names or synonyms listed at id\_types.

#### Value

A data frame (tibble) with columns 'From' and 'To'.

#### See Also

- translate\_ids
- id\_types
- hmdb\_table
- uniprot\_full\_id\_mapping\_table
- uniprot\_id\_mapping\_table
- ensembl\_id\_mapping\_table
- chalmers\_gem\_id\_mapping\_table

## **Examples**

```
hmdb_kegg <- hmdb_id_mapping_table("kegg", "hmdb")
hmdb_kegg</pre>
```

hmdb\_id\_type

HMDB identifier type label

### **Description**

HMDB identifier type label

## Usage

```
hmdb_id_type(label)
```

#### **Arguments**

label

Character: an ID type label, as shown in the table at translate\_ids

## Value

Character: the HMDB specific ID type label, or the input unchanged if it could not be translated (still might be a valid identifier name). These labels should be valid HMDB field names, as used in HMDB XML files.

### See Also

- chalmers\_gem\_id\_type
- uniprot\_id\_type
- ensembl\_id\_type
- uploadlists\_id\_type

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### **Examples**

```
hmdb_id_type("hmdb")
# [1] "accession"
```

hmdb\_metabolite\_fields

Field names for the HMDB metabolite dataset

# Description

Field names for the HMDB metabolite dataset

# Usage

```
hmdb_metabolite_fields()
```

# Value

Character vector of field names.

# See Also

- hmdb\_table
- hmdb\_protein\_fields

# **Examples**

```
hmdb_metabolite_fields()
```

hmdb\_protein\_fields

Field names for the HMDB proteins dataset

# Description

Field names for the HMDB proteins dataset

# Usage

```
hmdb_protein_fields()
```

#### Value

Character vector of field names.

### See Also

- hmdb\_table
- hmdb\_metabolite\_fields

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### **Examples**

hmdb\_protein\_fields()

hmdb\_table

Download a HMDB XML file and process it into a table

### **Description**

Download a HMDB XML file and process it into a table

# Usage

```
hmdb_table(dataset = "metabolites", fields = NULL)
```

# **Arguments**

dataset

Character: name of an HMDB XML dataset, such as "metabolites", "proteins",

"urine", "serum", "csf", "saliva", "feces", "sweat".

fields

Character: fields to extract from the XML. This is a very minimal parser that is able to extract the text content of simple fields and multiple value fields which contain a list of leaves within one container tag under the record tag. A full list of fields available in HMDB is available by the hmdb\_protein\_fields and hmdb\_metabolite\_fields functions. By default, all fields available in the dataset are extracted.

#### Value

A data frame (tibble) with each column corresponding to a field.

### See Also

- hmdb\_protein\_fields
- hmdb\_metabolite\_fields

## **Examples**

hmdb\_table()

homologene\_download

Orthology table for a pair of organisms

### **Description**

Orthologous pairs of genes for a pair of organisms from NCBI HomoloGene, using one identifier type.

## Usage

```
homologene_download(
  target = 10090L,
  source = 9606L,
  id_type = "genesymbol",
  hgroup_size = FALSE
)
```

### **Arguments**

target Character or integer: name or ID of the target organism.

source Character or integer: name or ID of the source organism.

id\_type Symbol or character: identifier type, possible values are "genesymbol", "entrez", "refseqp" or "gi".

hgroup\_size Logical: include a column with the size of the homology groups. This column distinguishes one-to-one and one-to-many or many-to-many mappings.

# Details

The operation of this function is symmetric, \*source\* and \*target\* are interchangeable but determine the column layout of the output. The column "hgroup" is a numberic identifier of the homology groups. Most of the groups consist of one pair of orthologous genes (one-to-one mapping), and a few of them multiple ones (one-to-many or many-to-many mappings).

### Value

A data frame with orthologous identifiers between the two organisms.

### See Also

- homologene\_raw
- homologene\_uniprot\_orthology

homologene\_organisms 73

```
# 3 6 NP_000010.1 XP_508738.2

# 4 7 NP_001096.1 XP_001145316.1

# 5 9 NP_000014.1 XP_523792.2

# # . with 17,732 more rows
```

homologene\_organisms

Organisms in NCBI HomoloGene

### **Description**

Organisms in NCBI HomoloGene

# Usage

```
homologene_organisms(name_type = "ncbi")
```

## **Arguments**

name\_type

Character: type of the returned name or identifier. Many synonyms are accepted, the shortest ones: "latin", "ncbi", "common", "ensembl". Case unsensitive.

### **Details**

Not all NCBI Taxonomy IDs can be translated to common or latin names. It means some organisms will be missing if translated to those name types. In the future we will address this issue, until then if you want to see all organisms use NCBI Taxonomy IDs.

### Value

A character vector of organism names.

homologene\_raw

Orthology data from NCBI HomoloGene

## Description

Retrieves NCBI HomoloGene data without any processing. Processed tables are more useful for most purposes, see below other functions that provide those. Genes of various organisms are grouped into homology groups ("hgroup" column). Organisms are identified by NCBI Taxonomy IDs, genes are identified by four different identifier types.

## Usage

```
homologene_raw()
```

## Value

A data frame as provided by NCBI HomoloGene.

### See Also

• homologene\_download

## **Examples**

```
hg <- homologene_raw()</pre>
hg
# # A tibble: 275,237 × 6
#
    hgroup ncbi_taxid entrez genesymbol gi
                                             refseqp
    #
#
 1
# 2
# 4
# 5
# # . with 275,232 more rows
# which organisms are available?
common_name(unique(hg$ncbi_taxid))
# [1] "Human" "Chimpanzee" "Macaque" "Dog" "Cow" "Mouse" "Rat" "Zebrafish"
# [9] "D. melanogaster" "Caenorhabditis elegans (PRJNA13758)"
# [11] "Tropical clawed frog" "Chicken"
# ...and 9 more organisms with missing English names.
```

```
homologene_uniprot_orthology
```

Orthology table with UniProt IDs

# Description

Orthologous pairs of UniProt IDs for a pair of organisms, based on NCBI HomoloGene data.

## Usage

```
homologene_uniprot_orthology(target = 10090L, source = 9606L, by = entrez, ...)
```

### **Arguments**

target	Character or integer: name or ID of the target organism.
source	Character or integer: name or ID of the source organism.
by	Symbol or character: the identifier type in NCBI HomoloGene to use. Possible values are "refseqp", "entrez", "genesymbol", "gi".
	Further arguments passed to translate_ids.

## Value

A data frame with orthologous pairs of UniProt IDs.

hpo\_download 75

### **Examples**

```
homologene_uniprot_orthology(by = genesymbol)
# # A tibble: 14,235 × 2
# source target
# <chr> <chr>
# 1 P11310 P45952
# 2 P49748 P50544
# 3 P24752 Q8QZT1
# 4 Q04771 P37172
# 5 Q16586 P82350
# # . with 14,230 more rows
```

hpo\_download

Downloads protein annotations from Human Phenotype Ontology

## **Description**

Human Phenotype Ontology (HPO) provides a standardized vocabulary of phenotypic abnormalities encountered in human disease. Each term in the HPO describes a phenotypic abnormality. HPO currently contains over 13,000 terms and over 156,000 annotations to hereditary diseases. See more at https://hpo.jax.org/app/.

## Usage

```
hpo_download()
```

# Value

A tibble (data frame) of annotations as it is provided by the database

```
hpo_data <- hpo_download()</pre>
hpo_data
# # A tibble: 231,738 x 9
#
     entrez_gene_id entrez_gene_symb. hpo_term_id hpo_term_name
#
              <dbl> <chr>
                                      <chr>
                                                  <chr>
#
               8192 CLPP
                                      HP:0000013 Hypoplasia of the ute.
  1
#
  2
               8192 CLPP
                                      HP:0004322
                                                  Short stature
#
  3
               8192 CLPP
                                      HP:0000786 Primary amenorrhea
#
  4
               8192 CLPP
                                      HP:0000007
                                                  Autosomal recessive i.
#
  5
               8192 CLPP
                                      HP:0000815 Hypergonadotropic hyp.
# # . with 231,733 more rows, and 5 more variables:
# #
      frequency_raw <chr>, frequency_hpo <chr>, info_gd_source <chr>,
      gd_source <chr>, disease_id <chr>
```

htridb\_download

Downloads TF-target interactions from HTRIdb

## **Description**

HTRIdb (https://www.lbbc.ibb.unesp.br/htri/) is a database of literature curated human TF-target interactions. As the database is recently offline, the data is distributed by the OmniPath rescued data repository (https://rescued.omnipathdb.org/).

### Usage

htridb\_download()

### Value

Data frame (tibble) with interactions.

## **Examples**

```
htridb_data <- htridb_download()</pre>
htridb_data
# # A tibble: 18,630 x 7
       OID GENEID_TF SYMBOL_TF GENEID_TG SYMBOL_TG TECHNIQUE
#
     <dbl>
            <dbl> <chr> <dbl> <chr> <dbl> <chr>
                                    675 BRCA2
# 1 32399
               142 PARP1
                                                    Electrophoretic Mobi.
                                    675 BRCA2 Chromatin Immunoprec.
1543 CYP1A1 Chromatin Immunoprec.
               142 PARP1
# 2 32399
# 3 28907
                 196 AHR
                                    1543 C...
1543 CYP1A1
# 4 29466
                 196 AHR
                                                    Electrophoretic Mobi.
                 196 AHR
# 5 28911
                                     1543 CYP1A1
                                                    Chromatin Immunoprec.
# # . with 18,620 more rows, and 1 more variable: PUBMED_ID <chr>
```

id\_translation\_resources

List available ID translation resources

### **Description**

List available ID translation resources

## Usage

```
id_translation_resources()
```

# Value

A character vector with the names of the available ID translation resources.

```
id_translation_resources()
```

id\_types 77

id\_types

ID types and synonyms in identifier translation

## **Description**

ID types and synonyms in identifier translation

## Usage

```
id_types()
```

## Value

Data frame with 4 columns: the ID type labels in the resource, their synonyms in OmniPath (this package), the name of the ID translation resource, and the entity type.

## See Also

- translate\_ids
- translate\_ids\_multi
- ensembl\_id\_mapping\_table
- uniprot\_id\_mapping\_table
- hmdb\_id\_mapping\_table
- chalmers\_gem\_id\_mapping\_table
- uniprot\_full\_id\_mapping\_table
- ensembl\_id\_type
- uniprot\_id\_type
- hmdb\_id\_type
- chalmers\_gem\_id\_type

## **Examples**

```
id_types()
```

inbiomap\_download

Downloads and preprocesses network data from InWeb InBioMap

# Description

Downloads the data by inbiomap\_raw, extracts the UniProt IDs, Gene Symbols and scores and removes the irrelevant columns.

### Usage

```
inbiomap\_download(...)
```

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### **Arguments**

```
... Passed to inbiomap_raw.
```

## Value

A data frame (tibble) of interactions.

### See Also

```
inbiomap_raw
```

# **Examples**

```
## Not run:
inbiomap_interactions <- inbiomap_download()</pre>
inbiomap_interactions
## End(Not run)
# # A tibble: 625,641 x 7
    uniprot_a uniprot_b genesymbol_a genesymbol_b inferred score1 score2
#
    <chr>
             <chr>
                        <chr>
                                    <chr>
                                                 <lgl>
                                                           <dbl> <dbl>
#
  1 A0A5B9
              P01892
                        TRBC2
                                    HLA-A
                                                 FALSE
                                                           0.417 0.458
            Q96CV9
  2 A0AUZ9
                        KANSL1L
                                    OPTN
                                                 FALSE
                                                           0.155 0.0761
# 3 A0AV02
              P24941
                        SLC12A8
                                    CDK2
                                                 TRUE
                                                           0.156 0.0783
  4 A0AV02
              Q00526
                        SLC12A8
                                     CDK3
                                                 TRUE
                                                           0.157 0.0821
            P0CG48
# 5 A0AV96
                        RBM47
                                     UBC
                                                 FALSE
                                                           0.144 0.0494
# # . with 625,631 more rows
```

inbiomap\_raw

Downloads network data from InWeb InBioMap

## Description

Downloads the data from <a href="https://inbio-discover.com/map.html#downloads">httml#downloads</a> in tar.gz format, extracts the PSI MITAB table and returns it as a data frame.

# Usage

```
inbiomap_raw(curl_verbose = FALSE)
```

## **Arguments**

curl\_verbose Logical. Perform CURL requests in verbose mode for debugging purposes.

### Value

A data frame (tibble) with the extracted interaction table.

## See Also

```
inbiomap_download
```

interaction\_datasets 79

## **Examples**

```
## Not run:
inbiomap_psimitab <- inbiomap_raw()
## End(Not run)</pre>
```

# Description

Datasets in the OmniPath Interactions database

# Usage

```
interaction_datasets()
```

## Value

Character: labels of interaction datasets.

## **Examples**

```
interaction_datasets()
```

interaction\_graph

Build Omnipath interaction graph

# Description

Transforms the interactions data frame to an igraph graph object.

## Usage

```
interaction_graph(interactions = interactions)
```

# Arguments

interactions data.frame created by

- enzyme\_substrate
- omnipath-interactions

## Value

An igraph graph object.

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### See Also

- graph\_interaction
- import\_omnipath\_interactions
- import\_pathwayextra\_interactions
- import\_kinaseextra\_interactions
- import\_ligrecextra\_interactions
- import\_dorothea\_interactions
- import\_mirnatarget\_interactions
- import\_all\_interactions
- giant\_component
- find\_all\_paths

### **Examples**

```
interactions <- import_omnipath_interactions(resources = c('SignaLink3'))
g <- interaction_graph(interactions)</pre>
```

# Description

Names of the resources available in https://omnipathdb.org/interactions.

### Usage

```
interaction_resources(dataset = NULL)
```

## **Arguments**

dataset

a dataset within the interactions query type. Currently available datasets are 'omnipath', 'kinaseextra', 'pathwayextra', 'ligrecextra', 'collectri', 'dorothea', 'tf\_target', 'tf\_mirna', 'mirnatarget', 'lncrna\_mrna' and 'small\_molecule\_protein'.

### Value

Character: names of the interaction resources.

## See Also

- resources
- omnipath
- pathwayextra
- kinaseextra
- ligrecextra
- post\_translational

interaction\_types 81

- dorothea
- collectri
- tf\_target
- transcriptional
- mirna\_target
- tf\_mirna
- small\_molecule
- all\_interactions

## **Examples**

interaction\_resources()

interaction\_types

Interaction types in the OmniPath Interactions database

# Description

Interaction types in the OmniPath Interactions database

# Usage

```
interaction_types()
```

## Value

Character: labels of interaction types.

## **Examples**

interaction\_types()

intercell

Cell-cell communication roles from OmniPath

# Description

Roles of proteins in intercellular communication from the <a href="https://omnipathdb.org/intercell">https://omnipathdb.org/intercell</a> endpoint of the OmniPath web service. It provides information on the roles in inter-cellular signaling. E.g. if a protein is a ligand, a receptor, an extracellular matrix (ECM) component, etc.

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### Usage

```
intercell(
  categories = NULL,
  parent = NULL,
  scope = NULL,
  aspect = NULL,
  source = NULL,
  transmitter = NULL,
  receiver = NULL,
  secreted = NULL,
 plasma_membrane_peripheral = NULL,
 plasma_membrane_transmembrane = NULL,
 proteins = NULL,
  topology = NULL,
  causality = NULL,
  consensus_percentile = NULL,
 loc_consensus_percentile = NULL,
)
```

## **Arguments**

categories vector containing the categories to be retrieved. All the genes belonging to those

categories will be returned. For further information about the categories see

get\_intercell\_categories.

parent vector containing the parent classes to be retrieved. All the genes belonging to

those classes will be returned. For furter information about the main classes see

get\_intercell\_categories.

scope either 'specific' or 'generic'
aspect either 'locational' or 'functional'

source either 'resource\_specific' or 'composite'

transmitter logical, include only transmitters i.e. proteins delivering signal from a cell to its

environment.

receiver logical, include only receivers i.e. proteins delivering signal to the cell from its

environment.

secreted logical, include only secreted proteins

plasma\_membrane\_peripheral

logical, include only plasma membrane peripheral membrane proteins.

plasma\_membrane\_transmembrane

logical, include only plasma membrane transmembrane proteins.

proteins limit the query to certain proteins

topology topology categories: one or more of 'secreted' (sec), 'plasma\_membrane\_peripheral'

(pmp), 'plasma\_membrane\_transmembrane' (pmtm) (both short or long nota-

tion can be used).

causality 'transmitter' (trans), 'receiver' (rec) or 'both' (both short or long notation can

be used).

consensus\_percentile

Numeric: a percentile cut off for the consensus score of generic categories. The consensus score is the number of resources supporting the classification of an

entity into a category based on combined information of many resources. Here you can apply a cut-off, keeping only the annotations supported by a higher number of resources than a certain percentile of each category. If NULL no filtering will be performed. The value is either in the 0-1 range, or will be divided by 100 if greater than 1. The percentiles will be calculated against the generic composite categories and then will be applied to their resource specific annotations and specific child categories.

### loc\_consensus\_percentile

Numeric: similar to consensus\_percentile for major localizations. For example, with a value of 50, the secreted, plasma membrane transmembrane or peripheral attributes will be true only where at least 50 percent of the resources support these.

Arguments passed on to omnipath\_query

organism Character or integer: name or NCBI Taxonomy ID of the organism. OmniPath is built of human data, and the web service provides orthology translated interactions and enzyme-substrate relationships for mouse and rat. For other organisms and query types, orthology translation will be called automatically on the downloaded human data before returning the result.

resources Character vector: name of one or more resources. Restrict the data to these resources. For a complete list of available resources, call the '<query\_type>\_resources' functions for the query type of interst.

fields Character vector: additional fields to include in the result. For a list of available fields, call 'query\_info("interactions")'.

default\_fields Logical: if TRUE, the default fields will be included.

silent Logical: if TRUE, no messages will be printed. By default a summary message is printed upon successful download.

logicals Character vector: fields to be cast to logical.

format Character: if "json", JSON will be retrieved and processed into a nested list; any other value will return data frame.

download\_args List: parameters to pass to the download function, which is 'readr::read\_tsv' by default, and 'jsonlite::safe\_load'.

license Character: license restrictions. By default, data from resources allowing "academic" use is returned by OmniPath. If you use the data for work in a company, you can provide "commercial" or "for-profit", which will restrict the data to those records which are supported by resources that allow for-profit use.

password Character: password for the OmniPath web service. You can provide a special password here which enables the use of 'license = "ignore" option, completely bypassing the license filter.

exclude Character vector: resource or dataset names to be excluded. The data will be filtered after download to remove records of the excluded datasets and resources

json\_param List: parameters to pass to the 'jsonlite::fromJSON' when processing JSON columns embedded in the downloaded data. Such columns are "extra\_attrs" and "evidences". These are optional columns which provide a lot of extra details about interactions.

strict\_evidences Logical: reconstruct the "sources" and "references" columns of interaction data frames based on the "evidences" column, strictly filtering them to the queried datasets and resources. Without this, the "sources" and

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"references" fields for each record might contain information for datasets and resources other than the queried ones, because the downloaded records are a result of a simple filtering of an already integrated data frame.

genesymbol\_resource Character: "uniprot" (default) or "ensembl". The OmniPath web service uses the primary gene symbols as provided by UniProt. By passing "ensembl" here, the UniProt gene symbols will be replaced by the ones used in Ensembl. This translation results in a loss of a few records, and multiplication of another few records due to ambiguous translation.

cache Logical: use caching, load data from and save to the. The cache directory by default belongs to the user, located in the user's default cache directory, and named "OmnipathR". Find out about it by getOption("omnipathr.cachedir"). Can be changed by omnipath\_set\_cachedir.

### Value

A data frame of intercellular communication roles.

#### See Also

- intercell\_network
- intercell\_consensus\_filter
- filter\_intercell
- intercell\_categories
- intercell\_generic\_categories
- intercell\_resources
- intercell\_summary
- intercell\_network

### **Examples**

```
ecm_proteins <- intercell(categories = "ecm")</pre>
```

 $intercell\_categories \quad \textit{Categories in the intercell database of OmniPath}$ 

# Description

Retrieves a list of categories from https://omnipathdb.org/intercell.

## Usage

```
intercell_categories()
```

## Value

character vector with the different intercell categories

#### See Also

- intercell
- intercell\_generic\_categories
- intercell\_summary

### **Examples**

```
intercell_categories()
```

```
intercell_consensus_filter
```

Quality filter for intercell annotations

### **Description**

Quality filter for intercell annotations

### Usage

```
intercell_consensus_filter(
  data,
 percentile = NULL,
  loc_percentile = NULL,
  topology = NULL
)
```

# Arguments

data

A data frame with intercell annotations, as provided by intercell.

percentile

Numeric: a percentile cut off for the consensus score of composite categories. The consensus score is the number of resources supporting the classification of an entity into a category based on combined information of many resources. Here you can apply a cut-off, keeping only the annotations supported by a higher number of resources than a certain percentile of each category. If NULL no filtering will be performed. The value is either in the 0-1 range, or will be divided by 100 if greater than 1. The percentiles will be calculated against the generic composite categories and then will be applied to their resource specific annotations and specific child categories.

loc\_percentile Numeric: similar to percentile for major localizations. For example, with a value of 50, the secreted, plasma membrane transmembrane or peripheral attributes will be TRUE only where at least 50 percent of the resources support these.

topology

Character vector: list of allowed topologies, possible values are \*"secreted"\*, \*"plasma\_membrane\_peripheral"\* and \*"plasma\_membrane\_transmembrane"\*.

### Value

The data frame in data filtered by the consensus scores.

### See Also

- resources
- intercell
- filter\_intercell
- intercell\_categories
- intercell\_generic\_categories
- intercell\_resources
- intercell\_summary
- intercell\_network

## **Examples**

```
ligand_receptor <- intercell(parent = c("ligand", "receptor"))
nrow(ligand_receptor)
# [1] 50174
lr_q50 <- intercell_consensus_filter(ligand_receptor, 50)
nrow(lr_q50)
# [1] 42863</pre>
```

intercell\_generic\_categories

Retrieves a list of the generic categories in the intercell database of OmniPath

# Description

Retrieves a list of the generic categories from https://omnipathdb.org/intercell.

### Usage

```
intercell_generic_categories()
```

### Value

character vector with the different intercell main classes

### See Also

- intercell
- intercell\_categories
- intercell\_summary

```
intercell_generic_categories()
```

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intercell\_network

Intercellular communication network

### **Description**

Imports an intercellular network by combining intercellular annotations and protein interactions. First imports a network of protein-protein interactions. Then, it retrieves annotations about the proteins intercellular communication roles, once for the transmitter (delivering information from the expressing cell) and second, the receiver (receiving signal and relaying it towards the expressing cell) side. These 3 queries can be customized by providing parameters in lists which will be passed to the respective methods (omnipath\_interactions for the network and intercell for the annotations). Finally the 3 data frames combined in a way that the source proteins in each interaction annotated by the transmitter, and the target proteins by the receiver categories. If undirected interactions present (these are disabled by default) they will be duplicated, i.e. both partners can be both receiver and transmitter.

### Usage

```
intercell_network(
  interactions_param = list(),
  transmitter_param = list(),
  receiver_param = list(),
  resources = NULL,
  entity_types = NULL,
  ligand_receptor = FALSE,
 high_confidence = FALSE,
  simplify = FALSE,
  unique_pairs = FALSE,
  consensus_percentile = NULL,
  loc_consensus_percentile = NULL,
  omnipath = TRUE,
  ligrecextra = TRUE,
  kinaseextra = !high_confidence,
 pathwayextra = !high_confidence,
)
```

### **Arguments**

interactions\_param

a list with arguments for an interactions query;  $\mbox{omnipath-interactions}$ .

transmitter\_param

a list with arguments for intercell, to define the transmitter side of intercellular connections

receiver\_param a list with arguments for intercell, to define the receiver side of intercellular connections

resources

A character vector of resources to be applied to both the interactions and the annotations. For example, resources = 'CellChatDB' will download the transmitters and receivers defined by CellChatDB, connected by connections from CellChatDB.

