# Package 'AMR'

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```
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ab\_from\_text

Retrieve Antimicrobial Drug Names and Doses from Clinical Text

# Description

Use this function on e.g. clinical texts from health care records. It returns a list with all antimicrobial drugs, doses and forms of administration found in the texts.

# Usage

```
ab_from_text(
  text,
  type = c("drug", "dose", "administration"),
  collapse = NULL,
  translate_ab = FALSE,
  thorough_search = NULL,
  info = interactive(),
  ...
)
```

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#### **Arguments**

text text to analyse

type type of property to search for, either "drug", "dose" or "administration",

see Examples

collapse a character to pass on to paste(, collapse = ...) to only return one character

per element of text, see Examples

translate\_ab if type = "drug": a column name of the antibiotics data set to translate the

antibiotic abbreviations to, using ab\_property(). The default is FALSE. Using

TRUE is equal to using "name".

thorough\_search

a logical to indicate whether the input must be extensively searched for misspelling and other faulty input values. Setting this to TRUE will take considerably more time than when using FALSE. At default, it will turn TRUE when all input

elements contain a maximum of three words.

info a logical to indicate whether a progress bar should be printed - the default is

TRUE only in interactive mode

... arguments passed on to as.ab()

#### Details

This function is also internally used by as.ab(), although it then only searches for the first drug name and will throw a note if more drug names could have been returned. Note: the as.ab() function may use very long regular expression to match brand names of antimicrobial drugs. This may fail on some systems.

# **Argument** type:

At default, the function will search for antimicrobial drug names. All text elements will be searched for official names, ATC codes and brand names. As it uses as.ab() internally, it will correct for misspelling.

With type = "dose" (or similar, like "dosing", "doses"), all text elements will be searched for numeric values that are higher than 100 and do not resemble years. The output will be numeric. It supports any unit (g, mg, IE, etc.) and multiple values in one clinical text, see *Examples*.

With type = "administration" (or abbreviations, like "admin", "adm"), all text elements will be searched for a form of drug administration. It supports the following forms (including common abbreviations): buccal, implant, inhalation, instillation, intravenous, nasal, oral, parenteral, rectal, sublingual, transdermal and vaginal. Abbreviations for oral (such as 'po', 'per os') will become "oral", all values for intravenous (such as 'iv', 'intraven') will become "iv". It supports multiple values in one clinical text, see *Examples*.

# **Argument** collapse:

Without using collapse, this function will return a list. This can be convenient to use e.g. inside a mutate()):

```
df %>% mutate(abx = ab_from_text(clinical_text))
```

The returned AB codes can be transformed to official names, groups, etc. with all ab\_\* functions such as ab\_name() and ab\_group(), or by using the translate\_ab argument.

```
With using collapse, this function will return a character:
```

```
df %>% mutate(abx = ab_from_text(clinical_text, collapse = "|"))
```

# Value

A list, or a character if collapse is not NULL

```
# mind the bad spelling of amoxicillin in this line,
# straight from a true health care record:
ab_from_text("28/03/2020 regular amoxicilliin 500mg po tid")
ab_from_text("500 mg amoxi po and 400mg cipro iv")
ab_from_text("500 mg amoxi po and 400mg cipro iv", type = "dose")
ab_from_text("500 mg amoxi po and 400mg cipro iv", type = "admin")
ab_from_text("500 mg amoxi po and 400mg cipro iv", collapse = ", ")
# if you want to know which antibiotic groups were administered, do e.g.:
abx <- ab_from_text("500 mg amoxi po and 400mg cipro iv")</pre>
ab_group(abx[[1]])
if (require("dplyr")) {
 tibble(clinical_text = c(
    "given 400mg cipro and 500 mg amox",
    "started on doxy iv today"
 )) %>%
    mutate(
      abx_codes = ab_from_text(clinical_text),
      abx_doses = ab_from_text(clinical_text, type = "doses"),
      abx_admin = ab_from_text(clinical_text, type = "admin"),
      abx_coll = ab_from_text(clinical_text, collapse = "|"),
      abx_coll_names = ab_from_text(clinical_text,
        collapse = "|",
        translate_ab = "name"
     ),
      abx_coll_doses = ab_from_text(clinical_text,
        type = "doses",
        collapse = "|"
      abx_coll_admin = ab_from_text(clinical_text,
        type = "admin",
        collapse = "|"
     )
   )
}
```

# **Description**

Use these functions to return a specific property of an antibiotic from the antibiotics data set. All input values will be evaluated internally with as.ab().

# Usage

```
ab_name(x, language = get_AMR_locale(), tolower = FALSE, ...)
ab_cid(x, ...)
ab_synonyms(x, ...)
ab_tradenames(x, ...)
ab_group(x, language = get_AMR_locale(), ...)
ab_atc(x, only_first = FALSE, ...)
ab_atc_group1(x, language = get_AMR_locale(), ...)
ab_atc_group2(x, language = get_AMR_locale(), ...)
ab_loinc(x, ...)
ab_ddd(x, administration = "oral", ...)
ab_ddd_units(x, administration = "oral", ...)
ab_info(x, language = get_AMR_locale(), ...)
ab_url(x, open = FALSE, ...)
ab_property(x, property = "name", language = get_AMR_locale(), ...)
set_ab_names(
 data,
  property = "name",
  language = get_AMR_locale(),
  snake_case = NULL
)
```

# **Arguments**

x any (vector of) text that can be coerced to a valid antibiotic drug code with as.ab()

language of the returned text - the default is the current system language (see get\_AMR\_locale()) and can also be set with the package option AMR\_locale.

Use language = NULL or language = "" to prevent translation.

tolower a logical to indicate whether the first character of every output should be trans-

formed to a lower case character. This will lead to e.g. "polymyxin B" and not

"polymyxin b".

... in case of set\_ab\_names() and data is a data.frame: columns to select (sup-

ports tidy selection such as column1: column4), otherwise other arguments passed

on to as.ab()

only\_first a logical to indicate whether only the first ATC code must be returned, with

giving preference to J0-codes (i.e., the antimicrobial drug group)

administration way of administration, either "oral" or "iv"

open browse the URL using utils::browseURL()

property one of the column names of one of the antibiotics data set: vector\_or(colnames(antibiotics),

sort = FALSE).

data a data frame of which the columns need to be renamed, or a character vector of

column names

snake\_case a logical to indicate whether the names should be in so-called snake case: in

lower case and all spaces/slashes replaced with an underscore (\_)

#### **Details**

All output will be translated where possible.

The function ab\_url() will return the direct URL to the official WHO website. A warning will be returned if the required ATC code is not available.

The function set\_ab\_names() is a special column renaming function for data.frames. It renames columns names that resemble antimicrobial drugs. It always makes sure that the new column names are unique. If property = "atc" is set, preference is given to ATC codes from the J-group.

### Value

- An integer in case of ab\_cid()
- A named list in case of ab\_info() and multiple ab\_atc()/ab\_synonyms()/ab\_tradenames()
- A double in case of ab\_ddd()
- A data.frame in case of set\_ab\_names()
- A character in all other cases

#### **Source**

World Health Organization (WHO) Collaborating Centre for Drug Statistics Methodology: https://www.whocc.no/atc\_ddd\_index/

European Commission Public Health PHARMACEUTICALS - COMMUNITY REGISTER: https://ec.europa.eu/health/documents/community-register/html/reg\_hum\_atc.htm

# Reference Data Publicly Available

All data sets in this AMR package (about microorganisms, antibiotics, SIR interpretation, EUCAST rules, etc.) are publicly and freely available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. We also provide tab-separated plain text files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

#### See Also

antibiotics

```
# all properties:
ab_name("AMX")
ab_atc("AMX")
ab_cid("AMX")
ab_synonyms("AMX")
ab_tradenames("AMX")
ab_group("AMX")
ab_atc_group1("AMX")
ab_atc_group2("AMX")
ab_url("AMX")
# smart lowercase transformation
ab_name(x = c("AMC", "PLB"))
ab_name(x = c("AMC", "PLB"), tolower = TRUE)
# defined daily doses (DDD)
ab_ddd("AMX", "oral")
ab_ddd_units("AMX", "oral")
ab_ddd("AMX", "iv")
ab_ddd_units("AMX", "iv")
ab_info("AMX") # all properties as a list
# all ab_* functions use as.ab() internally, so you can go from 'any' to 'any':
ab_atc("AMP")
ab_group("J01CA01")
ab_loinc("ampicillin")
ab_name("21066-6")
ab_name(6249)
ab_name("J01CA01")
# spelling from different languages and dyslexia are no problem
ab_atc("ceftriaxon")
ab_atc("cephtriaxone")
ab_atc("cephthriaxone")
ab_atc("seephthriaaksone")
```

```
# use set_ab_names() for renaming columns
colnames(example_isolates)
colnames(set_ab_names(example_isolates))
colnames(set_ab_names(example_isolates, NIT:VAN))
if (require("dplyr")) {
 example_isolates %>%
   set_ab_names()
 # this does the same:
 example_isolates %>%
    rename_with(set_ab_names)
 # set_ab_names() works with any AB property:
 example_isolates %>%
   set_ab_names(property = "atc")
 example_isolates %>%
    set_ab_names(where(is.sir)) %>%
   colnames()
 example_isolates %>%
    set_ab_names(NIT:VAN) %>%
    colnames()
}
```

add\_custom\_antimicrobials

Add Custom Antimicrobials

# **Description**

With add\_custom\_antimicrobials() you can add your own custom antimicrobial drug names and codes.

# Usage

```
add_custom_antimicrobials(x)
clear_custom_antimicrobials()
```

# **Arguments**

x a data.frame resembling the antibiotics data set, at least containing columns "ab" and "name"

#### **Details**

**Important:** Due to how R works, the add\_custom\_antimicrobials() function has to be run in every R session - added antimicrobials are not stored between sessions and are thus lost when R is exited.

There are two ways to circumvent this and automate the process of adding antimicrobials:

**Method 1:** Using the package option AMR\_custom\_ab, which is the preferred method. To use this method:

- 1. Create a data set in the structure of the antibiotics data set (containing at the very least columns "ab" and "name") and save it with saveRDS() to a location of choice, e.g. "~/my\_custom\_ab.rds", or any remote location.
- 2. Set the file location to the package option AMR\_custom\_ab: options(AMR\_custom\_ab = "~/my\_custom\_ab.rds"). This can even be a remote file location, such as an https URL. Since options are not saved between R sessions, it is best to save this option to the .Rprofile file so that it will be loaded on start-up of R. To do this, open the .Rprofile file using e.g. utils::file.edit("~/.Rprofile"), add this text and save the file:

```
# Add custom antimicrobial codes:
options(AMR_custom_ab = "~/my_custom_ab.rds")
```

Upon package load, this file will be loaded and run through the add\_custom\_antimicrobials() function.

**Method 2:** Loading the antimicrobial additions directly from your .Rprofile file. Note that the definitions will be stored in a user-specific  $\mathsf{R}$  file, which is a suboptimal workflow. To use this method:

- 1. Edit the .Rprofile file using e.g. utils::file.edit("~/.Rprofile").
- 2. Add a text like below and save the file:

Use clear\_custom\_antimicrobials() to clear the previously added antimicrobials.

#### See Also

add\_custom\_microorganisms() to add custom microorganisms.

```
# returns NA and throws a warning (which is suppressed here):
suppressWarnings(
   as.ab("testab")
```

```
)
# now add a custom entry - it will be considered by as.ab() and
# all ab_*() functions
add_custom_antimicrobials(
  data.frame(
   ab = "TESTAB",
   name = "Test Antibiotic",
    # you can add any property present in the
    # 'antibiotics' data set, such as 'group':
    group = "Test Group"
)
# "testab" is now a new antibiotic:
as.ab("testab")
ab_name("testab")
ab_group("testab")
ab_info("testab")
# Add Co-fluampicil, which is one of the many J01CR50 codes, see
# https://www.whocc.no/ddd/list_of_ddds_combined_products/
add_custom_antimicrobials(
  data.frame(
   ab = "COFLU",
   name = "Co-fluampicil",
   atc = "J01CR50",
    group = "Beta-lactams/penicillins"
  )
)
ab_atc("Co-fluampicil")
ab_name("J01CR50")
# even antibiotic selectors work
x <- data.frame(</pre>
  random_column = "some value",
  coflu = as.sir("S"),
  ampicillin = as.sir("R")
)
x[, betalactams()]
```

add\_custom\_microorganisms

# **Description**

With add\_custom\_microorganisms() you can add your own custom microorganisms, such the non-taxonomic outcome of laboratory analysis.

# **Usage**

```
add_custom_microorganisms(x)
clear_custom_microorganisms()
```

# **Arguments**

x a data.frame resembling the microorganisms data set, at least containing column "genus" (case-insensitive)

#### **Details**

This function will fill in missing taxonomy for you, if specific taxonomic columns are missing, see *Examples*.

**Important:** Due to how R works, the add\_custom\_microorganisms() function has to be run in every R session - added microorganisms are not stored between sessions and are thus lost when R is exited.

There are two ways to circumvent this and automate the process of adding microorganisms:

**Method 1:** Using the package option AMR\_custom\_mo, which is the preferred method. To use this method:

- 1. Create a data set in the structure of the microorganisms data set (containing at the very least column "genus") and save it with saveRDS() to a location of choice, e.g. "~/my\_custom\_mo.rds", or any remote location.
- 2. Set the file location to the package option AMR\_custom\_mo: options(AMR\_custom\_mo = "~/my\_custom\_mo.rds"). This can even be a remote file location, such as an https URL. Since options are not saved between R sessions, it is best to save this option to the .Rprofile file so that it will be loaded on start-up of R. To do this, open the .Rprofile file using e.g. utils::file.edit("~/.Rprofile"), add this text and save the file:

```
# Add custom microorganism codes:
options(AMR_custom_mo = "~/my_custom_mo.rds")
```

Upon package load, this file will be loaded and run through the add\_custom\_microorganisms() function.

**Method 2:** Loading the microorganism directly from your .Rprofile file. Note that the definitions will be stored in a user-specific R file, which is a suboptimal workflow. To use this method:

- 1. Edit the .Rprofile file using e.g. utils::file.edit("~/.Rprofile").
- 2. Add a text like below and save the file:

Use clear\_custom\_microorganisms() to clear the previously added microorganisms.

#### See Also

add\_custom\_antimicrobials() to add custom antimicrobials.

```
# a combination of species is not formal taxonomy, so
# this will result in "Enterobacter cloacae cloacae",
# since it resembles the input best:
mo_name("Enterobacter asburiae/cloacae")
# now add a custom entry - it will be considered by as.mo() and
# all mo_*() functions
add_custom_microorganisms(
  data.frame(
   genus = "Enterobacter",
    species = "asburiae/cloacae"
  )
)
# E. asburiae/cloacae is now a new microorganism:
mo_name("Enterobacter asburiae/cloacae")
# its code:
as.mo("Enterobacter asburiae/cloacae")
# all internal algorithms will work as well:
mo_name("Ent asburia cloacae")
# and even the taxonomy was added based on the genus!
mo_family("E. asburiae/cloacae")
mo_gramstain("Enterobacter asburiae/cloacae")
mo_info("Enterobacter asburiae/cloacae")
# the function tries to be forgiving:
add_custom_microorganisms(
  data.frame(
   GENUS = "BACTEROIDES / PARABACTEROIDES SLASHLINE",
    SPECIES = "SPECIES"
  )
)
mo_name("BACTEROIDES / PARABACTEROIDES")
```

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```
mo_rank("BACTEROIDES / PARABACTEROIDES")

# taxonomy still works, even though a slashline genus was given as input:
mo_family("Bacteroides/Parabacteroides")

# for groups and complexes, set them as species or subspecies:
add_custom_microorganisms(
    data.frame(
        genus = "Citrobacter",
            species = c("freundii", "braakii complex"),
            subspecies = c("complex", "")
    )
)
mo_name(c("C. freundii complex", "C. braakii complex"))
mo_species(c("C. freundii complex", "C. braakii complex"))
mo_gramstain(c("C. freundii complex", "C. braakii complex"))
```

age

Age in Years of Individuals

# **Description**

Calculates age in years based on a reference date, which is the system date at default.

# Usage

```
age(x, reference = Sys.Date(), exact = FALSE, na.rm = FALSE, ...)
```

# Arguments

X	date(s), character (vectors) will be coerced with as.POSIX1t()			
reference	reference date(s) (default is today), character (vectors) will be coerced with $as.POSIXlt()$			
exact	a logical to indicate whether age calculation should be exact, i.e. with decimals. It divides the number of days of year-to-date (YTD) of x by the number of days in the year of reference (either 365 or 366).			
na.rm	a logical to indicate whether missing values should be removed			
	arguments passed on to as.POSIX1t(), such as origin			

#### **Details**

Ages below 0 will be returned as NA with a warning. Ages above 120 will only give a warning.

This function vectorises over both x and reference, meaning that either can have a length of 1 while the other argument has a larger length.

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# Value

An integer (no decimals) if exact = FALSE, a double (with decimals) otherwise

# See Also

To split ages into groups, use the age\_groups() function.

# Examples

```
# 10 random pre-Y2K birth dates
df <- data.frame(birth_date = as.Date("2000-01-01") - runif(10) * 25000)
# add ages
df$age <- age(df$birth_date)
# add exact ages
df$age_exact <- age(df$birth_date, exact = TRUE)
# add age at millenium switch
df$age_at_y2k <- age(df$birth_date, "2000-01-01")</pre>
df
```

age\_groups

Split Ages into Age Groups

# Description

Split ages into age groups defined by the split argument. This allows for easier demographic (antimicrobial resistance) analysis.

# Usage

```
age\_groups(x, split\_at = c(12, 25, 55, 75), na.rm = FALSE)
```

# Arguments

```
x age, e.g. calculated with age()

split_at values to split x at - the default is age groups 0-11, 12-24, 25-54, 55-74 and 75+. See Details.

na.rm a logical to indicate whether missing values should be removed
```

age\_groups

#### **Details**

To split ages, the input for the split\_at argument can be:

• A numeric vector. A value of e.g. c(10, 20) will split x on 0-9, 10-19 and 20+. A value of only 50 will split x on 0-49 and 50+. The default is to split on young children (0-11), youth (12-24), young adults (25-54), middle-aged adults (55-74) and elderly (75+).

#### • A character:

- "children" or "kids", equivalent of: c(0, 1, 2, 4, 6, 13, 18). This will split on 0, 1, 2-3, 4-5, 6-12, 13-17 and 18+.
- "elderly" or "seniors", equivalent of: c(65, 75, 85). This will split on 0-64, 65-74, 75-84, 85+.
- "fives", equivalent of: 1:20 \* 5. This will split on 0-4, 5-9, ..., 95-99, 100+.
- "tens", equivalent of: 1:10 \* 10. This will split on 0-9, 10-19, ..., 90-99, 100+.

#### Value

Ordered factor

#### See Also

To determine ages, based on one or more reference dates, use the age() function.

```
ages <- c(3, 8, 16, 54, 31, 76, 101, 43, 21)
# split into 0-49 and 50+
age_groups(ages, 50)
# split into 0-19, 20-49 and 50+
age_groups(ages, c(20, 50))
# split into groups of ten years
age\_groups(ages, 1:10 * 10)
age_groups(ages, split_at = "tens")
# split into groups of five years
age\_groups(ages, 1:20 * 5)
age_groups(ages, split_at = "fives")
# split specifically for children
age_groups(ages, c(1, 2, 4, 6, 13, 18))
age_groups(ages, "children")
# resistance of ciprofloxacin per age group
if (require("dplyr") && require("ggplot2")) {
 example_isolates %>%
    filter_first_isolate() %>%
   filter(mo == as.mo("Escherichia coli")) %>%
```

```
group_by(age_group = age_groups(age)) %>%
select(age_group, CIP) %>%
ggplot_sir(
    x = "age_group",
    minimum = 0,
    x.title = "Age Group",
    title = "Ciprofloxacin resistance per age group"
)
}
```

antibiogram

Generate Antibiogram: Traditional, Combined, Syndromic, or Weighted-Incidence Syndromic Combination (WISCA)

# Description

Generate an antibiogram, and communicate the results in plots or tables. These functions follow the logic of Klinker *et al.* and Barbieri *et al.* (see *Source*), and allow reporting in e.g. R Markdown and Quarto as well.

# Usage

```
antibiogram(
 Х,
  antibiotics = where(is.sir),
 mo_transform = "shortname",
  ab_transform = NULL,
  syndromic_group = NULL,
  add_total_n = TRUE,
 only_all_tested = FALSE,
 digits = 0,
  col_mo = NULL,
 language = get_AMR_locale(),
 minimum = 30,
 combine_SI = TRUE,
  sep = " + ",
  info = interactive()
)
## S3 method for class 'antibiogram'
plot(x, ...)
## S3 method for class 'antibiogram'
autoplot(object, ...)
## S3 method for class 'antibiogram'
knit_print(
```

```
x,
italicise = TRUE,
na = getOption("knitr.kable.NA", default = ""),
...
)
```

# **Arguments**

a data.frame containing at least a column with microorganisms and columns with antibiotic results (class 'sir', see as.sir())

antibiotics vector of any antibiotic name or code (will be evaluated with as.ab(), column

name of x, or (any combinations of) antibiotic selectors such as aminoglycosides() or carbapenems(). For combination antibiograms, this can also be set to values separated with "+", such as "TZP+TOB" or "cipro + genta", given that columns

resembling such antibiotics exist in x. See Examples.

mo\_transform a character to transform microorganism input - must be "name", "shortname",

"gramstain", or one of the column names of the microorganisms data set: "mo", "fullname", "status", "kingdom", "phylum", "class", "order", "family", "genus", "species", "subspecies", "rank", "ref", "oxygen\_tolerance", "source", "lpsn", "lpsn\_parent", "lpsn\_renamed\_to", "gbif\_, "gbif\_parent", "gbif\_renamed\_to",

"prevalence", or "snomed". Can also be NULL to not transform the input.

ab\_transform a character to transform antibiotic input - must be one of the column names

of the antibiotics data set: "ab", "cid", "name", "group", "atc", "atc\_group1", "atc\_group2", "abbreviations", "synonyms", "oral\_ddd", "oral\_units", "iv\_ddd",

"iv\_units", or "loinc". Can also be NULL to not transform the input.

syndromic\_group

a column name of x, or values calculated to split rows of x, e.g. by using

ifelse() or case\_when(). See Examples.

add\_total\_n a logical to indicate whether total available numbers per pathogen should be

added to the table (default is TRUE). This will add the lowest and highest number of available isolate per antibiotic (e.g, if for *E. coli* 200 isolates are available for ciprofloxacin and 150 for amoxicillin, the returned number will be "150-200").

only\_all\_tested

(for combination antibiograms): a logical to indicate that isolates must be tested

for all antibiotics, see Details

digits number of digits to use for rounding

col\_mo column name of the names or codes of the microorganisms (see as.mo()) - the

default is the first column of class mo. Values will be coerced using as.mo().

language language to translate text, which defaults to the system language (see get\_AMR\_locale())

minimum the minimum allowed number of available (tested) isolates. Any isolate count

lower than minimum will return NA with a warning. The default number of 30 isolates is advised by the Clinical and Laboratory Standards Institute (CLSI) as

best practice, see Source.

combine\_SI a logical to indicate whether all susceptibility should be determined by results

of either S or I, instead of only S (default is TRUE)

sep	a separating character for antibiotic columns in combination antibiograms
info	a ${\color{blue} \log \text{ical}}$ to indicate info should be printed - the default is TRUE only in interactive mode
•••	when used in R Markdown or Quarto: arguments passed on to $knitr::kable()$ (otherwise, has no use)
object	an antibiogram() object
italicise	a logical to indicate whether the microorganism names in the knitr table should be made italic, using $italicise\_taxonomy()$ .
na	character to use for showing NA values

# **Details**

This function returns a table with values between 0 and 100 for susceptibility, not resistance.

Remember that you should filter your data to let it contain only first isolates! This is needed to exclude duplicates and to reduce selection bias. Use first\_isolate() to determine them in your data set with one of the four available algorithms.

All types of antibiograms as listed below can be plotted (using ggplot2::autoplot() or base R plot()/barplot()). The antibiogram object can also be used directly in R Markdown / Quarto (i.e., knitr) for reports. In this case, knitr::kable() will be applied automatically and microorganism names will even be printed in italics at default (see argument italicise). You can also use functions from specific 'table reporting' packages to transform the output of antibiogram() to your needs, e.g. with flextable::as\_flextable() or gt::gt().

# **Antibiogram Types:**

There are four antibiogram types, as proposed by Klinker *et al.* (2021, doi:10.1177/20499361211011373), and they are all supported by antibiogram():

# 1. Traditional Antibiogram

Case example: Susceptibility of *Pseudomonas aeruginosa* to piperacillin/tazobactam (TZP) Code example:

# 2. Combination Antibiogram

Case example: Additional susceptibility of *Pseudomonas aeruginosa* to TZP + tobramycin versus TZP alone

Code example:

# 3. Syndromic Antibiogram

Case example: Susceptibility of *Pseudomonas aeruginosa* to TZP among respiratory specimens (obtained among ICU patients only)

Code example:

# 4. Weighted-Incidence Syndromic Combination Antibiogram (WISCA)

Case example: Susceptibility of *Pseudomonas aeruginosa* to TZP among respiratory specimens (obtained among ICU patients only) for male patients age >=65 years with heart failure Code example:

Note that for combination antibiograms, it is important to realise that susceptibility can be calculated in two ways, which can be set with the only\_all\_tested argument (default is FALSE). See this example for two antibiotics, Drug A and Drug B, about how antibiogram() works to calculate the %SI:

		only_all_tested = FALSE		only_all_tested = TRUE	
Drug A	Drug B		include as denominator		include as denominator
S or I	S or I	Χ	X	Х	Χ
R	S or I	Χ	Χ	Χ	Χ
<na></na>	S or I	Χ	Χ	-	_
S or I	R	Χ	Χ	Χ	Χ
R	R	-	Χ	-	Χ
<na></na>	R	-	-	-	-
S or I	<na></na>	Χ	Χ	-	-
R	<na></na>	-	-	-	-
<na></na>	<na></na>	-	-	-	-

# Source

- Klinker KP *et al.* (2021). **Antimicrobial stewardship and antibiograms: importance of moving beyond traditional antibiograms**. *Therapeutic Advances in Infectious Disease*, May 5;8:20499361211011373; doi:10.1177/20499361211011373
- Barbieri E *et al.* (2021). **Development of a Weighted-Incidence Syndromic Combination Antibiogram (WISCA) to guide the choice of the empiric antibiotic treatment for urinary tract infection in paediatric patients: a Bayesian approach** *Antimicrobial Resistance & Infection Control* May 1;10(1):74; doi:10.1186/s13756021009392
- M39 Analysis and Presentation of Cumulative Antimicrobial Susceptibility Test Data, 5th Edition, 2022, Clinical and Laboratory Standards Institute (CLSI). https://clsi.org/standards/products/microbiology/documents/m39/.

```
# example_isolates is a data set available in the AMR package.
# run ?example_isolates for more info.
example_isolates
# Traditional antibiogram ------
antibiogram(example_isolates,
 antibiotics = c(aminoglycosides(), carbapenems())
antibiogram(example_isolates,
 antibiotics = aminoglycosides(),
 ab_transform = "atc",
 mo_transform = "gramstain"
)
antibiogram(example_isolates,
 antibiotics = carbapenems(),
 ab_transform = "name",
 mo_transform = "name"
)
# Combined antibiogram ------
# combined antibiotics yield higher empiric coverage
antibiogram(example_isolates,
 antibiotics = c("TZP", "TZP+TOB", "TZP+GEN"),
 mo_transform = "gramstain"
)
# names of antibiotics do not need to resemble columns exactly:
antibiogram(example_isolates,
 antibiotics = c("Cipro", "cipro + genta"),
 mo_transform = "gramstain",
 ab_transform = "name",
 sep = " & "
)
# Syndromic antibiogram ------
# the data set could contain a filter for e.g. respiratory specimens
antibiogram(example_isolates,
 antibiotics = c(aminoglycosides(), carbapenems()),
 syndromic_group = "ward"
)
# now define a data set with only E. coli
ex1 <- example_isolates[which(mo_genus() == "Escherichia"), ]</pre>
```

```
# with a custom language, though this will be determined automatically
# (i.e., this table will be in Spanish on Spanish systems)
antibiogram(ex1,
 antibiotics = aminoglycosides(),
 ab_transform = "name",
 syndromic_group = ifelse(ex1$ward == "ICU",
    "UCI", "No UCI"
 ),
 language = "es"
)
# Weighted-incidence syndromic combination antibiogram (WISCA) ------
# the data set could contain a filter for e.g. respiratory specimens/ICU
antibiogram(example_isolates,
 antibiotics = c("AMC", "AMC+CIP", "TZP", "TZP+TOB"),
 mo_transform = "gramstain",
 minimum = 10, # this should be >=30, but now just as example
 syndromic_group = ifelse(example_isolates$age >= 65 &
   example_isolates$gender == "M",
 "WISCA Group 1", "WISCA Group 2"
 )
)
# Print the output for R Markdown / Quarto ------
ureido <- antibiogram(example_isolates,</pre>
 antibiotics = ureidopenicillins(),
 ab_transform = "name"
)
# in an Rmd file, you would just need to return `ureido` in a chunk,
# but to be explicit here:
if (requireNamespace("knitr")) {
 cat(knitr::knit_print(ureido))
# Generate plots with ggplot2 or base R ------
ab1 <- antibiogram(example_isolates,</pre>
 antibiotics = c("AMC", "CIP", "TZP", "TZP+TOB"),
 mo_transform = "gramstain"
ab2 <- antibiogram(example_isolates,</pre>
 antibiotics = c("AMC", "CIP", "TZP", "TZP+TOB"),
 mo_transform = "gramstain",
 syndromic_group = "ward"
)
```

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```
if (requireNamespace("ggplot2")) {
   ggplot2::autoplot(ab1)
}
if (requireNamespace("ggplot2")) {
   ggplot2::autoplot(ab2)
}
plot(ab1)
plot(ab2)
```

antibiotics

Data Sets with 603 Antimicrobial Drugs

# Description

Two data sets containing all antibiotics/antimycotics and antivirals. Use as.ab() or one of the ab\_\* functions to retrieve values from the antibiotics data set. Three identifiers are included in this data set: an antibiotic ID (ab, primarily used in this package) as defined by WHONET/EARS-Net, an ATC code (atc) as defined by the WHO, and a Compound ID (cid) as found in PubChem. Other properties in this data set are derived from one or more of these codes. Note that some drugs have multiple ATC codes.

# Usage

antibiotics antivirals

#### **Format**

# For the antibiotics data set: a tibble with 483 observations and 14 variables::

ab

Antibiotic ID as used in this package (such as AMC), using the official EARS-Net (European Antimicrobial Resistance Surveillance Network) codes where available. *This is a unique identifier.* 

• cid

Compound ID as found in PubChem. This is a unique identifier.

• name

Official name as used by WHONET/EARS-Net or the WHO. This is a unique identifier.

group

A short and concise group name, based on WHONET and WHOCC definitions

• atc

ATC codes (Anatomical Therapeutic Chemical) as defined by the WHOCC, like J01CR02

• atc\_group1

Official pharmacological subgroup (3rd level ATC code) as defined by the WHOCC, like "Macrolides, lincosamides and streptogramins"

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• atc\_group2

Official chemical subgroup (4th level ATC code) as defined by the WHOCC, like "Macrolides"

abbr

List of abbreviations as used in many countries, also for antibiotic susceptibility testing (AST)

synonyms

Synonyms (often trade names) of a drug, as found in PubChem based on their compound ID

• oral\_ddd

Defined Daily Dose (DDD), oral treatment, currently available for 174 drugs

• oral\_units

Units of oral\_ddd

• iv\_ddd

Defined Daily Dose (DDD), parenteral (intravenous) treatment, currently available for 146 drugs

• iv\_units

Units of iv\_ddd

• loinc

All codes associated with the name of the antimicrobial drug from Logical Observation Identifiers Names and Codes (LOINC), Version 2.76 (18 September, 2023). Use ab\_loinc() to retrieve them quickly, see ab\_property().

#### For the antivirals data set: a tibble with 120 observations and 11 variables::

av

Antiviral ID as used in this package (such as ACI), using the official EARS-Net (European Antimicrobial Resistance Surveillance Network) codes where available. *This is a unique identifier*. Combinations are codes that contain a + to indicate this, such as ATA+COBI for atazanavir/cobicistat.

• name

Official name as used by WHONET/EARS-Net or the WHO. This is a unique identifier.

• atc

ATC codes (Anatomical Therapeutic Chemical) as defined by the WHOCC

• cid

Compound ID as found in PubChem. This is a unique identifier.

• atc\_group

Official pharmacological subgroup (3rd level ATC code) as defined by the WHOCC

svnonvms

Synonyms (often trade names) of a drug, as found in PubChem based on their compound ID

• oral\_ddd

Defined Daily Dose (DDD), oral treatment

• oral\_units

Units of oral\_ddd

• iv\_ddd

Defined Daily Dose (DDD), parenteral treatment

• iv\_units

Units of iv\_ddd

• loinc

All codes associated with the name of the antiviral drug from Logical Observation Identifiers

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Names and Codes (LOINC), Version 2.76 (18 September, 2023). Use av\_loinc() to retrieve them quickly, see av\_property().

An object of class tbl\_df (inherits from tbl, data.frame) with 120 rows and 11 columns.

#### **Details**

Properties that are based on an ATC code are only available when an ATC is available. These properties are: atc\_group1, atc\_group2, oral\_ddd, oral\_units, iv\_ddd and iv\_units.

Synonyms (i.e. trade names) were derived from the PubChem Compound ID (column cid) and consequently only available where a CID is available.

#### Direct download:

Like all data sets in this package, these data sets are publicly available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

# WHOCC

This package contains **all ~550 antibiotic**, **antimycotic and antiviral drugs** and their Anatomical Therapeutic Chemical (ATC) codes, ATC groups and Defined Daily Dose (DDD) from the World Health Organization Collaborating Centre for Drug Statistics Methodology (WHOCC, https://www.whocc.no) and the Pharmaceuticals Community Register of the European Commission (https://ec.europa.eu/health/documents/community-register/html/reg\_hum\_atc.htm).

These have become the gold standard for international drug utilisation monitoring and research.

The WHOCC is located in Oslo at the Norwegian Institute of Public Health and funded by the Norwegian government. The European Commission is the executive of the European Union and promotes its general interest.

NOTE: The WHOCC copyright does not allow use for commercial purposes, unlike any other info from this package. See <a href="https://www.whocc.no/copyright\_disclaimer/">https://www.whocc.no/copyright\_disclaimer/</a>.

# Source

- World Health Organization (WHO) Collaborating Centre for Drug Statistics Methodology (WHOCC): https://www.whocc.no/atc\_ddd\_index/
- Logical Observation Identifiers Names and Codes (LOINC), Version 2.76 (18 September, 2023). Accessed from https://loinc.org on October 19th, 2023.
- European Commission Public Health PHARMACEUTICALS COMMUNITY REGISTER: https://ec.europa.eu/health/documents/community-register/html/reg\_hum\_atc.htm

### See Also

microorganisms, intrinsic\_resistant

### **Examples**

antibiotics antivirals

```
antibiotic_class_selectors

Antibiotic Selectors
```

# **Description**

These functions allow for filtering rows and selecting columns based on antibiotic test results that are of a specific antibiotic class or group (according to the antibiotics data set), without the need to define the columns or antibiotic abbreviations.

In short, if you have a column name that resembles an antimicrobial drug, it will be picked up by any of these functions that matches its pharmaceutical class: "cefazolin", "kefzol", "CZO" and "J01DB04" will all be picked up by cephalosporins().

# Usage

```
ab_class(ab_class, only_sir_columns = FALSE, only_treatable = TRUE, ...)
ab_selector(filter, only_sir_columns = FALSE, only_treatable = TRUE, ...)
aminoglycosides(only_sir_columns = FALSE, only_treatable = TRUE, ...)
aminopenicillins(only_sir_columns = FALSE, ...)
antifungals(only_sir_columns = FALSE, ...)
antimycobacterials(only_sir_columns = FALSE, ...)
betalactams(only_sir_columns = FALSE, only_treatable = TRUE, ...)
carbapenems(only_sir_columns = FALSE, only_treatable = TRUE, ...)
cephalosporins(only_sir_columns = FALSE, ...)
cephalosporins_1st(only_sir_columns = FALSE, ...)
cephalosporins_2nd(only_sir_columns = FALSE, ...)
cephalosporins_3rd(only_sir_columns = FALSE, ...)
cephalosporins_4th(only_sir_columns = FALSE, ...)
cephalosporins_5th(only_sir_columns = FALSE, ...)
fluoroquinolones(only_sir_columns = FALSE, ...)
glycopeptides(only_sir_columns = FALSE, ...)
```

```
lincosamides(only_sir_columns = FALSE, ...)
lipoglycopeptides(only_sir_columns = FALSE, ...)
macrolides(only_sir_columns = FALSE, ...)
oxazolidinones(only_sir_columns = FALSE, ...)
penicillins(only_sir_columns = FALSE, ...)
polymyxins(only_sir_columns = FALSE, only_treatable = TRUE, ...)
streptogramins(only_sir_columns = FALSE, ...)
quinolones(only_sir_columns = FALSE, ...)
tetracyclines(only_sir_columns = FALSE, ...)
trimethoprims(only_sir_columns = FALSE, ...)
ureidopenicillins(only_sir_columns = FALSE, ...)
administrable_per_os(only_sir_columns = FALSE, ...)
administrable_iv(only_sir_columns = FALSE, ...)
not_intrinsic_resistant(
 only_sir_columns = FALSE,
 col_mo = NULL,
 version_expertrules = 3.3,
)
```

#### **Arguments**

ab\_class an antimicrobial class or a part of it, such as "carba" and "carbapenems". The columns group, atc\_group1 and atc\_group2 of the antibiotics data set will be searched (case-insensitive) for this value.

only\_sir\_columns

a logical to indicate whether only columns of class sir must be selected (default is FALSE), see as.sir()

only\_treatable a logical to indicate whether antimicrobial drugs should be excluded that are only for laboratory tests (default is TRUE), such as gentamicin-high (GEH) and imipenem/EDTA (IPE)

... ignored, only in place to allow future extensions

filter an expression to be evaluated in the antibiotics data set, such as name %like% "trim"

col\_mo column name of the names or codes of the microorganisms (see as.mo()) - the default is the first column of class mo. Values will be coerced using as.mo().

version\_expertrules

the version number to use for the EUCAST Expert Rules and Intrinsic Resistance guideline. Can be "3.3", "3.2", or "3.1".

#### **Details**

These functions can be used in data set calls for selecting columns and filtering rows. They work with base R, the Tidyverse, and data.table. They are heavily inspired by the Tidyverse selection helpers such as everything(), but are not limited to dplyr verbs. Nonetheless, they are very convenient to use with dplyr functions such as select(), filter() and summarise(), see Examples.

All columns in the data in which these functions are called will be searched for known antibiotic names, abbreviations, brand names, and codes (ATC, EARS-Net, WHO, etc.) according to the antibiotics data set. This means that a selector such as aminoglycosides() will pick up column names like 'gen', 'genta', 'J01GB03', 'tobra', 'Tobracin', etc.

The ab\_class() function can be used to filter/select on a manually defined antibiotic class. It searches for results in the antibiotics data set within the columns group, atc\_group1 and atc\_group2.

The ab\_selector() function can be used to internally filter the antibiotics data set on any results, see *Examples*. It allows for filtering on a (part of) a certain name, and/or a group name or even a minimum of DDDs for oral treatment. This function yields the highest flexibility, but is also the least user-friendly, since it requires a hard-coded filter to set.

The administrable\_per\_os() and administrable\_iv() functions also rely on the antibiotics data set - antibiotic columns will be matched where a DDD (defined daily dose) for resp. oral and IV treatment is available in the antibiotics data set.

The not\_intrinsic\_resistant() function can be used to only select antibiotic columns that pose no intrinsic resistance for the microorganisms in the data set. For example, if a data set contains only microorganism codes or names of *E. coli* and *K. pneumoniae* and contains a column "vancomycin", this column will be removed (or rather, unselected) using this function. It currently applies 'EUCAST Expert Rules' and 'EUCAST Intrinsic Resistance and Unusual Phenotypes' v3.3 (2021) to determine intrinsic resistance, using the eucast\_rules() function internally. Because of this determination, this function is quite slow in terms of performance.

#### Value

(internally) a character vector of column names, with additional class "ab\_selector"

# Full list of supported (antibiotic) classes

• aminoglycosides() can select: amikacin (AMK), amikacin/fosfomycin (AKF), amphotericin B-high (AMH), apramycin (APR), arbekacin (ARB), astromicin (AST), bekanamycin (BEK), dibekacin (DKB), framycetin (FRM), gentamicin (GEN), gentamicin-high (GEH), habekacin (HAB), hygromycin (HYG), isepamicin (ISE), kanamycin (KAN), kanamycin-high (KAH), kanamycin/cephalexin (KAC), micronomicin (MCR), neomycin (NEO), netilmicin (NET), pentisomicin (PIM), plazomicin (PLZ), propikacin (PKA), ribostamycin (RST), sisomicin (SIS), streptoduocin (STR), streptomycin (STR1), streptomycin-high (STH), tobramycin (TOB), and tobramycin-high (TOH)

- aminopenicillins() can select: amoxicillin (AMX) and ampicillin (AMP)
- antifungals() can select: amphotericin B (AMB), anidulafungin (ANI), butoconazole (BUT), caspofungin (CAS), ciclopirox (CIX), clotrimazole (CTR), econazole (ECO), fluconazole (FLU), flucytosine (FCT), fosfluconazole (FFL), griseofulvin (GRI), hachimycin (HCH), ibrexafungerp (IBX), isavuconazole (ISV), isoconazole (ISO), itraconazole (ITR), ketoconazole (KET), manogepix (MGX), micafungin (MIF), miconazole (MCZ), nystatin (NYS), oteseconazole (OTE), pimaricin (PMR), posaconazole (POS), rezafungin (RZF), ribociclib (RBC), sulconazole (SUC), terbinafine (TRB), terconazole (TRC), and voriconazole (VOR)
- antimycobacterials() can select:
   4-aminosalicylic acid (AMA), calcium aminosalicylate (CLA), capreomycin (CAP), clofazimine (CLF), delamanid (DLM), enviomycin (ENV), ethambutol (ETH), ethambutol/isoniazid (ETI), ethionamide (ETI1), isoniazid (INH), isoniazid/sulfamethoxazole/trimethoprim/pyridoxine (IST), morinamide (MRN), p-aminosalicylic acid (PAS), pretomanid (PMD), protionamide (PTH), pyrazinamide (PZA), rifabutin (RIB), rifampicin (RIF), rifampicin/ethambutol/isoniazid (REI), rifampicin/jyrazinamide/ethambutol/isoniazid (RPEI), rifampicin/pyrazinamide/isoniazid (RPI), rifamycin (RFM), rifapentine (RFP), simvastatin/fenofibrate (SMF), sodium aminosalicylate (SDA), streptomycin/isoniazid (STI), terizidone (TRZ), thioacetazone (TAT), thioacetazone/isoniazid (THII), tiocarlide (TCR), and viomycin (VIO)
- betalactams() can select: amoxicillin (AMX), amoxicillin/clavulanic acid (AMC), amoxicillin/sulbactam (AXS), ampicillin (AMP), ampicillin/sulbactam (SAM), apalcillin (APL), aspoxicillin (APX), avibactam (AVB), azidocillin (AZD), azlocillin (AZL), aztreonam (ATM), aztreonam/avibactam (AZA), aztreonam/nacubactam (ANC), bacampicillin (BAM), benzathine benzylpenicillin (BNB), benzathine phenoxymethylpenicillin (BNP), benzylpenicillin (PEN), biapenem (BIA), carbenicillin (CRB), carindacillin (CRN), cefacetrile (CAC), cefaclor (CEC), cefadroxil (CFR), cefalexin (LEX), cefaloridine (RID), cefalotin (CEP), cefamandole (MAN), cefapirin (HAP), cefatrizine (CTZ), cefazedone (CZD), cefazolin (CZO), cefcapene (CCP), cefcapene pivoxil (CCX), cefdinir (CDR), cefditoren (DIT), cefditoren pivoxil (DIX), cefepime (FEP), cefepime/clavulanic acid (CPC), cefepime/nacubactam (FNC), cefepime/tazobactam (FPT), cefetamet (CAT), cefetamet pivoxil (CPI), cefetecol (CCL), cefetrizole (CZL), cefixime (CFM), cefmenoxime (CMX), cefmetazole (CMZ), cefodizime (DIZ), cefonicid (CID), cefoperazone (CFP), cefoperazone/sulbactam (CSL), ceforanide (CND), cefoselis (CSE), cefotaxime (CTX), cefotaxime/clavulanic acid (CTC), cefotaxime/sulbactam (CTS), cefotetan (CTT), cefotiam (CTF), cefotiam hexetil (CHE), cefovecin (FOV), cefoxitin (FOX), cefoxitin screening (FOX1), cefozopran (ZOP), cefpimizole (CFZ), cefpiramide (CPM), cefpirome (CPO), cefpodoxime (CPD), cefpodoxime proxetil (CPX), cefpodoxime/clavulanic acid (CDC), cefprozil (CPR), cefquinome (CEQ), cefroxadine (CRD), cefsulodin (CFS), cefsumide (CSU), ceftaroline (CPT), ceftaroline/avibactam (CPA), ceftazidime (CAZ), ceftazidime/avibactam (CZA), ceftazidime/clavulanic acid (CCV), cefteram (CEM), cefteram pivoxil (CPL), ceftezole (CTL), ceftibuten (CTB), ceftiofur (TIO), ceftizoxime (CZX), ceftizoxime alapivoxil (CZP), ceftobiprole (BPR), ceftobiprole medocaril (CFM1), ceftolozane/tazobactam (CZT), ceftriaxone (CRO), ceftriaxone/betalactamase inhibitor (CEB), cefuroxime (CXM), cefuroxime axetil (CXA), cephradine (CED), ciclacillin (CIC), clometocillin (CLM), cloxacillin (CLO), dicloxacillin (DIC), doripenem (DOR), epicillin (EPC), ertapenem (ETP), flucloxacillin (FLC), hetacillin (HET), imipenem (IPM), imipenem/EDTA (IPE), imipenem/relebactam (IMR), latamoxef (LTM), lenampicillin (LEN), loracarbef (LOR), mecillinam (MEC), meropenem (MEM), meropenem/nacubactam

(MNC), meropenem/vaborbactam (MEV), metampicillin (MTM), meticillin (MET), mezlocillin (MEZ), mezlocillin/sulbactam (MSU), nacubactam (NAC), nafcillin (NAF), oxacillin (OXA), panipenem (PAN), penamecillin (PNM), penicillin/novobiocin (PNO), penicillin/sulbactam (PSU), pheneticillin (PHE), phenoxymethylpenicillin (PHN), piperacillin (PIP), piperacillin/sulbactam (PIS), piperacillin/tazobactam (TZP), piridicillin (PRC), pivampicillin (PVM), pivmecillinam (PME), procaine benzylpenicillin (PRB), propicillin (PRP), razupenem (RZM), ritipenem (RIT), ritipenem acoxil (RIA), sarmoxicillin (SRX), sulbactam (SUL), sulbenicillin (SBC), sultamicillin (SLT6), talampicillin (TAL), tazobactam (TAZ), tebipenem (TBP), temocillin (TEM), ticarcillin (TIC), and ticarcillin/clavulanic acid (TCC)

# • carbapenems() can select:

biapenem (BIA), doripenem (DOR), ertapenem (ETP), imipenem (IPM), imipenem/EDTA (IPE), imipenem/relebactam (IMR), meropenem (MEM), meropenem/nacubactam (MNC), meropenem/vaborbactam (MEV), panipenem (PAN), razupenem (RZM), ritipenem (RIT), ritipenem acoxil (RIA), and tebipenem (TBP)

#### • cephalosporins() can select:

cefacetrile (CAC), cefaclor (CEC), cefadroxil (CFR), cefalexin (LEX), cefaloridine (RID), cefalotin (CEP), cefamandole (MAN), cefapirin (HAP), cefatrizine (CTZ), cefazedone (CZD), cefazolin (CZO), cefcapene (CCP), cefcapene pivoxil (CCX), cefdinir (CDR), cefditoren (DIT), cefditoren pivoxil (DIX), cefepime (FEP), cefepime/clavulanic acid (CPC), cefepime/tazobactam (FPT), cefetamet (CAT), cefetamet pivoxil (CPI), cefetecol (CCL), cefetrizole (CZL), cefixime (CFM), cefmenoxime (CMX), cefmetazole (CMZ), cefodizime (DIZ), cefonicid (CID), cefoperazone (CFP), cefoperazone/sulbactam (CSL), ceforanide (CND), cefoselis (CSE), cefotaxime (CTX), cefotaxime/clavulanic acid (CTC), cefotaxime/sulbactam (CTS), cefotetan (CTT), cefotiam (CTF), cefotiam hexetil (CHE), cefovecin (FOV), cefoxitin (FOX), cefoxitin screening (FOX1), cefozopran (ZOP), cefpimizole (CFZ), cefpiramide (CPM), cefpirome (CPO), cefpodoxime (CPD), cefpodoxime proxetil (CPX), cefpodoxime/clavulanic acid (CDC), cefprozil (CPR), cefquinome (CEQ), cefroxadine (CRD), cefsulodin (CFS), cefsumide (CSU), ceftaroline (CPT), ceftaroline/avibactam (CPA), ceftazidime (CAZ), ceftazidime/avibactam (CZA), ceftazidime/clavulanic acid (CCV), cefteram (CEM), cefteram pivoxil (CPL), ceftezole (CTL), ceftibuten (CTB), ceftiofur (TIO), ceftizoxime (CZX), ceftizoxime alapivoxil (CZP), ceftobiprole (BPR), ceftobiprole medocaril (CFM1), ceftolozane/tazobactam (CZT), ceftriaxone (CRO), ceftriaxone/beta-lactamase inhibitor (CEB), cefuroxime (CXM), cefuroxime axetil (CXA), cephradine (CED), latamoxef (LTM), and loracarbef (LOR)

# cephalosporins\_1st() can select: cefacetrile (CAC), cefadroxil (CFR), cefalexin (LEX), cefaloridine (RID), cefalotin (CEP), cefapirin (HAP), cefatrizine (CTZ), cefazedone (CZD), cefazolin (CZO), cefroxadine (CRD), ceftezole (CTL), and cephradine (CED)

- cephalosporins\_2nd() can select: cefaclor (CEC), cefamandole (MAN), cefmetazole (CMZ), cefonicid (CID), ceforanide (CND), cefotetan (CTT), cefotiam (CTF), cefoxitin (FOX), cefoxitin screening (FOX1), cefprozil (CPR), cefuroxime (CXM), cefuroxime axetil (CXA), and loracarbef (LOR)
- cephalosporins\_3rd() can select:
   cefcapene (CCP), cefcapene pivoxil (CCX), cefdinir (CDR), cefditoren (DIT), cefditoren pivoxil (DIX), cefetamet (CAT), cefetamet pivoxil (CPI), cefixime (CFM), cefmenoxime (CMX), cefodizime (DIZ), cefoperazone (CFP), cefoperazone/sulbactam (CSL), cefotaxime (CTX), cefotaxime/clavulanic acid (CTC), cefotaxime/sulbactam (CTS), cefotiam hexetil (CHE), cefovecin (FOV), cefpimizole (CFZ), cefpiramide (CPM), cefpodoxime (CPD), cefpodoxime

proxetil (CPX), cefpodoxime/clavulanic acid (CDC), cefsulodin (CFS), ceftazidime (CAZ), ceftazidime/avibactam (CZA), ceftazidime/clavulanic acid (CCV), cefteram (CEM), cefteram pivoxil (CPL), ceftibuten (CTB), ceftiofur (TIO), ceftizoxime (CZX), ceftizoxime alapivoxil (CZP), ceftriaxone (CRO), ceftriaxone/beta-lactamase inhibitor (CEB), and latamoxef (LTM)