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Character, possible values are "protein", "complex" or both. entity\_types ligand\_receptor

> Logical. If TRUE, only *ligand* and *receptor* annotations will be used instead of the more generic transmitter and receiver categories.

high\_confidence

Logical: shortcut to do some filtering in order to include only higher confidence interactions. The intercell database of OmniPath covers a very broad range of possible ways of cell to cell communication, and the pieces of information, such as localization, topology, function and interaction, are combined from many, often independent sources. This unavoidably result some weird and unexpected combinations which are false positives in the context of intercellular communication. This option sets some minimum criteria to remove most (but definitely not all!) of the wrong connections. These criteria are the followings: 1) the receiver must be plasma membrane transmembrane; 2) the curation effort for interactions must be larger than one; 3) the consensus score for annotations must be larger than the 50 percentile within the generic category (you can override this by consensus\_percentile). 4) the transmitter must be secreted or exposed on the plasma membrane. 5) The major localizations have to be supported by at least 30 percent of the relevant resources (you can override this by loc\_consensus\_percentile). 6) The datasets with lower level of curation (kinaseextra and pathwayextra) will be disabled. These criteria are of medium stringency, you can always tune them to be more relaxed or stringent by filtering manually, using filter\_intercell\_network.

simplify

Logical: keep only the most often used columns. This function combines a network data frame with two copies of the intercell annotation data frames, all of them already having quite some columns. With this option we keep only the names of the interacting pair, their intercellular communication roles, and the minimal information of the origin of both the interaction and the annotations.

unique\_pairs

Logical: instead of having separate rows for each pair of annotations, drop the annotations and reduce the data frame to unique interacting pairs. See unique\_intercell\_network for details.

consensus\_percentile

Numeric: a percentile cut off for the consensus score of generic categories in intercell annotations. The consensus score is the number of resources supporting the classification of an entity into a category based on combined information of many resources. Here you can apply a cut-off, keeping only the annotations supported by a higher number of resources than a certain percentile of each category. If NULL no filtering will be performed. The value is either in the 0-1 range, or will be divided by 100 if greater than 1. The percentiles will be calculated against the generic composite categories and then will be applied to their resource specific annotations and specific child categories.

loc\_consensus\_percentile

Numeric: similar to consensus\_percentile for major localizations. For example, with a value of 50, the secreted, plasma membrane transmembrane or peripheral attributes will be TRUE only where at least 50 percent of the resources support these.

omnipath Logical: shortcut to include the *omnipath* dataset in the interactions query. Logical: shortcut to include the *ligrecextra* dataset in the interactions query. Logical: shortcut to include the *kinaseextra* dataset in the interactions query. kinaseextra Logical: shortcut to include the *pathwayextra* dataset in the interactions query.

ligrecextra

pathwayextra

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If simplify or unique\_pairs is TRUE, additional column names can be passed here to dplyr::select on the final data frame. Otherwise ignored.

#### **Details**

By default this function creates almost the largest possible network of intercellular interactions. However, this might contain a large number of false positives. Please refer to the documentation of the arguments, especially high\_confidence, and the filter\_intercell\_network function. Note: if you restrict the query to certain intercell annotation resources or small categories, it's not recommended to use the consensus\_percentile or high\_confidence options, instead filter the network with filter\_intercell\_network for more consistent results.

#### Value

A dataframe containing information about protein-protein interactions and the inter-cellular roles of the protiens involved in those interactions.

#### See Also

- intercell
- intercell\_summary
- intercell\_categories
- intercell\_generic\_categories
- intercell
- omnipath
- pathwayextra
- kinaseextra
- ligrecextra
- unique\_intercell\_network
- simplify\_intercell\_network
- filter\_intercell\_network

```
intercell_network <- intercell_network(
   interactions_param = list(datasets = 'ligrecextra'),
   receiver_param = list(categories = c('receptor', 'transporter')),
   transmitter_param = list(categories = c('ligand', 'secreted_enzyme'))
)</pre>
```

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intercell\_resources

Retrieves a list of intercellular communication resources available in OmniPath

## **Description**

Retrieves a list of the databases from https://omnipathdb.org/intercell.

## Usage

```
intercell_resources(dataset = NULL)
```

# Arguments

dataset

ignored at this query type

#### Value

character vector with the names of the databases

### See Also

- resources
- intercell
- filter\_intercell
- intercell\_categories
- intercell\_generic\_categories
- intercell\_summary
- intercell\_network

## **Examples**

```
intercell_resources()
```

intercell\_summary

Full list of intercell categories and resources

# Description

Full list of intercell categories and resources

# Usage

```
intercell_summary()
```

# Value

A data frame of categories and resources.

is\_ontology\_id 91

### **Examples**

```
ic_cat <- intercell_categories()</pre>
ic_cat
# # A tibble: 1,125 x 3
    category
                             parent
                                                     database
#
     <chr>
                             <chr>
                                                     <chr>
# 1 transmembrane
                            transmembrane
                                                     UniProt_location
# 2 transmembrane
                            transmembrane
                                                     UniProt_topology
# 3 transmembrane
                                                     UniProt_keyword
                            transmembrane
# 4 transmembrane
                             transmembrane_predicted Phobius
# 5 transmembrane_phobius transmembrane_predicted Almen2009
\# \# . with 1,120 more rows
```

is\_ontology\_id

Looks like an ontology ID

## **Description**

Tells if the input has the typical format of ontology IDs, i.e. a code of capital letters, a colon, followed by a numeric code.

### Usage

```
is_ontology_id(terms)
```

## **Arguments**

terms

Character vector with strings to check.

### Value

A logical vector with the same length as the input.

## **Examples**

```
is_ontology_id(c('GO:0000001', 'reproduction'))
# [1] TRUE FALSE
```

is\_swissprot

Check for SwissProt IDs

# Description

Check for SwissProt IDs

### Usage

```
is_swissprot(uniprots, organism = 9606)
```

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## **Arguments**

uniprots Character vector of UniProt IDs.

organism Character or integer: name or identifier of the organism.

## Value

Logical vector TRUE for SwissProt IDs and FALSE for any other element.

# **Examples**

```
is_swissprot(c("Q05BL1", "A0A654IBU3", "P00533"))
# [1] FALSE FALSE TRUE
```

is\_trembl

Check for TrEMBL IDs

# Description

Check for TrEMBL IDs

# Usage

```
is_trembl(uniprots, organism = 9606)
```

# Arguments

uniprots Character vector of UniProt IDs.

organism Character or integer: name or identifier of the organism.

## Value

Logical vector TRUE for TrEMBL IDs and FALSE for any other element.

```
is_trembl(c("Q05BL1", "A0A654IBU3", "P00533"))
# [1] TRUE TRUE FALSE
```

is\_uniprot 93

is\_uniprot

Looks like a UniProt ID?

## **Description**

This function checks only the format of the IDs, no guarantee that these IDs exist in UniProt.

## Usage

```
is_uniprot(identifiers)
```

### **Arguments**

identifiers

Character: one or more identifiers (typically a single string, a vector or a data frame column).

## Value

Logical: true if all elements in the input (except NAs) looks like valid UniProt IDs. If the input is not a character vector, 'FALSE' is returned.

## **Examples**

```
is_uniprot(all_uniprot_acs())
# [1] TRUE
is_uniprot("P00533")
# [1] TRUE
is_uniprot("pizza")
# [1] FALSE
```

kegg\_info

Information about a KEGG Pathway

# Description

Information about a KEGG Pathway

# Usage

```
kegg_info(pathway_id)
```

# Arguments

pathway\_id

Character: a KEGG Pathway identifier, e.g. "hsa04710". For a complete list of IDs see kegg\_pathway\_list.

## Value

List with the pathway information.

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### See Also

- kegg\_pathway\_list
- kegg\_picture
- kegg\_open

## **Examples**

```
kegg_info('map00563')
```

kegg\_open

Open a KEGG Pathway diagram in the browser

## **Description**

Open a KEGG Pathway diagram in the browser

# Usage

```
kegg_open(pathway_id)
```

# Arguments

pathway\_id

Character: a KEGG Pathway identifier, e.g. "hsa04710". For a complete list of IDs see kegg\_pathway\_list.

# **Details**

To open URLs in the web browser the "browser" option must to be set to a a valid executable. You can check the value of this option by getOption("browser"). If your browser is firefox and the executable is located in the system path, you can set the option to point to it: options(browser = "firefox"). To make it a permanent setting, you can also include this in your .Rprofile file.

### Value

Returns NULL.

## See Also

- kegg\_pathway\_list
- kegg\_picture
- kegg\_info

```
if(any(getOption('browser') != '')) kegg_open('hsa04710')
```

kegg\_pathways\_download

Download the KEGG Pathways database

## **Description**

Downloads all pathway diagrams in the KEGG Pathways database in KGML format and processes the XML to extract the interactions.

## Usage

```
kegg_pathways_download(max_expansion = NULL, simplify = FALSE)
```

#### **Arguments**

Numeric: the maximum number of relations derived from a single relation record. As one entry might represent more than one molecular entities, one relation might yield a large number of relations in the processing. This happens in a combinatorial way, e.g. if the two entries represent 3 and 4 entities, that results 12 relations. If NULL, all relations will be expanded.

Simplify

Logical: remove KEGG's internal identifiers and the pathway annotations, keep only unique interactions with direction and effect sign.

#### Value

A data frame (tibble) of interactions.

# See Also

- kegg\_pathway\_list
- kegg\_process
- kegg\_pathway\_download

```
## Not run:
kegg_pw <- kegg_pathways_download(simplify = TRUE)</pre>
kegg_pw
# # A tibble: 6,765 x 6
    uniprot_source uniprot_target type effect genesymbol_source
    <chr>
#
                   <chr>
                                  <chr> <chr> <chr>
  1 Q03113
                                  PPrel activ. GNA12
#
                   Q15283
  2 Q9Y4G8
                   P62070
                                 PPrel activ. RAPGEF2
# 3 Q13972
                                 PPrel activ. RASGRF1
                   P62070
# 4 095267
                   P62070
                                  PPrel activ. RASGRP1
# 5 P62834
                   P15056
                                  PPrel activ. RAP1A
# # . with 6,760 more rows, and 1 more variable: genesymbol_target <chr>
## End(Not run)
```

```
kegg_pathway_annotations
```

Protein pathway annotations

# Description

Downloads all KEGG pathways and creates a table of protein-pathway annotations.

# Usage

```
kegg_pathway_annotations(pathways = NULL)
```

### **Arguments**

pathways A table of KE

A table of KEGG pathways as produced by kegg\_pathways\_download.

#### Value

A data frame (tibble) with UniProt IDs and pathway names.

#### See Also

kegg\_pathways\_download

# **Examples**

## **Description**

Downloads one pathway diagram from the KEGG Pathways database in KGML format and processes the XML to extract the interactions.

### Usage

```
kegg_pathway_download(
  pathway_id,
  process = TRUE,
  max_expansion = NULL,
  simplify = FALSE
)
```

### **Arguments**

pathway\_id Character: a KEGG pathway identifier, for example "hsa04350".

process Logical: process the data or return it in raw format. processing means joining

the entries and relations into a single data frame and adding UniProt IDs.

max\_expansion Numeric: the maximum number of relations derived from a single relation

record. As one entry might represent more than one molecular entities, one relation might yield a large number of relations in the processing. This happens in a combinatorial way, e.g. if the two entries represent 3 and 4 entities, that

results 12 relations. If NULL, all relations will be expanded.

simplify Logical: remove KEGG's internal identifiers and the pathway annotations, keep

only unique interactions with direction and effect sign.

### Value

A data frame (tibble) of interactions if process is TRUE, otherwise a list with two data frames: "entries" is a raw table of the entries while "relations" is a table of relations extracted from the KGML file.

### See Also

- kegg\_process
- kegg\_pathways\_download
- kegg\_pathway\_list

```
tgf_pathway <- kegg_pathway_download('hsa04350')</pre>
tgf_pathway
# # A tibble: 50 x 12
    source target type effect arrow relation_id kegg_id_source
    <chr> <chr> <chr> <chr> <chr> <chr> <chr> <chr> 
                                                <chr>
 1 51
                  PPrel activ. --> hsa04350:1 hsa:7040 hsa:.
           49
                  PPrel activ. --> hsa04350:2 hsa:151449 hs.
#
 2 57
           55
                  PPrel activ. --> hsa04350:3 hsa:3624 hsa:.
#
 3 34
           32
                  PPrel activ. --> hsa04350:4 hsa:4838
# 4 20
           17
                  PPrel activ. --> hsa04350:5 hsa:4086 hsa:.
# 5 60
           46
# # . with 45 more rows, and 5 more variables: genesymbol_source <chr>,
# # uniprot_source <chr>, kegg_id_target <chr>,
# # genesymbol_target <chr>, uniprot_target <chr>
```

98 kegg\_pathway\_list

## **Description**

Retrieves a list of available KEGG pathways.

## Usage

```
kegg_pathway_list()
```

## Value

Data frame of pathway names and identifiers.

### See Also

- kegg\_process
- kegg\_pathway\_download
- kegg\_pathways\_download
- kegg\_open
- kegg\_picture
- kegg\_info

```
kegg_pws <- kegg_pathway_list()</pre>
kegg_pws
# # A tibble: 521 x 2
    id
             name
              <chr>
#
    <chr>
# 1 map01100 Metabolic pathways
# 2 map01110 Biosynthesis of secondary metabolites
# 3 map01120 Microbial metabolism in diverse environments
# 4 map01200 Carbon metabolism
# 5 map01210 2-Oxocarboxylic acid metabolism
# 6 map01212 Fatty acid metabolism
\# 7 map01230 Biosynthesis of amino acids
\# \# . with 514 more rows
```

kegg\_picture 99

kegg_p	oicture
--------	---------

Download a pathway diagram as a picture

## **Description**

Downloads a KEGG Pathway diagram as a PNG image.

### Usage

```
kegg_picture(pathway_id, path = NULL)
```

## **Arguments**

pathway\_id Character: a KEGG Pathway identifier, e.g. "hsa04710". For a complete list of

IDs see kegg\_pathway\_list.

path Character: save the image to this path. If NULL, the image will be saved in the

current directory under the name <pathway\_id>.png.

# Value

Invisibly returns the path to the downloaded file.

### See Also

kegg\_pathway\_list

- kegg\_pathway\_list
- kegg\_open
- kegg\_info

## **Examples**

```
kegg_picture('hsa04710')
kegg_picture('hsa04710', path = 'foo/bar')
kegg_picture('hsa04710', path = 'foo/bar/circadian.png')
```

kegg\_process

Interactions from KGML

### **Description**

Processes KEGG Pathways data extracted from a KGML file. Joins the entries and relations into a single data frame and translates the Gene Symbols to UniProt IDs.

## Usage

```
kegg_process(entries, relations, max_expansion = NULL, simplify = FALSE)
```

100 latin\_name

### **Arguments**

entries A data frames with entries extracted from a KGML file by kegg\_pathway\_download.

A data frames with relations extracted from a KGML file by kegg\_pathway\_download.

Numeric: the maximum number of relations derived from a single relation record. As one entry might represent more than one molecular entities, one relation might yield a large number of relations in the processing. This happens in a combinatorial way, e.g. if the two entries represent 3 and 4 entities, that results 12 relations. If NULL, all relations will be expanded.

Simplify Logical: remove KEGG's internal identifiers and the pathway annotations, keep only unique interactions with direction and effect sign.

Value

A data frame (tibble) of interactions. In rare cases when a pathway doesn't contain any relation, returns NULL.

#### See Also

- kegg\_pathway\_download
- kegg\_pathways\_download
- kegg\_pathway\_list

### **Examples**

```
hsa04350 <- kegg_pathway_download('hsa04350', process = FALSE)</pre>
tgf_pathway <- kegg_process(hsa04350$entries, hsa04350$relations)</pre>
tgf_pathway
# # A tibble: 50 x 12
                 source target type effect arrow relation_id kegg_id_source
                  <chr> <chr< <chr> <chr< <chr> <chr> <chr> <chr> <chr> <chr< <chr> <chr> <chr> <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr> <chr> <chr> <chr> <chr> <chr< <chr> <chr< <chr> <chr< <chr< <chr> <chr< <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr< <chr> <chr< <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr> <chr< <chr< <chr< <chr< <chr< <
                                                                                                                                                                                               <chr>
                                                                        PPrel activ. --> hsa04350:1 hsa:7040 hsa:.
       1 51
                                             49
                                                                        PPrel activ. --> hsa04350:2 hsa:151449 hs.
       2 57
                                             55
# 3 34
                                                                        PPrel activ. --> hsa04350:3 hsa:3624 hsa:.
                                             32
# 4 20
                                             17
                                                                        PPrel activ. --> hsa04350:4 hsa:4838
# 5 60
                                             46
                                                                        PPrel activ. --> hsa04350:5 hsa:4086 hsa:.
# # . with 45 more rows, and 5 more variables: genesymbol_source <chr>,
# # uniprot_source <chr>, kegg_id_target <chr>,
                      genesymbol_target <chr>, uniprot_target <chr>
```

latin\_name

Latin (scientific) names of organisms

## **Description**

Latin (scientific) names of organisms

### Usage

```
latin_name(name)
```

load\_db

## **Arguments**

name

Vector with any kind of organism name or identifier, can be also mixed type.

#### Value

Character vector with latin (scientific) names, NA if a name in the input could not be found.

### See Also

- ncbi\_taxid
- common\_name
- ensembl\_name

## **Examples**

```
latin_name(c(9606, "cat", "dog"))
# [1] "Homo sapiens" "Felis catus" "Canis lupus familiaris"
latin_name(c(9606, "cat", "doggy"))
# [1] "Homo sapiens" "Felis catus" NA
```

load\_db

Load a built in database

### **Description**

Load a built in database

### Usage

```
load_db(key, param = list())
```

# **Arguments**

key Character: the key of the database to load. For a list of available keys see

omnipath\_show\_db.

param List: override the defaults or pass further parameters to the database loader func-

tion. See the loader functions and their default parameters in omnipath\_show\_db.

### **Details**

This function loads a database which is stored within the package namespace until its expiry. The loaded database is accessible by get\_db and the loading process is typically initiated by get\_db, not by the users directly.

### Value

Returns NULL.

### See Also

```
omnipath_show_db, get_db
```

ncbi\_taxid

## **Examples**

```
load_db('go_slim')
omnipath_show_db()
```

ncbi\_taxid

NCBI Taxonomy IDs of organisms

# Description

NCBI Taxonomy IDs of organisms

# Usage

```
ncbi_taxid(name)
```

# Arguments

name

Vector with any kind of organism name or identifier, can be also mixed type.

### Value

Integer vector with NCBI Taxonomy IDs, NA if a name in the input could not be found.

## See Also

- latin\_name
- common\_name
- ensembl\_name

```
ncbi_taxid(c("Homo sapiens", "cat", "dog"))
# [1] 9606 9685 9615
ncbi_taxid(c(9606, "cat", "doggy"))
# [1] 9606 9685 NA
```

nichenet\_build\_model 103

### **Description**

Construct a NicheNet ligand-target model

## Usage

```
nichenet_build_model(optimization_results, networks, use_weights = TRUE)
```

## **Arguments**

optimization\_results

 $The \ outcome \ of \ NicheNet \ parameter \ optimization \ as \ produced \ by \ nichenet\_optimization.$ 

networks A list with NicheNet format signaling, ligand-receptor and gene regulatory net-

works as produced by nichenet\_networks.

use\_weights Logical: whether to use the optimized weights.

#### Value

A named list with two elements: 'weighted\_networks' and 'optimized\_parameters'.

### **Examples**

```
## Not run:
expression <- nichenet_expression_data()
networks <- nichenet_networks()
optimization_results <- nichenet_optimization(networks, expression)
nichenet_model <- nichenet_build_model(optimization_results, networks)
## End(Not run)</pre>
```

nichenet\_expression\_data

Expression data from ligand-receptor perturbation experiments used by NicheNet

## **Description**

NicheNet uses expression data from a collection of published ligand or receptor KO or perturbation experiments to build its model. This function retrieves the original expression data, deposited in Zenodo (https://zenodo.org/record/3260758).

## Usage

```
nichenet_expression_data()
```

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### Value

Nested list, each element contains a data frame of processed expression data and key variables about the experiment.

### **Examples**

```
exp_data <- nichenet_expression_data()</pre>
head(names(exp_data))
                      "tgfb_bmp4"
# [1] "bmp4_tgfb"
                                       "nodal_Nodal"
                                                        "spectrum_I14"
# [5] "spectrum_Tnf" "spectrum_Ifng"
purrr::map_chr(head(exp_data), 'from')
      \verb|bmp4_tgfb| & tgfb_bmp4 & nodal_Nodal & spectrum_Il4 & spectrum_Tnf| \\
        "BMP4"
                     "TGFB1"
                                    "NODAL"
                                                     "IL4"
                                                                    "TNF"
# spectrum_Ifng
       "IFNG"
```

nichenet\_gr\_network

Builds a NicheNet gene regulatory network

## **Description**

Builds gene regulatory network prior knowledge for NicheNet using multiple resources.

## Usage

```
nichenet_gr_network(
  omnipath = list(),
  harmonizome = list(),
  regnetwork = list(),
  htridb = list(),
  remap = list(),
  evex = list(),
  pathwaycommons = list(),
  trrust = list(),
  only_omnipath = FALSE
)
```

### **Arguments**

```
List with paramaters to be passed to nichenet_gr_network_omnipath.
omnipath
                 List with paramaters to be passed to nichenet\_gr\_network\_harmonizome.
harmonizome
                 List with paramaters to be passed to nichenet_gr_network_regnetwork.
regnetwork
                 List with paramaters to be passed to nichenet_gr_network_htridb.
htridb
remap
                 List with paramaters to be passed to nichenet_gr_network_remap.
                 List with paramaters to be passed to nichenet_gr_network_evex.
evex
pathwaycommons  List with paramaters to be passed to nichenet_gr_network_pathwaycommons.
                 List with paramaters to be passed to nichenet_gr_network_trrust.
trrust
only_omnipath
                Logical: a shortcut to use only OmniPath as network resource.
```

### Value

A network data frame (tibble) with gene regulatory interactions suitable for use with NicheNet.

### See Also

- nichenet\_gr\_network\_evex
- nichenet\_gr\_network\_harmonizome
- nichenet\_gr\_network\_htridb
- nichenet\_gr\_network\_omnipath
- nichenet\_gr\_network\_pathwaycommons
- nichenet\_gr\_network\_regnetwork
- nichenet\_gr\_network\_remap
- nichenet\_gr\_network\_trrust

## **Examples**

```
# load everything with the default parameters:
gr_network <- nichenet_gr_network()

# less targets from ReMap, not using RegNetwork:
gr_network <- nichenet_gr_network(
    # I needed to disable ReMap here due to some issues
    # of one of the Bioconductor build servers
    # remap = list(top_targets = 200),
    remap = NULL,
    regnetwork = NULL,
)

# use only OmniPath:
gr_network_omnipath <- nichenet_gr_network(only_omnipath = TRUE)</pre>
```

nichenet\_gr\_network\_evex

NicheNet gene regulatory network from EVEX

## **Description**

Builds a gene regulatory network using data from the EVEX database and converts it to a format suitable for NicheNet.

## Usage

```
nichenet_gr_network_evex(
  top_confidence = 0.75,
  indirect = FALSE,
  regulation_of_expression = FALSE
)
```

### **Arguments**

```
top_confidence Double, between 0 and 1. Threshold based on the quantile of the confidence score.

indirect Logical: whether to include indirect interactions.

regulation_of_expression

Logical: whether to include also the "regulation of expression" type interactions.
```

### Value

Data frame of interactions in NicheNet format.

Data frame with gene regulatory interactions in NicheNet format.

#### See Also

- nichenet\_gr\_network
- evex\_download

## **Examples**

```
# use only the 10% with the highest confidence:
evex_gr_network <- nichenet_gr_network_evex(top_confidence = .9)</pre>
```

```
\verb|nichenet_gr_network_harmonizome|\\
```

NicheNet gene regulatory network from Harmonizome

### **Description**

Builds gene regulatory network prior knowledge for NicheNet using Harmonizome

## Usage

```
nichenet_gr_network_harmonizome(
  datasets = c("cheappi", "encodetfppi", "jasparpwm", "transfac", "transfacpwm",
        "motifmap", "geotf", "geokinase", "geogene"),
        ...
)
```

## **Arguments**

```
datasets The datasets to use. For possible values please refer to default value and the Harmonizome webpage.

... Ignored.
```

## Value

Data frame with gene regulatory interactions in NicheNet format.

#### See Also

- nichenet\_gr\_network
- harmonizome\_download

## **Examples**

```
# use only JASPAR and TRANSFAC:
hz_gr_network <- nichenet_gr_network_harmonizome(
    datasets = c('jasparpwm', 'transfac', 'transfacpwm')
)</pre>
```

```
nichenet_gr_network_htridb
```

NicheNet gene regulatory network from HTRIdb

## **Description**

Builds a gene regulatory network using data from the HTRIdb database and converts it to a format suitable for NicheNet.

### Usage

```
nichenet_gr_network_htridb()
```

### Value

Data frame with gene regulatory interactions in NicheNet format.

# See Also

```
htridb_download, nichenet_gr_network
```

## **Examples**

```
htri_gr_network <- nichenet_gr_network_htridb()</pre>
```

```
nichenet_gr_network_omnipath
```

Builds gene regulatory network for NicheNet using OmniPath

# Description

Retrieves network prior knowledge from OmniPath and provides it in a format suitable for NicheNet. This method never downloads the 'ligrecextra' dataset because the ligand-receptor interactions are supposed to come from nichenet\_lr\_network\_omnipath.

### Usage

```
nichenet_gr_network_omnipath(min_curation_effort = 0, ...)
```

### **Arguments**

```
min_curation_effort

Lower threshold for curation effort

... Passed to import_transcriptional_interactions
```

#### Value

A network data frame (tibble) with gene regulatory interactions suitable for use with NicheNet.

## See Also

- nichenet\_gr\_network\_evex
- nichenet\_gr\_network\_harmonizome
- nichenet\_gr\_network\_htridb
- nichenet\_gr\_network\_omnipath
- nichenet\_gr\_network\_pathwaycommons
- nichenet\_gr\_network\_regnetwork
- nichenet\_gr\_network\_remap
- nichenet\_gr\_network\_trrust

## **Examples**

```
# use interactions up to confidence level "C" from DoRothEA:
op_gr_network <- nichenet_gr_network_omnipath(
    dorothea_levels = c('A', 'B', 'C')
)</pre>
```

nichenet\_gr\_network\_pathwaycommons

NicheNet gene regulatory network from PathwayCommons

## **Description**

Builds gene regulation prior knowledge for NicheNet using PathwayCommons.