- cephalosporins\_4th() can select: cefepime (FEP), cefepime/clavulanic acid (CPC), cefepime/tazobactam (FPT), cefetecol (CCL), cefoselis (CSE), cefozopran (ZOP), cefpirome (CPO), and cefquinome (CEQ)
- cephalosporins\_5th() can select: ceftaroline (CPT), ceftaroline/avibactam (CPA), ceftobiprole (BPR), ceftobiprole medocaril (CFM1), and ceftolozane/tazobactam (CZT)
- fluoroquinolones() can select: besifloxacin (BES), ciprofloxacin (CIP), clinafloxacin (CLX), danofloxacin (DAN), delafloxacin (DFX), difloxacin (DIF), enoxacin (ENX), enrofloxacin (ENR), finafloxacin (FIN), fleroxacin (FLE), garenoxacin (GRN), gatifloxacin (GAT), gemifloxacin (GEM), grepafloxacin (GRX), lascufloxacin (LSC), levofloxacin (LVX), levonadifloxacin (LND), lomefloxacin (LOM), marbofloxacin (MAR), metioxate (MXT), miloxacin (MIL), moxifloxacin (MFX), nadifloxacin (NAD), nifuroquine (NIF), norfloxacin (NOR), ofloxacin (OFX), orbifloxacin (ORB), pazufloxacin (PAZ), pefloxacin (PEF), pradofloxacin (PRA), premafloxacin (PRX), prulifloxacin (PRU), rufloxacin (RFL), sarafloxacin (SAR), sitafloxacin (SIT), sparfloxacin (SPX), temafloxacin (TMX), tilbroquinol (TBQ), tioxacin (TXC), tosufloxacin (TFX), and trovafloxacin (TVA)
- glycopeptides() can select: avoparcin (AVO), dalbavancin (DAL), norvancomycin (NVA), oritavancin (ORI), ramoplanin (RAM), teicoplanin (TEC), teicoplanin-macromethod (TCM), telavancin (TLV), vancomycin (VAN), and vancomycin-macromethod (VAM)
- lincosamides() can select: acetylmidecamycin (ACM), acetylspiramycin (ASP), clindamycin (CLI), gamithromycin (GAM), kitasamycin (KIT), lincomycin (LIN), meleumycin (MEL), nafithromycin (ZWK), pirlimycin (PRL), primycin (PRM), solithromycin (SOL), tildipirosin (TIP), tilmicosin (TIL), tulathromycin (TUL), tylosin (TYL), and tylvalosin (TYL1)
- lipoglycopeptides() can select: dalbavancin (DAL), oritavancin (ORI), and telavancin (TLV)
- macrolides() can select: acetylmidecamycin (ACM), acetylspiramycin (ASP), azithromycin (AZM), clarithromycin (CLR), dirithromycin (DIR), erythromycin (ERY), flurithromycin (FLR1), gamithromycin (GAM), josamycin (JOS), kitasamycin (KIT), meleumycin (MEL), midecamycin (MID), miocamycin (MCM), nafithromycin (ZWK), oleandomycin (OLE), pirlimycin (PRL), primycin (PRM), rokitamycin (ROK), roxithromycin (RXT), solithromycin (SOL), spiramycin (SPI), telithromycin (TLT), tildipirosin (TIP), tilmicosin (TIL), troleandomycin (TRL), tulathromycin (TUL), tylosin (TYL), and tylvalosin (TYL1)
- oxazolidinones() can select: cadazolid (CDZ), cycloserine (CYC), linezolid (LNZ), tedizolid (TZD), and thiacetazone (THA)
- penicillins() can select: amoxicillin (AMX), amoxicillin/clavulanic acid (AMC), amoxicillin/sulbactam (AXS), ampicillin (AMP), ampicillin/sulbactam (SAM), apalcillin (APL), aspoxicillin (APX), avibactam (AVB), azidocillin (AZD), azlocillin (AZL), aztreonam (ATM), aztreonam/avibactam (AZA),

aztreonam/nacubactam (ANC), bacampicillin (BAM), benzathine benzylpenicillin (BNB), benzathine phenoxymethylpenicillin (BNP), benzylpenicillin (PEN), carbenicillin (CRB), carindacillin (CRN), cefepime/nacubactam (FNC), ciclacillin (CIC), clometocillin (CLM), cloxacillin (CLO), dicloxacillin (DIC), epicillin (EPC), flucloxacillin (FLC), hetacillin (HET), lenampicillin (LEN), mecillinam (MEC), metampicillin (MTM), meticillin (MET), mezlocillin (MEZ), mezlocillin/sulbactam (MSU), nacubactam (NAC), nafcillin (NAF), oxacillin (OXA), penamecillin (PNM), penicillin/novobiocin (PNO), penicillin/sulbactam (PSU), pheneticillin (PHE), phenoxymethylpenicillin (PHN), piperacillin (PIP), piperacillin/sulbactam (PIS), piperacillin/tazobactam (TZP), piridicillin (PRC), pivampicillin (PVM), pivmecillinam (PME), procaine benzylpenicillin (PRB), propicillin (PRP), sarmoxicillin (SRX), sulbactam (SUL), sulbenicillin (SBC), sultamicillin (SLT6), talampicillin (TAL), tazobactam (TAZ), temocillin (TEM), ticarcillin (TIC), and ticarcillin/clavulanic acid (TCC)

- polymyxins() can select: colistin (COL), polymyxin B (PLB), and polymyxin B/polysorbate 80 (POP)
- quinolones() can select:
  besifloxacin (BES), cinoxacin (CIN), ciprofloxacin (CIP), clinafloxacin (CLX), danofloxacin
  (DAN), delafloxacin (DFX), difloxacin (DIF), enoxacin (ENX), enrofloxacin (ENR), finafloxacin
  (FIN), fleroxacin (FLE), flumequine (FLM), garenoxacin (GRN), gatifloxacin (GAT), gemifloxacin (GEM), grepafloxacin (GRX), lascufloxacin (LSC), levofloxacin (LVX), levonadifloxacin (LND), lomefloxacin (LOM), marbofloxacin (MAR), metioxate (MXT), miloxacin
  (MIL), moxifloxacin (MFX), nadifloxacin (NAD), nalidixic acid (NAL), nemonoxacin (NEM),
  nifuroquine (NIF), nitroxoline (NTR), norfloxacin (NOR), ofloxacin (OFX), orbifloxacin (ORB),
  oxolinic acid (OXO), pazufloxacin (PAZ), pefloxacin (PEF), pipemidic acid (PPA), piromidic
  acid (PIR), pradofloxacin (PRA), premafloxacin (PRX), prulifloxacin (PRU), rosoxacin (ROS),
  rufloxacin (RFL), sarafloxacin (SAR), sitafloxacin (SIT), sparfloxacin (SPX), temafloxacin
  (TMX), tilbroquinol (TBO), tioxacin (TXC), tosufloxacin (TFX), and trovafloxacin (TVA)
- streptogramins() can select: pristinamycin (PRI) and quinupristin/dalfopristin (QDA)
- tetracyclines() can select: cetocycline (CTO), chlortetracycline (CTE), clomocycline (CLM1), demeclocycline (DEM), doxycycline (DOX), eravacycline (ERV), lymecycline (LYM), metacycline (MTC), minocycline (MNO), omadacycline (OMC), oxytetracycline (OXY), penimepicycline (PNM1), rolitetracycline (RLT), sarecycline (SRC), tetracycline (TCY), and tigecycline (TGC)
- trimethoprims() can select: brodimoprim (BDP), sulfadiazine (SDI), sulfadiazine/tetroxoprim (SLT), sulfadiazine/trimethoprim (SLT1), sulfadimethoxine (SUD), sulfadimidine (SDM), sulfadimidine/trimethoprim (SLT2), sulfafurazole (SLF), sulfaisodimidine (SLF1), sulfalene (SLF2), sulfamazone (SZO), sulfamerazine (SLF3), sulfamerazine/trimethoprim (SLT3), sulfamethizole (SLF4), sulfamethoxazole (SMX), sulfamethoxypyridazine (SLF5), sulfametomidine (SLF6), sulfametoxydiazine (SLF7), sulfametrole/trimethoprim (SLT4), sulfamoxole (SLF8), sulfamoxole/trimethoprim (SLT5), sulfanilamide (SLF9), sulfaperin (SLF10), sulfaphenazole (SLF11), sulfapyridine (SLF12), sulfathiazole (SUT), sulfathiourea (SLF13), trimethoprim (TMP), and trimethoprim/sulfamethoxazole (SXT)
- ureidopenicillins() can select: azlocillin (AZL), mezlocillin (MEZ), piperacillin (PIP), and piperacillin/tazobactam (TZP)

#### Reference Data Publicly Available

All data sets in this AMR package (about microorganisms, antibiotics, SIR interpretation, EUCAST rules, etc.) are publicly and freely available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. We also provide tab-separated plain text files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

```
# `example_isolates` is a data set available in the AMR package.
# See ?example_isolates.
example_isolates
# Examples sections below are split into 'base R', 'dplyr', and 'data.table':
# base R -----
# select columns 'IPM' (imipenem) and 'MEM' (meropenem)
example_isolates[, carbapenems()]
# select columns 'mo', 'AMK', 'GEN', 'KAN' and 'TOB'
example_isolates[, c("mo", aminoglycosides())]
# select only antibiotic columns with DDDs for oral treatment
example_isolates[, administrable_per_os()]
# filter using any() or all()
example_isolates[any(carbapenems() == "R"), ]
subset(example_isolates, any(carbapenems() == "R"))
# filter on any or all results in the carbapenem columns (i.e., IPM, MEM):
example_isolates[any(carbapenems()), ]
example_isolates[all(carbapenems()), ]
# filter with multiple antibiotic selectors using c()
example_isolates[all(c(carbapenems(), aminoglycosides()) == "R"), ]
# filter + select in one go: get penicillins in carbapenem-resistant strains
example_isolates[any(carbapenems() == "R"), penicillins()]
# You can combine selectors with '&' to be more specific. For example,
# penicillins() would select benzylpenicillin ('peni G') and
# administrable_per_os() would select erythromycin. Yet, when combined these
# drugs are both omitted since benzylpenicillin is not administrable per os
# and erythromycin is not a penicillin:
example_isolates[, penicillins() & administrable_per_os()]
# ab_selector() applies a filter in the `antibiotics` data set and is thus
# very flexible. For instance, to select antibiotic columns with an oral DDD
```

```
# of at least 1 gram:
example_isolates[, ab_selector(oral_ddd > 1 & oral_units == "g")]
# dplyr -----
if (require("dplyr")) {
 tibble(kefzol = random_sir(5)) %>%
    select(cephalosporins())
}
if (require("dplyr")) {
 # get AMR for all aminoglycosides e.g., per ward:
 example_isolates %>%
    group_by(ward) %>%
    summarise(across(aminoglycosides(), resistance))
}
if (require("dplyr")) {
 # You can combine selectors with '&' to be more specific:
 example_isolates %>%
    select(penicillins() & administrable_per_os())
if (require("dplyr")) {
 # get AMR for only drugs that matter - no intrinsic resistance:
 example_isolates %>%
    filter(mo_genus() %in% c("Escherichia", "Klebsiella")) %>%
   group_by(ward) %>%
    summarise(across(not_intrinsic_resistant(), resistance))
if (require("dplyr")) {
 # get susceptibility for antibiotics whose name contains "trim":
 example_isolates %>%
    filter(first_isolate()) %>%
   group_by(ward) %>%
    summarise(across(ab_selector(name %like% "trim"), susceptibility))
if (require("dplyr")) {
 # this will select columns 'IPM' (imipenem) and 'MEM' (meropenem):
 example_isolates %>%
    select(carbapenems())
if (require("dplyr")) {
 \# this will select columns 'mo', 'AMK', 'GEN', 'KAN' and 'TOB':
 example_isolates %>%
    select(mo, aminoglycosides())
if (require("dplyr")) {
 # any() and all() work in dplyr's filter() too:
 example_isolates %>%
   filter(
     any(aminoglycosides() == "R"),
     all(cephalosporins_2nd() == "R")
   )
```

```
if (require("dplyr")) {
 # also works with c():
 example_isolates %>%
   filter(any(c(carbapenems(), aminoglycosides()) == "R"))
if (require("dplyr")) {
 # not setting any/all will automatically apply all():
 example_isolates %>%
    filter(aminoglycosides() == "R")
if (require("dplyr")) {
 # this will select columns 'mo' and all antimycobacterial drugs ('RIF'):
 example_isolates %>%
    select(mo, ab_class("mycobact"))
if (require("dplyr")) {
 # get bug/drug combinations for only glycopeptides in Gram-positives:
 example_isolates %>%
    filter(mo_is_gram_positive()) %>%
    select(mo, glycopeptides()) %>%
   bug_drug_combinations() %>%
    format()
if (require("dplyr")) {
 data.frame(
    some_column = "some_value",
    J01CA01 = "S"
 ) %>% # ATC code of ampicillin
    select(penicillins()) # only the 'J01CA01' column will be selected
if (require("dplyr")) {
 # with recent versions of dplyr, this is all equal:
 x <- example_isolates[carbapenems() == "R", ]</pre>
 y <- example_isolates %>% filter(carbapenems() == "R")
 z \leftarrow example_isolates \%\% filter(if_all(carbapenems(), ~ .x == "R"))
 identical(x, y) && identical(y, z)
}
\mbox{\#} data.table is supported as well, just use it in the same way as with
# base R, but add `with = FALSE` if using a single AB selector.
if (require("data.table")) {
 dt <- as.data.table(example_isolates)</pre>
 # this does not work, it returns column *names*
 dt[, carbapenems()]
if (require("data.table")) {
 # so `with = FALSE` is required
```

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```
dt[, carbapenems(), with = FALSE]
}

# for multiple selections or AB selectors, `with = FALSE` is not needed:
if (require("data.table")) {
   dt[, c("mo", aminoglycosides())]
}
if (require("data.table")) {
   dt[, c(carbapenems(), aminoglycosides())]
}

# row filters are also supported:
if (require("data.table")) {
   dt[any(carbapenems() == "S"), ]
}
if (require("data.table")) {
   dt[any(carbapenems() == "S"), penicillins(), with = FALSE]
}
```

as.ab

Transform Input to an Antibiotic ID

# Description

Use this function to determine the antibiotic drug code of one or more antibiotics. The data set antibiotics will be searched for abbreviations, official names and synonyms (brand names).

# Usage

```
as.ab(x, flag_multiple_results = TRUE, info = interactive(), ...)
is.ab(x)
```

# **Arguments**

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#### **Details**

All entries in the antibiotics data set have three different identifiers: a human readable EARS-Net code (column ab, used by ECDC and WHONET), an ATC code (column atc, used by WHO), and a CID code (column cid, Compound ID, used by PubChem). The data set contains more than 5,000 official brand names from many different countries, as found in PubChem. Not that some drugs contain multiple ATC codes.

All these properties will be searched for the user input. The as.ab() can correct for different forms of misspelling:

- Wrong spelling of drug names (such as "tobramicin" or "gentamycin"), which corrects for most audible similarities such as f/ph, x/ks, c/z/s, t/th, etc.
- Too few or too many vowels or consonants
- Switching two characters (such as "mreopenem", often the case in clinical data, when doctors typed too fast)
- Digitalised paper records, leaving artefacts like 0/o/O (zero and O's), B/8, n/r, etc.

Use the ab\_\* functions to get properties based on the returned antibiotic ID, see Examples.

Note: the as.ab() and ab\_\* functions may use very long regular expression to match brand names of antimicrobial drugs. This may fail on some systems.

You can add your own manual codes to be considered by as.ab() and all ab\_\* functions, see add\_custom\_antimicrobials().

#### Value

A character vector with additional class ab

#### Source

World Health Organization (WHO) Collaborating Centre for Drug Statistics Methodology: https://www.whocc.no/atc\_ddd\_index/

European Commission Public Health PHARMACEUTICALS - COMMUNITY REGISTER: https://ec.europa.eu/health/documents/community-register/html/reg\_hum\_atc.htm

## **WHOCC**

This package contains **all ~550 antibiotic**, **antimycotic and antiviral drugs** and their Anatomical Therapeutic Chemical (ATC) codes, ATC groups and Defined Daily Dose (DDD) from the World Health Organization Collaborating Centre for Drug Statistics Methodology (WHOCC, https://www.whocc.no) and the Pharmaceuticals Community Register of the European Commission (https://ec.europa.eu/health/documents/community-register/html/reg\_hum\_atc.htm).

These have become the gold standard for international drug utilisation monitoring and research.

The WHOCC is located in Oslo at the Norwegian Institute of Public Health and funded by the Norwegian government. The European Commission is the executive of the European Union and promotes its general interest.

NOTE: The WHOCC copyright does not allow use for commercial purposes, unlike any other info from this package. See https://www.whocc.no/copyright\_disclaimer/.

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## Reference Data Publicly Available

All data sets in this AMR package (about microorganisms, antibiotics, SIR interpretation, EUCAST rules, etc.) are publicly and freely available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. We also provide tab-separated plain text files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

#### See Also

- antibiotics for the data.frame that is being used to determine ATCs
- ab\_from\_text() for a function to retrieve antimicrobial drugs from clinical text (from health care records)

```
# these examples all return "ERY", the ID of erythromycin:
as.ab("J01FA01")
as.ab("J 01 FA 01")
as.ab("Erythromycin")
as.ab("eryt")
as.ab("
         eryt 123")
as.ab("ERYT")
as.ab("ERY")
as.ab("eritromicine") # spelled wrong, yet works
as.ab("Erythrocin") # trade name
as.ab("Romycin") # trade name
# spelling from different languages and dyslexia are no problem
ab_atc("ceftriaxon")
ab_atc("cephtriaxone") # small spelling error
ab_atc("cephthriaxone") # or a bit more severe
ab_atc("seephthriaaksone") # and even this works
# use ab_* functions to get a specific properties (see ?ab_property);
# they use as.ab() internally:
ab_name("J01FA01")
ab_name("eryt")
if (require("dplyr")) {
  # you can quickly rename 'sir' columns using set_ab_names() with dplyr:
  example_isolates %>%
    set_ab_names(where(is.sir), property = "atc")
}
```

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as.av

Transform Input to an Antiviral Drug ID

## **Description**

Use this function to determine the antiviral drug code of one or more antiviral drugs. The data set antivirals will be searched for abbreviations, official names and synonyms (brand names).

## Usage

```
as.av(x, flag_multiple_results = TRUE, info = interactive(), ...)
is.av(x)
```

### **Arguments**

## Details

All entries in the antivirals data set have three different identifiers: a human readable EARS-Net code (column ab, used by ECDC and WHONET), an ATC code (column atc, used by WHO), and a CID code (column cid, Compound ID, used by PubChem). The data set contains more than 5,000 official brand names from many different countries, as found in PubChem. Not that some drugs contain multiple ATC codes.

All these properties will be searched for the user input. The as.av() can correct for different forms of misspelling:

- Wrong spelling of drug names (such as "acyclovir"), which corrects for most audible similarities such as f/ph, x/ks, c/z/s, t/th, etc.
- Too few or too many vowels or consonants
- Switching two characters (such as "aycclovir", often the case in clinical data, when doctors typed too fast)
- Digitalised paper records, leaving artefacts like 0/o/O (zero and O's), B/8, n/r, etc.

Use the av\_\* functions to get properties based on the returned antiviral drug ID, see Examples.

Note: the as.av() and av\_\* functions may use very long regular expression to match brand names of antimicrobial drugs. This may fail on some systems.

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#### Value

A character vector with additional class ab

#### Source

```
World Health Organization (WHO) Collaborating Centre for Drug Statistics Methodology: https://www.whocc.no/atc_ddd_index/
```

European Commission Public Health PHARMACEUTICALS - COMMUNITY REGISTER: https://ec.europa.eu/health/documents/community-register/html/reg\_hum\_atc.htm

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#### See Also

- antivirals for the data.frame that is being used to determine ATCs
- av\_from\_text() for a function to retrieve antimicrobial drugs from clinical text (from health care records)

```
# these examples all return "ACI", the ID of aciclovir:
as.av("J05AB01")
as.av("J 05 AB 01")
as.av("Aciclovir")
as.av("aciclo")
as.av(" aciclo 123")
as.av("ACICL")
```

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```
as.av("ACI")
as.av("Virorax") # trade name
as.av("Zovirax") # trade name
as.av("acyklofir") # severe spelling error, yet works
# use av_* functions to get a specific properties (see ?av_property);
# they use as.av() internally:
av_name("J05AB01")
av_name("acicl")
```

as.disk

Transform Input to Disk Diffusion Diameters

# **Description**

This transforms a vector to a new class disk, which is a disk diffusion growth zone size (around an antibiotic disk) in millimetres between 6 and 50.

## Usage

```
as.disk(x, na.rm = FALSE)
NA_disk_
is.disk(x)
```

# **Arguments**

x vector
na.rm a logical indicating whether missing values should be removed

## Format

An object of class disk (inherits from integer) of length 1.

# **Details**

Interpret disk values as SIR values with as.sir(). It supports guidelines from EUCAST and CLSI.

Disk diffusion growth zone sizes must be between 6 and 50 millimetres. Values higher than 50 but lower than 100 will be maximised to 50. All others input values outside the 6-50 range will return NA.

NA\_disk\_ is a missing value of the new disk class.

## Value

An integer with additional class disk

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# See Also

```
as.sir()
```

## **Examples**

```
# transform existing disk zones to the `disk` class (using base R)
df <- data.frame(</pre>
 microorganism = "Escherichia coli",
  AMP = 20,
  CIP = 14,
  GEN = 18,
  TOB = 16
df[, 2:5] <- lapply(df[, 2:5], as.disk)</pre>
str(df)
# transforming is easier with dplyr:
if (require("dplyr")) {
  df %>% mutate(across(AMP:TOB, as.disk))
# interpret disk values, see ?as.sir
as.sir(
  x = as.disk(18),
 mo = "Strep pneu", # `mo` will be coerced with as.mo()
  ab = "ampicillin", # and `ab` with as.ab()
  guideline = "EUCAST"
)
# interpret whole data set, pretend to be all from urinary tract infections:
as.sir(df, uti = TRUE)
```

as.mic

Transform Input to Minimum Inhibitory Concentrations (MIC)

## **Description**

This transforms vectors to a new class mic, which treats the input as decimal numbers, while maintaining operators (such as ">=") and only allowing valid MIC values known to the field of (medical) microbiology.

## Usage

```
as.mic(x, na.rm = FALSE)
NA_mic_
```

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```
is.mic(x)
## S3 method for class 'mic'
droplevels(x, as.mic = FALSE, ...)
```

### **Arguments**

```
x a character or numeric vector

na.rm a logical indicating whether missing values should be removed

as.mic a logical to indicate whether the mic class should be kept - the default is FALSE

... arguments passed on to methods
```

## **Details**

To interpret MIC values as SIR values, use as.sir() on MIC values. It supports guidelines from EUCAST (2011-2023) and CLSI (2011-2023).

This class for MIC values is a quite a special data type: formally it is an ordered factor with valid MIC values as factor levels (to make sure only valid MIC values are retained), but for any mathematical operation it acts as decimal numbers:

```
x <- random_mic(10)
#> Class 'mic'
  [1] 16
               1
                      8
                              8
                                     64
                                            >=128 0.0625 32
                                                                  32
                                                                          16
is.factor(x)
#> [1] TRUE
x[1] * 2
#> [1] 32
median(x)
#> [1] 26
```

This makes it possible to maintain operators that often come with MIC values, such ">=" and "<=", even when filtering using numeric values in data analysis, e.g.:

```
x[x > 4]
#> Class 'mic'
#> [1] 16
                    8
                           64
                                 >=128 32
                                               32
                                                     16
df <- data.frame(x, hospital = "A")</pre>
subset(df, x > 4) \# or with dplyr: df %>% filter(x > 4)
#>
          x hospital
#> 1
         16
                    Α
#> 5
                    Α
#> 6 >=128
                    Α
```

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```
#> 8 32 A
#> 9 32 A
#> 10 16 A
```

The following generic functions are implemented for the MIC class: !, !=, %%, %/%, &, \*, +, -, /, <, <=, ==, >, >=, ^, |, abs(), acos(), acosh(), all(), any(), asin(), asinh(), atan(), atanh(), ceiling(), cos(), cosh(), cospi(), cummax(), cummin(), cumprod(), cumsum(), digamma(), exp(), expm1(), floor(), gamma(), lgamma(), log(), log1p(), log2(), log10(), max(), mean(), min(), prod(), range(), round(), sign(), signif(), sin(), sinh(), sinpi(), sqrt(), sum(), tanh(), tanpi(), trigamma() and trunc(). Some functions of the stats package are also implemented: median(), quantile(), mad(), IQR(), fivenum(). Also, boxplot.stats() is supported. Since sd() and var() are non-generic functions, these could not be extended. Use mad() as an alternative, or use e.g. sd(as.numeric(x)) where x is your vector of MIC values.

Using as.double() or as.numeric() on MIC values will remove the operators and return a numeric vector. Do **not** use as.integer() on MIC values as by the R convention on factors, it will return the index of the factor levels (which is often useless for regular users).

Use droplevels() to drop unused levels. At default, it will return a plain factor. Use droplevels(..., as.mic = TRUE) to maintain the mic class.

NA\_mic\_ is a missing value of the new mic class, analogous to e.g. base R's NA\_character\_.

#### Value

Ordered factor with additional class mic, that in mathematical operations acts as decimal numbers. Bare in mind that the outcome of any mathematical operation on MICs will return a numeric value.

## See Also

```
as.sir()
```

```
mic_data <- as.mic(c(">=32", "1.0", "1", "1.00", 8, "<=0.128", "8", "16", "16"))
mic_data
is.mic(mic_data)
# this can also coerce combined MIC/SIR values:
as.mic("<=0.002; S")
# mathematical processing treats MICs as numeric values
fivenum(mic_data)
quantile(mic_data)
all(mic_data < 512)
# interpret MIC values
as.sir(
 x = as.mic(2),
 mo = as.mo("Streptococcus pneumoniae"),
 ab = "AMX",
 guideline = "EUCAST"
)
```

```
as.sir(
  x = as.mic(c(0.01, 2, 4, 8)),
  mo = as.mo("Streptococcus pneumoniae"),
  ab = "AMX",
  guideline = "EUCAST"
)

# plot MIC values, see ?plot
plot(mic_data)
plot(mic_data, mo = "E. coli", ab = "cipro")

if (require("ggplot2")) {
  autoplot(mic_data, mo = "E. coli", ab = "cipro")
}
if (require("ggplot2")) {
  autoplot(mic_data, mo = "E. coli", ab = "cipro", language = "nl") # Dutch
}
```

as.mo

Transform Arbitrary Input to Valid Microbial Taxonomy

# **Description**

Use this function to get a valid microorganism code (mo) based on arbitrary user input. Determination is done using intelligent rules and the complete taxonomic tree of the kingdoms Animalia, Archaea, Bacteria, and Protozoa, and most microbial species from the kingdom Fungi (see *Source*). The input can be almost anything: a full name (like "Staphylococcus aureus"), an abbreviated name (such as "S. aureus"), an abbreviation known in the field (such as "MRSA"), or just a genus. See *Examples*.

# Usage

```
as.mo(
    x,
    Becker = FALSE,
    Lancefield = FALSE,
    minimum_matching_score = NULL,
    keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
    reference_df = get_mo_source(),
    ignore_pattern = getOption("AMR_ignore_pattern", NULL),
    cleaning_regex = getOption("AMR_cleaning_regex", mo_cleaning_regex()),
    language = get_AMR_locale(),
    info = interactive(),
    ...
)

is.mo(x)
```

```
mo_uncertainties()
mo_renamed()
mo_failures()
mo_reset_session()
mo_cleaning_regex()
```

#### **Arguments**

x a character vector or a data.frame with one or two columns

Becker a logical to indicate whether staphylococci should be categorised into coagulase-

negative staphylococci ("CoNS") and coagulase-positive staphylococci ("CoPS") instead of their own species, according to Karsten Becker *et al.* (see *Source*). Please see *Details* for a full list of staphylococcal species that will be converted. This excludes *Staphylococcus aureus* at default, use Becker = "al1" to also cat-

egorise S. aureus as "CoPS".

Lancefield a logical to indicate whether a beta-haemolytic Streptococcus should be cate-

gorised into Lancefield groups instead of their own species, according to Rebecca C. Lancefield (see *Source*). These streptococci will be categorised in their first group, e.g. *Streptococcus dysgalactiae* will be group C, although officially it was also categorised into groups G and L. . Please see *Details* for a full list of

streptococcal species that will be converted.

This excludes enterococci at default (who are in group D), use Lancefield =

"all" to also categorise all enterococci as group D.

minimum\_matching\_score

a numeric value to set as the lower limit for the MO matching score. When left blank, this will be determined automatically based on the character length of x,

its taxonomic kingdom and human pathogenicity.

keep\_synonyms a logical to indicate if old, previously valid taxonomic names must be preserved

and not be corrected to currently accepted names. The default is FALSE, which will return a note if old taxonomic names were processed. The default can be set with the package option AMR\_keep\_synonyms, i.e. options(AMR\_keep\_synonyms

= TRUE) or options(AMR\_keep\_synonyms = FALSE).

reference\_df a data.frame to be used for extra reference when translating x to a valid mo. See

set\_mo\_source() and get\_mo\_source() to automate the usage of your own

codes (e.g. used in your analysis or organisation).

ignore\_pattern a Perl-compatible regular expression (case-insensitive) of which all matches in

x must return NA. This can be convenient to exclude known non-relevant input and can also be set with the package option AMR\_ignore\_pattern, e.g. options(AMR\_ignore\_pattern = "(not reported|contaminated flora)").

cleaning\_regex a Perl-compatible regular expression (case-insensitive) to clean the input of x.

Every matched part in x will be removed. At default, this is the outcome of mo\_cleaning\_regex(), which removes texts between brackets and texts such

as "species" and "serovar". The default can be set with the package option AMR\_cleaning\_regex.

language language to translate text like "no growth", which defaults to the system language (see get\_AMR\_locale())

info a logical to indicate if a progress bar should be printed if more than 25 items are to be coerced - the default is TRUE only in interactive mode

... other arguments passed on to functions

#### **Details**

A microorganism (MO) code from this package (class: mo) is human readable and typically looks like these examples:

```
Code
                   Full name
B_KLBSL
                   Klebsiella
B_KLBSL_PNMN
                   Klebsiella pneumoniae
B_KLBSL_PNMN_RHNS Klebsiella pneumoniae rhinoscleromatis
         Ι
    1
              \---> subspecies, a 3-5 letter acronym
    1
         \----> species, a 3-6 letter acronym
    \---> genus, a 4-8 letter acronym
\----> taxonomic kingdom: A (Archaea), AN (Animalia), B (Bacteria),
                          F (Fungi), PL (Plantae), P (Protozoa)
```

Values that cannot be coerced will be considered 'unknown' and will be returned as the MO code UNKNOWN with a warning.

Use the mo\_\* functions to get properties based on the returned code, see *Examples*.

The as.mo() function uses a novel matching score algorithm (see *Matching Score for Microorganisms* below) to match input against the available microbial taxonomy in this package. This will lead to the effect that e.g. "E. coli" (a microorganism highly prevalent in humans) will return the microbial ID of *Escherichia coli* and not *Entamoeba coli* (a microorganism less prevalent in humans), although the latter would alphabetically come first.

With Becker = TRUE, the following 85 staphylococci will be converted to the **coagulase-negative group**: S. argensis, S. arlettae, S. auricularis, S. borealis, S. caeli, S. caledonicus, S. canis, S. capitis, S. capitis capitis, S. capitis urealyticus, S. capitis ureolyticus, S. caprae, S. carnosus, S. carnosus carnosus, S. carnosus utilis, S. casei, S. caseolyticus, S. chromogenes, S. cohnii, S. cohnii cohnii, S. cohnii urealyticum, S. cohnii urealyticus, S. condimenti, S. croceilyticus, S. debuckii, S. devriesei, S. durrellii, S. edaphicus, S. epidermidis, S. equorum, S. equorum equorum, S. equorum linens, S. felis, S. fleurettii, S. gallinarum, S. haemolyticus, S. hominis, S. hominis hominis, S. hominis novobiosepticus, S. jettensis, S. kloosii, S. lentus, S. lloydii, S. lugdunensis, S. massiliensis, S. microti, S. muscae, S. nepalensis, S. pasteuri, S. petrasii, S. petrasii croceilyticus, S. petrasii jettensis, S. petrasii petrasii, S. petrasii pragensis, S. pettenkoferi, S. piscifermentans, S. pragensis, S. pseudoxylosus, S. pulvereri, S. ratti, S. rostri, S. saccharolyticus, S. saprophyticus, S. sciuri carnaticus, S. sciuri lentus, S. sciuri rodentium, S. sciuri sciuri, S. simulans, S. stepanovicii, S.

succinus, S. succinus casei, S. succinus succinus, S. taiwanensis, S. urealyticus, S. ureilyticus, S. veratri, S. vitulinus, S. vitulus, S. warneri, and S. xylosus.

The following 16 staphylococci will be converted to the **coagulase-positive group**: S. agnetis, S. argenteus, S. coagulans, S. cornubiensis, S. delphini, S. hyicus, S. hyicus chromogenes, S. hyicus hyicus, S. intermedius, S. lutrae, S. pseudintermedius, S. roterodami, S. schleiferi coagulans, S. schweitzeri, S. simiae, and S. singaporensis.

With Lancefield = TRUE, the following streptococci will be converted to their corresponding Lancefield group: S. agalactiae (Group B), S. anginosus anginosus (Group F), S. anginosus whileyi (Group F), S. anginosus (Group F), S. canis (Group G), S. dysgalactiae dysgalactiae (Group C), S. dysgalactiae equisimilis (Group C), S. dysgalactiae (Group C), S. equi equi (Group C), S. equi ruminatorum (Group C), S. equi zooepidemicus (Group C), S. equi (Group C), S. pyogenes (Group A), S. salivarius salivarius (Group K), S. salivarius thermophilus (Group K), S. salivarius (Group K), and S. sanguinis (Group H).

## **Coping with Uncertain Results:**

Results of non-exact taxonomic input are based on their matching score. The lowest allowed score can be set with the minimum\_matching\_score argument. At default this will be determined based on the character length of the input, and the taxonomic kingdom and human pathogenicity of the taxonomic outcome. If values are matched with uncertainty, a message will be shown to suggest the user to evaluate the results with mo\_uncertainties(), which returns a data.frame with all specifications.

To increase the quality of matching, the cleaning\_regex argument can be used to clean the input (i.e., x). This must be a regular expression that matches parts of the input that should be removed before the input is matched against the available microbial taxonomy. It will be matched Perl-compatible and case-insensitive. The default value of cleaning\_regex is the outcome of the helper function mo\_cleaning\_regex().

There are three helper functions that can be run after using the as.mo() function:

- Use mo\_uncertainties() to get a data.frame that prints in a pretty format with all taxonomic names that were guessed. The output contains the matching score for all matches (see *Matching Score for Microorganisms* below).
- Use mo\_failures() to get a character vector with all values that could not be coerced to a valid value.
- Use mo\_renamed() to get a data.frame with all values that could be coerced based on old, previously accepted taxonomic names.

## **Microbial Prevalence of Pathogens in Humans:**

The coercion rules consider the prevalence of microorganisms in humans, which is available as the prevalence column in the microorganisms data set. The grouping into human pathogenic prevalence is explained in the section *Matching Score for Microorganisms* below.

#### Value

A character vector with additional class mo

#### Source

1. Berends MS *et al.* (2022). **AMR: An R Package for Working with Antimicrobial Resistance Data**. *Journal of Statistical Software*, 104(3), 1-31; doi:10.18637/jss.v104.i03

 Becker K et al. (2014). Coagulase-Negative Staphylococci. Clin Microbiol Rev. 27(4): 870-926; doi:10.1128/CMR.0010913

- 3. Becker K et al. (2019). Implications of identifying the recently defined members of the S. aureus complex, S. argenteus and S. schweitzeri: A position paper of members of the ESCMID Study Group for staphylococci and Staphylococcal Diseases (ESGS). Clin Microbiol Infect; doi:10.1016/j.cmi.2019.02.028
- 4. Becker K *et al.* (2020). **Emergence of coagulase-negative staphylococci.** *Expert Rev Anti Infect Ther.* 18(4):349-366; doi:10.1080/14787210.2020.1730813
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- Berends MS et al. (2022). Trends in Occurrence and Phenotypic Resistance of Coagulase-Negative Staphylococci (CoNS) Found in Human Blood in the Northern Netherlands between 2013 and 2019/ Micro.rganisms 10(9), 1801; doi:10.3390/microorganisms10091801
- Parte, AC et al. (2020). List of Prokaryotic names with Standing in Nomenclature (LPSN) moves to the DSMZ. International Journal of Systematic and Evolutionary Microbiology, 70, 5607-5612; doi:10.1099/ijsem.0.004332. Accessed from https://lpsn.dsmz.de on December 11th, 2022.
- 8. GBIF Secretariat (2022). GBIF Backbone Taxonomy. Checklist dataset doi:10.15468/39omei. Accessed from https://www.gbif.org on December 11th, 2022.
- 9. Reimer, LC *et al.* (2022). *BacDive* in 2022: the knowledge base for standardized bacterial and archaeal data. Nucleic Acids Res., 50(D1):D741-D74; doi:10.1093/nar/gkab961. Accessed from https://bacdive.dsmz.de on May 12th, 2023.
- Public Health Information Network Vocabulary Access and Distribution System (PHIN VADS).
   US Edition of SNOMED CT from 1 September 2020. Value Set Name 'Microorganism', OID 2.16.840.1.114222.4.11.1009 (v12). URL: https://phinvads.cdc.gov
- 11. Bartlett A *et al.* (2022). **A comprehensive list of bacterial pathogens infecting humans** *Microbiology* 168:001269; doi:10.1099/mic.0.001269

## **Matching Score for Microorganisms**

With ambiguous user input in as.mo() and all the mo\_\* functions, the returned results are chosen based on their matching score using mo\_matching\_score(). This matching score m, is calculated as:

$$m_{(x,n)} = \frac{l_n - 0.5 \cdot \min \begin{cases} l_n \\ \text{lev}(x,n) \end{cases}}{l_n \cdot p_n \cdot k_n}$$

where:

- x is the user input;
- n is a taxonomic name (genus, species, and subspecies);
- $l_n$  is the length of n;
- lev is the Levenshtein distance function (counting any insertion as 1, and any deletion or substitution as 2) that is needed to change x into n;

- $p_n$  is the human pathogenic prevalence group of n, as described below;
- $k_n$  is the taxonomic kingdom of n, set as Bacteria = 1, Fungi = 1.25, Protozoa = 1.5, Archaea = 2, others = 3.

The grouping into human pathogenic prevalence p is based on recent work from Bartlett  $et \, al.$  (2022, doi:10.1099/mic.0.001269) who extensively studied medical-scientific literature to categorise all bacterial species into these groups:

- **Established**, if a taxonomic species has infected at least three persons in three or more references. These records have prevalence = 1.0 in the microorganisms data set;
- **Putative**, if a taxonomic species has fewer than three known cases. These records have prevalence = 1.25 in the microorganisms data set.

#### Furthermore,

- Any genus present in the established list also has prevalence = 1.0 in the microorganisms data set;
- Any other genus present in the putative list has prevalence = 1.25 in the microorganisms data set:
- Any other species or subspecies of which the genus is present in the two aforementioned groups, has prevalence = 1.5 in the microorganisms data set;
- Any non-bacterial genus, species or subspecies of which the genus is present in the following list, has prevalence = 1.25 in the microorganisms data set: Absidia, Acanthamoeba, Acremonium, Aedes, Alternaria, Amoeba, Ancylostoma, Angiostrongylus, Anisakis, Anopheles, Apophysomyces, Aspergillus, Aureobasidium, Basidiobolus, Beauveria, Blastocystis, Blastomyces, Candida, Capillaria, Chaetomium, Chrysonilia, Cladophialophora, Cladosporium, Conidiobolus, Contracaecum, Cordylobia, Cryptococcus, Curvularia, Demodex, Dermatobia, Dientamoeba, Diphyllobothrium, Dirofilaria, Echinostoma, Entamoeba, Enterobius, Exophiala, Exserohilum, Fasciola, Fonsecaea, Fusarium, Giardia, Haloarcula, Halobacterium, Halococcus, Hendersonula, Heterophyes, Histomonas, Histoplasma, Hymenolepis, Hypomyces, Hysterothylacium, Leishmania, Malassezia, Malbranchea, Metagonimus, Meyerozyma, Microsporidium, Microsporum, Mortierella, Mucor, Mycocentrospora, Necator, Nectria, Ochroconis, Oesophagostomum, Oidiodendron, Opisthorchis, Pediculus, Penicillium, Phlebotomus, Phoma, Pichia, Piedraia, Pithomyces, Pityrosporum, Pneumocystis, Pseudallescheria, Pseudoterranova, Pulex, Rhizomucor, Rhizopus, Rhodotorula, Saccharomyces, Sarcoptes, Scolecobasidium, Scopulariopsis, Scytalidium, Spirometra, Sporobolomyces, Stachybotrys, Strongyloides, Syngamus, Taenia, Talaromyces, Toxocara, Trichinella, Trichobilharzia, Trichoderma, Trichomonas, Trichophyton, Trichosporon, Trichostrongylus, Trichuris, Tritirachium, Trombicula, Trypanosoma, Tunga, or Wuchereria;
- All other records have prevalence = 2.0 in the microorganisms data set.

When calculating the matching score, all characters in x and n are ignored that are other than A-Z, a-z, 0-9, spaces and parentheses.

All matches are sorted descending on their matching score and for all user input values, the top match will be returned. This will lead to the effect that e.g., "E. coli" will return the microbial ID of *Escherichia coli* (m=0.688, a highly prevalent microorganism found in humans) and not *Entamoeba coli* (m=0.381, a less prevalent microorganism in humans), although the latter would alphabetically come first.

## Reference Data Publicly Available

All data sets in this AMR package (about microorganisms, antibiotics, SIR interpretation, EUCAST rules, etc.) are publicly and freely available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. We also provide tab-separated plain text files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

#### See Also

microorganisms for the data.frame that is being used to determine ID's.

The mo\_\* functions (such as mo\_genus(), mo\_gramstain()) to get properties based on the returned code.

```
# These examples all return "B_STPHY_AURS", the ID of S. aureus:
  "sau", # WHONET code
  "stau",
  "STAU",
  "staaur",
 "S. aureus"
 "S aureus",
 "Sthafilokkockus aureus", # handles incorrect spelling
  "Staphylococcus aureus (MRSA)",
  "MRSA", # Methicillin Resistant S. aureus
  "VISA", # Vancomycin Intermediate S. aureus
 "VRSA", # Vancomycin Resistant S. aureus
 115329001 # SNOMED CT code
))
# Dyslexia is no problem - these all work:
as.mo(c(
  "Ureaplasma urealyticum",
  "Ureaplasma urealyticus",
  "Ureaplasmium urealytica",
  "Ureaplazma urealitycium"
))
# input will get cleaned up with the input given in the `cleaning_regex` argument,
# which defaults to `mo_cleaning_regex()`:
cat(mo_cleaning_regex(), "\n")
as.mo("Streptococcus group A")
as.mo("S. epidermidis") # will remain species: B_STPHY_EPDR
as.mo("S. epidermidis", Becker = TRUE) # will not remain species: B_STPHY_CONS
as.mo("S. pyogenes") # will remain species: B_STRPT_PYGN
```

```
as.mo("S. pyogenes", Lancefield = TRUE) # will not remain species: B_STRPT_GRPA
# All mo_* functions use as.mo() internally too (see ?mo_property):
mo_genus("E. coli")
mo_gramstain("ESCO")
mo_is_intrinsic_resistant("ESCCOL", ab = "vanco")
```

as.sir

Translate MIC and Disk Diffusion to SIR, or Clean Existing SIR Data

# **Description**

Interpret minimum inhibitory concentration (MIC) values and disk diffusion diameters according to EUCAST or CLSI, or clean up existing SIR values. This transforms the input to a new class sir, which is an ordered factor with levels S < I < R.

Currently available **breakpoint guidelines** are EUCAST 2011-2023 and CLSI 2011-2023, and available **breakpoint types** are "ECOFF", "animal", and "human".

All breakpoints used for interpretation are publicly available in the clinical\_breakpoints data set.

## Usage

```
as.sir(x, ...)
NA_sir_
is.sir(x)
is_sir_eligible(x, threshold = 0.05)
## S3 method for class 'mic'
as.sir(
  Х,
  mo = NULL,
  ab = deparse(substitute(x)),
  guideline = getOption("AMR_guideline", "EUCAST"),
  uti = NULL,
  conserve_capped_values = FALSE,
  add_intrinsic_resistance = FALSE,
  reference_data = AMR::clinical_breakpoints,
  include_screening = getOption("AMR_include_screening", FALSE),
  include_PKPD = getOption("AMR_include_PKPD", TRUE),
  breakpoint_type = getOption("AMR_breakpoint_type", "human"),
)
## S3 method for class 'disk'
```

```
as.sir(
  Х,
 mo = NULL,
  ab = deparse(substitute(x)),
  guideline = getOption("AMR_guideline", "EUCAST"),
  uti = NULL,
  add_intrinsic_resistance = FALSE,
  reference_data = AMR::clinical_breakpoints,
  include_screening = getOption("AMR_include_screening", FALSE),
  include_PKPD = getOption("AMR_include_PKPD", TRUE),
  breakpoint_type = getOption("AMR_breakpoint_type", "human"),
)
## S3 method for class 'data.frame'
as.sir(
 х,
  . . . ,
  col_mo = NULL,
  guideline = getOption("AMR_guideline", "EUCAST"),
  uti = NULL,
  conserve_capped_values = FALSE,
  add_intrinsic_resistance = FALSE,
  reference_data = AMR::clinical_breakpoints,
  include_screening = getOption("AMR_include_screening", FALSE),
  include_PKPD = getOption("AMR_include_PKPD", TRUE),
  breakpoint_type = getOption("AMR_breakpoint_type", "human")
)
sir_interpretation_history(clean = FALSE)
```

# **Arguments**

Х	vector of values (for class mic: MIC values in mg/L, for class disk: a disk diffusion radius in millimetres)
	for using on a data.frame: names of columns to apply as.sir() on (supports tidy selection such as column1:column4). Otherwise: arguments passed on to methods.
threshold	maximum fraction of invalid antimicrobial interpretations of x, see Examples
mo	any (vector of) text that can be coerced to valid microorganism codes with as.mo(), can be left empty to determine it automatically
ab	any (vector of) text that can be coerced to a valid antimicrobial drug code with as.ab()
guideline	defaults to EUCAST 2023 (the latest implemented EUCAST guideline in the clinical_breakpoints data set), but can be set with the package option AMR_guideline. Currently supports EUCAST (2011-2023) and CLSI (2011-2023), see <i>Details</i> .

uti

(Urinary Tract Infection) A vector with logicals (TRUE or FALSE) to specify whether a UTI specific interpretation from the guideline should be chosen. For using as.sir() on a data frame, this can also be a column containing logicals or when left blank, the data set will be searched for a column 'specimen', and rows within this column containing 'urin' (such as 'urine', 'urina') will be regarded isolates from a UTI. See Examples.

# conserve\_capped\_values

a logical to indicate that MIC values starting with ">" (but not ">=") must always return "R", and that MIC values starting with "<" (but not "<=") must always return "S"

# add\_intrinsic\_resistance

(only useful when using a EUCAST guideline) a logical to indicate whether intrinsic antibiotic resistance must also be considered for applicable bug-drug combinations, meaning that e.g. ampicillin will always return "R" in Klebsiella species. Determination is based on the intrinsic resistant data set, that itself is based on 'EUCAST Expert Rules' and 'EUCAST Intrinsic Resistance and Unusual Phenotypes' v3.3 (2021).

reference\_data a data.frame to be used for interpretation, which defaults to the clinical breakpoints data set. Changing this argument allows for using own interpretation guidelines. This argument must contain a data set that is equal in structure to the clinical breakpoints data set (same column names and column types). Please note that the guideline argument will be ignored when reference\_data is manually set.

## include\_screening

a logical to indicate that clinical breakpoints for screening are allowed - the default is FALSE. Can also be set with the package option AMR\_include\_screening.

include\_PKPD

a logical to indicate that PK/PD clinical breakpoints must be applied as a last resort - the default is TRUE. Can also be set with the package option AMR\_include\_PKPD.

## breakpoint\_type

the type of breakpoints to use, either "ECOFF", "animal", or "human". ECOFF stands for Epidemiological Cut-Off values. The default is "human", which can also be set with the package option AMR\_breakpoint\_type.

col\_mo

column name of the names or codes of the microorganisms (see as.mo()) - the default is the first column of class mo. Values will be coerced using as.mo().

clean

a logical to indicate whether previously stored results should be forgotten after returning the 'logbook' with results

#### **Format**

An object of class sir (inherits from ordered, factor) of length 1.

## **Details**

Note: The clinical breakpoints in this package were validated through and imported from WHONET and the public use of this AMR package has been endorsed by CLSI and EUCAST, please see clinical\_breakpoints for more information.

#### **How it Works:**

The as.sir() function works in four ways:

1. For **cleaning raw / untransformed data**. The data will be cleaned to only contain values S, I and R and will try its best to determine this with some intelligence. For example, mixed values with SIR interpretations and MIC values such as "<0.25; S" will be coerced to "S". Combined interpretations for multiple test methods (as seen in laboratory records) such as "S; S" will be coerced to "S", but a value like "S; I" will return NA with a warning that the input is unclear.

- 2. For interpreting minimum inhibitory concentration (MIC) values according to EUCAST or CLSI. You must clean your MIC values first using as .mic(), that also gives your columns the new data class mic. Also, be sure to have a column with microorganism names or codes. It will be found automatically, but can be set manually using the mo argument.
  - Using dplyr, SIR interpretation can be done very easily with either: your\_data %>% mutate\_if(is.mic, as.sir) your\_data %>% mutate(across(where(is.mic), as.sir))
  - Operators like "<=" will be stripped before interpretation. When using conserve\_capped\_values = TRUE, an MIC value of e.g. ">2" will always return "R", even if the breakpoint according to the chosen guideline is ">=4". This is to prevent that capped values from raw laboratory data would not be treated conservatively. The default behaviour (conserve\_capped\_values = FALSE) considers ">2" to be lower than ">=4" and might in this case return "S" or "I".
- 3. For interpreting disk diffusion diameters according to EUCAST or CLSI. You must clean your disk zones first using as.disk(), that also gives your columns the new data class disk. Also, be sure to have a column with microorganism names or codes. It will be found automatically, but can be set manually using the mo argument.
  - Using dplyr, SIR interpretation can be done very easily with either: your\_data %>% mutate\_if(is.disk, as.sir) your\_data %>% mutate(across(where(is.disk), as.sir))
- 4. For **interpreting a complete data set**, with automatic determination of MIC values, disk diffusion diameters, microorganism names or codes, and antimicrobial test results. This is done very simply by running as.sir(your\_data).

For points 2, 3 and 4: Use sir\_interpretation\_history() to retrieve a data.frame (or tibble if the tibble package is installed) with all results of the last as.sir() call.

# **Supported Guidelines:**

For interpreting MIC values as well as disk diffusion diameters, currently implemented guidelines are EUCAST (2011-2023) and CLSI (2011-2023).

Thus, the guideline argument must be set to e.g., "EUCAST 2023" or "CLSI 2023". By simply using "EUCAST" (the default) or "CLSI" as input, the latest included version of that guideline will automatically be selected. You can set your own data set using the reference\_data argument. The guideline argument will then be ignored.

You can set the default guideline with the package option AMR\_guideline (e.g. in your . Rprofile file), such as:

```
options(AMR_guideline = "CLSI")
options(AMR_guideline = "CLSI 2018")
options(AMR_guideline = "EUCAST 2020")
# or to reset:
options(AMR_guideline = NULL)
```

### **After Interpretation:**

After using as.sir(), you can use the eucast\_rules() defined by EUCAST to (1) apply inferred susceptibility and resistance based on results of other antimicrobials and (2) apply intrinsic resistance based on taxonomic properties of a microorganism.

### **Machine-Readable Clinical Breakpoints:**

The repository of this package contains a machine-readable version of all guidelines. This is a CSV file consisting of 29 747 rows and 12 columns. This file is machine-readable, since it contains one row for every unique combination of the test method (MIC or disk diffusion), the antimicrobial drug and the microorganism. **This allows for easy implementation of these rules in laboratory information systems (LIS)**. Note that it only contains interpretation guidelines for humans - interpretation guidelines from CLSI for animals were removed.

#### Other:

The function is.sir() detects if the input contains class sir. If the input is a data.frame, it iterates over all columns and returns a logical vector.

The function is\_sir\_eligible() returns TRUE when a columns contains at most 5% invalid antimicrobial interpretations (not S and/or I and/or R), and FALSE otherwise. The threshold of 5% can be set with the threshold argument. If the input is a data.frame, it iterates over all columns and returns a logical vector.

NA\_sir\_ is a missing value of the new sir class, analogous to e.g. base R's NA\_character\_.

#### Value

Ordered factor with new class sir

### **Interpretation of SIR**

In 2019, the European Committee on Antimicrobial Susceptibility Testing (EUCAST) has decided to change the definitions of susceptibility testing categories S, I, and R as shown below (https://www.eucast.org/newsiandr):

#### • S - Susceptible, standard dosing regimen

A microorganism is categorised as "Susceptible, standard dosing regimen", when there is a high likelihood of therapeutic success using a standard dosing regimen of the agent.

# • I - Susceptible, increased exposure

A microorganism is categorised as "Susceptible, Increased exposure" when there is a high likelihood of therapeutic success because exposure to the agent is increased by adjusting the dosing regimen or by its concentration at the site of infection.

## • R = Resistant

A microorganism is categorised as "Resistant" when there is a high likelihood of therapeutic failure even when there is increased exposure.

Exposure is a function of how the mode of administration, dose, dosing interval, infusion time, as well as distribution and excretion of the antimicrobial agent will influence the infecting organism at the site of infection.

This AMR package honours this insight. Use susceptibility() (equal to proportion\_SI()) to determine antimicrobial susceptibility and count\_susceptible() (equal to count\_SI()) to count susceptible isolates.

## Reference Data Publicly Available

All data sets in this AMR package (about microorganisms, antibiotics, SIR interpretation, EUCAST rules, etc.) are publicly and freely available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. We also provide tab-separated plain text files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

#### Source

For interpretations of minimum inhibitory concentration (MIC) values and disk diffusion diameters:

- M39 Analysis and Presentation of Cumulative Antimicrobial Susceptibility Test Data, 2011-2023, Clinical and Laboratory Standards Institute (CLSI). https://clsi.org/standards/products/microbiology/documents/m39/.
- M100 Performance Standard for Antimicrobial Susceptibility Testing, 2011-2023, Clinical and Laboratory Standards Institute (CLSI). https://clsi.org/standards/products/microbiology/documents/m100/.
- Breakpoint tables for interpretation of MICs and zone diameters, 2011-2023, European Committee on Antimicrobial Susceptibility Testing (EUCAST). https://www.eucast.org/clinical\_breakpoints.