## Usage

```
nichenet_gr_network_pathwaycommons(
  interaction_types = "controls-expression-of",
  ...
)
```

# **Arguments**

interaction\_types

Character vector with PathwayCommons interaction types. Please refer to the default value and the PathwayCommons webpage.

... Ignored.

#### Value

Data frame with gene regulatory interactions in NicheNet format.

### See Also

- nichenet\_gr\_network
- pathwaycommons\_download

### **Examples**

```
pc_gr_network <- nichenet_gr_network_pathwaycommons()</pre>
```

nichenet\_gr\_network\_regnetwork

NicheNet gene regulatory network from RegNetwork

# Description

Builds a gene regulatory network using data from the RegNetwork database and converts it to a format suitable for NicheNet.

## Usage

```
nichenet_gr_network_regnetwork()
```

## Value

Data frame with gene regulatory interactions in NicheNet format.

### See Also

- regnetwork\_download
- nichenet\_gr\_network

```
regn_gr_network <- nichenet_gr_network_regnetwork()</pre>
```

```
nichenet_gr_network_remap
```

NicheNet gene regulatory network from ReMap

# Description

Builds a gene regulatory network using data from the ReMap database and converts it to a format suitable for NicheNet.

## Usage

```
nichenet_gr_network_remap(
  score = 100,
  top_targets = 500,
  only_known_tfs = TRUE
)
```

## **Arguments**

score Numeric: a minimum score between 0 and 1000, records with lower scores will

be excluded. If NULL no filtering performed.

mum number of targets per TF. If NULL the number of targets is not restricted.

only\_known\_tfs Logical: whether to exclude TFs which are not in TF census.

### Value

Data frame with gene regulatory interactions in NicheNet format.

## See Also

- remap\_filtered
- nichenet\_gr\_network

```
# use only max. top 100 targets for each TF:
remap_gr_network <- nichenet_gr_network_remap(top_targets = 100)</pre>
```

```
nichenet_gr_network_trrust
```

NicheNet gene regulatory network from TRRUST

# Description

Builds a gene regulatory network using data from the TRRUST database and converts it to a format suitable for NicheNet.

## Usage

```
nichenet_gr_network_trrust()
```

#### Value

Data frame with gene regulatory interactions in NicheNet format.

### See Also

- trrust\_download
- nichenet\_gr\_network

## **Examples**

```
trrust_gr_network <- nichenet_gr_network_trrust()</pre>
```

```
nichenet_ligand_activities
```

Calls the NicheNet ligand activity analysis

# Description

Calls the NicheNet ligand activity analysis

## Usage

```
nichenet_ligand_activities(
   ligand_target_matrix,
   lr_network,
   expressed_genes_transmitter,
   expressed_genes_receiver,
   genes_of_interest,
   background_genes = NULL,
   n_top_ligands = 42,
   n_top_targets = 250
)
```

### **Arguments**

```
ligand_target_matrix
```

A matrix with rows and columns corresponding to ligands and targets, respectively. Produced by nichenet\_ligand\_target\_matrix or nichenetr::construct\_ligand\_target

1r\_network A data frame

A data frame with ligand-receptor interactions, as produced by  $\mbox{nichenet\_lr\_network}$ .

expressed\_genes\_transmitter

Character vector with the gene symbols of the genes expressed in the cells transmitting the signal.

expressed\_genes\_receiver

Character vector with the gene symbols of the genes expressed in the cells receiving the signal.

genes\_of\_interest

Character vector with the gene symbols of the genes of interest. These are the genes in the receiver cell population that are potentially affected by ligands expressed by interacting cells (e.g. genes differentially expressed upon cell-cell interaction).

background\_genes

Character vector with the gene symbols of the genes to be used as background.

n\_top\_ligands How many of the top ligands to include in the ligand-target table.

n\_top\_targets For each ligand, how many of the top targets to include in the ligand-target table.

#### Value

A named list with 'ligand\_activities' (a tibble giving several ligand activity scores; following columns in the tibble: \$test\_ligand, \$auroc, \$aupr and \$pearson) and 'ligand\_target\_links' (a tibble with columns ligand, target and weight (i.e. regulatory potential score)).

```
## Not run:
networks <- nichenet_networks()</pre>
expression <- nichenet_expression_data()</pre>
optimization_results <- nichenet_optimization(networks, expression)</pre>
nichenet_model <- nichenet_build_model(optimization_results, networks)</pre>
lt_matrix <- nichenet_ligand_target_matrix(</pre>
    \verb|nichenet_model| \verb|sweighted_networks|, \\
    networks$lr_network,
    nichenet_model$optimized_parameters
ligand_activities <- nichenet_ligand_activities(</pre>
    ligand_target_matrix = lt_matrix,
    lr_network = networks$lr_network,
    \ensuremath{\text{\#}} the rest of the parameters should come
    # from your transcriptomics data:
    expressed_genes_transmitter = expressed_genes_transmitter,
    expressed_genes_receiver = expressed_genes_receiver,
    genes_of_interest = genes_of_interest
)
## End(Not run)
```

```
nichenet_ligand_target_links
```

Compiles a table with weighted ligand-target links

# Description

A wrapper around nichenetr::get\_weighted\_ligand\_target\_links to compile a data frame with weighted links from the top ligands to their top targets.

## Usage

```
nichenet_ligand_target_links(
  ligand_activities,
  ligand_target_matrix,
  genes_of_interest,
  n_top_ligands = 42,
  n_top_targets = 250
)
```

### **Arguments**

ligand\_activities

Ligand activity table as produced by nichenetr::predict\_ligand\_activities.

ligand\_target\_matrix

Ligand-target matrix as produced by nichenetr::construct\_ligand\_target\_matrix or the wrapper around it in the current package: nichenet\_ligand\_target\_matrix.

genes\_of\_interest

Character vector with the gene symbols of the genes of interest. These are the genes in the receiver cell population that are potentially affected by ligands expressed by interacting cells (e.g. genes differentially expressed upon cell-cell interaction).

n\_top\_ligands How many of the top ligands to include in the ligand-target table.

n\_top\_targets For each ligand, how many of the top targets to include in the ligand-target table.

### Value

A tibble with columns ligand, target and weight (i.e. regulatory potential score).

```
## Not run:
networks <- nichenet_networks()
expression <- nichenet_expression_data()
optimization_results <- nichenet_optimization(networks, expression)
nichenet_model <- nichenet_build_model(optimization_results, networks)
lt_matrix <- nichenet_ligand_target_matrix(
    nichenet_model$weighted_networks,
    networks$lr_network,
    nichenet_model$optimized_parameters
)
ligand_activities <- nichenet_ligand_activities(</pre>
```

```
ligand_target_matrix = lt_matrix,
lr_network = networks$lr_network,
# the rest of the parameters should come
# from your transcriptomics data:
expressed_genes_transmitter = expressed_genes_transmitter,
expressed_genes_receiver = expressed_genes_receiver,
genes_of_interest = genes_of_interest
)
lt_links <- nichenet_ligand_target_links(
    ligand_activities = ligand_activities,
    ligand_target_matrix = lt_matrix,
    genes_of_interest = genes_of_interest,
    n_top_ligands = 20,
    n_top_targets = 100
)
## End(Not run)</pre>
```

nichenet\_ligand\_target\_matrix

Creates a NicheNet ligand-target matrix

## **Description**

Creates a NicheNet ligand-target matrix

### Usage

```
nichenet_ligand_target_matrix(
  weighted_networks,
  lr_network,
  optimized_parameters,
  use_weights = TRUE,
  construct_ligand_target_matrix_param = list()
)
```

## **Arguments**

weighted\_networks

Weighted networks as provided by nichenet\_build\_model.

The outcome of NicheNet parameter optimization as produced by  $\mbox{nichenet\_build\_model}.$ 

use\_weights Logical: wether the network sources are weighted. In this function it only affects the output file name.

construct\_ligand\_target\_matrix\_param

Override parameters for nichenetr::construct\_ligand\_target\_matrix.

## Value

A matrix containing ligand-target probability scores.

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#### **Examples**

```
## Not run:
networks <- nichenet_networks()
expression <- nichenet_expression_data()
optimization_results <- nichenet_optimization(networks, expression)
nichenet_model <- nichenet_build_model(optimization_results, networks)
lt_matrix <- nichenet_ligand_target_matrix(
    nichenet_model$weighted_networks,
    networks$lr_network,
    nichenet_model$optimized_parameters
)

## End(Not run)</pre>
```

nichenet\_lr\_network

Builds a NicheNet ligand-receptor network

### **Description**

Builds ligand-receptor network prior knowledge for NicheNet using multiple resources.

## Usage

```
nichenet_lr_network(
  omnipath = list(),
  guide2pharma = list(),
  ramilowski = list(),
  only_omnipath = FALSE,
  quality_filter_param = list()
)
```

### Arguments

omnipath List with paramaters to be passed to nichenet\_lr\_network\_omnipath.

guide2pharma List with paramaters to be passed to nichenet\_lr\_network\_guide2pharma.

ramilowski List with paramaters to be passed to nichenet\_lr\_network\_ramilowski.

only\_omnipath Logical: a shortcut to use only OmniPath as network resource.

quality\_filter\_param

Arguments for filter\_intercell\_network (quality filtering of the OmniPath ligand-receptor network). It is recommended to check these parameters and apply some quality filtering. The defaults already ensure certain filtering, but you might want more relaxed or stringent options.

### Value

A network data frame (tibble) with ligand-receptor interactions suitable for use with NicheNet.

### See Also

- nichenet\_lr\_network\_omnipath
- nichenet\_lr\_network\_guide2pharma
- nichenet\_lr\_network\_ramilowski
- filter\_intercell\_network

## **Examples**

```
# load everything with the default parameters:
lr_network <- nichenet_lr_network()

# don't use Ramilowski:
lr_network <- nichenet_lr_network(ramilowski = NULL)

# use only OmniPath:
lr_network_omnipath <- nichenet_lr_network(only_omnipath = TRUE)</pre>
```

```
nichenet_lr_network_guide2pharma
```

Ligand-receptor network from Guide to Pharmacology

# Description

Downloads ligand-receptor interactions from the Guide to Pharmacology database and converts it to a format suitable for NicheNet.

#### Usage

```
nichenet_lr_network_guide2pharma()
```

## Value

Data frame with ligand-receptor interactions in NicheNet format.

## See Also

```
nichenet_lr_network, guide2pharma_download
```

```
g2p_lr_network <- nichenet_lr_network_guide2pharma()</pre>
```

```
nichenet_lr_network_omnipath
```

Builds ligand-receptor network for NicheNet using OmniPath

### **Description**

Retrieves network prior knowledge from OmniPath and provides it in a format suitable for NicheNet. This method never downloads the 'ligrecextra' dataset because the ligand-receptor interactions are supposed to come from nichenet\_lr\_network\_omnipath.

### Usage

```
nichenet_lr_network_omnipath(quality_filter_param = list(), ...)
```

## **Arguments**

```
quality_filter_param
```

List with arguments for filter\_intercell\_network. It is recommended to check these parameters and apply some quality filtering. The defaults already ensure certain filtering, but you might want more relaxed or stringent options.

```
Passed to import_intercell_network
```

### Value

A network data frame (tibble) with ligand-receptor interactions suitable for use with NicheNet.

# See Also

- nichenet\_lr\_network
- import\_intercell\_network

```
# use only ligand-receptor interactions (not for example ECM-adhesion):
op_lr_network <- nichenet_lr_network_omnipath(ligand_receptor = TRUE)

# use only CellPhoneDB and Guide to Pharmacology:
op_lr_network <- nichenet_lr_network_omnipath(
    resources = c('CellPhoneDB', 'Guide2Pharma')
)

# only interactions where the receiver is a transporter:
op_lr_network <- nichenet_lr_network_omnipath(
    receiver_param = list(parent = 'transporter')
)</pre>
```

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```
nichenet_lr_network_ramilowski
```

Ligand-receptor network from Ramilowski 2015

## **Description**

Downloads ligand-receptor interactions from Supplementary Table 2 of the paper 'A draft network of ligand-receptor-mediated multicellular signalling in human' (Ramilowski et al. 2015, https://www.nature.com/articles/ncomms8866). It converts the downloaded table to a format suitable for NicheNet.

### Usage

```
nichenet_lr_network_ramilowski(
  evidences = c("literature supported", "putative")
)
```

### **Arguments**

evidences

Character: evidence types, "literature supported", "putative" or both.

#### Value

Data frame with ligand-receptor interactions in NicheNet format.

### See Also

- nichenet\_lr\_network
- ramilowski\_download

### **Examples**

```
# use only the literature supported data:
rami_lr_network <- nichenet_lr_network_ramilowski(
    evidences = 'literature supported'
)</pre>
```

nichenet\_main

Executes the full NicheNet pipeline

## **Description**

Builds all prior knowledge data required by NicheNet. For this it calls a multitude of methods to download and combine data from various databases according to the settings. The content of the prior knowledge data is highly customizable, see the documentation of the related functions. After the prior knowledge is ready, it performs parameter optimization to build a NicheNet model. This results a weighted ligand- target matrix. Then, considering the expressed genes from user provided data, a gene set of interest and background genes, it executes the NicheNet ligand activity analysis.

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#### Usage

```
nichenet_main(
  only_omnipath = FALSE,
  expressed_genes_transmitter = NULL,
  expressed_genes_receiver = NULL,
  genes_of_interest = NULL,
  background_genes = NULL,
  use_weights = TRUE,
  n_{top_ligands} = 42,
  n_{top_targets} = 250,
  signaling_network = list(),
  lr_network = list(),
  gr_network = list(),
  small = FALSE,
  tiny = FALSE,
  make_multi_objective_function_param = list(),
  objective_function_param = list(),
  mlrmbo_optimization_param = list(),
  construct_ligand_target_matrix_param = list(),
  results_dir = NULL,
  quality_filter_param = list()
)
```

#### **Arguments**

only\_omnipath

Logical: use only OmniPath for network knowledge. This is a simple switch for convenience, further options are available by the other arguments. By default we use all available resources. The networks can be customized on a resource by resource basis, as well as providing custom parameters for individual resources, using the parameters 'signaling\_network', 'lr\_network' and 'gr\_network'.

 ${\tt expressed\_genes\_transmitter}$ 

Character vector with the gene symbols of the genes expressed in the cells transmitting the signal.

expressed\_genes\_receiver

Character vector with the gene symbols of the genes expressed in the cells receiving the signal.

genes\_of\_interest

Character vector with the gene symbols of the genes of interest. These are the genes in the receiver cell population that are potentially affected by ligands expressed by interacting cells (e.g. genes differentially expressed upon cell-cell interaction).

 $background\_genes$ 

Character vector with the gene symbols of the genes to be used as background.

use\_weights

Logical: calculate and use optimized weights for resources (i.e. one resource seems to be better than another, hence the former is considered with a higher weight).

n\_top\_ligands How many of the top ligands to include in the ligand-target table.

n\_top\_targets How many of the top targets (for each of the top ligands) to consider in the ligand-target table.

signaling\_network

A list of parameters for building the signaling network, passed to nichenet\_signaling\_network.

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lr\_network A list of parameters for building the ligand-receptor network, passed to nichenet\_lr\_network.

gr\_network A list of parameters for building the gene regulatory network, passed to nichenet\_gr\_network.

small Logical: build a small network for testing purposes, using only OmniPath data.

It is also a high quality network, it is reasonable to try the analysis with this

small network.

tiny Logical: build an even smaller network for testing purposes. As this involves

random subsetting, it's not recommended to use this network for analysis.

make\_multi\_objective\_function\_param

Override parameters for smoof::makeMultiObjectiveFunction.

objective\_function\_param

Override additional arguments passed to the objective function.

mlrmbo\_optimization\_param

 $Override\ arguments\ for\ nichenetr:: {\tt mlrmbo\_optimization}.$ 

construct\_ligand\_target\_matrix\_param

Override parameters for nichenetr::construct\_ligand\_target\_matrix.

results\_dir Char

Character: path to the directory to save intermediate and final outputs from NicheNet methods.

quality\_filter\_param

Arguments for filter\_intercell\_network (quality filtering of the OmniPath ligand-receptor network). It is recommended to check these parameters and apply some quality filtering. The defaults already ensure certain filtering, but you might want more relaxed or stringent options.

### **Details**

About *small* and *tiny* networks: Building a NicheNet model is computationally demanding, taking several hours to run. As this is related to the enormous size of the networks, to speed up testing we can use smaller networks, around 1,000 times smaller, with few thousands of interactions instead of few millions. Random subsetting of the whole network would result disjunct fragments, instead we load only a few resources. To run the whole pipeline with tiny networks use nichenet\_test.

#### Value

A named list with the intermediate and final outputs of the pipeline: 'networks', 'expression', 'optimized\_parameters', 'weighted\_networks' and 'ligand\_target\_matrix'.

## See Also

- nichenet\_networks
- nichenet\_signaling\_network
- nichenet\_lr\_network
- nichenet\_gr\_network
- nichenet\_test
- nichenet\_workarounds
- nichenet\_results\_dir

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#### **Examples**

```
## Not run:
nichenet_results <- nichenet_main(
    # altering some network resource parameters, the rest
    # of the resources will be loaded according to the defaults
    signaling_network = list(
        cpdb = NULL, # this resource will be excluded
        inbiomap = NULL,
        evex = list(min_confidence = 1.0) # override some parameters
),
    gr_network = list(only_omnipath = TRUE),
    n_top_ligands = 20,
    # override the default number of CPU cores to use
    mlrmbo_optimization_param = list(ncores = 4)
)
## End(Not run)</pre>
```

nichenet\_networks

Builds NicheNet network prior knowledge

### **Description**

Builds network knowledge required by NicheNet. For this it calls a multitude of methods to download and combine data from various databases according to the settings. The content of the prior knowledge data is highly customizable, see the documentation of the related functions.

## Usage

```
nichenet_networks(
   signaling_network = list(),
   lr_network = list(),
   gr_network = list(),
   only_omnipath = FALSE,
   small = FALSE,
   tiny = FALSE,
   quality_filter_param = list()
)
```

# **Arguments**

signaling\_network

gr\_network

A list of parameters for building the signaling network, passed to nichenet\_signaling\_network

A list of parameters for building the gene regulatory network, passed to nichenet\_gr\_network

 ${\tt lr\_network} \qquad \qquad A\ list\ of\ parameters\ for\ building\ the\ ligand-receptor\ network,\ passed\ to\ \verb|nichenet_lr_network||$ 

only\_omnipath Logical: a shortcut to use only OmniPath as network resource.

small Logical: build a small network for testing purposes, using only OmniPath data.

It is also a high quality network, it is reasonable to try the analysis with this

small network.

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tiny

Logical: build an even smaller network for testing purposes. As this involves random subsetting, it's not recommended to use this network for analysis.

```
quality_filter_param
```

Arguments for filter\_intercell\_network (quality filtering of the OmniPath ligand-receptor network). It is recommended to check these parameters and apply some quality filtering. The defaults already ensure certain filtering, but you might want more relaxed or stringent options.

### Value

A named list with three network data frames (tibbles): the signaling, the ligand-receptor (lr) and the gene regulatory (gr) networks.

#### See Also

- nichenet\_signaling\_network
- nichenet\_lr\_network
- nichenet\_gr\_network

### **Examples**

```
## Not run:
networks <- nichenet_networks()</pre>
dplyr::sample_n(networks$gr_network, 10)
# # A tibble: 10 x 4
    from
                     source
                                          database
            to
#
    <chr>
            <chr>
                     <chr>
                                          <chr>
#
  1 MAX
            ALG3
                     harmonizome_ENCODE
                                          harmonizome
  2 MAX
            IMPDH1 harmonizome_ENCODE harmonizome
                                          Remap
  3 SMAD5
            LCP1
                     Remap_5
#
  4 HNF4A
            TNFRSF19 harmonizome_CHEA
                                          harmonizome
#
  5 SMC3
            FAP
                    harmonizome_ENCODE harmonizome
            HIST1H1B harmonizome_ENCODE
#
  6 E2F6
                                          harmonizome
  7 TFAP2C MAT2B harmonizome_ENCODE
                                          harmonizome
  8 USF1
            TBX4
                     harmonizome_TRANSFAC harmonizome
  9 MIR133B FETUB
                     harmonizome_TRANSFAC harmonizome
# 10 SP4
            HNRNPH2 harmonizome_ENCODE harmonizome
## End(Not run)
# use only OmniPath:
omnipath_networks <- nichenet_networks(only_omnipath = TRUE)</pre>
```

nichenet\_optimization Optimizes NicheNet model parameters

## **Description**

Optimize NicheNet method parameters, i.e. PageRank parameters and source weights, basedon a collection of experiments where the effect of a ligand on gene expression was measured.

#### Usage

```
nichenet_optimization(
  networks,
  expression,
  make_multi_objective_function_param = list(),
  objective_function_param = list(),
  mlrmbo_optimization_param = list()
)
```

## **Arguments**

networks A list with NicheNet format signaling, ligand-receptor and gene regulatory networks as produced by nichenet\_networks.

expression A list with expression data from ligand perturbation experiments, as produced

by nichenet\_expression\_data.

make\_multi\_objective\_function\_param

 $Override\ parameters\ for\ smoof:: \verb|makeMultiObjectiveFunction|.$ 

objective\_function\_param

Override additional arguments passed to the objective function.

mlrmbo\_optimization\_param

Override arguments for nichenetr::mlrmbo\_optimization.

#### Value

A result object from the function mlrMBO::mbo. Among other things, this contains the optimal parameter settings, the output corresponding to every input etc.

## **Examples**

```
## Not run:
networks <- nichenet_networks()
expression <- nichenet_expression_data()
optimization_results <- nichenet_optimization(networks, expression)
## End(Not run)</pre>
```

```
nichenet_remove_orphan_ligands
```

Removes experiments with orphan ligands

### **Description**

Removes from the expression data the perturbation experiments involving ligands without connections.

## Usage

```
nichenet_remove_orphan_ligands(expression, lr_network)
```

### **Arguments**

expression Expression data as returned by nichenet\_expression\_data.

#### Value

The same list as 'expression' with certain elements removed.

# **Examples**

```
lr_network <- nichenet_lr_network()
expression <- nichenet_expression_data()
expression <- nichenet_remove_orphan_ligands(expression, lr_network)</pre>
```

# Description

Path to the directory to save intermediate and final outputs from NicheNet methods.

# Usage

```
nichenet_results_dir()
```

### Value

Character: path to the NicheNet results directory.

# **Examples**

```
nichenet_results_dir()
# [1] "nichenet_results"
```

nichenet\_signaling\_network

Builds a NicheNet signaling network

## **Description**

Builds signaling network prior knowledge for NicheNet using multiple resources.

#### Usage

```
nichenet_signaling_network(
  omnipath = list(),
  pathwaycommons = list(),
  harmonizome = list(),
  vinayagam = list(),
  cpdb = list(),
  evex = list(),
  inbiomap = list(),
  only_omnipath = FALSE
)
```

### **Arguments**

```
omnipath List with paramaters to be passed to nichenet_signaling_network_omnipath.

List with paramaters to be passed to nichenet_signaling_network_pathwaycommons.

List with paramaters to be passed to nichenet_signaling_network_harmonizome.

List with paramaters to be passed to nichenet_signaling_network_vinayagam.

List with paramaters to be passed to nichenet_signaling_network_cpdb.

List with paramaters to be passed to nichenet_signaling_network_evex.

List with paramaters to be passed to nichenet_signaling_network_evex.

List with paramaters to be passed to nichenet_signaling_network_inbiomap.

List with paramaters to be passed to nichenet_signaling_network_inbiomap.

List with paramaters to be passed to nichenet_signaling_network_inbiomap.