#### See Also

```
as.mic(), as.disk(), as.mo()
```

```
example_isolates
summary(example_isolates) # see all SIR results at a glance
# For INTERPRETING disk diffusion and MIC values -------
# a whole data set, even with combined MIC values and disk zones
df <- data.frame(</pre>
 microorganism = "Escherichia coli",
 AMP = as.mic(8),
 CIP = as.mic(0.256),
 GEN = as.disk(18),
 TOB = as.disk(16),
 ERY = "R"
)
as.sir(df)
# return a 'logbook' about the results:
sir_interpretation_history()
# for single values
as.sir(
 x = as.mic(2),
```

```
mo = as.mo("S. pneumoniae"),
  ab = "AMP",
  guideline = "EUCAST"
)
as.sir(
  x = as.disk(18),
  mo = "Strep pneu", # `mo` will be coerced with as.mo()
 ab = "ampicillin", # and `ab` with as.ab()
  guideline = "EUCAST"
)
# the dplyr way
if (require("dplyr")) {
  df %>% mutate_if(is.mic, as.sir)
  df %>% mutate_if(function(x) is.mic(x) | is.disk(x), as.sir)
  df %>% mutate(across(where(is.mic), as.sir))
  df %>% mutate_at(vars(AMP:TOB), as.sir)
  df %>% mutate(across(AMP:TOB, as.sir))
   mutate_at(vars(AMP:TOB), as.sir, mo = .$microorganism)
  # to include information about urinary tract infections (UTI)
  data.frame(
   mo = "E. coli",
   NIT = c(" \le 2", 32),
    from_the_bladder = c(TRUE, FALSE)
  ) %>%
   as.sir(uti = "from_the_bladder")
  data.frame(
   mo = "E. coli",
   NIT = c(" \le 2", 32),
    specimen = c("urine", "blood")
    as.sir() # automatically determines urine isolates
  df %>%
    mutate_at(vars(AMP:TOB), as.sir, mo = "E. coli", uti = TRUE)
}
# For CLEANING existing SIR values ------
as.sir(c("S", "I", "R", "A", "B", "C"))
as.sir("<= 0.002; S") # will return "S"
sir_data <- as.sir(c(rep("S", 474), rep("I", 36), rep("R", 370)))</pre>
is.sir(sir_data)
plot(sir_data) # for percentages
barplot(sir_data) # for frequencies
# the dplyr way
```

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```
if (require("dplyr")) {
  example_isolates %>%
    mutate_at(vars(PEN:RIF), as.sir)
# same:
  example_isolates %>%
    as.sir(PEN:RIF)

# fastest way to transform all columns with already valid AMR results to class `sir`:
  example_isolates %>%
    mutate_if(is_sir_eligible, as.sir)

# since dplyr 1.0.0, this can also be:
  # example_isolates %>%
  # mutate(across(where(is_sir_eligible), as.sir))
}
```

atc\_online\_property

Get ATC Properties from WHOCC Website

# Description

Gets data from the WHOCC website to determine properties of an Anatomical Therapeutic Chemical (ATC) (e.g. an antibiotic), such as the name, defined daily dose (DDD) or standard unit.

## Usage

```
atc_online_property(
  atc_code,
  property,
  administration = "0",
  url = "https://www.whocc.no/atc_ddd_index/?code=%s&showdescription=no",
  url_vet = "https://www.whocc.no/atcvet/atcvet_index/?code=%s&showdescription=no")

atc_online_groups(atc_code, ...)

atc_online_ddd(atc_code, ...)

atc_online_ddd_units(atc_code, ...)
```

#### Arguments

atc\_code a character (vector) with ATC code(s) of antibiotics, will be coerced with as.ab() and ab\_atc() internally if not a valid ATC code

property property of an ATC code. Valid values are "ATC", "Name", "DDD", "U" ("unit"), "Adm.R", "Note" and groups. For this last option, all hierarchical groups of an ATC code will be returned, see *Examples*.

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administration type of administration when using property = "Adm.R", see *Details*url url of website of the WHOCC. The sign %s can be used as a placeholder for ATC codes.

url\_vet url of website of the WHOCC for veterinary medicine. The sign %s can be used as a placeholder for ATC\_vet codes (that all start with "Q").

... arguments to pass on to atc\_property

#### **Details**

Options for argument administration:

- "Implant" = Implant
- "Inhal" = Inhalation
- "Instill" = Instillation
- "N" = nasal
- "0" = oral
- "P" = parenteral
- "R" = rectal
- "SL" = sublingual/buccal
- "TD" = transdermal
- "V" = vaginal

Abbreviations of return values when using property = "U" (unit):

- "g" = gram
- "mg" = milligram
- "mcg" = microgram
- "U" = unit
- "TU" = thousand units
- "MU" = million units
- "mmol" = millimole
- "ml" = millilitre (e.g. eyedrops)

N.B. This function requires an internet connection and only works if the following packages are installed: curl, rvest, xml2.

#### **Source**

https://www.whocc.no/atc\_ddd\_alterations\_cumulative/ddd\_alterations/abbrevations/

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## **Examples**

```
if (requireNamespace("curl") && requireNamespace("rvest") && requireNamespace("xml2")) {
    # oral DDD (Defined Daily Dose) of amoxicillin
    atc_online_property("J01CA04", "DDD", "0")
    atc_online_ddd(ab_atc("amox"))

# parenteral DDD (Defined Daily Dose) of amoxicillin
    atc_online_property("J01CA04", "DDD", "P")

atc_online_property("J01CA04", property = "groups") # search hierarchical groups of amoxicillin
}
```

availability

Check Availability of Columns

# **Description**

Easy check for data availability of all columns in a data set. This makes it easy to get an idea of which antimicrobial combinations can be used for calculation with e.g. susceptibility() and resistance().

# Usage

```
availability(tbl, width = NULL)
```

# **Arguments**

tbl a data.frame or list

width number of characters to present the visual availability - the default is filling the

width of the console

## **Details**

The function returns a data.frame with columns "resistant" and "visual\_resistance". The values in that columns are calculated with resistance().

## Value

data.frame with column names of tbl as row names

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## **Examples**

```
availability(example_isolates)

if (require("dplyr")) {
   example_isolates %>%
    filter(mo == as.mo("Escherichia coli")) %>%
    select_if(is.sir) %>%
    availability()
}
```

av\_from\_text

Retrieve Antiviral Drug Names and Doses from Clinical Text

## Description

Use this function on e.g. clinical texts from health care records. It returns a list with all antiviral drugs, doses and forms of administration found in the texts.

# Usage

```
av_from_text(
   text,
   type = c("drug", "dose", "administration"),
   collapse = NULL,
   translate_av = FALSE,
   thorough_search = NULL,
   info = interactive(),
   ...
)
```

## **Arguments**

text text to analyse

type type of property to search for, either "drug", "dose" or "administration",

see Examples

collapse a character to pass on to paste(, collapse = ...) to only return one character

per element of text, see Examples

translate\_av if type = "drug": a column name of the antivirals data set to translate the antibi-

otic abbreviations to, using av\_property(). The default is FALSE. Using TRUE

is equal to using "name".

thorough\_search

a logical to indicate whether the input must be extensively searched for misspelling and other faulty input values. Setting this to TRUE will take considerably more time than when using FALSE. At default, it will turn TRUE when all input elements contain a maximum of three words.

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info a logical to indicate whether a progress bar should be printed - the default is TRUE only in interactive mode
... arguments passed on to as.av()

#### **Details**

This function is also internally used by as.av(), although it then only searches for the first drug name and will throw a note if more drug names could have been returned. Note: the as.av() function may use very long regular expression to match brand names of antiviral drugs. This may fail on some systems.

# **Argument** type:

At default, the function will search for antiviral drug names. All text elements will be searched for official names, ATC codes and brand names. As it uses as .av() internally, it will correct for misspelling.

With type = "dose" (or similar, like "dosing", "doses"), all text elements will be searched for numeric values that are higher than 100 and do not resemble years. The output will be numeric. It supports any unit (g, mg, IE, etc.) and multiple values in one clinical text, see *Examples*.

With type = "administration" (or abbreviations, like "admin", "adm"), all text elements will be searched for a form of drug administration. It supports the following forms (including common abbreviations): buccal, implant, inhalation, instillation, intravenous, nasal, oral, parenteral, rectal, sublingual, transdermal and vaginal. Abbreviations for oral (such as 'po', 'per os') will become "oral", all values for intravenous (such as 'iv', 'intraven') will become "iv". It supports multiple values in one clinical text, see *Examples*.

# **Argument** collapse:

Without using collapse, this function will return a list. This can be convenient to use e.g. inside a mutate()):

```
df %>% mutate(avx = av_from_text(clinical_text))
```

The returned AV codes can be transformed to official names, groups, etc. with all av\_\* functions such as av\_name() and av\_group(), or by using the translate\_av argument.

```
With using collapse, this function will return a character:

df %>% mutate(avx = av_from_text(clinical_text, collapse = "|"))
```

## Value

A list, or a character if collapse is not NULL

```
av_from_text("28/03/2020 valaciclovir po tid")
av_from_text("28/03/2020 valaciclovir po tid", type = "admin")
```

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av\_property

Get Properties of an Antiviral Drug

# **Description**

Use these functions to return a specific property of an antiviral drug from the antivirals data set. All input values will be evaluated internally with as.av().

### Usage

```
av_name(x, language = get_AMR_locale(), tolower = FALSE, ...)
av_cid(x, ...)
av_synonyms(x, ...)
av_tradenames(x, ...)
av_group(x, language = get_AMR_locale(), ...)
av_atc(x, ...)
av_loinc(x, ...)
av_ddd(x, administration = "oral", ...)
av_ddd_units(x, administration = "oral", ...)
av_info(x, language = get_AMR_locale(), ...)
av_url(x, open = FALSE, ...)
av_property(x, property = "name", language = get_AMR_locale(), ...)
```

# **Arguments**

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#### **Details**

All output will be translated where possible.

The function av\_url() will return the direct URL to the official WHO website. A warning will be returned if the required ATC code is not available.

### Value

- An integer in case of av\_cid()
- A named list in case of av\_info() and multiple av\_atc()/av\_synonyms()/av\_tradenames()
- A double in case of av\_ddd()
- A character in all other cases

#### Source

```
World Health Organization (WHO) Collaborating Centre for Drug Statistics Methodology: https://www.whocc.no/atc_ddd_index/
European Commission Public Health PHARMACEUTICALS - COMMUNITY REGISTER: https://ec.europa.eu/health/documents/community-register/html/reg_hum_atc.htm
```

## **Reference Data Publicly Available**

All data sets in this AMR package (about microorganisms, antibiotics, SIR interpretation, EUCAST rules, etc.) are publicly and freely available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. We also provide tab-separated plain text files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

#### See Also

antivirals

```
# all properties:
av_name("ACI")
av_atc("ACI")
av_cid("ACI")
av_synonyms("ACI")
av_tradenames("ACI")
av_group("ACI")
av_url("ACI")
# lowercase transformation
```

```
av_name(x = c("ACI", "VALA"))
av_name(x = c("ACI", "VALA"), tolower = TRUE)

# defined daily doses (DDD)
av_ddd("ACI", "oral")
av_ddd_units("ACI", "oral")
av_ddd("ACI", "iv")
av_ddd_units("ACI", "iv")
av_info("ACI") # all properties as a list

# all av_* functions use as.av() internally, so you can go from 'any' to 'any':
av_atc("ACI")
av_group("J05AB01")
av_loinc("abacavir")
av_name("29113-8")
av_name("J05AB01")
```

bug\_drug\_combinations Determine Bug-Drug Combinations

## **Description**

Determine antimicrobial resistance (AMR) of all bug-drug combinations in your data set where at least 30 (default) isolates are available per species. Use format() on the result to prettify it to a publishable/printable format, see *Examples*.

# Usage

```
bug_drug_combinations(x, col_mo = NULL, FUN = mo_shortname, ...)
## S3 method for class 'bug_drug_combinations'
format(
    x,
    translate_ab = "name (ab, atc)",
    language = get_AMR_locale(),
    minimum = 30,
    combine_SI = TRUE,
    add_ab_group = TRUE,
    remove_intrinsic_resistant = FALSE,
    decimal.mark = getOption("OutDec"),
    big.mark = ifelse(decimal.mark == ",", ".", ","),
    ...
)
```

## **Arguments**

	X	a data set with antibiotic columns, such as amox, AMX and AMC					
	col_mo	column name of the names or codes of the microorganisms (see $as.mo()$ ) - the default is the first column of class $mo$ . Values will be coerced using $as.mo()$ .					
	FUN	the function to call on the mo column to transform the microorganism codes - the default is $mo\_shortname()$					
		arguments passed on to FUN					
	translate_ab	a character of length 1 containing column names of the antibiotics data set					
	language	language of the returned text - the default is the current system language (see <pre>get_AMR_locale()</pre> ) and can also be set with the <pre>package</pre> option AMR_locale. Use language = NULL or language = "" to prevent translation.					
	minimum	the minimum allowed number of available (tested) isolates. Any isolate count lower than minimum will return NA with a warning. The default number of 30 isolates is advised by the Clinical and Laboratory Standards Institute (CLSI) as best practice, see <i>Source</i> .					
	combine_SI	a logical to indicate whether values S and I should be summed, so resistance will be based on only R - the default is $TRUE$					
	add_ab_group	a logical to indicate where the group of the antimicrobials must be included as a first column					
remove_intrinsic_resistant							
		logical to indicate that rows and columns with 100% resistance for all tested antimicrobials must be removed from the table					
	decimal.mark	the character to be used to indicate the numeric decimal point.					
	big.mark	character; if not empty used as mark between every big.interval decimals $\it before$ (hence big) the decimal point.					

# **Details**

The function format() calculates the resistance per bug-drug combination and returns a table ready for reporting/publishing. Use combine\_SI = TRUE (default) to test R vs. S+I and combine\_SI = FALSE to test R+I vs. S. This table can also directly be used in R Markdown / Quarto without the need for e.g. knitr::kable().

# Value

The function bug\_drug\_combinations() returns a data.frame with columns "mo", "ab", "S", "I", "R" and "total".

```
# example_isolates is a data set available in the AMR package.
# run ?example_isolates for more info.
example_isolates

x <- bug_drug_combinations(example_isolates)</pre>
```

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```
head(x)
format(x, translate_ab = "name (atc)")

# Use FUN to change to transformation of microorganism codes
bug_drug_combinations(example_isolates,
   FUN = mo_gramstain
)

bug_drug_combinations(example_isolates,
   FUN = function(x) {
    ifelse(x == as.mo("Escherichia coli"),
        "E. coli",
        "Others"
    )
   }
)
```

# Description

Data set containing clinical breakpoints to interpret MIC and disk diffusion to SIR values, according to international guidelines. Currently implemented guidelines are EUCAST (2011-2023) and CLSI (2011-2023). Use as.sir() to transform MICs or disks measurements to SIR values.

## Usage

clinical\_breakpoints

#### **Format**

A tibble with 29 747 observations and 12 variables:

- guideline Name of the guideline
- type
  Breakpoint type, either "ECOFF", "animal", or "human"
- method
   Testing method, either "DISK" or "MIC"
- site
   Body site for which the breakpoint must be applied, e.g. "Oral" or "Respiratory"
- mo
   Microbial ID, see as.mo()
- rank\_index
   Taxonomic rank index of mo from 1 (subspecies/infraspecies) to 5 (unknown microorganism)

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- ah
  - Antibiotic code as used by this package, EARS-Net and WHONET, see as.ab()
- ref\_tbl
  - Info about where the guideline rule can be found
- disk\_dose
  - Dose of the used disk diffusion method
- breakpoint\_S
  - Lowest MIC value or highest number of millimetres that leads to "S"
- breakpoint\_R
  - Highest MIC value or lowest number of millimetres that leads to "R"
- uti

A logical value (TRUE/FALSE) to indicate whether the rule applies to a urinary tract infection (UTI)

#### **Details**

### **Different types of breakpoints:**

Supported types of breakpoints are ECOFF, animal, and human. ECOFF (Epidemiological cutoff) values are used in antimicrobial susceptibility testing to differentiate between wild-type and non-wild-type strains of bacteria or fungi.

The default is "human", which can also be set with the package option AMR\_breakpoint\_type. Use as.sir(..., breakpoint\_type = ...) to interpret raw data using a specific breakpoint type, e.g. as.sir(..., breakpoint\_type = "ECOFF") to use ECOFFs.

### **Imported from WHONET:**

Clinical breakpoints in this package were validated through and imported from WHONET, a free desktop Windows application developed and supported by the WHO Collaborating Centre for Surveillance of Antimicrobial Resistance. More can be read on their website. The developers of WHONET and this AMR package have been in contact about sharing their work. We highly appreciate their development on the WHONET software.

# **Response from CLSI and EUCAST:**

The CEO of CLSI and the chairman of EUCAST have endorsed the work and public use of this AMR package (and consequently the use of their breakpoints) in June 2023, when future development of distributing clinical breakpoints was discussed in a meeting between CLSI, EUCAST, the WHO, and developers of WHONET and the AMR package.

### Download:

Like all data sets in this package, this data set is publicly available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. Please visit our website for the download links. The actual files are of course available on our GitHub repository. They allow for machine reading EUCAST and CLSI guidelines, which is almost impossible with the MS Excel and PDF files distributed by EUCAST and CLSI, though initiatives have started to overcome these burdens.

**NOTE:** this AMR package (and the WHONET software as well) contains internal methods to apply the guidelines, which is rather complex. For example, some breakpoints must be applied on certain species groups (which are in case of this package available through the microorganisms.groups data set). It is important that this is considered when using the breakpoints for own use.

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## See Also

intrinsic\_resistant

## **Examples**

clinical\_breakpoints

count

Count Available Isolates

## **Description**

These functions can be used to count resistant/susceptible microbial isolates. All functions support quasiquotation with pipes, can be used in summarise() from the dplyr package and also support grouped variables, see *Examples*.

count\_resistant() should be used to count resistant isolates, count\_susceptible() should be
used to count susceptible isolates.

## Usage

```
count_resistant(..., only_all_tested = FALSE)

count_susceptible(..., only_all_tested = FALSE)

count_R(..., only_all_tested = FALSE)

count_IR(..., only_all_tested = FALSE)

count_I(..., only_all_tested = FALSE)

count_SI(..., only_all_tested = FALSE)

count_S(..., only_all_tested = FALSE)

count_all(..., only_all_tested = FALSE)

n_sir(..., only_all_tested = FALSE)

count_df(
    data,
    translate_ab = "name",
    language = get_AMR_locale(),
    combine_SI = TRUE
)
```

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### **Arguments**

... one or more vectors (or columns) with antibiotic interpretations. They will be transformed internally with as.sir() if needed.

only\_all\_tested

(for combination therapies, i.e. using more than one variable for ...): a logical to indicate that isolates must be tested for all antibiotics, see section *Combina*-

tion Therapy below

data a data.frame containing columns with class sir (see as.sir())

translate\_ab a column name of the antibiotics data set to translate the antibiotic abbreviations

to, using ab\_property()

language language of the returned text - the default is the current system language (see

get\_AMR\_locale()) and can also be set with the package option AMR\_locale.

Use language = NULL or language = "" to prevent translation.

combine\_SI a logical to indicate whether all values of S and I must be merged into one, so

the output only consists of S+I vs. R (susceptible vs. resistant) - the default is

TRUE

#### **Details**

These functions are meant to count isolates. Use the resistance()/susceptibility() functions to calculate microbial resistance/susceptibility.

The function count\_resistant() is equal to the function count\_R(). The function count\_susceptible() is equal to the function count\_SI().

The function  $n_sir()$  is an alias of  $count_all()$ . They can be used to count all available isolates, i.e. where all input antibiotics have an available result (S, I or R). Their use is equal to  $n_distinct()$ . Their function is equal to  $count_susceptible(...) + count_resistant(...)$ .

The function count\_df() takes any variable from data that has an sir class (created with as.sir()) and counts the number of S's, I's and R's. It also supports grouped variables. The function sir\_df() works exactly like count\_df(), but adds the percentage of S, I and R.

#### Value

An integer

## **Interpretation of SIR**

In 2019, the European Committee on Antimicrobial Susceptibility Testing (EUCAST) has decided to change the definitions of susceptibility testing categories S, I, and R as shown below (https://www.eucast.org/newsiandr):

# • S - Susceptible, standard dosing regimen

A microorganism is categorised as "Susceptible, standard dosing regimen", when there is a high likelihood of therapeutic success using a standard dosing regimen of the agent.

#### • I - Susceptible, increased exposure

A microorganism is categorised as "Susceptible, Increased exposure" when there is a high likelihood of therapeutic success because exposure to the agent is increased by adjusting the dosing regimen or by its concentration at the site of infection.

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## • R = Resistant

A microorganism is categorised as "Resistant" when there is a high likelihood of therapeutic failure even when there is increased exposure.

Exposure is a function of how the mode of administration, dose, dosing interval, infusion time, as well as distribution and excretion of the antimicrobial agent will influence the infecting organism at the site of infection.

This AMR package honours this insight. Use susceptibility() (equal to proportion\_SI()) to determine antimicrobial susceptibility and count\_susceptible() (equal to count\_SI()) to count susceptible isolates.

# **Combination Therapy**

When using more than one variable for . . . (= combination therapy), use only\_all\_tested to only count isolates that are tested for all antibiotics/variables that you test them for. See this example for two antibiotics, Drug A and Drug B, about how susceptibility() works to calculate the %SI:

		only_all_tested = FALSE		only_all_tested = TRUE				
Drug A	Drug B		include as denominator	include as numerator	include as denominator			
S or I	S or I	X	Χ	Χ	X			
R	S or I	Χ	Χ	Χ	Χ			
<na></na>	S or I	Χ	Χ	-	-			
S or I	R	Χ	Χ	Χ	Χ			
R	R	-	Χ	-	Χ			
<na></na>	R	-	-	-	-			
S or I	<na></na>	Χ	Χ	-	-			
R	<na></na>	-	-	-	-			
<na></na>	<na></na>	-	-	-	-			

Please note that, in combination therapies, for only\_all\_tested = TRUE applies that:

```
count_S() + count_I() + count_R() = count_all()
proportion_S() + proportion_I() + proportion_R() = 1
```

and that, in combination therapies, for only\_all\_tested = FALSE applies that:

```
count_S() + count_I() + count_R() >= count_all()
proportion_S() + proportion_I() + proportion_R() >= 1
```

Using only\_all\_tested has no impact when only using one antibiotic as input.

## See Also

proportion\_\* to calculate microbial resistance and susceptibility.

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# **Examples**

```
# example_isolates is a data set available in the AMR package.
# run ?example_isolates for more info.
# base R -----
count_resistant(example_isolates$AMX) # counts "R"
count_susceptible(example_isolates$AMX) # counts "S" and "I"
count_all(example_isolates$AMX) # counts "S", "I" and "R"
# be more specific
count_S(example_isolates$AMX)
count_SI(example_isolates$AMX)
count_I(example_isolates$AMX)
count_IR(example_isolates$AMX)
count_R(example_isolates$AMX)
# Count all available isolates
count_all(example_isolates$AMX)
n_sir(example_isolates$AMX)
# n_sir() is an alias of count_all().
# Since it counts all available isolates, you can
# calculate back to count e.g. susceptible isolates.
# These results are the same:
count_susceptible(example_isolates$AMX)
susceptibility(example_isolates$AMX) * n_sir(example_isolates$AMX)
# dplyr -----
if (require("dplyr")) {
 example_isolates %>%
   group_by(ward) %>%
   summarise(
     R = count_R(CIP),
     I = count_I(CIP),
     S = count_S(CIP),
     n1 = count_all(CIP), # the actual total; sum of all three
     n2 = n_sir(CIP), # same - analogous to n_distinct
     total = n()
   ) # NOT the number of tested isolates!
 # Number of available isolates for a whole antibiotic class
 # (i.e., in this data set columns GEN, TOB, AMK, KAN)
 example_isolates %>%
   group_by(ward) %>%
   summarise(across(aminoglycosides(), n_sir))
 # Count co-resistance between amoxicillin/clav acid and gentamicin,
 # so we can see that combination therapy does a lot more than mono therapy.
 # Please mind that `susceptibility()` calculates percentages right away instead.
 example_isolates %>% count_susceptible(AMC) # 1433
 example_isolates %>% count_all(AMC) # 1879
```

```
example_isolates %>% count_susceptible(GEN) # 1399
example_isolates %>% count_all(GEN) # 1855

example_isolates %>% count_susceptible(AMC, GEN) # 1764
example_isolates %>% count_all(AMC, GEN) # 1936

# Get number of S+I vs. R immediately of selected columns
example_isolates %>%
    select(AMX, CIP) %>%
    count_df(translate = FALSE)

# It also supports grouping variables
example_isolates %>%
    select(ward, AMX, CIP) %>%
    group_by(ward) %>%
    count_df(translate = FALSE)
}
```

custom\_eucast\_rules

Define Custom EUCAST Rules

# **Description**

Define custom EUCAST rules for your organisation or specific analysis and use the output of this function in eucast\_rules().

# Usage

```
custom_eucast_rules(...)
```

# **Arguments**

... rules in formula notation, see *Examples* 

### **Details**

Some organisations have their own adoption of EUCAST rules. This function can be used to define custom EUCAST rules to be used in the eucast\_rules() function.

# Value

A list containing the custom rules

#### How it works

#### **Basics:**

If you are familiar with the case\_when() function of the dplyr package, you will recognise the input method to set your own rules. Rules must be set using what R considers to be the 'formula notation'. The rule itself is written *before* the tilde (~) and the consequence of the rule is written *after* the tilde:

These are two custom EUCAST rules: if TZP (piperacillin/tazobactam) is "S", all aminopenicillins (ampicillin and amoxicillin) must be made "S", and if TZP is "R", aminopenicillins must be made "R". These rules can also be printed to the console, so it is immediately clear how they work:

```
x
#> A set of custom EUCAST rules:
#>
#> 1. If TZP is "S" then set to S:
#> amoxicillin (AMX), ampicillin (AMP)
#>
#> 2. If TZP is "R" then set to R:
#> amoxicillin (AMX), ampicillin (AMP)
```

The rules (the part *before* the tilde, in above example TZP == "S" and TZP == "R") must be evaluable in your data set: it should be able to run as a filter in your data set without errors. This means for the above example that the column TZP must exist. We will create a sample data set and test the rules set:

```
df <- data.frame(mo = c("Escherichia coli", "Klebsiella pneumoniae"),</pre>
                  TZP = as.sir("R"),
                  ampi = as.sir("S"),
                  cipro = as.sir("S"))
df
#>
                         mo TZP ampi cipro
#> 1
          Escherichia coli
                                    S
                                          S
                                    S
                                          S
#> 2 Klebsiella pneumoniae
eucast_rules(df, rules = "custom", custom_rules = x, info = FALSE)
#>
                         mo TZP ampi cipro
#> 1
          Escherichia coli
                              R
                                    R
                                          S
#> 2 Klebsiella pneumoniae
                                    R
                                          S
```

# Using taxonomic properties in rules:

There is one exception in columns used for the rules: all column names of the microorganisms data set can also be used, but do not have to exist in the data set. These column names are: "mo", "fullname", "status", "kingdom", "phylum", "class", "order", "family", "genus", "species", "subspecies", "rank", "ref", "oxygen\_tolerance", "source", "lpsn", "lpsn\_parent", "lpsn\_renamed\_to", "gbif", "gbif\_parent", "gbif\_renamed\_to", "prevalence", and "snomed". Thus, this next example will work as well, despite the fact that the df data set does not contain a column genus:

#### Usage of antibiotic group names:

It is possible to define antibiotic groups instead of single antibiotics for the rule consequence, the part *after* the tilde. In above examples, the antibiotic group aminopenicillins is used to include ampicillin and amoxicillin. The following groups are allowed (case-insensitive). Within parentheses are the drugs that will be matched when running the rule.

- "aminoglycosides"

   (amikacin, amikacin/fosfomycin, amphotericin B-high, apramycin, arbekacin, astromicin, bekanamycin, dibekacin, framycetin, gentamicin, gentamicin-high, habekacin, hygromycin, isepamicin, kanamycin, kanamycin-high, kanamycin/cephalexin, micronomicin, neomycin, netilmicin, pentisomicin, plazomicin, propikacin, ribostamycin, sisomicin, streptoduocin, streptomycin, streptomycin-high, tobramycin, and tobramycin-high)
- "aminopenicillins" (amoxicillin and ampicillin)
- "antifungals"
   (amphotericin B, anidulafungin, butoconazole, caspofungin, ciclopirox, clotrimazole, econazole, fluconazole, flucytosine, fosfluconazole, griseofulvin, hachimycin, ibrexafungerp, isavuconazole, isoconazole, itraconazole, ketoconazole, manogepix, micafungin, miconazole, nystatin, oteseconazole, pimaricin, posaconazole, rezafungin, ribociclib, sulconazole, terbinafine, terconazole, and voriconazole)
- "antimycobacterials"
   (4-aminosalicylic acid, calcium aminosalicylate, capreomycin, clofazimine, delamanid, enviomycin, ethambutol, ethambutol/isoniazid, ethionamide, isoniazid, isoniazid/sulfamethoxazole/trimethoprim/pyridomorinamide, p-aminosalicylic acid, pretomanid, protionamide, pyrazinamide, rifabutin, rifampicin, rifampicin/ethambutol/isoniazid, rifampicin/isoniazid, rifampicin/pyrazinamide/ethambutol/isoniazid, rifampicin/pyrazinamide/isoniazid, rifampicin, rifapentine, simvastatin/fenofibrate, sodium aminosalicylate, streptomycin/isoniazid, terizidone, thioacetazone, thioacetazone/isoniazid, tiocarlide, and viomycin)
- "betalactams" (amoxicillin, amoxicillin/clavulanic acid, amoxicillin/sulbactam, ampicillin, ampicillin/sulbactam, apalcillin, aspoxicillin, avibactam, azidocillin, azlocillin, aztreonam, aztreonam/avibactam, aztreonam/nacubactam, bacampicillin, benzathine benzylpenicillin, benzathine phenoxymethylpenicillin, benzylpenicillin, biapenem, carbenicillin, carindacillin, cefacetrile, cefaclor, cefadroxil, cefalexin, cefaloridine, cefalotin, cefamandole, cefapirin, cefatrizine, cefazedone, cefazolin, cefcapene, cefcapene pivoxil, cefdinir, cefditoren, cefditoren pivoxil, cefepime, cefepime/clavulanic acid, cefepime/nacubactam, cefepime/tazobactam, cefetamet, cefetamet pivoxil, cefetecol, cefetrizole, cefixime, cefmenoxime, cefmetazole, cefodizime, cefonicid, cefoperazone, cefoperazone/sulbactam, ceforanide, cefoselis, cefotaxime, cefotaxime/clavulanic acid, cefotaxime/sulbactam, cefotetan, cefotiam, cefotiam hexetil, cefovecin, cefoxitin, cefoxitin screening, cefozopran, cefpimizole, cefpiramide, cefpirome, cefpodoxime, cefpodoxime proxetil,

cefpodoxime/clavulanic acid, cefprozil, cefquinome, cefroxadine, cefsulodin, cefsumide, ceftaroline, ceftaroline/avibactam, ceftazidime, ceftazidime/avibactam, ceftazidime/clavulanic acid, cefteram, cefteram pivoxil, ceftezole, ceftibuten, ceftiofur, ceftizoxime, ceftizoxime alapivoxil, ceftobiprole, ceftobiprole medocaril, ceftolozane/tazobactam, ceftriaxone, ceftriaxone/beta-lactamase inhibitor, cefuroxime, cefuroxime axetil, cephradine, ciclacillin, clometocillin, cloxacillin, dicloxacillin, doripenem, epicillin, ertapenem, flucloxacillin, hetacillin, imipenem, imipenem/EDTA, imipenem/relebactam, latamoxef, lenampicillin, loracarbef, mecillinam, meropenem, meropenem/nacubactam, meropenem/vaborbactam, metampicillin, meticillin, mezlocillin, sulbactam, nacubactam, nafcillin, oxacillin, panipenem, pename-cillin, penicillin/novobiocin, penicillin/sulbactam, pheneticillin, phenoxymethylpenicillin, piperacillin, piperacillin, piperacillin, pivencillin, pivencillin, propicillin, propicillin, razupenem, ritipenem, ritipenem acoxil, sarmoxicillin, sulbactam, sulbactam, sulbenicillin, talampicillin, tazobactam, tebipenem, temocillin, ticarcillin, and ticarcillin/clavulanic acid)

# · "carbapenems"

(biapenem, doripenem, ertapenem, imipenem, imipenem/EDTA, imipenem/relebactam, meropenem, meropenem/nacubactam, meropenem/vaborbactam, panipenem, razupenem, ritipenem, ritipenem acoxil, and tebipenem)

# · "cephalosporins"

lactamase inhibitor, and latamoxef)

(cefacetrile, cefaclor, cefadroxil, cefalexin, cefaloridine, cefalotin, cefamandole, cefapirin, cefatrizine, cefazedone, cefazolin, cefcapene, cefcapene pivoxil, cefdinir, cefditoren, cefditoren pivoxil, cefepime, cefepime/clavulanic acid, cefepime/tazobactam, cefetamet, cefetamet pivoxil, cefetecol, cefetrizole, cefixime, cefmenoxime, cefmetazole, cefodizime, cefonicid, cefoperazone, cefoperazone/sulbactam, ceforanide, cefoselis, cefotaxime, cefotaxime/clavulanic acid, cefotaxime/sulbactam, cefotetan, cefotiam, cefotiam hexetil, cefovecin, cefoxitin, cefoxitin screening, cefozopran, cefpimizole, cefpiramide, cefpirome, cefpodoxime, cefpodoxime proxetil, cefpodoxime/clavulanic acid, cefprozil, cefquinome, cefroxadine, cefsulodin, cefsumide, ceftaroline, ceftaroline/avibactam, ceftazidime, ceftazidime/avibactam, ceftazidime/clavulanic acid, cefteram pivoxil, ceftezole, ceftibuten, ceftiofur, ceftizoxime, ceftizoxime alapivoxil, ceftobiprole, ceftobiprole medocaril, ceftolozane/tazobactam, ceftriaxone, ceftriaxone/beta-lactamase inhibitor, cefuroxime, cefuroxime axetil, cephradine, latamoxef, and loracarbef)

- "cephalosporins\_1st"
   (cefacetrile, cefadroxil, cefalexin, cefaloridine, cefalotin, cefapirin, cefatrizine, cefazedone, cefazolin, cefroxadine, ceftezole, and cephradine)
- "cephalosporins\_2nd" (cefaclor, cefamandole, cefmetazole, cefonicid, ceforanide, cefotetan, cefotiam, cefoxitin, cefoxitin screening, cefprozil, cefuroxime, cefuroxime axetil, and loracarbef)
- "cephalosporins\_3rd"
   (cefcapene, cefcapene pivoxil, cefdinir, cefditoren, cefditoren pivoxil, cefetamet, cefetamet pivoxil, cefixime, cefmenoxime, cefodizime, cefoperazone, cefoperazone/sulbactam, cefotaxime, cefotaxime/clavulanic acid, cefotaxime/sulbactam, cefotiam hexetil, cefovecin, cefpimizole, cefpiramide, cefpodoxime, cefpodoxime proxetil, cefpodoxime/clavulanic acid, cefsulodin, ceftazidime, ceftazidime/avibactam, ceftazidime/clavulanic acid, cefteram pivoxil, ceftibuten, ceftiofur, ceftizoxime, ceftizoxime alapivoxil, ceftriaxone, ceftriaxone/beta-
- "cephalosporins\_4th" (cefepime, cefepime/clavulanic acid, cefepime/tazobactam, cefetecol, cefoselis, cefozopran,

cefpirome, and cefquinome)

• "cephalosporins\_5th" (ceftaroline, ceftaroline/avibactam, ceftobiprole, ceftobiprole medocaril, and ceftolozane/tazobactam)

"cephalosporins\_except\_caz"
 (cefacetrile, cefaclor, cefadroxil, cefalexin, cefaloridine, cefalotin, cefamandole, cefapirin, cefatrizine, cefazedone, cefazolin, cefcapene, cefcapene pivoxil, cefdinir, cefditoren, cefditoren pivoxil, cefepime, cefepime/clavulanic acid, cefepime/tazobactam, cefetamet, cefetamet pivoxil, cefetecol, cefetrizole, cefixime, cefmenoxime, cefmetazole, cefodizime, cefonicid, cefoperazone, cefoperazone/sulbactam, ceforanide, cefoselis, cefotaxime, cefotaxime/clavulanic acid, cefotaxime/sulbactam, cefotetan, cefotiam hexetil, cefovecin, cefoxitin, cefoxitin screening, cefozopran, cefpimizole, cefpiramide, cefpirome, cefpodoxime, cefpodoxime proxetil, cefpodoxime/clavulanic acid, cefprozil, cefquinome, cefroxadine, cefsulodin, cefsumide, ceftaroline, ceftaroline/avibactam, ceftazidime/avibactam, ceftazidime/clavulanic

alapiyoxil, ceftobiprole, ceftobiprole medocaril, ceftolozane/tazobactam, ceftriaxone, ceftriaxone/beta-

• "fluoroquinolones"

(besifloxacin, ciprofloxacin, clinafloxacin, danofloxacin, delafloxacin, difloxacin, enoxacin, enrofloxacin, finafloxacin, fleroxacin, garenoxacin, gatifloxacin, gemifloxacin, grepafloxacin, lascufloxacin, levofloxacin, levonadifloxacin, lomefloxacin, marbofloxacin, metioxate, miloxacin, moxifloxacin, nadifloxacin, nifuroquine, norfloxacin, ofloxacin, orbifloxacin, pazufloxacin, pefloxacin, pradofloxacin, premafloxacin, prulifloxacin, rufloxacin, sarafloxacin, sitafloxacin, sparfloxacin, temafloxacin, tilbroquinol, tioxacin, tosufloxacin, and trovafloxacin)

acid, cefteram, cefteram pivoxil, ceftezole, ceftibuten, ceftiofur, ceftizoxime, ceftizoxime

lactamase inhibitor, cefuroxime, cefuroxime axetil, cephradine, latamoxef, and loracarbef)

- "glycopeptides" (avoparcin, dalbavancin, norvancomycin, oritavancin, ramoplanin, teicoplanin, teicoplanin-macromethod, telavancin, vancomycin, and vancomycin-macromethod)
- "glycopeptides\_except\_lipo"

   (avoparcin, norvancomycin, ramoplanin, teicoplanin, teicoplanin-macromethod, vancomycin, and vancomycin-macromethod)
- "lincosamides" (acetylmidecamycin, acetylspiramycin, clindamycin, gamithromycin, kitasamycin, lincomycin, meleumycin, nafithromycin, pirlimycin, primycin, solithromycin, tildipirosin, tilmicosin, tulathromycin, tylosin, and tylvalosin)
- "lipoglycopeptides" (dalbavancin, oritavancin, and telavancin)
- · "macrolides"

(acetylmidecamycin, acetylspiramycin, azithromycin, clarithromycin, dirithromycin, erythromycin, flurithromycin, gamithromycin, josamycin, kitasamycin, meleumycin, midecamycin, miocamycin, nafithromycin, oleandomycin, pirlimycin, primycin, rokitamycin, roxithromycin, solithromycin, spiramycin, telithromycin, tildipirosin, tilmicosin, troleandomycin, tulathromycin, tylosin, and tylvalosin)

- "oxazolidinones" (cadazolid, cycloserine, linezolid, tedizolid, and thiacetazone)
- "penicillins"

   (amoxicillin, amoxicillin/clavulanic acid, amoxicillin/sulbactam, ampicillin, ampicillin, sulbactam, apalcillin, aspoxicillin, avibactam, azidocillin, azlocillin, aztreonam, aztreonam/avibactam, aztreonam/nacubactam, bacampicillin, benzathine benzylpenicillin, benzathine phenoxymethylpenicillin,

cillin, benzylpenicillin, carbenicillin, carindacillin, cefepime/nacubactam, ciclacillin, clometocillin, cloxacillin, dicloxacillin, epicillin, flucloxacillin, hetacillin, lenampicillin, mecillinam, metampicillin, meticillin, mezlocillin, mezlocillin/sulbactam, nacubactam, nafcillin, oxacillin, penamecillin, penicillin/novobiocin, penicillin/sulbactam, pheneticillin, phenoxymethylpenicillin, piperacillin, piperacillin/sulbactam, piperacillin/tazobactam, piridicillin, pivampicillin, pivmecillinam, procaine benzylpenicillin, propicillin, sarmoxicillin, sulbactam, sulbenicillin, sultamicillin, talampicillin, tazobactam, temocillin, ticarcillin, and ticarcillin/clavulanic acid)

- "polymyxins" (colistin, polymyxin B, and polymyxin B/polysorbate 80)
- "quinolones"

(besifloxacin, cinoxacin, ciprofloxacin, clinafloxacin, danofloxacin, delafloxacin, difloxacin, enoxacin, enrofloxacin, finafloxacin, fleroxacin, flumequine, garenoxacin, gatifloxacin, gemifloxacin, grepafloxacin, lascufloxacin, levofloxacin, levonadifloxacin, lomefloxacin, marbofloxacin, metioxate, miloxacin, moxifloxacin, nadifloxacin, nalidixic acid, nemonoxacin, nifuroquine, nitroxoline, norfloxacin, ofloxacin, orbifloxacin, oxolinic acid, pazufloxacin, pefloxacin, pipemidic acid, piromidic acid, pradofloxacin, premafloxacin, prulifloxacin, rosoxacin, rufloxacin, sarafloxacin, sitafloxacin, sparfloxacin, temafloxacin, tilbroquinol, tioxacin, tosufloxacin, and trovafloxacin)

- "streptogramins" (pristinamycin and quinupristin/dalfopristin)
- "tetracyclines"
   (cetocycline, chlortetracycline, clomocycline, demeclocycline, doxycycline, eravacycline, lymecycline, metacycline, minocycline, omadacycline, oxytetracycline, penimepicycline, rolitetracycline, sarecycline, tetracycline, and tigecycline)
- "tetracyclines\_except\_tgc" (cetocycline, chlortetracycline, clomocycline, demeclocycline, doxycycline, eravacycline, lymecycline, metacycline, minocycline, omadacycline, oxytetracycline, penimepicycline, rolitetracycline, sarecycline, and tetracycline)
- "trimethoprims"

(brodimoprim, sulfadiazine, sulfadiazine/tetroxoprim, sulfadiazine/trimethoprim, sulfadimethoxine, sulfadimidine, sulfadimidine/trimethoprim, sulfafurazole, sulfaisodimidine, sulfalene, sulfamazone, sulfamerazine, sulfamerazine/trimethoprim, sulfamethizole, sulfamethoxazole, sulfamethoxypyridazine, sulfametomidine, sulfametoxydiazine, sulfametrole/trimethoprim, sulfamoxole, sulfamoxole/trimethoprim, sulfanilamide, sulfaperin, sulfaphenazole, sulfapyridine, sulfathiourea, trimethoprim, and trimethoprim/sulfamethoxazole)

• "ureidopenicillins" (azlocillin, mezlocillin, piperacillin, and piperacillin/tazobactam)

### **Examples**

```
x <- custom_eucast_rules(
  AMC == "R" & genus == "Klebsiella" ~ aminopenicillins == "R",
  AMC == "I" & genus == "Klebsiella" ~ aminopenicillins == "I"
)
x
# run the custom rule set (verbose = TRUE will return a logbook instead of the data set):
eucast_rules(example_isolates,</pre>
```

80 dosage

```
rules = "custom",
  custom_rules = x,
  info = FALSE,
  verbose = TRUE
)

# combine rule sets
x2 <- c(
    x,
    custom_eucast_rules(TZP == "R" ~ carbapenems == "R")
)
x2</pre>
```

dosage

Data Set with Treatment Dosages as Defined by EUCAST

# Description

EUCAST breakpoints used in this package are based on the dosages in this data set. They can be retrieved with eucast\_dosage().

# Usage

dosage

#### **Format**

A tibble with 503 observations and 9 variables:

• ab

Antibiotic ID as used in this package (such as AMC), using the official EARS-Net (European Antimicrobial Resistance Surveillance Network) codes where available

• name

Official name of the antimicrobial drug as used by WHONET/EARS-Net or the WHO

type

Type of the dosage, either "high\_dosage", "standard\_dosage", or "uncomplicated\_uti"

dose

Dose, such as "2 g" or "25 mg/kg"

• dose\_times

Number of times a dose must be administered

• administration

Route of administration, either "im", "iv", or "oral"

notes

Additional dosage notes

• original\_txt

Original text in the PDF file of EUCAST

• eucast\_version

Version number of the EUCAST Clinical Breakpoints guideline to which these dosages apply, either 13, 12, or 11

# **Details**

Like all data sets in this package, this data set is publicly available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

# **Examples**

dosage

eucast\_rules

Apply EUCAST Rules

# **Description**

Apply rules for clinical breakpoints and intrinsic resistance as defined by the European Committee on Antimicrobial Susceptibility Testing (EUCAST, https://www.eucast.org), see *Source*. Use eucast\_dosage() to get a data.frame with advised dosages of a certain bug-drug combination, which is based on the dosage data set.

To improve the interpretation of the antibiogram before EUCAST rules are applied, some non-EUCAST rules can applied at default, see *Details*.

# Usage

```
eucast_rules(
    x,
    col_mo = NULL,
    info = interactive(),
    rules = getOption("AMR_eucastrules", default = c("breakpoints", "expert")),
    verbose = FALSE,
    version_breakpoints = 12,
    version_expertrules = 3.3,
    ampc_cephalosporin_resistance = NA,
    only_sir_columns = FALSE,
    custom_rules = NULL,
    ...
)

eucast_dosage(ab, administration = "iv", version_breakpoints = 12)
```

# **Arguments**

X	a data set with antibiotic columns, such as amox, AMX and AMC
col_mo	column name of the names or codes of the microorganisms (see as.mo()) - the default is the first column of class mo. Values will be coerced using as.mo().
info	a logical to indicate whether progress should be printed to the console - the default is only print while in interactive sessions

rules

a character vector that specifies which rules should be applied. Must be one or more of "breakpoints", "expert", "other", "custom", "all", and defaults to c("breakpoints", "expert"). The default value can be set to another value using the package option AMR\_eucastrules: options(AMR\_eucastrules = "all"). If using "custom", be sure to fill in argument custom\_rules too. Custom rules can be created with custom\_eucast\_rules().

verbose

a logical to turn Verbose mode on and off (default is off). In Verbose mode, the function does not apply rules to the data, but instead returns a data set in logbook form with extensive info about which rows and columns would be effected and in which way. Using Verbose mode takes a lot more time.

version\_breakpoints

the version number to use for the EUCAST Clinical Breakpoints guideline. Can be "12.0", "11.0", or "10.0".

version\_expertrules

the version number to use for the EUCAST Expert Rules and Intrinsic Resistance guideline. Can be "3.3", "3.2", or "3.1".

ampc\_cephalosporin\_resistance

a character value that should be applied to cefotaxime, ceftriaxone and ceftazidime for AmpC de-repressed cephalosporin-resistant mutants - the default is NA. Currently only works when version\_expertrules is 3.2 and higher; these version of 'EUCAST Expert Rules on Enterobacterales' state that results of cefotaxime, ceftriaxone and ceftazidime should be reported with a note, or results should be suppressed (emptied) for these three drugs. A value of NA (the default) for this argument will remove results for these three drugs, while e.g. a value of "R" will make the results for these drugs resistant. Use NULL or FALSE to not alter results for these three drugs of AmpC de-repressed cephalosporin-resistant mutants. Using TRUE is equal to using "R".

For EUCAST Expert Rules v3.2, this rule applies to: Citrobacter braakii, Citrobacter freundii, Citrobacter gillenii, Citrobacter murliniae, Citrobacter rodenticum, Citrobacter sedlakii, Citrobacter werkmanii, Citrobacter youngae, Enterobacter, Hafnia alvei, Klebsiella aerogenes, Morganella morganii, Providencia, and Serratia.

only\_sir\_columns

a logical to indicate whether only antibiotic columns must be detected that were transformed to class sir (see as.sir()) on beforehand (default is FALSE)

custom\_rules custom rules to apply, created with custom\_eucast\_rules()
... column name of an antibiotic, see section *Antibiotics* below

ab any (vector of) text that can be coerced to a valid antibiotic drug code with as.ab()

administration route of administration, either "im", "iv", or "oral"

### **Details**

**Note:** This function does not translate MIC values to SIR values. Use as .sir() for that.

Note: When ampicillin (AMP, J01CA01) is not available but amoxicillin (AMX, J01CA04) is, the

latter will be used for all rules where there is a dependency on ampicillin. These drugs are interchangeable when it comes to expression of antimicrobial resistance.

The file containing all EUCAST rules is located here: https://github.com/msberends/AMR/blob/main/data-raw/eucast\_rules.tsv. Note: Old taxonomic names are replaced with the current taxonomy where applicable. For example, *Ochrobactrum anthropi* was renamed to *Brucella anthropi* in 2020; the original EUCAST rules v3.1 and v3.2 did not yet contain this new taxonomic name. The AMR package contains the full microbial taxonomy updated until December 11th, 2022, see microorganisms.

### **Custom Rules:**

Custom rules can be created using custom\_eucast\_rules(), e.g.:

#### 'Other' Rules:

Before further processing, two non-EUCAST rules about drug combinations can be applied to improve the efficacy of the EUCAST rules, and the reliability of your data (analysis). These rules are:

- 1. A drug with enzyme inhibitor will be set to S if the same drug without enzyme inhibitor is S
- 2. A drug **without** enzyme inhibitor will be set to R if the same drug **with** enzyme inhibitor is R

Important examples include amoxicillin and amoxicillin/clavulanic acid, and trimethoprim and trimethoprim/sulfamethoxazole. Needless to say, for these rules to work, both drugs must be available in the data set.

Since these rules are not officially approved by EUCAST, they are not applied at default. To use these rules, include "other" to the rules argument, or use eucast\_rules(..., rules = "all"). You can also set the package option AMR\_eucastrules, i.e. run options(AMR\_eucastrules = "all").

# Value

The input of x, possibly with edited values of antibiotics. Or, if verbose = TRUE, a data.frame with all original and new values of the affected bug-drug combinations.

#### **Antibiotics**

To define antibiotics column names, leave as it is to determine it automatically with guess\_ab\_col() or input a text (case-insensitive), or use NULL to skip a column (e.g. TIC = NULL to skip ticarcillin). Manually defined but non-existing columns will be skipped with a warning.

The following antibiotics are eligible for the functions eucast\_rules() and mdro(). These are shown below in the format 'name (antimicrobial ID, ATC code)', sorted alphabetically:

Amikacin (AMK, J01GB06), amoxicillin (AMX, J01CA04), amoxicillin/clavulanic acid (AMC, J01CR02), ampicillin (AMP, J01CA01), ampicillin/sulbactam (SAM, J01CR01), arbekacin (ARB, J01GB12), aspoxicillin (APX, J01CA19), azidocillin (AZD, J01CE04), azithromycin (AZM, J01FA10), azlocillin

(AZL, J01CA09), aztreonam (ATM, J01DF01), bacampicillin (BAM, J01CA06), bekanamycin (BEK, J01GB13), benzathine benzylpenicillin (BNB, J01CE08), benzathine phenoxymethylpenicillin (BNP, J01CE10), benzylpenicillin (PEN, J01CE01), besifloxacin (BES, S01AE08), biapenem (BIA, J01DH05), carbenicillin (CRB, J01CA03), carindacillin (CRN, J01CA05), cefacetrile (CAC, J01DB10), cefaclor (CEC, J01DC04), cefadroxil (CFR, J01DB05), cefalexin (LEX, J01DB01), cefaloridine (RID, J01DB02), cefalotin (CEP, J01DB03), cefamandole (MAN, J01DC03), cefapirin (HAP, J01DB08), cefatrizine (CTZ, J01DB07), cefazedone (CZD, J01DB06), cefazolin (CZO, J01DB04), cefcapene (CCP, J01DD17), cefdinir (CDR, J01DD15), cefditoren (DIT, J01DD16), cefepime (FEP, J01DE01), cefetamet (CAT, J01DD10), cefixime (CFM, J01DD08), cefmenoxime (CMX, J01DD05), cefmetazole (CMZ, J01DC09), cefodizime (DIZ, J01DD09), cefonicid (CID, J01DC06), cefoperazone (CFP, J01DD12), cefoperazone/sulbactam (CSL, J01DD62), ceforanide (CND, J01DC11), cefotaxime (CTX, J01DD01), cefotaxime/clavulanic acid (CTC, J01DD51), cefotetan (CTT, J01DC05), cefotiam (CTF, J01DC07), cefoxitin (F0X, J01DC01), cefozopran (Z0P, J01DE03), cefpiramide (CPM, J01DD11), cefpirome (CPO, J01DE02), cefpodoxime (CPD, J01DD13), cefprozil (CPR, J01DC10), cefroxadine (CRD, J01DB11), cefsulodin (CFS, J01DD03), ceftaroline (CPT, J01DI02), ceftazidime (CAZ, J01DD02), ceftazidime/clavulanic acid (CCV, J01DD52), cefteram (CEM, J01DD18), ceftezole (CTL, J01DB12), ceftibuten (CTB, J01DD14), ceftizoxime (CZX, J01DD07), ceftobiprole medocaril (CFM1, J01DI01), ceftolozane/tazobactam (CZT, J01DI54), ceftriaxone (CRO, J01DD04), ceftriaxone/betalactamase inhibitor (CEB, J01DD63), cefuroxime (CXM, J01DC02), cephradine (CED, J01DB09), chloramphenicol (CHL, J01BA01), ciprofloxacin (CIP, J01MA02), clarithromycin (CLR, J01FA09), clindamycin (CLI, J01FF01), clometocillin (CLM, J01CE07), cloxacillin (CLO, J01CF02), colistin (COL, J01XB01), cycloserine (CYC, J04AB01), dalbavancin (DAL, J01XA04), daptomycin (DAP, J01XX09), delafloxacin (DFX, J01MA23), dibekacin (DKB, J01GB09), dicloxacillin (DIC, J01CF01), dirithromycin (DIR, J01FA13), doripenem (DOR, J01DH04), doxycycline (DOX, J01AA02), enoxacin (ENX, J01MA04), epicillin (EPC, J01CA07), ertapenem (ETP, J01DH03), erythromycin (ERY, J01FA01), fleroxacin (FLE, J01MA08), flucloxacillin (FLC, J01CF05), flurithromycin (FLR1, J01FA14), fosfomycin (FOS, J01XX01), framycetin (FRM, D09AA01), fusidic acid (FUS, J01XC01), garenoxacin (GRN, J01MA19), gatifloxacin (GAT, J01MA16), gemifloxacin (GEM, J01MA15), gentamicin (GEN, J01GB03), grepafloxacin (GRX, J01MA11), hetacillin (HET, J01CA18), imipenem (IPM, J01DH51), imipenem/relebactam (IMR, J01DH56), isepamicin (ISE, J01GB11), josamycin (JOS, J01FA07), kanamycin (KAN, J01GB04), lascufloxacin (LSC, J01MA25), latamoxef (LTM, J01DD06), levofloxacin (LVX, J01MA12), levonadifloxacin (LND, J01MA24), lincomycin (LIN, J01FF02), linezolid (LNZ, J01XX08), lomefloxacin (LOM, J01MA07), loracarbef (LOR, J01DC08), mecillinam (MEC, J01CA11), meropenem (MEM, J01DH02), meropenem/vaborbactam (MEV, J01DH52), metampicillin (MTM, J01CA14), meticillin (MET, J01CF03), mezlocillin (MEZ, J01CA10), micronomicin (MCR, S01AA22), midecamycin (MID, J01FA03), minocycline (MNO, J01AA08), miocamycin (MCM, J01FA11), moxifloxacin (MFX, J01MA14), nadifloxacin (NAD, D10AF05), nafcillin (NAF, J01CF06), nalidixic acid (NAL, J01MB02), neomycin (NEO, J01GB05), netilmicin (NET, J01GB07), nitrofurantoin (NIT, J01XE01), norfloxacin (NOR, J01MA06), ofloxacin (OFX, J01MA01), oleandomycin (OLE, J01FA05), oritavancin (ORI, J01XA05), oxacillin (OXA, J01CF04), panipenem (PAN, J01DH55), pazufloxacin (PAZ, J01MA18), pefloxacin (PEF, J01MA03), penamecillin (PNM, J01CE06), pheneticillin (PHE, J01CE05), phenoxymethylpenicillin (PHN, J01CE02), piperacillin (PIP, J01CA12), piperacillin/tazobactam (TZP, J01CR05), pivampicillin (PVM, J01CA02), pivmecillinam (PME, J01CA08), plazomicin (PLZ, J01GB14), polymyxin B (PLB, J01XB02), pristinamycin (PRI, J01FG01), procaine benzylpenicillin (PRB, J01CE09), propicillin (PRP, J01CE03), prulifloxacin (PRU, J01MA17), quinupristin/dalfopristin (QDA, J01FG02), ribostamycin (RST, J01GB10), rifampicin (RIF, J04AB02), rokitamycin (R0K, J01FA12), roxithromycin (RXT, J01FA06), rufloxacin (RFL, J01MA10), sisomicin (SIS, J01GB08), sitafloxacin (SIT, J01MA21), solithromycin (SOL, J01FA16), sparfloxacin (SPX, J01MA09), spiramycin (SPI, J01FA02), strepto-

duocin (STR, J01GA02), streptomycin (STR1, J01GA01), sulbactam (SUL, J01CG01), sulbenicillin (SBC, J01CA16), sulfadiazine (SDI, J01EC02), sulfadiazine/trimethoprim (SLT1, J01EE02), sulfadimethoxine (SUD, J01ED01), sulfadimidine (SDM, J01EB03), sulfadimidine/trimethoprim (SLT2, J01EE05), sulfafurazole (SLF, J01EB05), sulfaisodimidine (SLF1, J01EB01), sulfalene (SLF2, J01ED02), sulfamazone (SZO, J01ED09), sulfamerazine (SLF3, J01ED07), sulfamerazine/trimethoprim (SLT3, J01EE07), sulfamethizole (SLF4, J01EB02), sulfamethoxazole (SMX, J01EC01), sulfamethoxypyridazine (SLF5, J01ED05), sulfametomidine (SLF6, J01ED03), sulfametoxydiazine (SLF7, J01ED04), sulfametrole/trimethoprim (SLT4, J01EE03), sulfamoxole (SLF8, J01EC03), sulfamoxole/trimethoprim (SLT5, J01EE04), sulfanilamide (SLF9, J01EB06), sulfaperin (SLF10, J01ED06), sulfaphenazole (SLF11, J01ED08), sulfapyridine (SLF12, J01EB04), sulfathiazole (SUT, J01EB07), sulfathiourea (SLF13, J01EB08), sultamicillin (SLT6, J01CR04), talampicillin (TAL, J01CA15), tazobactam (TAZ, J01CG02), tebipenem (TBP, J01DH06), tedizolid (TZD, J01XX11), teicoplanin (TEC, J01XA02), telavancin (TLV, J01XA03), telithromycin (TLT, J01FA15), temafloxacin (TMX, J01MA05), temocillin (TEM, J01CA17), tetracycline (TCY, J01AA07), ticarcillin (TIC, J01CA13), ticarcillin/clavulanic acid (TCC, J01CR03), tigecycline (TGC, J01AA12), tilbroquinol (TBQ, P01AA05), tobramycin (T0B, J01GB01), tosufloxacin (TFX, J01MA22), trimethoprim (TMP, J01EA01), trimethoprim/sulfamethoxazole (SXT, J01EE01), troleandomycin (TRL, J01FA08), trovafloxacin (TVA, J01MA13), vancomycin (VAN, J01XA01)