List with paramaters to be passed to nichenet_signaling_network_inbiomap.
```

#### Value

A network data frame (tibble) with signaling interactions suitable for use with NicheNet.

# See Also

- nichenet\_signaling\_network\_omnipath
- nichenet\_signaling\_network\_pathwaycommons
- nichenet\_signaling\_network\_harmonizome
- nichenet\_signaling\_network\_vinayagam
- nichenet\_signaling\_network\_cpdb
- nichenet\_signaling\_network\_evex
- nichenet\_signaling\_network\_inbiomap

```
# load everything with the default parameters:
# we don't load inBio Map due to the - hopefully
# temporary - issues of their server
sig_network <- nichenet_signaling_network(inbiomap = NULL, cpdb = NULL)
# override parameters for some resources:
sig_network <- nichenet_signaling_network(
    omnipath = list(resources = c('SIGNOR', 'SignaLink3', 'SPIKE')),
    pathwaycommons = NULL,
    harmonizome = list(datasets = c('phosphositeplus', 'depod')),
    # we can not include this in everyday tests as it takes too long:</pre>
```

```
# cpdb = list(complex_max_size = 1, min_score = .98),
    cpdb = NULL,
    evex = list(min_confidence = 1.5),
    inbiomap = NULL
)

# use only OmniPath:
sig_network_omnipath <- nichenet_signaling_network(only_omnipath = TRUE)</pre>
```

nichenet\_signaling\_network\_cpdb

Builds signaling network for NicheNet using ConsensusPathDB

## **Description**

Builds signaling network prior knowledge using ConsensusPathDB (CPDB) data. Note, the interactions from CPDB are not directed and many of them comes from complex expansion. Find out more at http://cpdb.molgen.mpg.de/.

## Usage

```
nichenet_signaling_network_cpdb(...)
```

# Arguments

.. Passed to consensuspathdb\_download.

### Value

A network data frame (tibble) with signaling interactions suitable for use with NicheNet.

### See Also

- nichenet\_signaling\_network
- consensuspathdb\_download

```
# use some parameters stricter than default:
cpdb_signaling_network <- nichenet_signaling_network_cpdb(
    complex_max_size = 2,
    min_score = .99
)</pre>
```

```
nichenet_signaling_network_evex
                        NicheNet signaling network from EVEX
```

# Description

Builds signaling network prior knowledge for NicheNet from the EVEX database.

## Usage

```
nichenet_signaling_network_evex(top_confidence = 0.75, indirect = FALSE, ...)
```

## **Arguments**

```
top_confidence Double, between 0 and 1. Threshold based on the quantile of the confidence
indirect
                  Logical: whether to include indirect interactions.
                  Ignored.
```

## Value

A network data frame (tibble) with signaling interactions suitable for use with NicheNet.

### See Also

- evex\_download
- nichenet\_signaling\_network

## **Examples**

```
ev_signaling_network <- nichenet_signaling_network_evex(</pre>
    top_confidence = .9
```

```
\verb|nichenet_signaling_network_harmonizome|\\
                          NicheNet signaling network from Harmonizome
```

# **Description**

Builds signaling network prior knowledge for NicheNet using Harmonizome

# Usage

```
nichenet_signaling_network_harmonizome(
 datasets = c("phosphositeplus", "kea", "depod"),
)
```

#### **Arguments**

datasets The datasets to use. For possible values please refer to default value and the Harmonizome webpage.

... Ignored.

#### Value

A network data frame (tibble) with signaling interactions suitable for use with NicheNet.

# **Examples**

```
# use only KEA and PhosphoSite:
hz_signaling_network <- nichenet_signaling_network_harmonizome(
    datasets = c('kea', 'phosphositeplus')
)</pre>
```

# Description

Builds signaling network prior knowledge for NicheNet from the InWeb InBioMap database.

# Usage

```
\verb|nichenet_signaling_network_inbiomap(...)|
```

# **Arguments**

```
... Ignored.
```

# Value

A network data frame (tibble) with signaling interactions suitable for use with NicheNet.

## See Also

```
nichenet_signaling_network, inbiomap_download
```

```
## Not run:
ib_signaling_network <- nichenet_signaling_network_inbiomap()
## End(Not run)</pre>
```

```
nichenet_signaling_network_omnipath

Builds signaling network for NicheNet using OmniPath
```

## **Description**

Retrieves network prior knowledge from OmniPath and provides it in a format suitable for NicheNet. This method never downloads the 'ligrecextra' dataset because the ligand-receptor interactions are supposed to come from nichenet\_lr\_network\_omnipath.

### Usage

```
nichenet_signaling_network_omnipath(min_curation_effort = 0, ...)
```

### **Arguments**

```
min_curation_effort

Lower threshold for curation effort

... Passed to import_post_translational_interactions
```

#### Value

A network data frame (tibble) with signaling interactions suitable for use with NicheNet.

#### See Also

• nichenet\_signaling\_network

# **Examples**

```
# use interactions with at least 2 evidences (reference or database)
op_signaling_network <- nichenet_signaling_network_omnipath(
    min_curation_effort = 2
)</pre>
```

```
nichenet_signaling_network_pathwaycommons

NicheNet signaling network from PathwayCommons
```

### **Description**

Builds signaling network prior knowledge for NicheNet using PathwayCommons.

### Usage

```
nichenet_signaling_network_pathwaycommons(
  interaction_types = c("catalysis-precedes", "controls-phosphorylation-of",
    "controls-state-change-of", "controls-transport-of", "in-complex-with",
    "interacts-with"),
    ...
)
```

### **Arguments**

```
interaction_types
```

Character vector with PathwayCommons interaction types. Please refer to the default value and the PathwayCommons webpage.

... Ignored.

#### Value

A network data frame (tibble) with signaling interactions suitable for use with NicheNet.

# **Examples**

```
# use only the "controls-transport-of" interactions:
pc_signaling_network <- nichenet_signaling_network_pathwaycommons(
   interaction_types = 'controls-transport-of'
)</pre>
```

nichenet\_signaling\_network\_vinayagam

NicheNet signaling network from Vinayagam

## **Description**

Builds signaling network prior knowledge for NicheNet using Vinayagam 2011 Supplementary Table S6. Find out more at https://doi.org/10.1126/scisignal.2001699.

# Usage

```
nichenet_signaling_network_vinayagam(...)
```

## **Arguments**

... Ignored.

## Value

A network data frame (tibble) with signaling interactions suitable for use with NicheNet.

```
vi_signaling_network <- nichenet_signaling_network_vinayagam()</pre>
```

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nichenet\_test

Run the NicheNet pipeline with a little dummy network

### **Description**

Loads a tiny network and runs the NicheNet pipeline with low number of iterations in the optimization process. This way the pipeline runs in a reasonable time in order to test the code. Due to the random subsampling disconnected networks might be produced sometimes. If you see an error like "Error in if (sd(prediction\_vector) == 0) ... missing value where TRUE/FALSE needed", the random subsampled input is not appropriate. In this case just interrupt and call again. This test ensures the computational integrity of the pipeline. If it fails during the optimization process, try to start it over several times, even restarting R. The unpredictability is related to mlrMBO and nichenetr not being prepared to handle certain conditions, and it's also difficult to find out which conditions lead to which errors. At least 3 different errors appear time to time, depending on the input. It also seems like restarting R sometimes helps, suggesting that the entire system might be somehow stateful. You can ignore the Parallelization was not stopped warnings on repeated runs.

### Usage

```
nichenet_test(...)
```

## **Arguments**

... Passed to nichenet\_main.

# Value

A named list with the intermediate and final outputs of the pipeline: 'networks', 'expression', 'optimized\_parameters', 'weighted\_networks' and 'ligand\_target\_matrix'.

# **Examples**

```
## Not run:
nnt <- nichenet_test()
## End(Not run)</pre>
```

nichenet\_workarounds

Workarounds using NicheNet without attaching the package

### **Description**

NicheNet requires the availability of some lazy loaded external data which are not available if the package is not loaded and attached. Also, the BBmisc::convertToShortString used for error reporting in mlrMBO::evalTargetFun.OptState is patched here to print longer error messages. Maybe it's a better solution to attach nichenetr before running the NicheNet pipeline. Alternatively you can try to call this function in the beginning. Why we don't call this automatically is just because we don't want to load datasets from another package without the user knowing about it.

obo\_parser

#### Usage

```
nichenet_workarounds()
```

#### Value

Returns NULL.

### **Examples**

```
## Not run:
nichenet_workarounds()
## End(Not run)
```

obo\_parser

Generic OBO parser

#### **Description**

Reads the contents of an OBO file and processes it into data frames or a list based data structure.

## Usage

## **Arguments**

path Path to the OBO file.

relations Character vector: process only these relations.

shorten\_namespace

Logical: shorten the namespace to a single letter code (as usual for Gene Ontol-

ogy, e.g. cellular\_component = "C").

tables Logical: return data frames (tibbles) instead of nested lists.

### Value

A list with the following elements: 1) "names" a list with terms as names and names as values; 2) "namespaces" a list with terms as names and namespaces as values; 3) "relations" a list with relations between terms: terms are keys, values are lists with relations as names and character vectors of related terms as values; 4) "subsets" a list with terms as keys and character vectors of subset names as values (or NULL if the term does not belong to any subset); 5) "obsolete" character vector with all the terms labeled as obsolete. If the tables parameter is TRUE, "names", "namespaces", "relations" and "subsets" will be data frames (tibbles).

oma\_code 133

### See Also

```
• relations_list_to_table
```

- relations\_table\_to\_list
- swap\_relations

## **Examples**

```
goslim_url <-</pre>
    "http://current.geneontology.org/ontology/subsets/goslim_generic.obo"
path <- tempfile()</pre>
httr::GET(goslim_url, httr::write_disk(path, overwrite = TRUE))
obo <- obo_parser(path, tables = FALSE)</pre>
unlink(path)
names(obo)
# [1] "names"
                   "namespaces" "relations" "subsets"
                                                             "obsolete"
head(obo\$relations, n = 2)
# $`GO:0000001`
# $`GO:0000001`$is_a
# [1] "GO:0048308" "GO:0048311"
# $`GO:0000002`
# $`GO:0000002`$is_a
# [1] "GO:0007005"
```

oma\_code

Orthologous Matrix (OMA) codes of organisms

# Description

Note: OMA species codes are whenever possible identical to UniProt codes.

# Usage

```
oma_code(name)
```

## **Arguments**

name

Vector with any kind of organism name or identifier, can be also mixed type.

## Value

A character vector with the Orthologous Matrix (OMA) codes of the organisms.

### See Also

- ncbi\_taxid
- latin\_name
- ensembl\_name
- common\_name

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## **Examples**

```
oma_code(c(10090, "cjacchus", "Vicugna pacos"))
# [1] "MOUSE" "CALJA" "VICPA"
```

oma\_organisms

Organism identifiers from the Orthologous Matrix

# **Description**

Organism identifiers from the Orthologous Matrix

# Usage

```
oma_organisms()
```

### Value

A data frame with organism identifiers.

### See Also

```
ensembl_organisms
```

# **Examples**

```
oma_organisms()
```

oma\_pairwise

Orthologous gene pairs between two organisms

# Description

From the web API of Orthologous Matrix (OMA). Items which could not be translated to 'id\_type' (but present in the data with their internal OMA IDs) are removed.

## Usage

```
oma_pairwise(
  organism_a = "human",
  organism_b = "mouse",
  id_type = "uniprot",
  mappings = c("1:1", "1:m", "n:1", "n:m"),
  only_ids = TRUE
)
```

### **Arguments**

organism_a	Name or identifier of an organism.
organism_b	Name or identifier of another organism.
id_type	The gene or protein identifier to use in the table. For a list of supported ID types see 'omnipathr.env\$id_types\$oma'. In addition, "genesymbol" is supported, in this case oma_pairwise_genesymbols will be called automatically.
mappings	Character vector: control ambiguous mappings:
	• 1:1 - unambiguous
	• 1:m - one-to-many
	• n:1 - many-to-one
	• n:m - many-to-many
only_ids	Logical: include only the two identifier columns, not the mapping type and the orthology group columns.

### Value

A data frame with orthologous gene pairs.

## **Examples**

```
oma_pairwise("human", "mouse", "uniprot")
# # A tibble: 21,753 × 4
#
     id_organism_a id_organism_b mapping oma_group
#
    <chr>
                  <chr>
                                <chr>
                                            <dbl>
#
  1 015326
                  Q8R5C8
                                1:1
                                          1129380
  2 Q9Y2E4
                  B2RQ71
                                1:1
                                          681224
  3 Q92615
                  06A0A2
                                1:1
                                          1135087
  4 Q9BZE4
                  Q99ME9
                                1:1
                                          1176239
# 5 Q9BXS1
                  Q8BFZ6
                                1:m
                                               NA
# # . . . with 21,743 more rows
```

```
oma_pairwise_genesymbols
```

Orthologous pairs of gene symbols between two organisms

# Description

The Orthologous Matrix (OMA), a resource of orthologous relationships between genes, doesn't provide gene symbols, the identifier preferred in many bioinformatics pipelines. Hence this function wraps oma\_pairwise by translating the identifiers used in OMA to gene symbols. Items that can not be translated to 'id\_type' (but present in the data with their internal OMA IDs) will be removed. Then, in this function we translate the identifiers to gene symbols.

#### **Usage**

```
oma_pairwise_genesymbols(
  organism_a = "human",
  organism_b = "mouse",
  oma_id_type = "uniprot_entry",
  mappings = c("1:1", "1:m", "n:1", "n:m"),
  only_ids = TRUE
)
```

### **Arguments**

organism\_a Name or identifier of an organism.

organism\_b Name or identifier of another organism.

oma\_id\_type Character: the gene or protein identifier to be queried from OMA. These IDs will be translated to 'id\_type'.

mappings Character vector: control ambiguous mappings:

• 1:1 - unambiguous

• 1:m - one-to-many

• n:1 - many-to-one

• n:m - many-to-many

only\_ids Logical: include only the two identifier columns, not the mapping type and the

### Value

A data frame with orthologous gene pairs.

orthology group columns.

#### **Examples**

```
oma_pairwise_genesymbols("human", "mouse")
```

```
oma_pairwise_translated
```

Orthologous pairs between two organisms for ID types not supported by OMA

# Description

The Orthologous Matrix (OMA), a resource of orthologous relationships between genes, doesn't provide gene symbols, the identifier preferred in many bioinformatics pipelines. Hence this function wraps oma\_pairwise by translating the identifiers used in OMA to gene symbols. Items that can not be translated to 'id\_type' (but present in the data with their internal OMA IDs) will be removed. Then, in this function we translate the identifiers to the desired ID type.

#### Usage

```
oma_pairwise_translated(
  organism_a = "human",
  organism_b = "mouse",
  id_type = "uniprot",
  oma_id_type = "uniprot_entry",
  mappings = c("1:1", "1:m", "n:1", "n:m"),
  only_ids = TRUE
)
```

# **Arguments**

Name or identifier of an organism. organism\_a organism\_b Name or identifier of another organism. id\_type The gene or protein identifier to use in the table. For a list of supported ID types see 'omnipathr.env\$id\_types\$oma'. These are the identifiers that will be translated to gene symbols. Character: the gene or protein identifier to be queried from OMA. These IDs oma\_id\_type will be translated to 'id\_type'. mappings Character vector: control ambiguous mappings: • 1:1 - unambiguous • 1:m - one-to-many • n:1 - many-to-one • n:m - many-to-many only\_ids Logical: include only the two identifier columns, not the mapping type and the orthology group columns.

#### Value

A data frame with orthologous gene pairs.

### **Examples**

```
oma_pairwise_translated("human", "mouse")
```

omnipath-interactions Molecular interactions from OmniPath

### **Description**

The functions listed here all download pairwise, causal molecular interactions from the <a href="https://omnipathdb.org/interactions">https://omnipathdb.org/interactions</a> endpoint of the OmniPath web service. They are different only in the type of interactions and the kind of resources and data they have been compiled from. A complete list of these functions is available below, these cover the interaction datasets and types currently available in OmniPath:

Interactions from the <a href="https://omnipathdb.org/interactions">https://omnipathdb.org/interactions</a> endpoint of the OmniPath web service. By default, it downloads only the "omnipath" dataset, which corresponds to the curated causal interactions described in Turei et al. 2016.

Imports interactions from the 'omnipath' dataset of OmniPath, a dataset that inherits most of its design and contents from the original OmniPath core from the 2016 publication. This dataset consists of about 40k interactions.

Imports the dataset from: https://omnipathdb.org/interactions?datasets=pathwayextra, which contains activity flow interactions without literature reference. The activity flow interactions supported by literature references are part of the 'omnipath' dataset.

Imports the dataset from: https://omnipathdb.org/interactions?datasets=kinaseextra, which contains enzyme-substrate interactions without literature reference. The enzyme-substrate interactions supported by literature references are part of the 'omnipath' dataset.

Imports the dataset from: https://omnipathdb.org/interactions?datasets=ligrecextra, which contains ligand-receptor interactions without literature reference. The ligand-receptor interactions supported by literature references are part of the 'omnipath' dataset.

Imports interactions from all post-translational datasets of OmniPath. The datasets are "omnipath", "kinaseextra", "pathwayextra" and "ligrecextra".

Imports the dataset from: https://omnipathdb.org/interactions?datasets=dorothea which contains transcription factor (TF)-target interactions from DoRothEA https://github.com/saezlab/DoRothEA DoRothEA is a comprehensive resource of transcriptional regulation, consisting of 16 original resources, in silico TFBS prediction, gene expression signatures and ChIP-Seq binding site analysis.

Imports the dataset from: https://omnipathdb.org/interactions?datasets=tf\_target, which contains transcription factor-target protein coding gene interactions. Note: this is not the only TF-target dataset in OmniPath, 'dorothea' is the other one and the 'tf\_mirna' dataset provides TF-miRNA gene interactions.

Imports the dataset from: https://omnipathdb.org/interactions?datasets=tf\_target, dorothea, which contains transcription factor-target protein coding gene interactions.

CollecTRI is a comprehensive resource of transcriptional regulation, published in 2023, consisting of 14 resources and original literature curation.

Imports the dataset from: https://omnipathdb.org/interactions?datasets=mirnatarget, which contains miRNA-mRNA interactions.

Imports the dataset from: https://omnipathdb.org/interactions?datasets=tf\_mirna, which contains transcription factor-miRNA gene interactions

Imports the dataset from: https://omnipathdb.org/interactions?datasets=lncrna\_mrna, which contains lncRNA-mRNA interactions

Imports the dataset from: https://omnipathdb.org/interactions?datasets=small\_molecule, which contains small molecule-protein interactions. Small molecules can be metabolites, intrinsic ligands or drug compounds.

#### Usage

```
omnipath_interactions(...)
omnipath(...)
pathwayextra(...)
kinaseextra(...)
ligrecextra(...)
```

```
post_translational(...)

dorothea(dorothea_levels = c("A", "B"), ...)

tf_target(...)

transcriptional(dorothea_levels = c("A", "B"), ...)

collectri(...)

mirna_target(...)

tf_mirna(...)

lncrna_mrna(...)

small_molecule(...)

all_interactions(
  dorothea_levels = c("A", "B"),
  types = NULL,
  fields = NULL,
  exclude = NULL,
  ...
)
```

#### **Arguments**

. . .

Arguments passed on to omnipath\_query, omnipath\_query

organism Character or integer: name or NCBI Taxonomy ID of the organism. OmniPath is built of human data, and the web service provides orthology translated interactions and enzyme-substrate relationships for mouse and rat. For other organisms and query types, orthology translation will be called automatically on the downloaded human data before returning the result.

resources Character vector: name of one or more resources. Restrict the data to these resources. For a complete list of available resources, call the '<query\_type>\_resources' functions for the query type of interst.

datasets Character vector: name of one or more datasets. In the interactions query type a number of datasets are available. The default is caled "omnipath", and corresponds to the curated causal signaling network published in the 2016 OmniPath paper.

genesymbols Character or logical: TRUE or FALS or "yes" or "no". Include the 'genesymbols' column in the results. OmniPath uses UniProt IDs as the primary identifiers, gene symbols are optional.

default\_fields Logical: if TRUE, the default fields will be included.

silent Logical: if TRUE, no messages will be printed. By default a summary message is printed upon successful download.

logicals Character vector: fields to be cast to logical.

> format Character: if "json", JSON will be retrieved and processed into a nested list; any other value will return data frame.

- download\_args List: parameters to pass to the download function, which is 'readr::read\_tsv' by default, and 'jsonlite::safe\_load'.
- references\_by\_resource Logical: if TRUE,, in the 'references' column the PubMed IDs will be prefixed with the names of the resources they are coming from. If FALSE, the 'references' column will be a list of unique PubMed IDs.
- add\_counts Logical: if TRUE, the number of references and number of resources for each record will be added to the result.
- license Character: license restrictions. By default, data from resources allowing "academic" use is returned by OmniPath. If you use the data for work in a company, you can provide "commercial" or "for-profit", which will restrict the data to those records which are supported by resources that allow for-profit use.
- password Character: password for the OmniPath web service. You can provide a special password here which enables the use of 'license = "ignore"' option, completely bypassing the license filter.
- json\_param List: parameters to pass to the 'jsonlite::fromJSON' when processing JSON columns embedded in the downloaded data. Such columns are "extra attrs" and "evidences". These are optional columns which provide a lot of extra details about interactions.
- strict\_evidences Logical: reconstruct the "sources" and "references" columns of interaction data frames based on the "evidences" column, strictly filtering them to the gueried datasets and resources. Without this, the "sources" and "references" fields for each record might contain information for datasets and resources other than the queried ones, because the downloaded records are a result of a simple filtering of an already integrated data frame.
- genesymbol\_resource Character: "uniprot" (default) or "ensembl". The OmniPath web service uses the primary gene symbols as provided by UniProt. By passing "ensembl" here, the UniProt gene symbols will be replaced by the ones used in Ensembl. This translation results in a loss of a few records, and multiplication of another few records due to ambiguous translation.
- cache Logical: use caching, load data from and save to the. The cache directory by default belongs to the user, located in the user's default cache directory, and named "OmnipathR". Find out about it by getOption("omnipathr.cachedir"). Can be changed by omnipath\_set\_cachedir.

## dorothea\_levels

The confidence levels of the dorothea interactions (TF-target) which range from A to D. Set to A and B by default.

Character: interaction types, such as "transcriptional", "post\_transcriptional", "post translational", etc.

Character: additional fields (columns) to be included in the result. For a list of available fields, see query\_info.

Character: names of datasets or resource to be excluded from the result. By deafult, the records supported by only these resources or datasets will be removed from the output. If strict\_evidences = TRUE, the resource, reference and causality information in the data frame will be reconstructed to remove all information coming from the excluded resources.

types

fields

exclude

#### **Details**

# Post-translational (protein-protein, PPI) interactions

• omnipath: the OmniPath data as defined in the 2016 paper, an arbitrary optimum between coverage and quality. This dataset contains almost entirely causal (stimulatory or inhibitory; i.e. activity flow, according to the SBGN standard), physical interactions between pairs of proteins, curated by experts from the literature.

- pathwayextra: activity flow interactions without literature references.
- kinaseextra: enzyme-substrate interactions without literature references.
- ligrecextra: ligand-receptor interactions without literature references.
- post\_translational: all post-translational (protein-protein, PPI) interactions; this is the combination of the *omnipath*, *pathwayextra*, *kinaseextra* and *ligrecextra* datasets.

## TF-target (gene regulatory, GRN) interactions

- collectri: transcription factor (TF)-target interactions from CollecTRI.
- dorothea: transcription factor (TF)-target interactions from DoRothEA
- tf\_target: transcription factor (TF)-target interactions from other resources
- transcriptional: all transcription factor (TF)-target interactions; this is the combination of the *collectri*, *dorothea* and *tf\_target* datasets.

### Post-transcriptional (miRNA-target) and other RNA related interactions

In these datasets we intend to collect the literature curated resources, hence we don't include some of the most well known large databases if those are based on predictions or high-throughput assays.

- mirna\_target: miRNA-mRNA interactions
- tf\_mirna: TF-miRNA interactions
- lncrna\_mrna: lncRNA-mRNA interactions

#### Other interaction access functions

- small\_molecule: interactions between small molecules and proteins. Currently this is a small, experimental dataset that includes drug-target, ligand-receptor, enzyme-metabolite and other interactions. In the future this will be largely expanded and divided into multiple datasets.
- all\_interactions: all the interaction datasets combined.

### Value

A dataframe of molecular interactions.

A dataframe of literature curated, post-translational signaling interactions.

A dataframe containing activity flow interactions between proteins without literature reference

A dataframe containing enzyme-substrate interactions without literature reference

A dataframe containing ligand-receptor interactions including the ones without literature references

A dataframe containing post-translational interactions

A data frame of TF-target interactions from DoRothEA.

A dataframe containing TF-target interactions

A dataframe containing TF-target interactions.

A dataframe of TF-target interactions.

A dataframe containing miRNA-mRNA interactions

A dataframe containing TF-miRNA interactions

A dataframe containing lncRNA-mRNA interactions

A dataframe of small molecule-protein interactions

A dataframe containing all the datasets in the interactions query

## See Also

- interaction\_resources
- interaction\_graph
- print\_interactions
- annotated\_network
- omnipath\_interactions
- post\_translational
- interaction\_resources
- all\_interactions
- interaction\_graph
- print\_interactions

```
op <- omnipath(resources = c("CA1", "SIGNOR", "SignaLink3"))</pre>
interactions = omnipath_interactions(
    resources = "SignaLink3",
    organism = 9606
)
pathways <- omnipath()</pre>
pathways
interactions <-
    pathwayextra(
        resources = c("BioGRID", "IntAct"),
        organism = 9606
    )
kinase_substrate <-
   kinaseextra(
       resources = c('PhosphoPoint', 'PhosphoSite'),
       organism = 9606
   )
ligand_receptor <- ligrecextra(</pre>
    resources = c('HPRD', 'Guide2Pharma'),
    organism = 9606
)
```

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```
interactions <- post_translational(resources = "BioGRID")</pre>
dorothea_grn <- dorothea(</pre>
    resources = c('DoRothEA', 'ARACNe-GTEx_DoRothEA'),
    organism = 9606,
    dorothea_levels = c('A', 'B', 'C')
dorothea_grn
interactions <- tf_target(resources = c("DoRothEA", "SIGNOR"))</pre>
grn <- transcriptional(resources = c("PAZAR", "ORegAnno", "DoRothEA"))</pre>
collectri_grn <- collectri()</pre>
collectri_grn
interactions <- mirna_target( resources = c("miRTarBase", "miRecords"))</pre>
interactions <- tf_mirna(resources = "TransmiR")</pre>
interactions <- lncrna_mrna(resources = c("ncRDeathDB"))</pre>
# What are the targets of aspirin?
interactions <- small_molecule(sources = "ASPIRIN")</pre>
# The prostaglandin synthases:
interactions
interactions <- all_interactions(</pre>
    resources = c("HPRD", "BioGRID"),
    organism = 9606
```

OmnipathR

The OmnipathR package

#### **Description**

OmnipathR is an R package built to provide easy access to the data stored in the OmniPath web service:

```
https://omnipathdb.org/
```

And a number of other resources, such as BioPlex, ConsensusPathDB, EVEX, Guide to Pharmacology (IUPHAR/BPS), Harmonizome, HTRIdb, InWeb InBioMap, KEGG Pathway, Pathway Commons, Ramilowski et al. 2015, RegNetwork, ReMap, TF census, TRRUST and Vinayagam et al. 2011.

The OmniPath web service implements a very simple REST style API. This package make requests by the HTTP protocol to retreive the data. Hence, fast Internet access is required for a propser use of OmnipathR.

The package also provides some utility functions to filter, analyse and visualize the data. Furthermore, OmnipathR features a close integration with the NicheNet method for ligand activity prediction from transcriptomics data, and its R implementation nichenetr (available in CRAN).

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#### Author(s)

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### See Also

Useful links:

- https://r.omnipathdb.org/
- Report bugs at https://github.com/saezlab/OmnipathR/issues

```
## Not run:
# Download post-translational modifications:
enzsub <- enzyme_substrate(resources = c("PhosphoSite", "SIGNOR"))</pre>
# Download protein-protein interactions
interactions <- omnipath(resources = "SignaLink3")</pre>
# Convert to igraph objects:
enzsub_g <- enzsub_graph(enzsub = enzsub)</pre>
OPI_g <- interaction_graph(interactions = interactions)</pre>
# Print some interactions:
print_interactions(head(enzsub))
# interactions with references:
print_interactions(tail(enzsub), writeRefs = TRUE)
# find interactions between kinase and substrate:
print_interactions(dplyr::filter(ptms,enzyme_genesymbol=="MAP2K1",
   substrate_genesymbol=="MAPK3"))
# find shortest paths on the directed network between proteins
print_path_es(shortest_paths(OPI_g, from = "TYRO3", to = "STAT3",
   output = 'epath')$epath[[1]], OPI_g)
# find all shortest paths between proteins
print_path_vs(
    all\_shortest\_paths(
        enzsub_g,
        from = "SRC",
        to = "STAT1"
    )$res,
    enzsub_g
)
## End(Not run)
```

```
omnipath_cache_autoclean
```

Keeps only the latest versions of complete downloads

# Description

Removes the old versions, the failed downloads and the files in the cache directory which are missing from the database. For more flexible operations use omnipath\_cache\_remove and omnipath\_cache\_clean.

# Usage

```
omnipath_cache_autoclean()
```

### Value

Invisibl returns the cache database (list of cache records).

# **Examples**

```
## Not run:
omnipath_cache_autoclean()
## End(Not run)
```

omnipath\_cache\_clean

Removes the items from the cache directory which are unknown by the cache database

### **Description**

Removes the items from the cache directory which are unknown by the cache database

### Usage

```
omnipath_cache_clean()
```

#### Value

Returns 'NULL'.

```
omnipath_cache_clean()
```

```
omnipath_cache_clean_db
```

Removes the cache database entries without existing files

# Description

Removes the cache database entries without existing files

# Usage

```
omnipath_cache_clean_db(...)
```

# **Arguments**

```
... Ignored.
```

#### Value

```
Returns 'NULL'.
```

# **Examples**

```
omnipath_cache_clean_db()
```

```
omnipath_cache_download_ready
```

Sets the download status to ready for a cache item

# Description

Sets the download status to ready for a cache item

# Usage

```
omnipath_cache_download_ready(version, key = NULL)
```

# **Arguments**

version Version of the cache item. If does not exist a new version item will be created

key Key of the cache item

### Value

Character: invisibly returns the version number of the cache version item.

### **Examples**

```
bioc_url <- 'https://bioconductor.org/'</pre>
# request a new version item (or retrieve the latest)
new_version <- omnipath_cache_latest_or_new(url = bioc_url)</pre>
# check if the version item is not a finished download
new_version$status
# [1] "unknown"
# download the file
httr::GET(bioc_url, httr::write_disk(new_version$path, overwrite = TRUE))
# report to the cache database that the download is ready
omnipath_cache_download_ready(new_version)
# now the status is ready:
version <- omnipath_cache_latest_or_new(url = bioc_url)</pre>
version$status
# "ready"
version$dl_finished
# [1] "2021-03-09 16:48:38 CET"
omnipath_cache_remove(url = bioc_url) # cleaning up
```

```
omnipath_cache_filter_versions
```

Filters the versions from one cache record

#### **Description**

Filters the versions based on multiple conditions: their age and status

# Usage

```
omnipath_cache_filter_versions(
  record,
  latest = FALSE,
  max_age = NULL,
  min_age = NULL,
  status = CACHE_STATUS$READY
)
```

### **Arguments**

record	A cache record
latest	Return the most recent version
max_age	The maximum age in days (e.g. 5: 5 days old or more recent)
min_age	The minimum age in days (e.g. 5: 5 days old or older)
status	Character vector with status codes. By default only the versions with 'ready' (completed download) status are selected

### Value

Character vector with version IDs, NA if no version satisfies the conditions.

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### **Examples**

```
# creating an example cache record
bioc_url <- 'https://bioconductor.org/'
version <- omnipath_cache_latest_or_new(url = bioc_url)
httr::GET(bioc_url, httr::write_disk(version$path, overwrite = TRUE))
omnipath_cache_download_ready(version)
record <- dplyr::first(omnipath_cache_search('biocond'))
# only the versions with status "ready"
version_numbers <- omnipath_cache_filter_versions(record, status = 'ready')
omnipath_cache_remove(url = bioc_url) # cleaning up</pre>
```

omnipath\_cache\_get

Retrieves one item from the cache directory

# Description

Retrieves one item from the cache directory

# Usage

```
omnipath_cache_get(
  key = NULL,
  url = NULL,
  post = NULL,
  payload = NULL,
  create = TRUE,
  ...
)
```

### **Arguments**

key	The key of the cache record
url	URL pointing to the resource
post	HTTP POST parameters as a list
payload	HTTP data payload
create	Create a new entry if doesn't exist yet
	Passed to omnipath_cache_record (internal function)

# Value

Cache record: an existing record if the entry already exists, otherwise a newly created and inserted record

omnipath\_cache\_key 149

### **Examples**

```
# create an example cache record
bioc_url <- 'https://bioconductor.org/'
version <- omnipath_cache_latest_or_new(url = bioc_url)
omnipath_cache_remove(url = bioc_url) # cleaning up

# retrieve the cache record
record <- omnipath_cache_get(url = bioc_url)
record$key
# [1] "41346a00fb20d2a9df03aa70cf4d50bf88ab154a"
record$url
# [1] "https://bioconductor.org/"</pre>
```

omnipath\_cache\_key

Generates a hash which identifies an element in the cache database

# Description

Generates a hash which identifies an element in the cache database

# Usage

```
omnipath_cache_key(url, post = NULL, payload = NULL)
```

### Arguments

url	Character vector with URLs
post	List with the HTTP POST parameters or a list of lists if the url vector is longer than 1. NULL for queries without POST parameters.
payload	HTTP data payload. List with multiple items if the url vector is longer than 1. NULL for queries without data.

#### Value

Character vector of cache record keys.

```
bioc_url <- 'https://bioconductor.org/'
omnipath_cache_key(bioc_url)
# [1] "41346a00fb20d2a9df03aa70cf4d50bf88ab154a"</pre>
```

```
omnipath_cache_latest_or_new
```

The latest or a new version of a cache record

# Description

Looks up a record in the cache and returns its latest valid version. If the record doesn't exist or no valid version available, creates a new one.

#### Usage

```
omnipath_cache_latest_or_new(
  key = NULL,
  url = NULL,
  post = NULL,
  payload = NULL,
  create = TRUE,
)
```

### **Arguments**

key	The key of the cache record
url	URL pointing to the resource
post	HTTP POST parameters as a list
payload	HTTP data payload
create	Logical: whether to create and return a new version. If FALSE only the latest existing valid version is returned, if available.
	Passed to omnipath_cache_get

# Value

A cache version item.

# **Examples**

```
## Not run:
# retrieve the latest version of the first cache record
# found by the search keyword "bioplex"
latest_bioplex <-</pre>
    omnipath_cache_latest_or_new(
        names(omnipath_cache_search('bioplex'))[1]
    )
latest\_bioplex\$dl\_finished
# [1] "2021-03-09 14:28:50 CET"
latest_bioplex$path
# [1] "/home/denes/.cache/OmnipathR/378e0def2ac97985f629-1.rds"
## End(Not run)
```

```
# create an example cache record
bioc_url <- 'https://bioconductor.org/'
version <- omnipath_cache_latest_or_new(url = bioc_url)
omnipath_cache_remove(url = bioc_url) # cleaning up</pre>
```

```
omnipath_cache_latest_version
```

Finds the most recent version in a cache record

#### **Description**

Finds the most recent version in a cache record

#### Usage

```
omnipath_cache_latest_version(record)
```

#### **Arguments**

record

A cache record

#### Value

Character: the version ID with the most recent download finished time

omnipath\_cache\_load

Loads an R object from the cache

### **Description**

Loads the object from RDS format.

# Usage

```
omnipath_cache_load(
  key = NULL,
  version = NULL,
  url = NULL,
  post = NULL,
  payload = NULL
)
```

### **Arguments**

key Key of the cache item

version Version of the cache item. If does not exist or NULL, the latest version will be

retrieved

url URL of the downloaded resource post HTTP POST parameters as a list

payload HTTP data payload

#### Value

Object loaded from the cache RDS file.

#### See Also

```
omnipath_cache_save
```

### **Examples**

```
url <- paste0(
    'https://omnipathdb.org/intercell?resources=Adhesome,Almen2009,',
    'Baccin2019, CSPA, CellChatDB&license=academic'
result <- read.delim(url, sep = '\t')
omnipath_cache_save(result, url = url)
# works only if you have already this item in the cache
intercell_data <- omnipath_cache_load(url = url)</pre>
class(intercell_data)
# [1] "data.frame"
nrow(intercell_data)
# [1] 16622
attr(intercell_data, 'origin')
# [1] "cache"
# basic example of saving and loading to and from the cache:
bioc_url <- 'https://bioconductor.org/'</pre>
bioc_html <- readChar(url(bioc_url), nchars = 99999)</pre>
omnipath_cache_save(bioc_html, url = bioc_url)
bioc_html <- omnipath_cache_load(url = bioc_url)</pre>
```

```
omnipath_cache_move_in
```

Moves an existing file into the cache

### **Description**

Either the key or the URL (with POST and payload) must be provided.

# Usage

```
omnipath_cache_move_in(
  path,
  key = NULL,
  version = NULL,
  url = NULL,
  post = NULL,
  payload = NULL,
  keep_original = FALSE
)
```

#### **Arguments**

path	Path to the source file
key	Key of the cache item

version Version of the cache item. If does not exist a new version item will be created

url URL of the downloaded resource post HTTP POST parameters as a list

payload HTTP data payload

keep\_original Whether to keep or remove the original file

#### Value

Character: invisibly returns the version number of the cache version item.

#### See Also

```
omnipath_cache_save
```

#### **Examples**

```
path <- tempfile()
saveRDS(rnorm(100), file = path)
omnipath_cache_move_in(path, url = 'the_download_address')

# basic example of moving a file to the cache:

bioc_url <- 'https://bioconductor.org/'
html_file <- tempfile(fileext = '.html')
httr::GET(bioc_url, httr::write_disk(html_file, overwrite = TRUE))
omnipath_cache_move_in(path = html_file, url = bioc_url)
omnipath_cache_remove(url = bioc_url) # cleaning up</pre>
```

omnipath\_cache\_remove Removes contents from the cache directory

# Description

According to the parameters, it can remove contents older than a certain age, or contents having a more recent version, one specific item, or wipe the entire cache.

#### Usage

```
omnipath_cache_remove(key = NULL, url = NULL, post = NULL,
    payload = NULL, max_age = NULL, min_age = NULL, status = NULL,
    only_latest = FALSE, wipe = FALSE, autoclean = TRUE)
```

### **Arguments**

key	The key of the cache record
url	URL pointing to the resource
post	HTTP POST parameters as a list
payload	HTTP data payload
max_age	Age of cache items in days. Remove everything that is older than this age
min_age	Age of cache items in days. Remove everything more recent than this age
status	Remove items having any of the states listed here
only_latest	Keep only the latest version
wipe	Logical: if TRUE, removes all files from the cache and the cache database. Same as calling omnipath_cache_wipe.
autoclean	Remove the entries about failed downloads, the files in the cache directory which are missing from the cache database, and the entries without existing files in the

# Value

Invisibly returns the cache database (list of cache records).

cache directory

### See Also

- omnipath\_cache\_wipe
- omnipath\_cache\_clean
- omnipath\_cache\_autoclean

```
# remove all cache data from the BioPlex database
cache_records <- omnipath_cache_search(</pre>
    'bioplex',
    ignore.case = TRUE
)
omnipath_cache_remove(names(cache_records))
# remove a record by its URL
regnetwork_url <- 'http://www.regnetworkweb.org/download/human.zip'</pre>
omnipath_cache_remove(url = regnetwork_url)
# remove all records older than 30 days
omnipath_cache_remove(max_age = 30)
\ensuremath{\text{\#}} for each record, remove all versions except the latest
omnipath_cache_remove(only_latest = TRUE)
## End(Not run)
bioc_url <- 'https://bioconductor.org/'</pre>
version <- omnipath_cache_latest_or_new(url = bioc_url)</pre>
httr::GET(bioc_url, httr::write_disk(version$path, overwrite = TRUE))
omnipath_cache_download_ready(version)
```

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```
key <- omnipath_cache_key(bioc_url)
omnipath_cache_remove(key = key)</pre>
```

omnipath\_cache\_save

Saves an R object to the cache

### **Description**

Exports the object in RDS format, creates new cache record if necessary.

### Usage

```
omnipath_cache_save(
  data,
  key = NULL,
  version = NULL,
  url = NULL,
  post = NULL,
  payload = NULL
)
```

### **Arguments**

data An object

key Key of the cache item

version Version of the cache item. If does not exist a new version item will be created

url URL of the downloaded resource post HTTP POST parameters as a list

payload HTTP data payload

# Value

Returns invisibly the data itself.

Invisibly returns the 'data'.

#### See Also

```
omnipath_cache_move_in
```

```
mydata <- data.frame(a = c(1, 2, 3), b = c('a', 'b', 'c'))
omnipath_cache_save(mydata, url = 'some_dummy_address')
from_cache <- omnipath_cache_load(url = 'some_dummy_address')
from_cache
# a b
# 1 1 a
# 2 2 b
# 3 3 c</pre>
```

```
attr(from_cache, 'origin')
# [1] "cache"

# basic example of saving and loading to and from the cache:
bioc_url <- 'https://bioconductor.org/'
bioc_html <- readChar(url(bioc_url), nchars = 99999)
omnipath_cache_save(bioc_html, url = bioc_url)
bioc_html <- omnipath_cache_load(url = bioc_url)</pre>
```

# Description

Searches the cache records by matching the URL against a string or regexp.

#### Usage

```
omnipath_cache_search(pattern, ...)
```

# Arguments

```
pattern String or regular expression.
... Passed to grep
```

#### Value

List of cache records matching the pattern.

### **Examples**

```
# find all cache records from the BioPlex database
bioplex_cache_records <- omnipath_cache_search(
   'bioplex',
   ignore.case = TRUE
)
```

```
omnipath_cache_set_ext
```

Sets the file extension for a cache record

### **Description**

Sets the file extension for a cache record

#### Usage

```
omnipath_cache_set_ext(key, ext)
```

### **Arguments**

key Character: key for a cache item, alternatively a version entry.

ext Character: the file extension, e.g. "zip".

#### Value

Returns 'NULL'.

#### **Examples**

```
bioc_url <- 'https://bioconductor.org/'
version <- omnipath_cache_latest_or_new(url = bioc_url)
version$path
# [1] "/home/denes/.cache/OmnipathR/41346a00fb20d2a9df03-1"
httr::GET(bioc_url, httr::write_disk(version$path, overwrite = TRUE))
key <- omnipath_cache_key(url = bioc_url)
omnipath_cache_set_ext(key = key, ext = 'html')
version <- omnipath_cache_latest_or_new(url = bioc_url)
version$path
# [1] "/home/denes/.cache/OmnipathR/41346a00fb20d2a9df03-1.html"
record <- omnipath_cache_get(url = bioc_url)
record$ext
# [1] "html"
omnipath_cache_remove(url = bioc_url) # cleaning up</pre>
```

omnipath\_cache\_update\_status

Updates the status of an existing cache record

#### **Description**

Updates the status of an existing cache record

# Usage

```
omnipath_cache_update_status(key, version, status,
    dl_finished = NULL)
```

#### **Arguments**

key Key of the cache item

version Version of the cache item. If does not exist a new version item will be created

status The updated status value

dl\_finished Timestamp for the time when download was finished, if 'NULL' the value re-

mains unchanged

### Value

Character: invisibly returns the version number of the cache version item.

### **Examples**

```
bioc_url <- 'https://bioconductor.org/'
latest_version <- omnipath_cache_latest_or_new(url = bioc_url)
key <- omnipath_cache_key(bioc_url)
omnipath_cache_update_status(
    key = key,
    version = latest_version$number,
    status = 'ready',
    dl_finished = Sys.time()
)
omnipath_cache_remove(url = bioc_url) # cleaning up</pre>
```

omnipath\_cache\_wipe

Permanently removes all the cache contents

### **Description**

After this operation the cache directory will be completely empty, except an empty cache database file.

### Usage

```
omnipath_cache_wipe(...)
```

### **Arguments**

... Ignored.

#### Value

Returns 'NULL'.

# See Also

```
omnipath_cache_remove
```

```
## Not run:
omnipath_cache_wipe()
# the cache is completely empty:
print(omnipathr.env$cache)
# list()
list.files(omnipath_get_cachedir())
# [1] "cache.json"
## End(Not run)
```

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### **Description**

Current config file path of OmnipathR

Current config file path for a certain package

### Usage

```
omnipath_config_path(user = FALSE)
config_path(user = FALSE, pkg = "OmnipathR")
```

# **Arguments**

user Logical: prioritize the user level config even if a config in the current working

directory is available.

pkg Character: name of the package.

### Value

Character: path to the config file.

# **Examples**

```
omnipath_config_path()
```

omnipath\_for\_cosmos

OmniPath PPI for the COSMOS PKN

# Description

OmniPath PPI for the COSMOS PKN

### Usage

```
omnipath_for_cosmos(
  organism = 9606L,
  resources = NULL,
  datasets = NULL,
  interaction_types = NULL,
  id_types = c("uniprot", "genesymbol"),
  ...
)
```

#### **Arguments**

organism Character or integer: name or NCBI Taxonomy ID of the organism.

resources Character: names of one or more resources. Correct spelling is important.

datasets Character: one or more network datasets in OmniPath.

interaction\_types

Character: one or more interaction type

id\_types Character: translate the protein identifi

Character: translate the protein identifiers to these ID types. Each ID type results two extra columns in the output, for the "source" and "target" sides of the interaction, respectively. The default ID type for proteins is Esembl Gene ID, and by default UniProt IDs and Gene Symbols are included. The UniProt IDs returned by the web service are left intact, while the Gene Symbols are queried from Ensembl. These Gene Symbols are different from the ones returned from the web service, and match the Ensembl Gene Symbols used by other compo-

nents of the COSMOS PKN.

... Further parameters to omnipath\_interactions.

#### Value

Data frame with the columns source, target and sign.

### See Also

- cosmos\_pkn
- omnipath-interactions

# Examples

```
op_cosmos <- omnipath_for_cosmos()
op_cosmos</pre>
```

### **Description**

Load the package configuration from a config file Load the coniguration of a certain package

## Usage

```
omnipath_load_config(path = NULL, title = "default", user = FALSE, ...)

load_config(
  path = NULL,
  title = "default",
  user = FALSE,
  pkg = "OmnipathR",
  ...
)
```

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# **Arguments**

path	Path to the config file.
title	Load the config under this title. One config file might contain multple configurations, each identified by a title. If the title is not available the first section of the config file will be used.
user	Force to use the user level config even if a config file exists in the current directory. By default, the local config files have prioroty over the user level config.
	Passed to yaml::yaml.load_file.
pkg	Character: name of the package

#### Value

Invisibly returns the config as a list.

# **Examples**

```
## Not run:
# load the config from a custom config file:
omnipath_load_config(path = 'my_custom_omnipath_config.yml')
## End(Not run)
```

 $omnipath_log$ 

Browse the current OmnipathR log file

# Description

Browse the current OmnipathR log file Browse the latest log from a package

# Usage

```
omnipath_log()
read_log(pkg = "OmnipathR")
```

# **Arguments**

pkg Character: name of a package.

### Value

Returns 'NULL'.

### See Also

```
omnipath_logfile
```

omnipath\_logfile

### **Examples**

```
## Not run:
omnipath_log()
# then you can browse the log file, and exit with `q`
## End(Not run)
```

omnipath\_logfile

Path to the current OmnipathR log file

# Description

Path to the current OmnipathR log file

Path to the current logfile of a package

# Usage

```
omnipath_logfile()
logfile(pkg = "OmnipathR")
```

# Arguments

pkg

Character: name of a package.

### Value

Character: path to the current logfile, or NULL if no logfile is available.

# See Also

```
omnipath_log
```

```
omnipath_logfile()
# [1] "/home/denes/omnipathr/omnipathr-log/omnipathr-20210309-1642.log"
```

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omnipath\_msg

Dispatch a message to the OmnipathR logger

### **Description**

Any package or script can easily send log messages and establish a logging facility with the fantastic 'logger' package. This function serves the only purpose if you want to inject messages into the logger of OmnipathR. Otherwise we recommend to use the 'logger' package directly.

#### Usage

```
omnipath_msg(level, ...)
```

### **Arguments**

```
Character, numeric or class loglevel. A log level, if character one of the followings: "fatal", "error", "warn", "success", "info", "trace".
Arguments for string formatting, passed sprintf or str_glue.
```

#### Value

Returns 'NULL'.

# **Examples**

```
omnipath_msg(
    level = 'success',
    'Talking to you in the name of OmnipathR, my favourite number is %d',
    round(runif(1, 1, 10))
)
```

omnipath\_query

Download data from the OmniPath web service

# Description

This is the most generic method for accessing data from the OmniPath web service. All other functions retrieving data from OmniPath call this function with various parameters. In general, every query can retrieve data in tabular or JSON format, the tabular (data frame) being the default.

### Usage

```
omnipath_query(
  query_type,
  organism = 9606L,
  resources = NULL,
  datasets = NULL,
  types = NULL,
  genesymbols = "yes",
```

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```
fields = NULL,
 default_fields = TRUE.
 silent = FALSE,
 logicals = NULL,
 download_args = list(),
  format = "data.frame",
 references_by_resource = TRUE,
 add_counts = TRUE,
 license = NULL,
 password = NULL,
 exclude = NULL,
  json_param = list(),
  strict_evidences = FALSE,
 genesymbol_resource = "UniProt",
 cache = NULL,
)
```

#### **Arguments**

query\_type Character: "interactions", "enzsub", "complexes", "annotations", or "intercell".

organism Character or integer: name or NCBI Taxonomy ID of the organism. Omni-

Path is built of human data, and the web service provides orthology translated interactions and enzyme-substrate relationships for mouse and rat. For other organisms and query types, orthology translation will be called automatically on

the downloaded human data before returning the result.

resources Character vector: name of one or more resources. Restrict the data to these re-

sources. For a complete list of available resources, call the '<query\_type>\_resources'

functions for the query type of interst.

datasets Character vector: name of one or more datasets. In the interactions query type

a number of datasets are available. The default is called "omnipath", and corresponds to the curated causal signaling network published in the 2016~OmniPath

paper.

types Character vector: one or more interaction types, such as "transcriptional" or

"post\_translational". For a full list of interaction types see 'query\_info("interaction")\$types'.

genesymbols Character or logical: TRUE or FALS or "yes" or "no". Include the 'genesym-

bols' column in the results. OmniPath uses UniProt IDs as the primary identi-

fiers, gene symbols are optional.

fields Character vector: additional fields to include in the result. For a list of available

fields, call 'query\_info("interactions")'.

default\_fields Logical: if TRUE, the default fields will be included.

silent Logical: if TRUE, no messages will be printed. By default a summary message

is printed upon successful download.

logicals Character vector: fields to be cast to logical.

download\_args List: parameters to pass to the download function, which is 'readr::read tsv' by

default, and 'jsonlite::safe\_load'.

format Character: if "json", JSON will be retrieved and processed into a nested list; any

other value will return data frame.

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references\_by\_resource

Logical: if TRUE,, in the 'references' column the PubMed IDs will be prefixed with the names of the resources they are coming from. If FALSE, the 'references' column will be a list of unique PubMed IDs.

Logical: if TRUE, the number of references and number of resources for each add\_counts

record will be added to the result.

license Character: license restrictions. By default, data from resources allowing "aca-

demic" use is returned by OmniPath. If you use the data for work in a company, you can provide "commercial" or "for-profit", which will restrict the data to

those records which are supported by resources that allow for-profit use.