#### Reference Data Publicly Available

All data sets in this AMR package (about microorganisms, antibiotics, SIR interpretation, EUCAST rules, etc.) are publicly and freely available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. We also provide tab-separated plain text files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

# Source

- EUCAST Expert Rules. Version 2.0, 2012.
   Leclercq et al. EUCAST expert rules in antimicrobial susceptibility testing. Clin Microbiol Infect. 2013;19(2):141-60; doi:10.1111/j.14690691.2011.03703.x
- EUCAST Expert Rules, Intrinsic Resistance and Exceptional Phenotypes Tables. Version 3.1, 2016. (link)
- EUCAST Intrinsic Resistance and Unusual Phenotypes. Version 3.2, 2020. (link)
- EUCAST Intrinsic Resistance and Unusual Phenotypes. Version 3.3, 2021. (link)
- EUCAST Breakpoint tables for interpretation of MICs and zone diameters. Version 9.0, 2019. (link)
- EUCAST Breakpoint tables for interpretation of MICs and zone diameters. Version 10.0, 2020. (link)
- EUCAST Breakpoint tables for interpretation of MICs and zone diameters. Version 11.0, 2021. (link)
- EUCAST Breakpoint tables for interpretation of MICs and zone diameters. Version 12.0, 2022. (link)

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# **Examples**

```
a <- data.frame(
  mo = c(
    "Staphylococcus aureus",
    "Enterococcus faecalis",
    "Escherichia coli",
    "Klebsiella pneumoniae",
    "Pseudomonas aeruginosa"
 ),
  VAN = "-", # Vancomycin
  AMX = "-", # Amoxicillin
  COL = "-", # Colistin
  CAZ = "-", # Ceftazidime
  CXM = "-", # Cefuroxime
  PEN = "S", # Benzylpenicillin
  FOX = "S", # Cefoxitin
  stringsAsFactors = FALSE
)
head(a)
# apply EUCAST rules: some results wil be changed
b <- eucast_rules(a)</pre>
head(b)
# do not apply EUCAST rules, but rather get a data.frame
# containing all details about the transformations:
c <- eucast_rules(a, verbose = TRUE)</pre>
head(c)
# Dosage guidelines:
eucast_dosage(c("tobra", "genta", "cipro"), "iv")
eucast_dosage(c("tobra", "genta", "cipro"), "iv", version_breakpoints = 10)
```

example\_isolates

Data Set with 2 000 Example Isolates

# **Description**

A data set containing 2 000 microbial isolates with their full antibiograms. This data set contains randomised fictitious data, but reflects reality and can be used to practise AMR data analysis. For examples, please read the tutorial on our website.

# Usage

```
example_isolates
```

#### **Format**

A tibble with 2 000 observations and 46 variables:

- date
   Date of receipt at the laboratory
- patient ID of the patient
- age
   Age of the patient
- gender
  Gender of the patient, either "F" or "M"
- ward
   Ward type where the patient was admitted, either "Clinical", "ICU", or "Outpatient"
- mo
  ID of microorganism created with as . mo(), see also the microorganisms data set
- PEN:RIF
   40 different antibiotics with class sir (see as.sir()); these column names occur in the antibiotics data set and can be translated with set\_ab\_names() or ab\_name()

### **Details**

Like all data sets in this package, this data set is publicly available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

#### **Examples**

```
example_isolates
```

example\_isolates\_unclean

Data Set with Unclean Data

# Description

A data set containing 3 000 microbial isolates that are not cleaned up and consequently not ready for AMR data analysis. This data set can be used for practice.

# Usage

```
example_isolates_unclean
```

#### **Format**

A tibble with 3 000 observations and 8 variables:

- patient\_idID of the patient
- date date of receipt at the laboratory
- hospital ID of the hospital, from A to C
- bacteria info about microorganism that can be transformed with as.mo(), see also microorganisms
- AMX:GEN
   4 different antibiotics that have to be transformed with as.sir()

### **Details**

Like all data sets in this package, this data set is publicly available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

# **Examples**

```
example_isolates_unclean
```

first\_isolate

Determine First Isolates

# **Description**

Determine first isolates of all microorganisms of every patient per episode and (if needed) per specimen type. These functions support all four methods as summarised by Hindler *et al.* in 2007 (doi:10.1086/511864). To determine patient episodes not necessarily based on microorganisms, use is\_new\_episode() that also supports grouping with the dplyr package.

# Usage

```
first_isolate(
  x = NULL,
  col_date = NULL,
  col_patient_id = NULL,
  col_mo = NULL,
  col_testcode = NULL,
  col_specimen = NULL,
  col_icu = NULL,
  col_keyantimicrobials = NULL,
  episode_days = 365,
```

```
testcodes_exclude = NULL,
  icu_exclude = FALSE,
  specimen_group = NULL,
  type = "points",
 method = c("phenotype-based", "episode-based", "patient-based", "isolate-based"),
  ignore_I = TRUE,
 points_threshold = 2,
  info = interactive(),
  include_unknown = FALSE,
  include_untested_sir = TRUE,
)
filter_first_isolate(
  x = NULL,
  col_date = NULL,
  col_patient_id = NULL,
  col_mo = NULL,
  episode_days = 365,
 method = c("phenotype-based", "episode-based", "patient-based", "isolate-based"),
)
```

### **Arguments**

x a data.frame containing isolates. Can be left blank for automatic determination,

see Examples.

col\_date column name of the result date (or date that is was received on the lab) - the

default is the first column with a date class

col\_patient\_id column name of the unique IDs of the patients - the default is the first column

that starts with 'patient' or 'patid' (case insensitive)

col\_mo column name of the names or codes of the microorganisms (see as.mo()) - the

default is the first column of class mo. Values will be coerced using as.mo().

col\_testcode column name of the test codes. Use col\_testcode = NULL to **not** exclude cer-

tain test codes (such as test codes for screening). In that case testcodes\_exclude

will be ignored.

col\_specimen column name of the specimen type or group

col\_icu column name of the logicals (TRUE/FALSE) whether a ward or department is an

Intensive Care Unit (ICU). This can also be a logical vector with the same length

as rows in x.

col\_keyantimicrobials

(only useful when method = "phenotype-based") column name of the key antimicrobials to determine first isolates, see key\_antimicrobials(). The default is the first column that starts with 'key' followed by 'ab' or 'antibiotics' or 'antimicrobials' (case insensitive). Use col\_keyantimicrobials = FALSE to prevent this. Can also be the output of key\_antimicrobials().

episode\_days episode in days after which a genus/species combination will be determined as

'first isolate' again. The default of 365 days is based on the guideline by CLSI,

see Source.

testcodes\_exclude

a character vector with test codes that should be excluded (case-insensitive)

icu\_exclude a logical to indicate whether ICU isolates should be excluded (rows with value

TRUE in the column set with col\_icu)

specimen\_group value in the column set with col\_specimen to filter on

type type to determine weighed isolates; can be "keyantimicrobials" or "points",

see Details

method the method to apply, either "phenotype-based", "episode-based", "patient-based"

or "isolate-based" (can be abbreviated), see Details. The default is "phenotype-based"

if antimicrobial test results are present in the data, and "episode-based" oth-

erwise.

ignore\_I logical to indicate whether antibiotic interpretations with "I" will be ignored

when type = "keyantimicrobials", see *Details* 

points\_threshold

minimum number of points to require before differences in the antibiogram will

lead to inclusion of an isolate when type = "points", see *Details* 

info a logical to indicate info should be printed - the default is TRUE only in interactive

mode

include\_unknown

a logical to indicate whether 'unknown' microorganisms should be included too, i.e. microbial code "UNKNOWN", which defaults to FALSE. For WHONET users, this means that all records with organism code "con" (contamination) will be excluded at default. Isolates with a microbial ID of NA will always be excluded

as first isolate.

include\_untested\_sir

a logical to indicate whether also rows without antibiotic results are still eligible for becoming a first isolate. Use include\_untested\_sir = FALSE to always return FALSE for such rows. This checks the data set for columns of class sir and consequently requires transforming columns with antibiotic results using

as.sir() first.

arguments passed on to first\_isolate() when using filter\_first\_isolate(), otherwise arguments passed on to key\_antimicrobials() (such as universal,

gram\_negative, gram\_positive)

# **Details**

To conduct epidemiological analyses on antimicrobial resistance data, only so-called first isolates should be included to prevent overestimation and underestimation of antimicrobial resistance. Different methods can be used to do so, see below.

These functions are context-aware. This means that the x argument can be left blank if used inside a data.frame call, see *Examples*.

The first\_isolate() function is a wrapper around the is\_new\_episode() function, but more efficient for data sets containing microorganism codes or names.

All isolates with a microbial ID of NA will be excluded as first isolate.

#### **Different methods:**

According to Hindler *et al.* (2007, doi:10.1086/511864), there are different methods (algorithms) to select first isolates with increasing reliability: isolate-based, patient-based, episode-based and phenotype-based. All methods select on a combination of the taxonomic genus and species (not subspecies).

All mentioned methods are covered in the first\_isolate() function:

```
Method
                                            Function to apply
Isolate-based
                                            first_isolate(x, method = "isolate-based")
(= all isolates)
Patient-based
                                            first_isolate(x, method = "patient-based")
(= first isolate per patient)
Episode-based
                                            first_isolate(x, method = "episode-based"), or:
(= first isolate per episode)
- 7-Day interval from initial isolate
                                            - first_isolate(x, method = "e", episode_days = 7)
- 30-Day interval from initial isolate
                                            - first_isolate(x, method = "e", episode_days = 30)
Phenotype-based
                                            first_isolate(x, method = "phenotype-based"), or:
(= first isolate per phenotype)
- Major difference in any antimicrobial result - first_isolate(x, type = "points")
- Any difference in key antimicrobial results
                                           - first_isolate(x, type = "keyantimicrobials")
```

### **Isolate-based:**

This method does not require any selection, as all isolates should be included. It does, however, respect all arguments set in the first\_isolate() function. For example, the default setting for include\_unknown (FALSE) will omit selection of rows without a microbial ID.

### Patient-based:

To include every genus-species combination per patient once, set the episode\_days to Inf. Although often inappropriate, this method makes sure that no duplicate isolates are selected from the same patient. In a large longitudinal data set, this could mean that isolates are *excluded* that were found years after the initial isolate.

# **Episode-based:**

To include every genus-species combination per patient episode once, set the episode\_days to a sensible number of days. Depending on the type of analysis, this could be 14, 30, 60 or 365. Short episodes are common for analysing specific hospital or ward data, long episodes are common for analysing regional and national data.

This is the most common method to correct for duplicate isolates. Patients are categorised into episodes based on their ID and dates (e.g., the date of specimen receipt or laboratory result). While this is a common method, it does not take into account antimicrobial test results. This means that e.g. a methicillin-resistant *Staphylococcus aureus* (MRSA) isolate cannot be differentiated from a wildtype *Staphylococcus aureus* isolate.

# Phenotype-based:

This is a more reliable method, since it also *weighs* the antibiogram (antimicrobial test results) yielding so-called 'first weighted isolates'. There are two different methods to weigh the antibiogram:

- 1. Using type = "points" and argument points\_threshold (default)
   This method weighs all antimicrobial drugs available in the data set. Any difference from I to S or R (or vice versa) counts as 0.5 points, a difference from S to R (or vice versa) counts as 1 point. When the sum of points exceeds points\_threshold, which defaults to 2, an isolate will be selected as a first weighted isolate.
   All antimicrobials are internally selected using the all\_antimicrobials() function. The output of this function does not need to be passed to the first\_isolate() function.
- 2. Using type = "keyantimicrobials" and argument ignore\_I This method only weighs specific antimicrobial drugs, called key antimicrobials. Any difference from S to R (or vice versa) in these key antimicrobials will select an isolate as a first weighted isolate. With ignore\_I = FALSE, also differences from I to S or R (or vice versa) will lead to this.

Key antimicrobials are internally selected using the key\_antimicrobials() function, but can also be added manually as a variable to the data and set in the col\_keyantimicrobials argument. Another option is to pass the output of the key\_antimicrobials() function directly to the col\_keyantimicrobials argument.

The default method is phenotype-based (using type = "points") and episode-based (using episode\_days = 365). This makes sure that every genus-species combination is selected per patient once per year, while taking into account all antimicrobial test results. If no antimicrobial test results are available in the data set, only the episode-based method is applied at default.

### Value

A logical vector

### Source

Methodology of this function is strictly based on:

- M39 Analysis and Presentation of Cumulative Antimicrobial Susceptibility Test Data, 5th Edition, 2022, Clinical and Laboratory Standards Institute (CLSI). https://clsi.org/standards/products/microbiology/documents/m39/.
- Hindler JF and Stelling J (2007). Analysis and Presentation of Cumulative Antibiograms:
   A New Consensus Guideline from the Clinical and Laboratory Standards Institute. Clinical Infectious Diseases, 44(6), 867-873. doi:10.1086/511864

#### See Also

key\_antimicrobials()

# **Examples**

- # `example\_isolates` is a data set available in the AMR package.
- # See ?example\_isolates.

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```
example_isolates[first_isolate(info = TRUE), ]
# get all first Gram-negatives
example_isolates[which(first_isolate(info = FALSE) & mo_is_gram_negative()), ]
if (require("dplyr")) {
 # filter on first isolates using dplyr:
 example_isolates %>%
    filter(first_isolate(info = TRUE))
}
if (require("dplyr")) {
 # short-hand version:
 example_isolates %>%
    filter_first_isolate(info = FALSE)
}
if (require("dplyr")) {
 # flag the first isolates per group:
 example_isolates %>%
   group_by(ward) %>%
   mutate(first = first_isolate(info = TRUE)) %>%
   select(ward, date, patient, mo, first)
}
```

g.test

G-test for Count Data

# **Description**

g.test() performs chi-squared contingency table tests and goodness-of-fit tests, just like chisq.test() but is more reliable (1). A *G*-test can be used to see whether the number of observations in each category fits a theoretical expectation (called a *G*-test of goodness-of-fit), or to see whether the proportions of one variable are different for different values of the other variable (called a *G*-test of independence).

# Usage

```
g.test(x, y = NULL, p = rep(1/length(x), length(x)), rescale.p = FALSE)
```

# **Arguments**

X	a numeric vector or matrix. x and y can also both be factors.
У	a numeric vector; ignored if $x$ is a matrix. If $x$ is a factor, $y$ should be a factor of the same length.
p	a vector of probabilities of the same length as $x$ . An error is given if any entry of $p$ is negative.
rescale.p	a logical scalar; if TRUE then p is rescaled (if necessary) to sum to 1. If rescale.p is FALSE, and p does not sum to 1, an error is given.

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#### **Details**

If x is a matrix with one row or column, or if x is a vector and y is not given, then a *goodness-of-fit* test is performed (x is treated as a one-dimensional contingency table). The entries of x must be non-negative integers. In this case, the hypothesis tested is whether the population probabilities equal those in p, or are all equal if p is not given.

If x is a matrix with at least two rows and columns, it is taken as a two-dimensional contingency table: the entries of x must be non-negative integers. Otherwise, x and y must be vectors or factors of the same length; cases with missing values are removed, the objects are coerced to factors, and the contingency table is computed from these. Then Pearson's chi-squared test is performed of the null hypothesis that the joint distribution of the cell counts in a 2-dimensional contingency table is the product of the row and column marginals.

The p-value is computed from the asymptotic chi-squared distribution of the test statistic.

In the contingency table case simulation is done by random sampling from the set of all contingency tables with given marginals, and works only if the marginals are strictly positive. Note that this is not the usual sampling situation assumed for a chi-squared test (such as the G-test) but rather that for Fisher's exact test.

In the goodness-of-fit case simulation is done by random sampling from the discrete distribution specified by p, each sample being of size n = sum(x). This simulation is done in R and may be slow.

### G-test Of Goodness-of-Fit (Likelihood Ratio Test):

Use the *G*-test of goodness-of-fit when you have one nominal variable with two or more values (such as male and female, or red, pink and white flowers). You compare the observed counts of numbers of observations in each category with the expected counts, which you calculate using some kind of theoretical expectation (such as a 1:1 sex ratio or a 1:2:1 ratio in a genetic cross).

If the expected number of observations in any category is too small, the G-test may give inaccurate results, and you should use an exact test instead (fisher.test()).

The *G*-test of goodness-of-fit is an alternative to the chi-square test of goodness-of-fit (chisq.test()); each of these tests has some advantages and some disadvantages, and the results of the two tests are usually very similar.

### **G**-test of Independence:

Use the *G*-test of independence when you have two nominal variables, each with two or more possible values. You want to know whether the proportions for one variable are different among values of the other variable.

It is also possible to do a *G*-test of independence with more than two nominal variables. For example, Jackson et al. (2013) also had data for children under 3, so you could do an analysis of old vs. young, thigh vs. arm, and reaction vs. no reaction, all analyzed together.

Fisher's exact test (fisher.test()) is an **exact** test, where the *G*-test is still only an **approximation**. For any 2x2 table, Fisher's Exact test may be slower but will still run in seconds, even if the sum of your observations is multiple millions.

The *G*-test of independence is an alternative to the chi-square test of independence (chisq.test()), and they will give approximately the same results.

### **How the Test Works:**

Unlike the exact test of goodness-of-fit (fisher.test()), the *G*-test does not directly calculate the probability of obtaining the observed results or something more extreme. Instead, like almost

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all statistical tests, the G-test has an intermediate step; it uses the data to calculate a test statistic that measures how far the observed data are from the null expectation. You then use a mathematical relationship, in this case the chi-square distribution, to estimate the probability of obtaining that value of the test statistic.

The G-test uses the log of the ratio of two likelihoods as the test statistic, which is why it is also called a likelihood ratio test or log-likelihood ratio test. The formula to calculate a G-statistic is:

```
G = 2 * sum(x * log(x/E))
```

where E are the expected values. Since this is chi-square distributed, the p value can be calculated in R with:

```
p <- stats::pchisq(G, df, lower.tail = FALSE)</pre>
```

where df are the degrees of freedom.

If there are more than two categories and you want to find out which ones are significantly different from their null expectation, you can use the same method of testing each category vs. the sum of all categories, with the Bonferroni correction. You use *G*-tests for each category, of course.

#### Value

A list with class "htest" containing the following components:

statistic	the value the chi-squared test statistic.
parameter	the degrees of freedom of the approximate chi-squared distribution of the test statistic, NA if the p-value is computed by Monte Carlo simulation.
p.value	the p-value for the test.
method	a character string indicating the type of test performed, and whether Monte Carlo simulation or continuity correction was used.
data.name	a character string giving the name(s) of the data.
observed	the observed counts.
expected	the expected counts under the null hypothesis.
residuals	the Pearson residuals, (observed - expected) / sqrt(expected).
stdres	standardized residuals, (observed - expected) / sqrt(V), where V is the residual cell variance (Agresti, 2007, section 2.4.5 for the case where x is a matrix, n

# Source

The code for this function is identical to that of chisq.test(), except that:

\*p\*(1-p) otherwise).

- The calculation of the statistic was changed to 2 \* sum(x \* log(x/E))
- Yates' continuity correction was removed as it does not apply to a G-test
- The possibility to simulate p values with simulate.p.value was removed

#### References

1. McDonald, J.H. 2014. **Handbook of Biological Statistics (3rd ed.)**. Sparky House Publishing, Baltimore, Maryland. http://www.biostathandbook.com/gtestgof.html.

# See Also

```
chisq.test()
```

# Examples

```
# = EXAMPLE 1 =
# Shivrain et al. (2006) crossed clearfield rice (which are resistant
# to the herbicide imazethapyr) with red rice (which are susceptible to
# imazethapyr). They then crossed the hybrid offspring and examined the
# F2 generation, where they found 772 resistant plants, 1611 moderately
# resistant plants, and 737 susceptible plants. If resistance is controlled
# by a single gene with two co-dominant alleles, you would expect a 1:2:1
# ratio.
x < -c(772, 1611, 737)
g.test(x, p = c(1, 2, 1) / 4)
# There is no significant difference from a 1:2:1 ratio.
# Meaning: resistance controlled by a single gene with two co-dominant
# alleles, is plausible.
# = EXAMPLE 2 =
# Red crossbills (Loxia curvirostra) have the tip of the upper bill either
# right or left of the lower bill, which helps them extract seeds from pine
# cones. Some have hypothesized that frequency-dependent selection would
# keep the number of right and left-billed birds at a 1:1 ratio. Groth (1992)
# observed 1752 right-billed and 1895 left-billed crossbills.
x < -c(1752, 1895)
g.test(x)
# There is a significant difference from a 1:1 ratio.
# Meaning: there are significantly more left-billed birds.
```

get\_episode

Determine Clinical or Epidemic Episodes

# Description

These functions determine which items in a vector can be considered (the start of) a new episode. This can be used to determine clinical episodes for any epidemiological analysis. The get\_episode() function returns the index number of the episode per group, while the is\_new\_episode() function returns TRUE for every new get\_episode() index. Both absolute and relative episode determination are supported.

# Usage

```
get_episode(x, episode_days = NULL, case_free_days = NULL, ...)
is_new_episode(x, episode_days = NULL, case_free_days = NULL, ...)
```

#### **Arguments**

X	vector of dates (class Date or POSIXt), will be sorted internally to determine episodes
episode_days	episode length in days to specify the time period after which a new episode begins, can also be less than a day or Inf, see <i>Details</i>
case_free_days	(inter-epidemic) interval length in days after which a new episode will start, can also be less than a day or Inf, see <i>Details</i>
	ignored, only in place to allow future extensions

# **Details**

Episodes can be determined in two ways: absolute and relative.

# 1. Absolute

This method uses episode\_days to define an episode length in days, after which a new episode will start. A common use case in AMR data analysis is microbial epidemiology: episodes of *S. aureus* bacteraemia in ICU patients for example. The episode length could then be 30 days, so that new *S. aureus* isolates after an ICU episode of 30 days will be considered a different (or new) episode.

Thus, this method counts since the start of the previous episode.

### 2. Relative

This method uses case\_free\_days to quantify the duration of case-free days (the interepidemic interval), after which a new episode will start. A common use case is infectious disease epidemiology: episodes of norovirus outbreaks in a hospital for example. The case-free period could then be 14 days, so that new norovirus cases after that time will be considered a different (or new) episode.

Thus, this methods counts since the last case in the previous episode.

# In a table:

Date	Using episode_days = 7	Using case_free_days = 7
2023-01-01	1	1
2023-01-02	1	1
2023-01-05	1	1
2023-01-08	2**	1
2023-02-21	3	2***
2023-02-22	3	2
2023-02-23	3	2
2023-02-24	3	2
2023-03-01	4	2

\*\* This marks the start of a new episode, because 8 January 2023 is more than 7 days since the start of the previous episode (1 January 2023).

\*\*\* This marks the start of a new episode, because 21 January 2023 is more than 7 days since the last case in the previous episode (8 January 2023).

Either episode\_days or case\_free\_days must be provided in the function.

# Difference between get\_episode() and is\_new\_episode():

The get\_episode() function returns the index number of the episode, so all cases/patients/isolates in the first episode will have the number 1, all cases/patients/isolates in the second episode will have the number 2, etc.

The is\_new\_episode() function on the other hand, returns TRUE for every new get\_episode() index.

To specify, when setting episode\_days = 365 (using method 1 as explained above), this is how the two functions differ:

patient	date	<pre>get_episode()</pre>	is_new_episode()
A	2019-01-01	1	TRUE
A	2019-03-01	1	FALSE
A	2021-01-01	2	TRUE
В	2008-01-01	1	TRUE
В	2008-01-01	1	FALSE
C	2020-01-01	1	TRUE

### Other:

The first\_isolate() function is a wrapper around the is\_new\_episode() function, but is more efficient for data sets containing microorganism codes or names and allows for different isolate selection methods.

The dplyr package is not required for these functions to work, but these episode functions do support variable grouping and work conveniently inside dplyr verbs such as filter(), mutate() and summarise().

### Value

```
• get_episode(): an integer vector
```

```
• is_new_episode(): a logical vector
```

# See Also

```
first_isolate()
```

# **Examples**

```
# difference between absolute and relative determination of episodes: x <- data.frame(dates = as.Date(c("2021-01-01", "2021-01-02", "2021-01-05", "2021-01-05", "2021-01-08",
```

```
"2021-02-21",
  "2021-02-22"
  "2021-02-23",
 "2021-02-24",
 "2021-03-01",
 "2021-03-01"
x$absolute <- get_episode(x$dates, episode_days = 7)</pre>
x$relative <- get_episode(x$dates, case_free_days = 7)</pre>
# `example_isolates` is a data set available in the AMR package.
# See ?example_isolates
df <- example_isolates[sample(seq_len(2000), size = 100), ]</pre>
get_episode(df$date, episode_days = 60) # indices
is_new_episode(df$date, episode_days = 60) # TRUE/FALSE
# filter on results from the third 60-day episode only, using base R
df[which(get_episode(df$date, 60) == 3), ]
# the functions also work for less than a day, e.g. to include one per hour:
get_episode(
 c(
   Sys.time(),
   Sys.time() + 60 \times 60
 episode_days = 1 / 24
)
if (require("dplyr")) {
 # is_new_episode() can also be used in dplyr verbs to determine patient
 # episodes based on any (combination of) grouping variables:
 df %>%
   mutate(condition = sample(
      x = c("A", "B", "C"),
      size = 100,
      replace = TRUE
   )) %>%
   group_by(patient, condition) %>%
   mutate(new_episode = is_new_episode(date, 365)) %>%
   select(patient, date, condition, new_episode) %>%
   arrange(patient, condition, date)
}
if (require("dplyr")) {
 df %>%
    group_by(ward, patient) %>%
    transmute(date,
      patient,
      new_index = get_episode(date, 60),
```

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```
new_logical = is_new_episode(date, 60)
   arrange(patient, ward, date)
}
if (require("dplyr")) {
 df %>%
   group_by(ward) %>%
   summarise(
      n_patients = n_distinct(patient),
      n_episodes_365 = sum(is_new_episode(date, episode_days = 365)),
      n_episodes_60 = sum(is_new_episode(date, episode_days = 60)),
      n_episodes_30 = sum(is_new_episode(date, episode_days = 30))
}
\ensuremath{\text{\#}} grouping on patients and microorganisms leads to the same
# results as first_isolate() when using 'episode-based':
if (require("dplyr")) {
 x <- df %>%
   filter_first_isolate(
      include_unknown = TRUE,
      method = "episode-based"
   )
 y <- df %>%
    group_by(patient, mo) %>%
    filter(is_new_episode(date, 365)) %>%
   ungroup()
 identical(x, y)
}
# but is_new_episode() has a lot more flexibility than first_isolate(),
# since you can now group on anything that seems relevant:
if (require("dplyr")) {
 df %>%
    group_by(patient, mo, ward) %>%
   mutate(flag_episode = is_new_episode(date, 365)) %>%
    select(group_vars(.), flag_episode)
}
```

ggplot\_pca

PCA Biplot with ggplot2

# **Description**

Produces a ggplot2 variant of a so-called biplot for PCA (principal component analysis), but is more flexible and more appealing than the base R biplot() function.