Character: password for the OmniPath web service. You can provide a special password here which enables the use of 'license = "ignore" option, completely

bypassing the license filter.

exclude Character vector: resource or dataset names to be excluded. The data will be

filtered after download to remove records of the excluded datasets and resources.

json\_param List: parameters to pass to the 'isonlite::fromJSON' when processing JSON

> columns embedded in the downloaded data. Such columns are "extra attrs" and "evidences". These are optional columns which provide a lot of extra details

about interactions.

strict\_evidences

password

Logical: reconstruct the "sources" and "references" columns of interaction data frames based on the "evidences" column, strictly filtering them to the queried datasets and resources. Without this, the "sources" and "references" fields for each record might contain information for datasets and resources other than the queried ones, because the downloaded records are a result of a simple filtering of an already integrated data frame.

genesymbol\_resource

Character: "uniprot" (default) or "ensembl". The OmniPath web service uses the primary gene symbols as provided by UniProt. By passing "ensembl" here, the UniProt gene symbols will be replaced by the ones used in Ensembl. This translation results in a loss of a few records, and multiplication of another few

records due to ambiguous translation.

cache Logical: use caching, load data from and save to the. The cache directory by default belongs to the user, located in the user's default cache directory, and named

"OmnipathR". Find out about it by getOption("omnipathr.cachedir"). Can

be changed by omnipath\_set\_cachedir.

Additional parameters for the OmniPath web service. These parameters will be processed, validated and included in the query string. Many parameters are already explicitly set by the arguments above. A number of query type specific parameters are also available, learn more about these by the query\_info function. For functions more specific than omnipath\_query, arguments for all

downstream functions are also passed here.

### Value

Data frame (tibble) or list: the data returned by the OmniPath web service (or loaded from cache), after processing. Nested list if the "format" parameter is "json", otherwise a tibble.

#### **Examples**

interaction\_data <- omnipath\_query("interaction", datasets = "omnipath")</pre>

omnipath\_save\_config

```
interaction_data
```

# Description

Save the current package configuration

Save the configuration of a certain package

# Usage

```
omnipath_save_config(path = NULL, title = "default", local = FALSE)
save_config(path = NULL, title = "default", local = FALSE, pkg = "OmnipathR")
```

# Arguments

path	Path to the config file. Directories and the file will be created if don't exist.
title	Save the config under this title. One config file might contain multiple configurations, each identified by a title.
local	Save into a config file in the current directory instead of a user level config file. When loading, the config in the current directory has priority over the user level config.
pkg	Character: name of the package

# Value

Returns 'NULL'.

```
## Not run:
# after this, all downloads will default to commercial licenses
# i.e. the resources that allow only academic use will be excluded:
options(omnipathr.license = 'commercial')
omnipath_save_config()
## End(Not run)
```

omnipath\_set\_cachedir Change the cache directory

# Description

Change the cache directory

# Usage

```
omnipath_set_cachedir(path = NULL)
```

### **Arguments**

path

Character: path to the new cache directory. If don't exist, the directories will be created. If the path is an existing cache directory, the package's cache database for the current session will be loaded from the database in the directory. If NULL, the cache directory will be set to its default path.

#### Value

Returns NULL.

# **Examples**

```
tmp_cache <- tempdir()
omnipath_set_cachedir(tmp_cache)
# restore the default cache directory:
omnipath_set_cachedir()</pre>
```

```
omnipath_set_console_loglevel
```

Sets the log level for the console

# Description

Use this method to change during a session which messages you want to be printed on the console. Before loading the package, you can set it also by the config file, with the omnipathr.console\_loglevel key.

# Usage

```
omnipath_set_console_loglevel(level)
```

### **Arguments**

level

Character or class 'loglevel'. The desired log level.

#### Value

Returns 'NULL'.

#### See Also

```
omnipath_set_logfile_loglevel
```

# **Examples**

```
omnipath_set_console_loglevel('warn')
# or:
omnipath_set_console_loglevel(logger::WARN)
```

```
omnipath_set_logfile_loglevel

Sets the log level for the logfile
```

# Description

Use this method to change during a session which messages you want to be written into the logfile. Before loading the package, you can set it also by the config file, with the "omnipathr.loglevel" key.

# Usage

```
omnipath_set_logfile_loglevel(level)
```

# Arguments

level

Character or class 'loglevel'. The desired log level.

#### Value

Returns 'NULL'.

# See Also

```
omnipath_set_console_loglevel
```

```
omnipath_set_logfile_loglevel('info')
# or:
omnipath_set_logfile_loglevel(logger::INFO)
```

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```
omnipath_set_loglevel Sets the log level for the package logger
```

# Description

```
Sets the log level for the package logger
Sets the log level for any package
```

### Usage

```
omnipath_set_loglevel(level, target = "logfile")
set_loglevel(level, target = "logfile", pkg = "OmnipathR")
```

### **Arguments**

level Character or class 'loglevel'. The desired log level.

target Character, either 'logfile' or 'console'

pkg Character: name of the package.

#### Value

```
Returns 'NULL'.
```

# **Examples**

```
omnipath_set_loglevel(logger::FATAL, target = 'console')
```

omnipath\_show\_db

Built in database definitions

# Description

Databases are resources which might be costly to load but can be used many times by functions which usually automatically load and retrieve them from the database manager. Each database has a lifetime and will be unloaded automatically upon expiry.

### Usage

```
omnipath_show_db()
```

### Value

A data frame with the built in database definitions.

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#### **Examples**

```
database_definitions <- omnipath_show_db()</pre>
{\tt database\_definitions}
# # A tibble: 14 x 10
            last_used lifetime package loader
    name
                                                              loader_p.
#
    <chr>
               <dttm>
                                     <dbl> <chr> <chr>
                                                              st>
# 1 Gene Onto. 2021-04-04 20:19:15
                                       300 Omnipat. go_ontol. <named 1.
  2 Gene Onto. NA
                                       300 Omnipat. go_ontol. <named 1.
  3 Gene Onto. NA
                                       300 Omnipat. go_ontol. <named 1.
# 4 Gene Onto. NA
                                       300 Omnipat. go_ontol. <named 1.
# 5 Gene Onto. NA
                                       300 Omnipat. go_ontol. <named 1.
# ... (truncated)
# # . with 4 more variables: latest_param <list>, loaded <lgl>, db <list>,
# # key <chr>
```

```
omnipath_unlock_cache_db
```

Removes the lock file from the cache directory

# Description

A lock file in the cache directory avoids simulatneous write and read. It's supposed to be removed after each read and write operation. This might not happen if the process crashes during such an operation. In this case you can manually call this function.

#### Usage

```
omnipath_unlock_cache_db()
```

#### Value

Logical: returns TRUE if the cache was locked and now is unlocked; FALSE if it was not locked.

# Examples

```
omnipath_unlock_cache_db()
```

 $\verb"only_from"$ 

Recreate interaction data frame based on certain datasets and resources

# Description

Recreate interaction data frame based on certain datasets and resources

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#### Usage

```
only_from(
  data,
  datasets = NULL,
  resources = NULL,
  exclude = NULL,
  .keep = FALSE
)
```

#### **Arguments**

data An interaction data frame from the OmniPath web service with evidences col-

umn.

datasets Character: a vector of dataset labels. Only evidences from these datasets will be

used.

resources Character: a vector of resource labels. Only evidences from these resources will

be used

exclude Character vector of resource names to be excluded.

.keep Logical: keep the "evidences" column.

#### Details

The OmniPath interactions database fully integrates all attributes from all resources for each interaction. This comes with the advantage that interaction data frames are ready for use in most of the applications; however, it makes it impossible to know which of the resources and references support the direction or effect sign of the interaction. This information can be recovered from the "evidences" column. The "evidences" column preserves all the details about interaction provenances. In cases when you want to use a faithful copy of a certain resource or dataset, this function will help you do so. Still, in most of the applications the best is to use the interaction data as it is returned by the web service.

**Note:** This function is automatically applied if the 'strict\_evidences' argument is passed to any function querying interactions (e.g. omnipath-interactions).

### Value

A copy of the interaction data frame restricted to the given datasets and resources.

#### See Also

- omnipath-interactions
- filter\_evidences
- unnest\_evidences
- from\_evidences

```
## Not run:
ci <- collectri(evidences = TRUE)
ci <- only_from(ci, datasets = 'collectri')
## End(Not run)</pre>
```

172 ontology\_ensure\_name

ontology\_ensure\_id Only ontology IDs

### **Description**

Converts a mixture of ontology IDs and names to only IDs. If an element of the input is missing from the chosen ontology it will be dropped. This can happen if the ontology is a subset (slim) version, but also if the input is not a valid ID or name.

### Usage

```
ontology_ensure_id(terms, db_key = "go_basic")
```

### **Arguments**

terms Character: ontology IDs or term names.

db\_key Character: key to identify the ontology database. For the available keys see

omnipath\_show\_db.

#### Value

Character vector of ontology IDs.

### **Examples**

```
ontology_ensure_id(c('mitochondrion inheritance', 'GO:0001754'))
# [1] "GO:0000001" "GO:0001754"
```

### **Description**

Converts a mixture of ontology IDs and names to only names. If an element of the input is missing from the chosen ontology it will be dropped. This can happen if the ontology is a subset (slim) version, but also if the input is not a valid ID or name.

# Usage

```
ontology_ensure_name(terms, db_key = "go_basic")
```

# Arguments

terms Character: ontology IDs or term names.

db\_key Character: key to identify the ontology database. For the available keys see

omnipath\_show\_db.

ontology\_name\_id 173

#### Value

Character vector of ontology term names.

### **Examples**

```
ontology_ensure_name(c('reproduction', 'GO:0001754', 'foo bar')) # [1] "eye photoreceptor cell differentiation" "reproduction"
```

ontology\_name\_id

Translate between ontology IDs and names

### **Description**

Makes sure that the output contains only valid IDs or term names. The input can be a mixture of IDs and names. The order of the input won't be preserved in the output.

### Usage

```
ontology_name_id(terms, ids = TRUE, db_key = "go_basic")
```

# Arguments

terms	Character: ontology IDs or term names.
ids	Logical: the output should contain IDs or term names.
db_key	Character: key to identify the ontology database. For the available keys see omnipath show db.

### Value

Character vector of ontology IDs or term names.

organism\_for

Make sure the resource supports the organism and it has the ID

# Description

Make sure the resource supports the organism and it has the ID

### Usage

```
organism_for(organism, resource, error = TRUE)
```

#### **Arguments**

organism Character or integer: name or NCBI Taxonomy ID of the organism.

resource Charater: name of the resource.

error Logical: raise an error if the organism is not supported in the resource. Other-

wise it only emits a warning.

#### Value

Character: the ID of the organism as it is used by the resource. NA if the organism can not be translated to the required identifier type.

# **Examples**

```
organism_for(10116, 'chalmers-gem')
# [1] "Rat"
organism_for(6239, 'chalmers-gem')
# [1] "Worm"
# organism_for('foobar', 'chalmers-gem')
# Error in organism_for("foobar", "chalmers-gem") :
# Organism `foobar` (common_name: `NA`; common_name: `NA`)
# is not supported by resource `chalmers-gem`. Supported organisms:
# Human, Mouse, Rat, Zebrafish, Drosophila melanogaster (Fruit fly),
# Caenorhabditis elegans (PRJNA13758).
```

orthology\_translate\_column

Translate a column of identifiers by orthologous gene pairs

### **Description**

Translate a column of identifiers by orthologous gene pairs

#### Usage

```
orthology_translate_column(
   data,
   column,
   id_type = NULL,
   target_organism = "mouse",
   source_organism = "human",
   resource = "oma",
   replace = FALSE,
   one_to_many = NULL,
   keep_untranslated = FALSE,
   translate_complexes = FALSE,
   uniprot_by_id_type = "entrez"
)
```

#### **Arguments**

data A data frame with the column to be translated.

column Name of a character column with identifiers of the source organism of type

'id\_type'.

id\_type Type of identifiers in 'column'. Available ID types include "uniprot", "entrez",

"ensg", "refseq" and "swissprot" for OMA, and "uniprot", "entrez", "genesymbol", "refseq" and "gi" for NCBI HomoloGene. If you want to translate an ID type not directly available in your preferred resource, use first translate\_ids to translate to an ID type directly available in the orthology resource. If not

provided, it is assumed the column name is the ID type.

target\_organism

Name or NCBI Taxonomy ID of the target organism.

source\_organism

Name or NCBI Taxonomy ID of the source organism.

resource Character: source of the orthology mapping. Currently Orthologous Matrix

(OMA) and NCBI HomoloGene are available, refer to them by "oma" and "ho-

mologene", respectively.

replace Logical or character: replace the column with the translated identifiers, or create

a new column. If it is character, it will be used as the name of the new column.

ganism. Genes mapping to higher number of orthologues will be dropped.

keep\_untranslated

Logical: keep records without orthologous pairs. If 'replace' is TRUE, this option is ignored, and untranslated records will be dropped. Genes with more

than 'one\_to\_many' orthologues will always be dropped.

translate\_complexes

Logical: translate the complexes by translating their components.

uniprot\_by\_id\_type

Character: translate NCBI HomoloGene to UniProt by this ID type. One of "genesymbol", "entrez", "refseq" or "gi".

#### Value

The data frame with identifiers translated to other organism.

pivot\_annotations

```
pathwaycommons_download
```

Interactions from PathwayCommons

### **Description**

PathwayCommons (http://www.pathwaycommons.org/) provides molecular interactions from a number of databases, in either BioPAX or SIF (simple interaction format). This function retrieves all interactions in SIF format. The data is limited to the interacting pair and the type of the interaction.

# Usage

```
pathwaycommons_download()
```

#### Value

A data frame (tibble) with interactions.

### **Examples**

```
pc_interactions <- pathwaycommons_download()</pre>
pc_interactions
# # A tibble: 1,884,849 x 3
    from type
                                       to
     <chr> <chr>
                                       <chr>
  1 A1BG controls-expression-of
                                       A2M
  2 A1BG interacts-with
                                       ABCC6
  3 A1BG interacts-with
                                       ACE2
  4 A1BG interacts-with
                                       ADAM10
# 5 A1BG interacts-with
                                       ADAM17
# # . with 1,884,839 more rows
```

pivot\_annotations

Converts annotation tables to a wide format

### **Description**

Use this method to reconstitute the annotation tables into the format of the original resources. With the 'wide=TRUE' option annotations applies this function to the downloaded data.

### Usage

```
pivot_annotations(annotations)
```

### **Arguments**

annotations

A data frame of annotations downloaded from the OmniPath web service by annotations.

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#### Value

A wide format data frame (tibble) if the provided data contains annotations from one resource, otherwise a list of wide format tibbles.

#### See Also

annotations

### **Examples**

```
# single resource: the result is a data frame
disgenet <- annotations(resources = "DisGeNet")</pre>
disgenet <- pivot_annotations(disgenet)</pre>
disgenet
# # A tibble: 126,588 × 11
     uniprot genesymbol entity_type disease
                                                    type score dsi
                                                     <chr> <dbl> <dbl> <dbl>
     <chr> <chr> <chr> <chr>
  1 P04217 A1BG
                        protein
                                      Schizophren. dise. 0.3 0.7 0.538
# 2 P04217 A1BG protein Hepatomegaly phen. 0.3 0.7 0.538
# 3 P01023 A2M protein Fibrosis, L. dise. 0.3 0.529 0.769
# 4 P01023 A2M protein Acute kidne. dise. 0.3 0.529 0.769
# 5 P01023 A2M protein Mental Depr. dise. 0.3 0.529 0.769
# # . with 126,583 more rows, and 3 more variables: nof_pmids <dbl>,
# # nof_snps <dbl>, source <chr>
# multiple resources: the result is a list
annot_long <- annotations(</pre>
    resources = c("DisGeNet", "SignaLink_function", "DGIdb", "kinase.com")
annot_wide <- pivot_annotations(annot_long)</pre>
names(annot_wide)
# [1] "DGIdb"
                             "DisGeNet"
                                                    "kinase.com"
# [4] "SignaLink_function"
annot_wide$kinase.com
# # A tibble: 825 x 6
     uniprot genesymbol entity_type group family subfamily
     <chr> <chr> <chr> <chr>
  1 P31749 AKT1
                         protein
                                      AGC Akt
                                                     NA
  2 P31751 AKT2
                                      AGC Akt
                        protein
                                                     NA
  3 Q9Y243 AKT3
                                      AGC Akt
                        protein
                                                     NA
                     protein
protein
  4 014578 CIT
                                      AGC
                                             DMPK
                                                     CRIK
# 5 Q09013 DMPK
                                      AGC DMPK
                                                    GEK
\# # . with 815 more rows
```

preppi\_download

Interactions from PrePPI

# Description

Retrieves predicted protein-protein interactions from the PrePPI database (http://honig.c2b2.columbia.edu/preppi). The interactions in this table are supposed to be correct with a > 0.5 probability.

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### Usage

```
preppi_download(...)
```

#### **Arguments**

. . .

Minimum values for the scores. The available scores are: str, protpep, str\_max, red, ort, phy, coexp, go, total, exp and final. Furthermore, an operator can be passed, either .op = '&' or .op = '|', which is then used for combined filtering by multiple scores.

### **Details**

PrePPI is a combination of many prediction methods, each resulting a score. For an explanation of the scores see <a href="https://honiglab.c2b2.columbia.edu/hfpd/help/Manual.html">https://honiglab.c2b2.columbia.edu/hfpd/help/Manual.html</a>. The minimum, median and maximum values of the scores:

	Score	Minimum	-	Median		Maximum	1
1							-
1	str	0		5.5		6,495	-
1	protpep	0		3.53	1	38,138	-
1	str_max	0		17.9		38,138	-
1	red	0		1.25		24.4	-
1	ort	0		0		5,000	-
1	phy	0		2.42		2.42	-
1	coexp	0		2.77	1	45.3	-
1	go	0		5.86		181	-
1	total	0		1,292	1	106,197,000,000	-
	exp	1		958		4,626	
Ι	final	600	-	1,778	Ι	4.91e14	Ι

#### Value

A data frame (tibble) of interactions with scores, databases and literature references.

### See Also

```
preppi_filter
```

```
preppi <- preppi_download()</pre>
# # A tibble: 1,545,710 x 15
    prot1 prot2 str_score protpep_score str_max_score red_score ort_score
    <chr> <chr> <dbl> <dbl>
                                        <dbl>
                                                    <dbl>
                                                              <dbl>
                               6.45
# 1 Q131. P146.
                 18.6
                                            18.6
                                                     4.25
                                                              0.615
                 1.83
4.57
# 2 P064. Q96N.
                                                     4.25
                             14.3
                                           14.3
                                                              0
# 3 Q7Z6. Q8NC.
                              0
                                            4.57
                                                     0
                                                              0
                               0
# 4 P370. P154.
                485.
                                           485.
                                                     1.77
                                                              0.615
# 5 0004. Q9NR.
                  34.0
                               0
                                           34.0
                                                     0.512
# # . with 1,545,700 more rows, and 8 more variables: phy_score <dbl>,
    coexp_score <dbl>, go_score <dbl>, total_score <dbl>, dbs <chr>,
     pubs <chr>, exp_score <dbl>, final_score <dbl>
```

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Filter PrePPI interactions by scores

# Description

Filter PrePPI interactions by scores

# Usage

```
preppi_filter(data, ..., .op = "&")
```

# **Arguments**

data	A data frame of PrePPI interactions as provided by preppi_download.
	Minimum values for the scores. The available scores are: str, protpep, str_max, red, ort, phy, coexp, go, total, exp and final. See more about the scores at preppi_download.
. op	The operator to combine the scores with: either '&' or ' '. With the former, only records where all scores are above the threshold will be kept; with the latter, records where at least one score is above its threshold will be kept.

### Value

The input data frame (tibble) filtered by the score thresholds.

# See Also

```
preppi_download
```

# Examples

```
preppi <- preppi_download()
preppi_filtered <- preppi_filter(preppi, red = 10, str = 4.5, ort = 1)
nrow(preppi_filtered)
# [1] 8443</pre>
```

print\_bma\_motif\_es

Prints BMA motifs to the screen from a sequence of edges

# Description

The motifs can be copy-pasted into a BMA canvas.

# Usage

```
print_bma_motif_es(edge_seq, G, granularity = 2)
```

print\_bma\_motif\_vs

### **Arguments**

```
edge_seq An igraph edge sequence.

G An igraph graph object.

granularity Numeric: granularity value.
```

# Value

Returns 'NULL'.

# **Examples**

```
interactions <- omnipath(resources = "ARN")</pre>
graph <- interaction_graph(interactions)</pre>
print_bma_motif_es(igraph::E(graph)[1], graph)
# {"Model": {
      "Name": "Omnipath motif",
#
      "Variables":[{
#
          "Name": "ULK1",
#
          "Id":1,
#
          "RangeFrom":0,
#
          "RangeTo":2,
#
          "Formula":""
#
#
      },
#
          "Name": "ATG13",
#
#
#
      }],
# ... (truncated)
# }}
```

print\_bma\_motif\_vs

Prints BMA motifs to the screen from a sequence of nodes

### **Description**

The motifs can be copy-pasted into a BMA canvas.

### Usage

```
print_bma_motif_vs(node_seq, G)
```

#### **Arguments**

```
node_seq An igraph node sequence.

G An igraph graph object.
```

### Value

Returns 'NULL'.

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### **Examples**

```
interactions <- omnipath(resources = "ARN")
graph <- interaction_graph(interactions)
print_bma_motif_vs(
    igraph::all_shortest_paths(
        graph,
        from = 'ULK1',
        to = 'ATG13'
    )$res,
    graph
)</pre>
```

print\_interactions

Print OmniPath interactions

### **Description**

Prints the interactions or enzyme-substrate relationships in a nice format.

#### Usage

```
print_interactions(interactions, refs = FALSE)
```

## **Arguments**

interactions Data frame with the interactions generated by any of the functions in omnipath-interactions. refs Logical: include PubMed IDs where available.

#### Value

Returns 'NULL'.

```
enzsub <- enzyme_substrate()</pre>
print_interactions(head(enzsub))
print_interactions(tail(enzsub), refs = TRUE)
print_interactions(
   dplyr::filter(
       enzsub,
       enzyme_genesymbol == 'MAP2K1',
        substrate_genesymbol == 'MAPK3'
   )
)
signor <- omnipath(resources = "SIGNOR")</pre>
print_interactions(head(signor))
            source interaction
                                          target n_resources
# 6 MAPK14 (Q16539) ==( + )==> MAPKAPK2 (P49137)
                                                         23
# 4 TRPM7 (Q96QT4) ==( + )==> ANXA1 (P04083)
                                                         10
# 1 PRKG1 (Q13976) ==( - )==> TRPC3 (Q13507)
                                                          8
# 2 PTPN1 (P18031) ==( - )==> TRPV6 (Q9H1D0)
                                                          6
# 5 PRKACA (P17612) ==( - )==> MCOLN1 (Q9GZU1)
# 3 RACK1 (P63244) ==( - )==> TRPM6 (Q9BX84)
                                                           2
```

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print\_path\_es

Prints network paths in an edge sequence

## Description

Pretty prints the interactions in a path.

## Usage

```
print_path_es(edges, G)
```

## **Arguments**

edges An igraph edge sequence object.

G igraph object (from ptms or any interaction dataset)

#### Value

Returns 'NULL'.

#### See Also

• print\_path\_vs

## **Examples**

print\_path\_vs

Print networks paths given by node sequence

# Description

Prints the interactions in the path in a nice format.

## Usage

```
print_path_vs(nodes, G)
```

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## **Arguments**

nodes An igraph node sequence object.

G An igraph graph object (from ptms or interactions)

#### Value

Returns 'NULL'.

## See Also

```
print_path_es
```

## **Examples**

```
interactions <- omnipath(resources = "SignaLink3")</pre>
{\tt OPI\_g} \, \mathrel{<\!\!\!\text{--}} \, \text{interaction\_graph(interactions} \, = \, \text{interactions})
print_path_vs(
    igraph::all_shortest_paths(
         OPI_g,
         from = 'TYRO3',
         to = 'STAT3'
    )$vpath,
    OPI_g
enzsub <- enzyme_substrate(resources=c("PhosphoSite", "SIGNOR"))</pre>
enzsub_g <- enzsub_graph(enzsub)</pre>
print_path_vs(
    igraph::all_shortest_paths(
         enzsub_g,
         from = 'SRC',
         to = 'STAT1'
    )$res,
    {\sf enzsub\_g}
)
```

pubmed\_open

Open one or more PubMed articles

# Description

Open one or more PubMed articles

## Usage

```
pubmed_open(pmids, browser = NULL, sep = ";", max_pages = 25L)
```

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### **Arguments**

pmids Character or numberic vector of one or more PubMed IDs.

browser Character: name of the web browser executable. If 'NULL', the default web

browser will be used.

sep Character: split the PubMed IDs by this separator.

max\_pages Numeric: largest number of pages to open. This is to prevent opening hundreds

or thousands of pages at once.

#### Value

Returns 'NULL'.

# **Examples**

```
interactions <- omnipath()
pubmed_open(interactions$references[1])</pre>
```

query\_info

OmniPath query parameters

## **Description**

All parameter names and their possible values for a query type. Note: parameters with 'NULL' values have too many possible values to list them.

## Usage

```
query_info(query_type)
```

# Arguments

query\_type

Character: interactions, annotations, complexes, enz\_sub or intercell.

## Value

A named list with the parameter names and their possible values.

```
ia_param <- query_info('interactions')
ia_param$datasets[1:5]
# [1] "dorothea" "kinaseextra" "ligrecextra" "lncrna_mrna" "mirnatarget"</pre>
```

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ramilowski\_download

Downloads ligand-receptor interactions from Ramilowski et al. 2015

### **Description**

Curated ligand-receptor pairs from Supplementary Table 2 of the article "A draft network of ligand-receptor mediated multicellular signaling in human" (https://www.nature.com/articles/ncomms8866).

## Usage

```
ramilowski_download()
```

#### Value

A data frame (tibble) with interactions.

### **Examples**

```
rami_interactions <- ramilowski_download()</pre>
rami_interactions
# # A tibble: 2,557 x 16
    Pair.Name Ligand.Approved. Ligand.Name Receptor.Approv.
    <chr>
                                          <chr>
           <chr>
                              <chr>
# 1 A2M_LRP1 A2M
                             alpha-2-ma. LRP1
# 2 AANAT_MT. AANAT
                             aralkylami. MTNR1A
                             aralkylami. MTNR1B
# 3 AANAT_MT. AANAT
# 4 ACE_AGTR2 ACE
                             angiotensi. AGTR2
# 5 ACE_BDKR. ACE
                              angiotensi. BDKRB2
# # . with 2,547 more rows, and 12 more variables: Receptor.Name <chr>,
# # DLRP <chr>, HPMR <chr>, IUPHAR <chr>, HPRD <chr>,
# # STRING.binding <chr>, STRING.experiment <chr>, HPMR.Ligand <chr>,
# # HPMR.Receptor <chr>, PMID.Manual <chr>, Pair.Source <chr>,
    Pair.Evidence <chr>
```

ramp\_id\_mapping\_table Pairwise ID translation table from RaMP database

#### **Description**

Pairwise ID translation table from RaMP database

### Usage

```
ramp_id_mapping_table(from, to, version = "2.5.4")
```

## **Arguments**

from	Character or Symbol. Name of an identifier type.
to	Character or Symbol. Name of an identifier type.
version	Character. The version of RaMP to download.

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#### Value

Dataframe of pairs of identifiers.

#### See Also

- ramp\_sqlite
- ramp\_tables
- ramp\_table
- translate\_ids
- id\_types
- hmdb\_table
- uniprot\_full\_id\_mapping\_table
- uniprot\_id\_mapping\_table
- ensembl\_id\_mapping\_table
- chalmers\_gem\_id\_mapping\_table

## **Examples**

```
ramp_id_mapping_table('hmdb', 'kegg')
```

ramp\_id\_type

RaMP identifier type label

# Description

RaMP identifier type label

## Usage

```
ramp_id_type(label)
```

## **Arguments**

label

Character: an ID type label, as shown in the table returned by id\_types

#### Value

Character: the RaMP specific ID type label, or the input unchanged if it could not be translated (still might be a valid identifier name). These labels should be valid value names, as used in RaMP SQL database.

### See Also

- chalmers\_gem\_id\_type
- uniprot\_id\_type
- ensembl\_id\_type
- uploadlists\_id\_type

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## **Examples**

```
ramp_id_type("rhea")
# [1] "rhea-comp"
```

ramp\_sqlite

Download and open RaMP database SQLite

## Description

Download and open RaMP database SQLite

## Usage

```
ramp_sqlite(version = "2.5.4")
```

## **Arguments**

version

Character. The version of RaMP to download.

## Value

SQLite connection.

## See Also

```
• ramp_tables
```

## **Examples**

```
sqlite_con <- ramp_sqlite()</pre>
```

ramp\_table

Return table from RaMP database

# Description

Return table from RaMP database

## Usage

```
ramp_table(name, version = "2.5.4")
```

## **Arguments**

name Character. The name of the RaMP table to fetch.

version Character. The version of RaMP to download.

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## Value

Character vector of table names in the RaMP SQLite database.

#### See Also

- ramp\_sqlite
- ramp\_tables

## **Examples**

```
ramp_table('source')
```

ramp\_tables

List tables in RaMP database

# Description

List tables in RaMP database

## Usage

```
ramp_tables(version = "2.5.4")
```

# **Arguments**

version

Character. The version of RaMP to download.

## Value

Character vector of table names in the RaMP SQLite database.

# See Also

• ramp\_sqlite

```
ramp_tables()
```

regnetwork\_directions 189

regnetwork\_directions Transcription factor effects from RegNetwork

## **Description**

Transcription factor effects from RegNetwork

### Usage

```
regnetwork_directions(organism = "human")
```

## **Arguments**

organism

Character: either human or mouse.

#### Value

A data frame (tibble) of TF-target interactions with effect signs.

## **Examples**

```
regn_dir <- regnetwork_directions()</pre>
regn_dir
# # A tibble: 3,954 x 5
    \verb|source_genesymb.| | source_entrez | target_genesymb.| | target_entrez|
#
                    <chr> <chr>
    <chr>
                                                    <chr>
                     196
                                   CDKN1B
                                                    1027
#
 1 AHR
  2 APLNR
                     187
                                   PIK3C3
                                                    5289
  3 APLNR
                     187
                                   PIK3R4
                                                    30849
# 4 AR
                     367
                                   KLK3
                                                    354
            405
# 5 ARNT
                                   ALDOA
                                                    226
# # . with 3,944 more rows, and 1 more variable: effect <dbl>
```

 $regnetwork\_download$ 

Interactions from RegNetwork

## **Description**

Downloads transcriptional and post-transcriptional regulatory interactions from the RegNetwork database (http://www.regnetworkweb.org/). The information about effect signs (stimulation or inhibition), provided by regnetwork\_directions are included in the result.

## Usage

```
regnetwork_download(organism = "human")
```

### **Arguments**

organism

Character: either human or mouse.

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#### Value

Data frame with interactions.

#### **Examples**

```
regn_interactions <- regnetwork_download()</pre>
regn_interactions
# # A tibble: 372,778 x 7
     source_genesymb. source_entrez target_genesymb. target_entrez
#
                      <chr>
                                    <chr>
                                                      <chr>
  1 USF1
#
                      7391
                                    S100A6
                                                     6277
  2 USF1
                      7391
                                    DUSP1
                                                     1843
# 3 USF1
                      7391
                                                     720
                                    C4A
# 4 USF1
                      7391
                                    ABCA1
                                                     19
# 5 TP53
                      7157
                                    TP73
                                                     7161
# # . with 372,768 more rows, and 3 more variables: effect <dbl>,
# # source_type <chr>, target_type <chr>
```

```
relations_list_to_table
```

Table from a nested list of ontology relations

## **Description**

Converting the nested list to a table is a more costly operation, it takes a few seconds. Best to do it only once, or pass tables = TRUE to obo\_parser, and convert the data frame to list, if you also need it in list format.

#### Usage

```
relations_list_to_table(relations, direction = NULL)
```

## Arguments

relations A nested list of ontology relations (the "relations" element of the list returned by

obo\_parser in case its argument 'tables' is FALSE).

direction Override the direction (i.e. child -> parents or parent -> children). The nested

lists produced by functions in the current package add an attribute "direction" thus no need to pass this value. If the attribute and the argument are both missing, the column will be named simply "side2" and it won't be clear whether the relations point from "term" to "side2" or the other way around. The direction should be a character vector of length 2 with the values "parents" and "children".

#### Value

The relations converted to a data frame (tibble).

### See Also

- swap\_relations
- relations\_table\_to\_list
- obo\_parser

## **Examples**

```
goslim_url <-
    "http://current.geneontology.org/ontology/subsets/goslim_generic.obo"
path <- tempfile()
httr::GET(goslim_url, httr::write_disk(path, overwrite = TRUE))
obo <- obo_parser(path, tables = FALSE)
unlink(path)
rel_tbl <- relations_list_to_table(obo$relations)</pre>
```

```
relations_table_to_graph
```

Graph from a table of ontology relations

## Description

Graph from a table of ontology relations

## Usage

```
relations_table_to_graph(relations)
```

## **Arguments**

relations

A data frame of ontology relations (the "relations" element of the list returned by obo\_parser in case its argument 'tables' is TRUE).

## **Details**

By default the relations point from child to parents, the edges in the graph will be of the same direction. Use swap\_relations on the data frame to reverse the direction.

## Value

The relations converted to an igraph graph object.

```
## Not run:
go <- get_db('go_basic')
go_graph <- relations_table_to_graph(go$relations)
## End(Not run)</pre>
```

```
relations_table_to_list
```

Nested list from a table of ontology relations

## Description

Nested list from a table of ontology relations

#### Usage

```
relations_table_to_list(relations)
```

## **Arguments**

relations

A data frame of ontology relations (the "relations" element of the list returned by obo\_parser in case its argument 'tables' is TRUE).

#### Value

The relations converted to a nested list.

## See Also

- relations\_list\_to\_table
- swap\_relations
- obo\_parser

## **Examples**

```
goslim_url <-
    "http://current.geneontology.org/ontology/subsets/goslim_generic.obo"
path <- tempfile()
httr::GET(goslim_url, httr::write_disk(path, overwrite = TRUE))
obo <- obo_parser(path, tables = TRUE)
unlink(path)
rel_list <- relations_table_to_list(obo$relations)</pre>
```

remap\_dorothea\_download

Downloads TF-target interactions from ReMap

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### **Description**

ReMap (http://remap.univ-amu.fr/) is a database of ChIP-Seq experiments. It provides raw and merged peaks and CRMs (cis regulatory motifs) with their associations to regulators (TFs). TF-target relationships can be derived as it is written in Garcia-Alonso et al. 2019: "For ChIP-seq, we downloaded the binding peaks from ReMap and scored the interactions between each TF and each gene according to the distance between the TFBSs and the genes' transcription start sites. We evaluated different filtering strategies that consisted of selecting only the top-scoring 100, 200, 500, and 1000 target genes for each TF." (https://www.ncbi.nlm.nih.gov/pmc/articles/PMC6673718/#s1title). This function returns the top TF-target relationships as used in DoRothEA: https://github.com/saezlab/dorothea/blob/master/inst/scripts/02\_chip\_seq.R).

### Usage

```
remap_dorothea_download()
```

#### Value

Data frame with TF-target relationships.

#### See Also

```
remap_tf_target_download
```

#### **Examples**

remap\_filtered

Downloads TF-target interactions from ReMap

## Description

Downloads the ReMap TF-target interactions as processed by Garcia-Alonso et al. (https://www.ncbi.nlm.nih.gov/pmc/articles/PMC6673718/#s1title) and filters them based on a score threshold, the top targets and whether the TF is included in the TF census (Vaquerizas et al. 2009). The code for filtering is adapted from DoRothEA, written by Christian Holland.

## Usage

```
remap_filtered(score = 100, top_targets = 500, only_known_tfs = TRUE)
```

### Arguments

score Numeric: a minimum score between 0 and 1000, records with lower scores will

be excluded. If NULL no filtering performed.

mum number of targets per TF. If NULL the number of targets is not restricted.

only\_known\_tfs Logical: whether to exclude TFs which are not in TF census.

#### Value

Data frame with TF-target relationships.

#### See Also

- remap\_tf\_target\_download
- remap\_filtered
- tfcensus download

### **Examples**

```
## Not run:
remap_interactions <- remap_filtered()</pre>
nrow(remap_interactions)
# [1] 145680
remap_interactions <- remap_filtered(top_targets = 100)</pre>
remap_interactions
# # A tibble: 30,330 x 2
    source_genesymbol target_genesymbol
#
#
                        <chr>
     <chr>>
  1 ADNP
                       ABCC1
 2 ADNP
                       ABT1
# 3 ADNP
                       AC006076.1
# 4 ADNP
                       AC007792.1
# 5 ADNP
                       AC011288.2
# # . with 30,320 more rows
## End(Not run)
```

remap\_tf\_target\_download

Downloads TF-target interactions from ReMap

### **Description**

ReMap (http://remap.univ-amu.fr/) is a database of ChIP-Seq experiments. It provides raw and merged peaks and CRMs (cis regulatory motifs) with their associations to regulators (TFs). TF-target relationships can be derived as it is written in Garcia-Alonso et al. 2019: "For ChIP-seq, we downloaded the binding peaks from ReMap and scored the interactions between each TF and each gene according to the distance between the TFBSs and the genes' transcription start sites. We evaluated different filtering strategies that consisted of selecting only the top-scoring 100, 200, 500, and

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1000 target genes for each TF." (https://www.ncbi.nlm.nih.gov/pmc/articles/PMC6673718/#s1title). This function retrieves the full processed TF-target list from the data deposited in https://zenodo.org/record/3713238.

## Usage

```
remap_tf_target_download()
```

#### Value

Data frame with TF-target relationships.

#### See Also

- remap\_dorothea\_download
- remap\_filtered

## **Examples**

```
## Not run:
remap_interactions <- remap_tf_target_download()</pre>
remap_interactions
# # A tibble: 9,546,470 \times 4
     source_genesymbol target_genesymbol target_ensembl
                                                             score
                       <chr>
                                         <chr>
                                                             <dbl>
# 1 ADNP
                       PTPRS
                                         ENSG00000105426.16 1000
# 2 AFF4
                       PRKCH
                                         ENSG00000027075.14 1000
# 3 AHR
                                         ENSG00000169862.18 1000
                       CTNND2
# 4 AR
                       PDE4D
                                         ENSG00000113448.18 1000
# 5 ARID1A
                                         ENSG00000178209.14 1000
                       PLEC
# # . with 9,546,460 more rows
## End(Not run)
```

reset\_config

Restore the built-in default values of all config parameters of a package

# Description

Restore the built-in default values of all config parameters of a package Restore the built-in default values of all config parameters of OmnipathR

# Usage

```
reset_config(save = NULL, reset_all = FALSE, pkg = "OmnipathR")
omnipath_reset_config(...)
```

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### Arguments

save If a path, the restored config will be also saved to this file. If TRUE, the config

will be saved to the current default config path (see omnipath\_config\_path).

reset\_all Reset to their defaults also the options already set in the R options.

pkg Character: name of a package

... Ignored.

#### Value

The config as a list.

### See Also

```
omnipath_load_config, omnipath_save_config
```

## **Examples**

```
## Not run:
# restore the defaults and write them to the default config file:
omnipath_reset_config()
omnipath_save_config()
## End(Not run)
```

resources

Retrieve the available resources for a given query type

## **Description**

Collects the names of the resources available in OmniPath for a certain query type and optionally for a dataset within that.

### Usage

```
resources(query_type, datasets = NULL, generic_categories = NULL)
```

#### Arguments

query\_type one of the query types 'interactions', 'enz\_sub', 'complexes', 'annotations' or

'intercell'

datasets currently within the 'interactions' query type only, multiple datasets are avail-

able: 'omnipath', 'kinaseextra', 'pathwayextra', 'ligrecextra', 'dorothea', 'tf\_target',

'tf\_mirna', 'mirnatarget' and 'lncrna\_mrna'.

generic\_categories

for the 'intercell' query type, restrict the search for some generic categories e.g.

'ligand' or 'receptor'.

## Value

a character vector with resource names

resources\_colname 197

## **Examples**

```
resources(query_type = "interactions")
```

resources\_colname

Name of the column with the resources

## **Description**

Unfortunately the column title is different across the various query types in the OmniPath web service, so we need to guess.

## Usage

```
resources_colname(data)
```

### **Arguments**

data

A data frame downloaded by any import\_... function in the current package.

#### Value

Character: the name of the column, if any of the column names matches.

## **Examples**

```
co <- complexes()
resources_colname(co)
# [1] "sources"</pre>
```

resources\_in

Collect resource names from a data frame

## **Description**

Collect resource names from a data frame

## Usage

```
resources_in(data)
```

## Arguments

data

A data frame from an OmniPath query.

## Value

Character: resource names occuring in the data frame.

show\_network

### **Examples**

```
pathways <- omnipath_interactions()
resources_in(pathways)</pre>
```

resource\_info

OmniPath resource information

## Description

The 'resources' query type provides resource metadata in JSON format. Here we retrieve this JSON and return it as a nested list structure.

## Usage

```
resource_info()
```

#### Value

A nested list structure with resource metadata.

## **Examples**

```
resource_info()
```

show\_network

Visualize node neighborhood with SigmaJS

## **Description**

This function takes an OmniPath interaction data frame as input and returns a sigmaJS object for the subgraph formed by the neighbors of a node of interest.

### Usage

```
show_network(interactions, node = NULL)
```

## **Arguments**

interactions An OmniPath interaction data frame.

node The node of interest.

## Value

A sigmaJS object, check http://sigmajs.john-coene.com/index.html for further details and customization options.

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## **Examples**

```
## Not run:
# get interactions from omnipath
interactions <- omnipath()
# create and plot the network containing ATM neighbors
viz_sigmajs_neighborhood(interactions_df = interactions, int_node = "ATM")
## End(Not run)</pre>
```

signed\_ptms

Causal effect enzyme-PTM interactions

## **Description**

Enzyme-substrate data does not contain sign (activation/inhibition), we generate this information based on the interaction network.

## Usage

```
signed_ptms(
  enzsub = enzyme_substrate(),
  interactions = omnipath_interactions()
)
```

## **Arguments**

enzsub Enzyme-substrate data frame generated by enzyme\_substrate

interactions interaction data frame generated by an OmniPath interactions query: omnipath-interactions

## Value

Data frame of enzyme-substrate relationships with is\_inhibition and is\_stimulation columns.

## See Also

- enzyme\_substrate
- omnipath-interactions

```
enzsub <- enzyme_substrate(resources = c("PhosphoSite", "SIGNOR"))
interactions <- omnipath_interactions()
enzsub <- signed_ptms(enzsub, interactions)</pre>
```

```
simplify_intercell_network

Simplify an intercell network
```

## **Description**

The intercellular communication network data frames, created by intercell\_network, are combinations of a network data frame with two copies of the intercell annotation data frames, all of them already having quite some columns. Here we keep only the names of the interacting pair, their intercellular communication roles, and the minimal information of the origin of both the interaction and the annotations. Optionally further columns can be selected.

## Usage

```
simplify_intercell_network(network, ...)
```

## Arguments

```
network An intercell network data frame, as provided by intercell_network.
... Optional, further columns to select.
```

## Value

An intercell network data frame with some columns removed.

## See Also

- intercell\_network
- filter\_intercell\_network
- unique\_intercell\_network
- intercell
- intercell\_categories
- intercell\_generic\_categories
- intercell\_summary

```
icn <- intercell_network()
icn_s <- simplify_intercell_network(icn)</pre>
```

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static\_table

Retrieve a static table from OmniPath

#### **Description**

A few resources and datasets are available also as plain TSV files and can be accessed without TLS. The purpose of these tables is to make the most often used OmniPath data available on computers with configuration issues. These tables are not the recommended way to access OmniPath data, and a warning is issued each time they are accessed.

#### Usage

```
static_table(
  query,
  resource,
  organism = 9606L,
  strict_evidences = TRUE,
  wide = TRUE,
  dorothea_levels = c("A", "B", "C")
)
```

#### **Arguments**

query Character: a query type such as "annotations" or "interactions".

resource Character: name of the resource or dataset, such as "CollecTRI" or "PROGENy".

organism Integer: NCBI Taxonomy of the organism: 9606 for human, 10090 for mouse

and 10116 for rat.

strict\_evidences

Logical: restrict the evidences to the queried datasets and resources. If set to FALSE, the directions and effect signs and references might be based on other

datasets and resources.

wide

Convert the annotation table to wide format, which corresponds more or less to the original resource. If the data comes from more than one resource a list of wide tables will be returned. See examples at pivot\_annotations.

dorothea\_levels

Vector detailing the confidence levels of the interactions to be downloaded. In dorothea, every TF-target interaction has a confidence score ranging from A to E, being A the most reliable interactions. By default here we take A, B and C level interactions (c("A", "B", "C")). It is to note that E interactions are not available in OmnipathR.

## Value

A data frame (tibble) with the requested resource.

## See Also

```
static_tables
```

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#### **Examples**

```
static_table("annotations", "PROGENy")
```

static\_tables

List the static tables available from OmniPath

### **Description**

A few resources and datasets are available also as plain TSV files and can be accessed without TLS. The purpose of these tables is to make the most often used OmniPath data available on computers with configuration issues. These tables are not the recommended way to access OmniPath data, and a warning is issued each time they are accessed.

## Usage

```
static_tables()
```

#### Value

A data frame listing the available tables.

### See Also

```
static_table
```

## **Examples**

static\_tables()

stitch\_actions

Retrieve the STITCH actions dataset

## **Description**

Retrieve the STITCH actions dataset

### Usage

```
stitch_actions(organism = "human", prefixes = FALSE)
```

## Arguments

organism Character or integer: name or NCBI Taxonomy ID of an organism. STITCH

supports many organisms, please refer to their web site at https://stitch.

embl.de/.

prefixes Logical: include the prefixes in front of identifiers.

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#### Value

Data frame of STITCH actions.

#### See Also

- stitch\_actions
- stitch\_links
- stitch\_network

## **Examples**

```
sta <- stitch_actions(organism = 'mouse')</pre>
```

stitch\_links

Retrieve the STITCH links dataset

## Description

Retrieve the STITCH links dataset

## Usage

```
stitch_links(organism = "human", prefixes = FALSE)
```

# Arguments

organism Character or integer: name or NCBI Taxonomy ID of an organism. STITCH

supports many organisms, please refer to their web site at https://stitch.

embl.de/.

prefixes Logical: include the prefixes in front of identifiers.

# Value

Data frame: organism specific STITCH links dataset.

## See Also

- stitch\_actions
- stitch\_links
- stitch\_network

```
stl <- stitch_links()</pre>
```

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stitch\_network

Chemical-protein interactions from STITCH

### **Description**

Chemical-protein interactions from STITCH

#### Usage

```
stitch_network(
  organism = "human",
  min_score = 700L,
  protein_ids = c("uniprot", "genesymbol"),
  metabolite_ids = c("hmdb", "kegg"),
  cosmos = FALSE
)
```

#### Arguments

organism Character or integer: name or NCBI Taxonomy ID of an organism. STITCH supports many organisms, please refer to their web site at https://stitch.

embl.de/.

min\_score Confidence cutoff used for STITCH connections (700 by default).

protein\_ids Character: translate the protein identifiers to these ID types. Each ID type results

two extra columns in the output, for the "a" and "b" sides of the interaction, respectively. The default ID type for proteins is Esembl Protein ID, and by

default UniProt IDs and Gene Symbols are included.

metabolite\_ids Character: translate the protein identifiers to these ID types. Each ID type results

two extra columns in the output, for the "a" and "b" sides of the interaction, respectively. The default ID type for metabolites is PubChem CID, and HMDB

IDs and KEGG IDs are included.

cosmos Logical: use COSMOS format?

### Value

A data frame of STITCH chemical-protein and protein-chemical interactions with their effect signs, and optionally with identifiers translated.

#### See Also

```
• stitch_actions
```

- stitch\_links
- stitch\_remove\_prefixes

```
stn <- stitch_network(protein_ids = 'genesymbol', metabolite_ids = 'hmdb')</pre>
```

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```
stitch_remove_prefixes
```

Remove the prefixes from STITCH identifiers

## Description

STITCH adds the NCBI Taxonomy ID as a prefix to Ensembl protein identifiers, e.g. "9606.ENSP00000170630", and "CID" followed by "s" or "m" (stereospecific or merged, respectively) in front of PubChem Compound Identifiers. It also pads the CID with zeros. This function removes these prefixes, leaving only the identifiers.

## Usage

```
stitch_remove_prefixes(d, ..., remove = TRUE)
```

## Arguments

d Data frame, typically the output of stitch\_links or stitch\_actions.
... Names of columns to remove prefixes from. NSE is supported.
remove Logical: remove the prefixes? If FALSE, this function does nothing.

#### Value

Data frame with prefixes removed in the specified columns.

#### See Also

- stitch\_actions
- stitch\_links
- stitch\_network

# **Examples**

```
stitch_remove_prefixes(
    data.frame(a = c('9606.ENSP00000170630', 'CIDs00012345')),
    a
)
```

subnetwork

Extract a custom subnetwork from a large network

## Description

Extract a custom subnetwork from a large network

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#### Usage

```
subnetwork(
  network,
  nodes = NULL,
  order = 1L,
  mode = "all",
  mindist = 0L,
  return_df = TRUE
)
```

## **Arguments**

network Either an OmniPath interaction data frame, or an igraph graph object.

nodes Character or integer vector: names, identifiers or indices of the nodes to build

the subnetwork around.

order Integer: order of neighbourhood around nodes; i.e., number of steps starting

from the provided nodes.

mode Character: "all", "out" or "in". Follow directed edges from the provided nodes

in any, outbound or inbound direction, respectively.

mindist Integer: The minimum distance to include the vertex in the result.

return\_df Logical: return an interaction data frame instead of an igraph object.

#### Value

A network data frame or an igraph object, depending on the "return\_df" parameter.

#### See Also

- interaction\_graph
- graph\_interaction
- show\_network

swap\_relations

Reverse the direction of ontology relations

## **Description**

Reverse the direction of ontology relations

# Usage

```
swap_relations(relations)
```

## **Arguments**

relations

The 'relations' component of the data returned by obo\_parser or any '...ontology\_download' function such as go\_ontology\_download. Depending on the tables argument of those functions the 'relations' can be a data frame or a nested list.

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#### Value

Same type as the input, but the relations swapped: if in the input these pointed from each child to the parents, in the output they point from each parent to their children, and vice versa.

## See Also

```
• relations_list_to_table
```

- relations\_table\_to\_list
- obo\_parser

## **Examples**

```
goslim_url <-
    "http://current.geneontology.org/ontology/subsets/goslim_generic.obo"
path <- tempfile()
httr::GET(goslim_url, httr::write_disk(path, overwrite = TRUE))
obo <- obo_parser(path)
unlink(path)
rel_swapped <- swap_relations(obo$relations)</pre>
```

swissprots\_only

Retain only SwissProt IDs

# Description

Retain only SwissProt IDs

# Usage

```
swissprots_only(uniprots, organism = 9606)
```

# Arguments

uniprots Character vector of UniProt IDs.

organism Character or integer: name or identifier of the organism.

## Value

Character vector with only SwissProt IDs.

```
swissprots_only(c("Q05BL1", "A0A654IBU3", "P00533"))
# [1] "P00533"
```

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tfcensus\_download

Downloads the list of transcription factors from TF census

## **Description**

Vaquerizas et al. published in 2009 a list of transcription factors. This function retrieves Supplementary Table 2 from the article (http://www.nature.com/nrg/journal/v10/n4/index.html).

#### Usage

```
tfcensus_download()
```

#### Value

A data frame (tibble) listing transcription factors.

#### **Examples**

```
tfcensus <- tfcensus_download()</pre>
tfcensus
# # A tibble: 1,987 x 7
    Class `Ensembl ID` `IPI ID` `Interpro DBD` `Interpro DNA-b.
    <chr> <chr>
                        <chr>
                                 <chr>
                                                 <chr>
  1 a
          ENSG0000000. IPI0021. NA
                                                 IPR001289
#
  2 a
          ENSG0000000. IPI0004. IPR000047; IPR. NA
          ENSG0000000. IPI0001. IPR001356; IPR. NA
  3 a
 4 a
          ENSG0000000. IPI0029. IPR000910; IPR. NA
# 5 a
          ENSG0000000. IPI0001. IPR007087; IPR. IPR006794
# # . with 1,977 more rows, and 2 more variables: `HGNC symbol` <chr>,
# # `Tissue-specificity` <chr>
```

translate\_ids

Translate gene, protein and small molecule identifiers

## **Description**

Translates a vector of identifiers, resulting a new vector, or a column of identifiers in a data frame by creating another column with the target identifiers.

## Usage

```
translate_ids(
    d,
    ...,
    uploadlists = FALSE,
    ensembl = FALSE,
    hmdb = FALSE,
    ramp = FALSE,
    chalmers = FALSE,
    entity_type = NULL,
```

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```
keep_untranslated = TRUE,
return_df = FALSE,
organism = 9606,
reviewed = TRUE,
complexes = NULL,
complexes_one_to_many = NULL)
```

## **Arguments**

d Character vector or data frame.

At least two arguments, with or without names. The first of these arguments describes the source identifier, the rest of them describe the target identifier(s). The values of all these arguments must be valid identifier types as shown in Details. The names of the arguments are column names. In case of the first (source) ID the column must exist. For the rest of the IDs new columns will be created with the desired names. For ID types provided as arguments without

names, the name of the ID type will be used for column name.

uploadlists Force using the uploadlists service from UniProt. By default the plain query

interface is used (implemented in uniprot\_full\_id\_mapping\_table in this package). If any of the provided ID types is only available in the uploadlists service, it will be automatically selected. The plain query interface is preferred because in the long term, with caching, it requires less download and data stor-

age.

ensembl Logical: use data from Ensembl BioMart instead of UniProt.

hmdb Logical: use HMDB ID translation data.
ramp Logical: use RaMP ID translation data.

chalmers Logical: use ID translation data from Chalmers Sysbio GEM.

entity\_type Character: "gene" and "smol" are short symbols for proteins, genes and small

molecules respectively. Several other synonyms are also accepted.

keep\_untranslated

In case the output is a data frame, keep the records where the source identifier could not be translated. At these records the target identifier will be NA.

return\_df Return a data frame even if the input is a vector.

organism Character or integer, name or NCBI Taxonomy ID of the organism (by default

9606 for human). Matters only if uploadlists is FALSE.

reviewed Translate only reviewed (TRUE), only unreviewed (FALSE) or both (NULL) UniProt

records. Matters only if uploadlists is FALSE.

complexes Logical: translate complexes by their members. Only complexes where all

members can be translated will be included in the result. If NULL, the option

omnipathr.complex\_translation will be used.

complexes\_one\_to\_many

Logical: allow combinatorial expansion or use only the first target identifier for each member of each complex. If NULL, the option omnipathr.complex\_translation\_one\_to\_many will be used.

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#### **Details**

This function, depending on the uploadlists parameter, uses either the uploadlists service of UniProt or plain UniProt queries to obtain identifier translation tables. The possible values for from and to are the identifier type abbreviations used in the UniProt API, please refer to the table here: <a href="https://www.uniprot.org/help/api\_idmapping">https://www.uniprot.org/help/api\_idmapping</a>. In addition, simple synonyms are available which realize a uniform API for the uploadlists and UniProt query based backends. These are the followings:

OmnipathR	Uploadlists	UniProt query	Ensembl BioMart
uniprot	ACC	id	uniprotswissprot
uniprot_entry	ID	entry name	
trembl	reviewed = FALSE	reviewed = FALSE	uniprotsptrembl
genesymbol	GENENAME	genes(PREFERRED)	external_gene_name
genesymbol_syn		genes(ALTERNATIVE)	external_synonym
hgnc	HGNC_ID	database(HGNC)	hgnc_symbol
entrez	P_ENTREZGENEID	database(GeneID)	
ensembl	ENSEMBL_ID		ensembl_gene_id
ensg	ENSEMBL_ID		ensembl_gene_id
enst	ENSEMBL_TRS_ID	database(Ensembl)	ensembl_transcript_id
ensp	ENSEMBL_PRO_ID		ensembl_peptide_id
ensgg	ENSEMBLGENOME_ID		
ensgt	ENSEMBLGENOME_TRS_ID		
ensgp	ENSEMBLGENOME_PRO_ID		
protein_name		protein names	
pir	PIR	database(PIR)	
ccds		database(CCDS)	
refseqp	P_REFSEQ_AC	database(refseq)	
ipro			interpro
ipro_desc			interpro_description
ipro_sdesc			interpro_short_description
wikigene			wikigene_name
rnacentral			rnacentral
gene_desc			description
wormbase		database(WormBase)	
flybase		database(FlyBase)	
xenbase		database(Xenbase)	
zfin		database(ZFIN)	
pbd	PBD_ID	database(PDB)	pbd

For a complete list of ID types and their synonyms, including metabolite and chemical ID types which are not shown here, see id\_types.

The mapping between identifiers can be ambiguous. In this case one row in the original data frame yields multiple rows or elements in the returned data frame or vector(s).

## Value

- Data frame: if the input is a data frame or the input is a vector and return\_df is TRUE.
- Vector: if the input is a vector, there is only one target ID type and return\_df is FALSE.
- List of vectors: if the input is a vector, there are more than one target ID types and return\_df is FALSE. The names of the list will be ID types (as they were column names, see the description of the . . . argument), and the list will also include the source IDs.

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#### See Also

```
translate_ids_multi
uniprot_id_mapping_table
uniprot_full_id_mapping_table
ensembl_id_mapping_table
hmdb_id_mapping_table
id_types
ensembl_id_type
uniprot_id_type
uploadlists_id_type
hmdb_id_type
chalmers_gem_id_type
```

### **Examples**

```
d <- data.frame(uniprot_id = c('P00533', 'Q9ULV1', 'P43897', 'Q9Y2P5'))</pre>
d <- translate_ids(d, uniprot_id = uniprot, genesymbol)</pre>
d
#
    uniprot_id genesymbol
# 1
        P00533
                       FGFR
# 2
        Q9ULV1
                       FZD4
# 3
        P43897
                       TSFM
        Q9Y2P5
                   SLC27A5
```

translate\_ids\_multi Translate gene, protein and small molecule identifiers from multiple columns

## **Description**

Especially when translating network interactions, where two ID columns exist (source and target), it is convenient to call the same ID translation on multiple columns. The translate\_ids function is already able to translate to multiple ID types in one call, but is able to work only from one source column. Here too, multiple target IDs are supported. The source columns can be listed explicitely, or they might share a common stem, in this case the first element of . . . will be used as stem, and the column names will be created by adding the suffixes. The suffixes are also used to name the target columns. If no suffixes are provided, the name of the source columns will be added to the name of the target columns. ID types can be defined the same way as for translate\_ids. The only limitation is that, if the source columns are provided as stem+suffixes, they must be the same ID type.

#### Usage

```
translate_ids_multi(
   d,
    ...,
   suffixes = NULL,
```

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```
suffix_sep = "_",
  uploadlists = FALSE.
  ensembl = FALSE,
 hmdb = FALSE,
  chalmers = FALSE,
  entity_type = NULL,
  keep_untranslated = TRUE,
  organism = 9606,
  reviewed = TRUE
)
```

## **Arguments**

A data frame.

At least two arguments, with or without names. These arguments describe identifier columns, either the ones we translate from (source), or the ones we translate to (target). Columns existing in the data frame will be used as source columns. All the rest will be considered target columns. Alternatively, the source columns can be defined as a stem and a vector of suffixes, plus a separator between the stem and suffix. In this case, the source columns will be the ones that exist in the data frame with the suffixes added. The values of all these arguments must be valid identifier types as shown at translate\_ids. If ID type is provided only for the first source column, the rest of the source columns will be assumed to have the same ID type. For the target identifiers new columns will be created with the desired names, with the suffixes added. If no suffixes provided, the names of the source columns will be used instead.

suffixes Column name suffixes in case the names should be composed of stem and suffix.

suffix\_sep Character: separator between the stem and suffixes.

Force using the 'uploadlists' service from UniProt. By default the plain query uploadlists interface is used (implemented in uniprot\_full\_id\_mapping\_table in this

package). If any of the provided ID types is only available in the uploadlists service, it will be automatically selected. The plain query interface is preferred because in the long term, with caching, it requires less download and data stor-

ensemb1 Logical: use data from Ensembl BioMart instead of UniProt.

hmdb Logical: use HMDB ID translation data.

Logical: use ID translation data from Chalmers Sysbio GEM. chalmers

Character: "gene" and "smol" are short symbols for proteins, genes and small entity\_type

molecules respectively. Several other synonyms are also accepted.

keep\_untranslated

In case the output is a data frame, keep the records where the source identifier could not be translated. At these records the target identifier will be NA.

Character or integer, name or NCBI Taxonomy ID of the organism (by default organism

9606 for human). Matters only if uploadlists is FALSE.

Translate only reviewed (TRUE), only unreviewed (FALSE) or both (NULL) UniProt

records. Matters only if uploadlists is FALSE.

d

. . .

reviewed

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#### Value

A data frame with all source columns translated to all target identifiers. The number of new columns is the product of source and target columns. The target columns are distinguished by the suffexes added to their names.

#### See Also

```
translate_ids
```

## **Examples**

```
ia <- omnipath()
translate_ids_multi(ia, source = uniprot, target, ensp, ensembl = TRUE)</pre>
```

trembls\_only

Retain only TrEMBL IDs

## Description

Retain only TrEMBL IDs

# Usage

```
trembls_only(uniprots, organism = 9606)
```

## **Arguments**

uniprots Character vector of UniProt IDs.

organism Character or integer: name or identifier of the organism.

## Value

Character vector with only TrEMBL IDs.

```
trembls_only(c("Q05BL1", "A0A654IBU3", "P00533"))
# [1] "Q05BL1" "A0A654IBU3"
```

trrust\_download

Downloads TF-target interactions from TRRUST

## **Description**

TRRUST v2 (https://www.grnpedia.org/trrust/) is a database of literature mined TF-target interactions for human and mouse.

## Usage

```
trrust_download(organism = "human")
```

# **Arguments**

organism

Character: either "human" or "mouse".

## Value

A data frame of TF-target interactions.

## **Examples**

```
trrust_interactions <- trrust_download()</pre>
trrust_interactions
# # A tibble: 11,698 x 4
    source_genesymbol target_genesymbol effect reference
#
                      <chr>
                                         <dbl> <chr>
# 1 AATF
                      BAX
                                            -1 22909821
# 2 AATF
                      CDKN1A
                                             0 17157788
# 3 AATF
                      KLK3
                                             0 23146908
                                             1 20549547
 4 AATF
                      MYC
# 5 AATF
                      TP53
                                             0 17157788
# 6 ABL1
                      BAX
                                             1 11753601
  7 ABL1
                      BCL2
                                            -1 11753601
# # . with 11,688 more rows
```

```
uniprot_full_id_mapping_table
```

Creates an ID translation table from UniProt data

### **Description**

Creates an ID translation table from UniProt data

### Usage

```
uniprot_full_id_mapping_table(
  to,
  from = "accession",
  reviewed = TRUE,
  organism = 9606
)
```

## **Arguments**

to	Character or symbol: target ID type. See Details for possible values.
from	Character or symbol: source ID type. See Details for possible values.

reviewed (TRUE), only unreviewed (FALSE) or both (NULL) UniProt

records.

organism Integer, NCBI Taxonomy ID of the organism (by default 9606 for human).

#### **Details**

For both source and target ID type, this function accepts column codes used by UniProt and some simple shortcuts defined here. For the UniProt codes please refer to https://www.uniprot.org/help/uniprotkb The shortcuts are entrez, genesymbol, genesymbol\_syn (synonym gene symbols), hgnc, embl, ref-seqp (RefSeq protein), enst (Ensembl transcript), uniprot\_entry (UniProtKB AC, e.g. EGFR\_HUMAN), protein\_name (full name of the protein), uniprot (UniProtKB ID, e.g. P00533). For a complete table please refer to translate\_ids.

#### Value

A data frame (tibble) with columns 'From' and 'To', UniProt IDs and the corresponding foreign IDs, respectively.

#### See Also

- translate\_ids
- ensembl\_id\_mapping\_table
- uniprot\_id\_mapping\_table

### **Examples**

```
uniprot_entrez <- uniprot_full_id_mapping_table(to = 'entrez')
uniprot_entrez
# # A tibble: 20,723 x 2
# From To
# <chr> <chr> # 1 Q96R72 NA
# 2 Q9UKL2 23538
# 3 Q9H205 144125
# 4 Q8NGN2 219873
# 5 Q8NGC1 390439
# # . with 20,713 more rows
```

```
uniprot_genesymbol_cleanup
```

TrEMBL to SwissProt by gene names

## **Description**

TrEMBL to SwissProt by gene names

#### Usage

```
uniprot_genesymbol_cleanup(uniprots, organism = 9606, only_trembls = TRUE)
```

#### **Arguments**

uniprots Character vector possibly containing TrEMBL IDs.

organism Character or integer: organism name or identifier.

only\_trembls Attempt to convert only known TrEMBL IDs of the organism. This is the rec-

ommended practice.

#### **Details**

Sometimes one gene or protein is represented by multiple identifiers in UniProt. These are typically slightly different isoforms, some of them having TrEMBL IDs, some of the SwissProt. For the purposes of most systems biology application, the most important is to identify the protein or gene in a way that we can recognize it in other datasets. Unfortunately UniProt or Ensembl do not seem to offer solution for this issue. Hence, if we find that a TrEMBL ID has a gene name which is also associated with a SwissProt ID, we replace this TrEMBL ID by that SwissProt. There might be a minor difference in their sequence, but most of the omics analyses do not even consider isoforms. And it is quite possible that later UniProt will convert the TrEMBL record to an isoform within the SwissProt record. Typically this translation is not so important (but still beneficial) for human, but for other organisms it is critical especially when translating from foreign identifiers.

This function accepts a mixed input of UniProt IDs and provides a distinct translation table that you can use to translate your data.

#### Value

Data frame with two columns: "input" and "output". The first one contains all identifiers from the input vector 'uniprots'. The second one has the corresponding identifiers which are either SwissProt IDs with gene names identical to the TrEMBL IDs in the input, or if no such records are available, the output has the input items unchanged.

## Examples

```
uniprot_idmapping_id_types
```

ID types available in the UniProt ID Mapping service

### **Description**

ID types available in the UniProt ID Mapping service

#### Usage

```
uniprot_idmapping_id_types()
```

#### Value

A data frame listing the ID types.

#### **Examples**

```
uniprot_idmapping_id_types()
```

```
uniprot_id_mapping_table
```

ID translation data from UniProt ID Mapping

# Description

Retrieves an identifier translation table from the UniProt ID Mapping service (https://www.uniprot.org/help/id\_mapping).

#### Usage

```
uniprot_id_mapping_table(identifiers, from, to, chunk_size = NULL)
```

download failures, try lower values.

## **Arguments**

identifiers	Character vector of identifiers
from	Character or symbol: type of the identifiers provided. See Details for possible values.
to	Character or symbol: identifier type to be retrieved from UniProt. See Details for possible values.
chunk_size	Integer: query the identifiers in chunks of this size. If you are experiencing

#### Details

This function uses the uploadlists service of UniProt to obtain identifier translation tables. The possible values for 'from' and 'to' are the identifier type abbreviations used in the UniProt API, please refer to the table here: uniprot\_idmapping\_id\_types or the table of synonyms supported by the current package: translate\_ids. Note: if the number of identifiers is larger than the chunk size the log message about the cache origin is not guaranteed to be correct (most of the times it is still correct).

# Value

A data frame (tibble) with columns 'From' and 'To', the identifiers provided and the corresponding target IDs, respectively.

#### See Also

```
translate_ids
```

218 uniprot\_id\_type

## **Examples**

```
uniprot_genesymbol <- uniprot_id_mapping_table(
    c('P00533', 'P23771'), uniprot, genesymbol
)
uniprot_genesymbol
# # A tibble: 2 x 2
# From To
# <chr> <chr> # 1 P00533 EGFR
# 2 P23771 GATA3
```

uniprot\_id\_type

UniProt identifier type label

# **Description**

UniProt identifier type label

# Usage

```
uniprot_id_type(label)
```

## **Arguments**

label

Character: an ID type label, as shown in the table at translate\_ids

#### Value

Character: the UniProt specific ID type label, or the input unchanged if it could not be translated (still might be a valid identifier name). This is the label that one can use in UniProt REST queries.

## See Also

- ensembl\_id\_type
- uploadlists\_id\_type

```
ensembl_id_type("entrez")
# [1] "database(GeneID)"
```

```
unique_intercell_network
```

Unique intercellular interactions

# Description

In the intercellular network data frames produced by intercell\_network, by default each pair of annotations for an interaction is represented in a separate row. This function drops the annotations and keeps only the distinct interacting pairs.

# Usage

```
unique_intercell_network(network, ...)
```

## Arguments

network An intercellular network data frame as produced by intercell\_network.

Additional columns to keep. Note: if these have multiple values for an interacting pair, only the first row will be preserved.

#### Value

A data frame with interacting pairs and interaction attributes.

#### See Also

- intercell\_network
- simplify\_intercell\_network
- filter\_intercell\_network
- intercell
- intercell\_categories
- intercell\_generic\_categories
- intercell\_summary

```
icn <- intercell_network()
icn_unique <- unique_intercell_network(icn)</pre>
```

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unnest\_evidences

Separate evidences by direction and effect sign

#### **Description**

Separate evidences by direction and effect sign

#### Usage

```
unnest_evidences(data, longer = FALSE, .keep = FALSE)
```

# Arguments

data An interaction data frame with "evidences" column.

longer Logical: If TRUE, the "evidences" column is split into rows.

.keep Logical: keep the "evidences" column. When unnesting to longer data frame,

the "evidences" column will contain the unnested evidences, while the original column will be retained under the "all\_evidences" name (if '.keep = TRUE').

#### Value

The data frame with new columns or new rows by direction and sign.

## See Also

- only\_from
- filter\_evidences
- from\_evidences

# **Examples**

```
## Not run:
op <- omnipath_interactions(fields = "evidences")
op <- unnest_evidences(op)
colnames(op)
## End(Not run)</pre>
```

uploadlists\_id\_type

UniProt Uploadlists identifier type label

## **Description**

UniProt Uploadlists identifier type label

#### Usage

```
uploadlists_id_type(label, side = "from")
```

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#### **Arguments**

label	Character: an ID type label, as shown in the table at translate_ids
side	Character: either "from" or "to": direction of the mapping.

#### Value

Character: the UniProt Uploadlists specific ID type label, or the input unchanged if it could not be translated (still might be a valid identifier name). This is the label that one can use in UniProt Uploadlists (ID Mapping) queries.

## See Also

```
• ensembl_id_type
```

- uniprot\_id\_type
- hmdb\_id\_type
- chalmers\_gem\_id\_type

#### **Examples**

```
ensembl_id_type("entrez")
# [1] "GeneID"
```

vinayagam\_download

Protein-protein interactions from Vinayagam 2011

## **Description**

Retrieves the Supplementary Table S6 from Vinayagam et al. 2011. Find out more at https://doi.org/10.1126/scisignal.2001699.

# Usage

```
vinayagam_download()
```

## Value

A data frame (tibble) with interactions.

```
vinayagam_interactions <- vinayagam_download()</pre>
{\tt vinayagam\_interactions}
# # A tibble: 34,814 x 5
     `Input-node Gen. `Input-node Gen. `Output-node Ge. `Output-node Ge.
    <chr>
                                 <dbl> <chr>
                                                                    <dbl>
#
# 1 C1orf103
                                 55791 MNAT1
                                                                     4331
# 2 MAST2
                                 23139 DYNLL1
                                                                     8655
# 3 RAB22A
                                 57403 APPL2
                                                                    55198
# 4 TRAP1
                                 10131 EXT2
                                                                     2132
# 5 STAT2
                                  6773 COPS4
                                                                    51138
```

222 walk\_ontology\_tree

```
# # . with 34,804 more rows, and 1 more variable:
# # `Edge direction score` <dbl>
```

walk\_ontology\_tree

All nodes of a subtree starting from the selected nodes

## Description

Starting from the selected nodes, recursively walks the ontology tree until it reaches either the root or leaf nodes. Collects all visited nodes.

# Usage

## **Arguments**

terms	Character vector of ontology term IDs or names. A mixture of IDs and names can be provided.
ancestors	Logical: if FALSE the ontology tree is traversed towards the leaf nodes; if TRUE, the tree is traversed until the root. The former returns the ancestors (parents), the latter the descendants (children).
db_key	Character: key to identify the ontology database. For the available keys see omnipath_show_db.
ids	Logical: whether to return IDs or term names.
method	Character: either "gra" or "lst". The implementation to use for traversing the ontology tree. The graph based implementation is faster than the list based, the latter will be removed in the future.
relations	Character vector of ontology relation types. Only these relations will be used.

# **Details**

Note: this function relies on the database manager, the first call might take long because of the database load process. Subsequent calls within a short period should be faster. See <a href="mailto:get\_ontology\_db">get\_ontology\_db</a>.

## Value

Character vector of ontology IDs. If the input terms are all leaves or roots NULL is returned. The starting nodes won't be included in the result unless they fall onto the traversal path from other nodes.

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#### See Also

- omnipath\_show\_db
- get\_ontology\_db

## **Examples**

with\_extra\_attrs

Interaction records having certain extra attributes

## **Description**

Interaction records having certain extra attributes

## Usage

```
with_extra_attrs(data, ...)
```

## **Arguments**

data An interaction data frame.

... The name(s) of the extra attributes; NSE is supported.

#### Value

The data frame filtered to the records having the extra attribute.

#### See Also

- extra\_attrs
- has\_extra\_attrs
- extra\_attrs\_to\_cols
- filter\_extra\_attrs
- extra\_attr\_values

```
i <- omnipath(fields = "extra_attrs")
with_extra_attrs(i, Macrophage_type)</pre>
```

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with\_references

Interactions having references

## **Description**

Interactions having references

#### Usage

```
with_references(data, resources = NULL)
```

## **Arguments**

data An interaction data frame.

resources Character: consider only these resources. If 'NULL', records with any reference

will be accepted.

#### Value

A subset of the input interaction data frame.

# **Examples**

```
cc <- import_post_translational_interactions(resources = 'CellChatDB')
with_references(cc, 'CellChatDB')</pre>
```

zenodo\_download

Retrieves data from Zenodo

# Description

Zenodo is a repository of large scientific datasets. Many projects and publications make their datasets available at Zenodo. This function downloads an archive from Zenodo and extracts the requested file.

# Usage

```
zenodo_download(
  path,
  reader = NULL,
  reader_param = list(),
  url_key = NULL,
  zenodo_record = NULL,
  zenodo_fname = NULL,
  url_param = list(),
  url_key_param = list(),
  ...
)
```

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## **Arguments**

path Character: path to the file within the archive. reader Optional, a function to read the connection. reader\_param List: arguments for the reader function. url\_key Character: name of the option containing the URL The Zenodo record ID, either integer or character. zenodo\_record zenodo\_fname The file name within the record. url\_param List: variables to insert into the URL string (which is returned from the options). url\_key\_param List: variables to insert into the 'url\_key'. Passed to archive\_extractor . . .

#### Value

A connection

```
# an example from the OmnipathR::remap_tf_target_download function:
remap_dorothea <- zenodo_download(</pre>
    zenodo_record = 3713238,
    zenodo_fname = 'tf_target_sources.zip',
    path = (
        'tf_target_sources/chip_seq/remap/gene_tf_pairs_genesymbol.txt'
    ),
    reader = read_tsv,
    reader_param = list(
        col_names = c(
            'source_genesymbol',
            'target_genesymbol',
            'target_ensembl',
            'score'
        col_types = cols(),
        progress = FALSE
    ),
  resource = 'ReMap'
```

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