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# Usage

```
ggplot_pca(
  Х,
  choices = 1:2,
  scale = 1,
  pc.biplot = TRUE,
  labels = NULL,
  labels_textsize = 3,
  labels_text_placement = 1.5,
  groups = NULL,
  ellipse = TRUE,
  ellipse_prob = 0.68,
  ellipse_size = 0.5,
  ellipse_alpha = 0.5,
  points_size = 2,
  points_alpha = 0.25,
  arrows = TRUE,
  arrows_colour = "darkblue",
  arrows_size = 0.5,
  arrows_textsize = 3,
  arrows_textangled = TRUE,
  arrows_alpha = 0.75,
  base_textsize = 10,
)
```

# **Arguments**

x an object returned by pca(), prcomp() or princomp()

choices length 2 vector specifying the components to plot. Only the default is a biplot in

the strict sense.

scale The variables are scaled by lambda ^ scale and the observations are scaled

by lambda  $^$  (1-scale) where lambda are the singular values as computed by princomp. Normally  $0 \le$ scale  $\le 1$ , and a warning will be issued if the spec-

ified scale is outside this range.

pc.biplot If true, use what Gabriel (1971) refers to as a "principal component biplot", with

lambda = 1 and observations scaled up by sqrt(n) and variables scaled down by sqrt(n). Then inner products between variables approximate covariances and

distances between observations approximate Mahalanobis distance.

labels an optional vector of labels for the observations. If set, the labels will be placed

below their respective points. When using the pca() function as input for x, this will be determined automatically based on the attribute non\_numeric\_cols, see

pca().

labels\_textsize

the size of the text used for the labels

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labels\_text\_placement

adjustment factor the placement of the variable names (>=1 means further away

from the arrow head)

groups an optional vector of groups for the labels, with the same length as labels. If

set, the points and labels will be coloured according to these groups. When using the pca() function as input for x, this will be determined automatically

based on the attribute non\_numeric\_cols, see pca().

ellipse a logical to indicate whether a normal data ellipse should be drawn for each

group (set with groups)

ellipse\_prob statistical size of the ellipse in normal probability

ellipse\_size the size of the ellipse line

ellipse\_alpha the alpha (transparency) of the ellipse line

points\_size the size of the points

points\_alpha the alpha (transparency) of the points

arrows a logical to indicate whether arrows should be drawn

arrows\_colour the colour of the arrow and their text the size (thickness) of the arrow lines

arrows\_textsize

the size of the text at the end of the arrows

arrows\_textangled

a logical whether the text at the end of the arrows should be angled

arrows\_alpha the alpha (transparency) of the arrows and their text

base\_textsize the text size for all plot elements except the labels and arrows

... arguments passed on to functions

### Details

The colours for labels and points can be changed by adding another scale layer for colour, such as scale\_colour\_viridis\_d() and scale\_colour\_brewer().

# Source

The ggplot\_pca() function is based on the ggbiplot() function from the ggbiplot package by Vince Vu, as found on GitHub: https://github.com/vqv/ggbiplot (retrieved: 2 March 2020, their latest commit: 7325e88; 12 February 2015).

As per their GPL-2 licence that demands documentation of code changes, the changes made based on the source code were:

- 1. Rewritten code to remove the dependency on packages plyr, scales and grid
- 2. Parametrised more options, like arrow and ellipse settings
- 3. Hardened all input possibilities by defining the exact type of user input for every argument
- 4. Added total amount of explained variance as a caption in the plot
- Cleaned all syntax based on the lintr package, fixed grammatical errors and added integrity checks
- 6. Updated documentation

# **Examples**

```
# `example_isolates` is a data set available in the AMR package.
# See ?example_isolates.
if (require("dplyr")) {
 # calculate the resistance per group first
 resistance_data <- example_isolates %>%
   group_by(
     order = mo_order(mo), # group on anything, like order
     genus = mo_genus(mo)
    ) %>% # and genus as we do here;
    filter(n() >= 30) %>% # filter on only 30 results per group
   summarise_if(is.sir, resistance) # then get resistance of all drugs
 # now conduct PCA for certain antimicrobial drugs
 pca_result <- resistance_data %>%
   pca(AMC, CXM, CTX, CAZ, GEN, TOB, TMP, SXT)
 summary(pca_result)
 # old base R plotting method:
 biplot(pca_result)
 # new ggplot2 plotting method using this package:
 if (require("ggplot2")) {
    ggplot_pca(pca_result)
    # still extendible with any ggplot2 function
   ggplot_pca(pca_result) +
     scale_colour_viridis_d() +
     labs(title = "Title here")
 }
}
```

ggplot\_sir

AMR Plots with ggplot2

# **Description**

Use these functions to create bar plots for AMR data analysis. All functions rely on ggplot2 functions.

# Usage

```
ggplot_sir(
  data,
  position = NULL,
```

```
x = "antibiotic",
  fill = "interpretation",
  facet = NULL,
  breaks = seq(0, 1, 0.1),
  limits = NULL,
  translate_ab = "name",
  combine_SI = TRUE,
  minimum = 30,
  language = get_AMR_locale(),
  nrow = NULL,
  colours = c(S = "#3CAEA3", SI = "#3CAEA3", I = "#F6D55C", IR = "#ED553B", R =
    "#ED553B"),
  datalabels = TRUE,
  datalabels.size = 2.5,
  datalabels.colour = "grey15",
  title = NULL,
  subtitle = NULL,
  caption = NULL,
  x.title = "Antimicrobial",
 y.title = "Proportion",
)
geom_sir(
 position = NULL,
 x = c("antibiotic", "interpretation"),
  fill = "interpretation",
  translate_ab = "name",
 minimum = 30,
  language = get_AMR_locale(),
  combine_SI = TRUE,
)
facet_sir(facet = c("interpretation", "antibiotic"), nrow = NULL)
scale_y_percent(breaks = seq(0, 1, 0.1), limits = NULL)
scale_sir_colours(..., aesthetics = "fill")
theme_sir()
labels_sir_count(
  position = NULL,
  x = "antibiotic",
  translate_ab = "name",
  minimum = 30,
  language = get_AMR_locale(),
```

```
combine_SI = TRUE,
datalabels.size = 3,
datalabels.colour = "grey15"
)
```

#### **Arguments**

y.title

data a data.frame with column(s) of class sir (see as.sir()) position position adjustment of bars, either "fill", "stack" or "dodge" variable to show on x axis, either "antibiotic" (default) or "interpretation" Х or a grouping variable fill variable to categorise using the plots legend, either "antibiotic" (default) or "interpretation" or a grouping variable facet variable to split plots by, either "interpretation" (default) or "antibiotic" or a grouping variable breaks a numeric vector of positions limits a numeric vector of length two providing limits of the scale, use NA to refer to the existing minimum or maximum translate\_ab a column name of the antibiotics data set to translate the antibiotic abbreviations to, using ab\_property() combine SI a logical to indicate whether all values of S and I must be merged into one, so the output only consists of S+I vs. R (susceptible vs. resistant) - the default is minimum the minimum allowed number of available (tested) isolates. Any isolate count lower than minimum will return NA with a warning. The default number of 30 isolates is advised by the Clinical and Laboratory Standards Institute (CLSI) as best practice, see Source. language language of the returned text - the default is the current system language (see get\_AMR\_locale()) and can also be set with the package option AMR\_locale. Use language = NULL or language = "" to prevent translation. (when using facet) number of rows nrow colours a named vactor with colour to be used for filling. The default colours are colourblind friendly. datalabels show datalabels using labels\_sir\_count() datalabels.size size of the datalabels datalabels.colour colour of the datalabels title text to show as title of the plot text to show as subtitle of the plot subtitle caption text to show as caption of the plot x.title text to show as x axis description

text to show as y axis description

```
other arguments passed on to <code>geom_sir()</code> or, in case of <code>scale_sir_colours()</code>, named values to set colours. The default colours are colour-blind friendly, while maintaining the convention that e.g. 'susceptible' should be green and 'resistant' should be red. See <code>Examples</code>.

aesthetics aesthetics to apply the colours to - the default is "fill" but can also be (a combination of) "alpha", "colour", "fill", "linetype", "shape" or "size"
```

#### **Details**

At default, the names of antibiotics will be shown on the plots using ab\_name(). This can be set with the translate\_ab argument. See count\_df().

#### The Functions:

```
geom_sir() will take any variable from the data that has an sir class (created with as.sir())
using sir_df() and will plot bars with the percentage S, I, and R. The default behaviour is to
have the bars stacked and to have the different antibiotics on the x axis.
facet_sir() creates 2d plots (at default based on S/I/R) using ggplot2::facet_wrap().
scale_y_percent() transforms the y axis to a 0 to 100% range using ggplot2::scale_y_continuous().
scale_sir_colours() sets colours to the bars (green for S, yellow for I, and red for R). with multilingual support. The default colours are colour-blind friendly, while maintaining the convention
that e.g. 'susceptible' should be green and 'resistant' should be red.
theme_sir() is a [ggplot2 theme][ggplot2::theme() with minimal distraction.
labels_sir_count() print datalabels on the bars with percentage and amount of isolates using
ggplot2::geom_text().
ggplot_sir() is a wrapper around all above functions that uses data as first input. This makes it
possible to use this function after a pipe (%>%). See Examples.
```

# **Examples**

```
if (require("ggplot2") && require("dplyr")) {
 # get antimicrobial results for drugs against a UTI:
 ggplot(example_isolates %>% select(AMX, NIT, FOS, TMP, CIP)) +
   geom_sir()
}
if (require("ggplot2") && require("dplyr")) {
 # prettify the plot using some additional functions:
 df <- example_isolates %>% select(AMX, NIT, FOS, TMP, CIP)
 ggplot(df) +
   geom_sir() +
   scale_y_percent() +
   scale_sir_colours() +
   labels_sir_count() +
   theme_sir()
if (require("ggplot2") && require("dplyr")) {
 # or better yet, simplify this using the wrapper function - a single command:
 example_isolates %>%
   select(AMX, NIT, FOS, TMP, CIP) %>%
   ggplot_sir()
```

```
if (require("ggplot2") && require("dplyr")) {
 # get only proportions and no counts:
 example_isolates %>%
   select(AMX, NIT, FOS, TMP, CIP) %>%
   ggplot_sir(datalabels = FALSE)
if (require("ggplot2") && require("dplyr")) {
 # add other ggplot2 arguments as you like:
 example_isolates %>%
    select(AMX, NIT, FOS, TMP, CIP) %>%
   ggplot_sir(
     width = 0.5,
     colour = "black",
     size = 1,
     linetype = 2,
     alpha = 0.25
   )
}
if (require("ggplot2") && require("dplyr")) {
  # you can alter the colours with colour names:
 example_isolates %>%
    select(AMX) %>%
   ggplot_sir(colours = c(SI = "yellow"))
}
if (require("ggplot2") && require("dplyr")) {
 # but you can also use the built-in colour-blind friendly colours for
 \# your plots, where "S" is green, "I" is yellow and "R" is red:
 data.frame(
   x = c("Value1", "Value2", "Value3"),
   y = c(1, 2, 3),
    z = c("Value4", "Value5", "Value6")
 ) %>%
   ggplot() +
   geom\_col(aes(x = x, y = y, fill = z)) +
   scale_sir_colours(Value4 = "S", Value5 = "I", Value6 = "R")
if (require("ggplot2") && require("dplyr")) {
 # resistance of ciprofloxacine per age group
 example_isolates %>%
   mutate(first_isolate = first_isolate()) %>%
   filter(
     first_isolate == TRUE,
     mo == as.mo("Escherichia coli")
    ) %>%
    # age_groups() is also a function in this AMR package:
   group_by(age_group = age_groups(age)) %>%
   select(age_group, CIP) %>%
   ggplot_sir(x = "age_group")
if (require("ggplot2") && require("dplyr")) {
 # a shorter version which also adjusts data label colours:
 example_isolates %>%
```

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```
select(AMX, NIT, FOS, TMP, CIP) %>%
   ggplot_sir(colours = FALSE)
if (require("ggplot2") && require("dplyr")) {
 # it also supports groups (don't forget to use the group var on `x` or `facet`):
 example_isolates %>%
    filter(mo_is_gram_negative(), ward != "Outpatient") %>%
    # select only UTI-specific drugs
    select(ward, AMX, NIT, FOS, TMP, CIP) %>%
   group_by(ward) %>%
   ggplot_sir(
     x = "ward",
     facet = "antibiotic",
     nrow = 1,
     title = "AMR of Anti-UTI Drugs Per Ward",
     x.title = "Ward",
     datalabels = FALSE
}
```

guess\_ab\_col

Guess Antibiotic Column

# Description

This tries to find a column name in a data set based on information from the antibiotics data set. Also supports WHONET abbreviations.

# Usage

```
guess_ab_col(
  x = NULL,
  search_string = NULL,
  verbose = FALSE,
  only_sir_columns = FALSE
)
```

# Arguments

```
x a data.frame

search_string a text to search x for, will be checked with as.ab() if this value is not a column in x

verbose a logical to indicate whether additional info should be printed only_sir_columns
```

a logical to indicate whether only antibiotic columns must be detected that were transformed to class sir (see as.sir()) on beforehand (default is FALSE)

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### **Details**

You can look for an antibiotic (trade) name or abbreviation and it will search x and the antibiotics data set for any column containing a name or code of that antibiotic.

#### Value

A column name of x, or NULL when no result is found.

### **Examples**

```
df <- data.frame(
   amox = "S",
   tetr = "R"
)

guess_ab_col(df, "amoxicillin")
guess_ab_col(df, "J01AA07") # ATC code of tetracycline

guess_ab_col(df, "J01AA07", verbose = TRUE)
# NOTE: Using column 'tetr' as input for J01AA07 (tetracycline).

# WHONET codes
df <- data.frame(
   AMP_ND10 = "R",
   AMC_ED20 = "S"
)
guess_ab_col(df, "ampicillin")
guess_ab_col(df, "J01CR02")
guess_ab_col(df, as.ab("augmentin"))</pre>
```

intrinsic\_resistant

Data Set with Bacterial Intrinsic Resistance

# Description

Data set containing defined intrinsic resistance by EUCAST of all bug-drug combinations.

### Usage

```
intrinsic_resistant
```

## **Format**

A tibble with 134 634 observations and 2 variables:

- mo Microorganism ID
- ab
   Antibiotic ID

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#### **Details**

This data set is based on 'EUCAST Expert Rules' and 'EUCAST Intrinsic Resistance and Unusual Phenotypes' v3.3 (2021).

#### Direct download:

Like all data sets in this package, this data set is publicly available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

They allow for machine reading EUCAST and CLSI guidelines, which is almost impossible with the MS Excel and PDF files distributed by EUCAST and CLSI.

# **Examples**

intrinsic\_resistant

italicise\_taxonomy

Italicise Taxonomic Families, Genera, Species, Subspecies

# Description

According to the binomial nomenclature, the lowest four taxonomic levels (family, genus, species, subspecies) should be printed in italics. This function finds taxonomic names within strings and makes them italic.

## Usage

```
italicise_taxonomy(string, type = c("markdown", "ansi"))
italicize_taxonomy(string, type = c("markdown", "ansi"))
```

#### Arguments

string a character (vector)

type type of conversion of the taxonomic names, either "markdown" or "ansi", see

Details

## **Details**

This function finds the taxonomic names and makes them italic based on the microorganisms data set.

The taxonomic names can be italicised using markdown (the default) by adding \* before and after the taxonomic names, or using ANSI colours by adding \033[3m before and \033[23m after the taxonomic names. If multiple ANSI colours are not available, no conversion will occur.

This function also supports abbreviation of the genus if it is followed by a species, such as "E. coli" and "K. pneumoniae ozaenae".

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## **Examples**

```
italicise_taxonomy("An overview of Staphylococcus aureus isolates")
italicise_taxonomy("An overview of S. aureus isolates")
cat(italicise_taxonomy("An overview of S. aureus isolates", type = "ansi"))
```

join

Join microorganisms to a Data Set

# **Description**

Join the data set microorganisms easily to an existing data set or to a character vector.

### Usage

```
inner_join_microorganisms(x, by = NULL, suffix = c("2", ""), \ldots)

left_join_microorganisms(x, by = NULL, suffix = c("2", ""), \ldots)

right_join_microorganisms(x, by = NULL, suffix = c("2", ""), \ldots)

full_join_microorganisms(x, by = NULL, suffix = c("2", ""), \ldots)

semi_join_microorganisms(x, by = NULL, ...)

anti_join_microorganisms(x, by = NULL, ...)
```

## **Arguments**

X	existing data set to join, or character vector. In case of a character vector, the resulting data.frame will contain a column 'x' with these values.
by	a variable to join by - if left empty will search for a column with class mo (created with as.mo()) or will be "mo" if that column name exists in x, could otherwise be a column name of x with values that exist in microorganisms\$mo (such as by = "bacteria_id"), or another column in microorganisms (but then it should be named, like by = c("bacteria_id" = "fullname"))
suffix	if there are non-joined duplicate variables in x and y, these suffixes will be added to the output to disambiguate them. Should be a character vector of length 2.
	ignored, only in place to allow future extensions

# **Details**

**Note:** As opposed to the join() functions of dplyr, character vectors are supported and at default existing columns will get a suffix "2" and the newly joined columns will not get a suffix.

If the dplyr package is installed, their join functions will be used. Otherwise, the much slower merge() and interaction() functions from base R will be used.

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### Value

a data.frame

### **Examples**

```
left_join_microorganisms(as.mo("K. pneumoniae"))
left_join_microorganisms("B_KLBSL_PNMN")
df <- data.frame(</pre>
 date = seq(
   from = as.Date("2018-01-01"),
   to = as.Date("2018-01-07"),
 ),
 bacteria = as.mo(c(
    "S. aureus", "MRSA", "MSSA", "STAAUR",
    "E. coli", "E. coli", "E. coli"
 stringsAsFactors = FALSE
colnames(df)
df_joined <- left_join_microorganisms(df, "bacteria")</pre>
colnames(df_joined)
if (require("dplyr")) {
 example_isolates %>%
    left_join_microorganisms() %>%
    colnames()
}
```

key\_antimicrobials

(Key) Antimicrobials for First Weighted Isolates

### **Description**

These functions can be used to determine first weighted isolates by considering the phenotype for isolate selection (see first\_isolate()). Using a phenotype-based method to determine first isolates is more reliable than methods that disregard phenotypes.

# Usage

```
key_antimicrobials(
    x = NULL,
    col_mo = NULL,
    universal = c("ampicillin", "amoxicillin/clavulanic acid", "cefuroxime",
    "piperacillin/tazobactam", "ciprofloxacin", "trimethoprim/sulfamethoxazole"),
```

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```
gram_negative = c("gentamicin", "tobramycin", "colistin", "cefotaxime", "ceftazidime",
    "meropenem"),
 gram_positive = c("vancomycin", "teicoplanin", "tetracycline", "erythromycin",
    "oxacillin", "rifampin"),
 antifungal = c("anidulafungin", "caspofungin", "fluconazole", "miconazole", "nystatin",
    "voriconazole"),
 only_sir_columns = FALSE,
)
all_antimicrobials(x = NULL, only_sir_columns = FALSE, ...)
antimicrobials_equal(
 у,
  Ζ,
  type = c("points", "keyantimicrobials"),
  ignore_I = TRUE,
  points_threshold = 2,
)
```

#### **Arguments**

a data.frame with antibiotics columns, like AMX or amox. Can be left blank to Х determine automatically column name of the names or codes of the microorganisms (see as.mo()) - the col\_mo default is the first column of class mo. Values will be coerced using as.mo(). names of broad-spectrum antimicrobial drugs, case-insensitive. Set to NULL to universal ignore. See *Details* for the default antimicrobial drugs names of antibiotic drugs for Gram-positives, case-insensitive. Set to NULL to gram\_negative ignore. See Details for the default antibiotic drugs gram\_positive names of antibiotic drugs for Gram-negatives, case-insensitive. Set to NULL to ignore. See Details for the default antibiotic drugs antifungal names of antifungal drugs for **fungi**, case-insensitive. Set to NULL to ignore. See Details for the default antifungal drugs only\_sir\_columns a logical to indicate whether only columns must be included that were transformed to class sir (see as.sir()) on beforehand (default is FALSE) ignored, only in place to allow future extensions . . . character vectors to compare y,z type to determine weighed isolates; can be "keyantimicrobials" or "points", type see Details ignore\_I logical to indicate whether antibiotic interpretations with "I" will be ignored when type = "keyantimicrobials", see *Details* points\_threshold minimum number of points to require before differences in the antibiogram will

lead to inclusion of an isolate when type = "points", see Details

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#### **Details**

The key\_antimicrobials() and all\_antimicrobials() functions are context-aware. This means that the x argument can be left blank if used inside a data.frame call, see *Examples*.

The function key\_antimicrobials() returns a character vector with 12 antimicrobial results for every isolate. The function all\_antimicrobials() returns a character vector with all antimicrobial drug results for every isolate. These vectors can then be compared using antimicrobials\_equal(), to check if two isolates have generally the same antibiogram. Missing and invalid values are replaced with a dot (".") by key\_antimicrobials() and ignored by antimicrobials\_equal().

Please see the first\_isolate() function how these important functions enable the 'phenotype-based' method for determination of first isolates.

The default antimicrobial drugs used for all rows (set in universal) are:

- Ampicillin
- · Amoxicillin/clavulanic acid
- Cefuroxime
- Ciprofloxacin
- Piperacillin/tazobactam
- Trimethoprim/sulfamethoxazole

The default antimicrobial drugs used for **Gram-negative bacteria** (set in gram\_negative) are:

- · Cefotaxime
- Ceftazidime
- Colistin
- Gentamicin
- Meropenem
- Tobramycin

The default antimicrobial drugs used for **Gram-positive bacteria** (set in gram\_positive) are:

- Erythromycin
- Oxacillin
- Rifampin
- Teicoplanin
- Tetracycline
- · Vancomycin

The default antimicrobial drugs used for **fungi** (set in antifungal) are:

- · Anidulafungin
- Caspofungin
- Fluconazole
- Miconazole
- Nystatin
- Voriconazole

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### See Also

```
first_isolate()
```

### **Examples**

```
# `example_isolates` is a data set available in the AMR package.
# See ?example_isolates.
# output of the `key_antimicrobials()` function could be like this:
strainA <- "SSSRR.S.R..S"</pre>
strainB <- "SSSIRSSSRSSS"</pre>
# those strings can be compared with:
antimicrobials_equal(strainA, strainB, type = "keyantimicrobials")
# TRUE, because I is ignored (as well as missing values)
antimicrobials_equal(strainA, strainB, type = "keyantimicrobials", ignore_I = FALSE)
# FALSE, because I is not ignored and so the 4th [character] differs
if (require("dplyr")) {
 # set key antibiotics to a new variable
 my_patients <- example_isolates %>%
   mutate(keyab = key_antimicrobials(antifungal = NULL)) %>% # no need to define `x`
   mutate(
     # now calculate first isolates
     first_regular = first_isolate(col_keyantimicrobials = FALSE),
     # and first WEIGHTED isolates
     first_weighted = first_isolate(col_keyantimicrobials = "keyab")
   )
 # Check the difference in this data set, 'weighted' results in more isolates:
 sum(my_patients$first_regular, na.rm = TRUE)
 sum(my_patients$first_weighted, na.rm = TRUE)
}
```

kurtosis

Kurtosis of the Sample

### **Description**

Kurtosis is a measure of the "tailedness" of the probability distribution of a real-valued random variable. A normal distribution has a kurtosis of 3 and a excess kurtosis of 0.

#### Usage

```
kurtosis(x, na.rm = FALSE, excess = FALSE)
```

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```
## Default S3 method:
kurtosis(x, na.rm = FALSE, excess = FALSE)
## S3 method for class 'matrix'
kurtosis(x, na.rm = FALSE, excess = FALSE)
## S3 method for class 'data.frame'
kurtosis(x, na.rm = FALSE, excess = FALSE)
```

### **Arguments**

x a vector of values, a matrix or a data.frame

na.rm a logical to indicate whether NA values should be stripped before the computation

proceeds

excess a logical to indicate whether the excess kurtosis should be returned, defined as

the kurtosis minus 3.

#### See Also

skewness()

# **Examples**

```
kurtosis(rnorm(10000))
kurtosis(rnorm(10000), excess = TRUE)
```

like

Vectorised Pattern Matching with Keyboard Shortcut

# Description

Convenient wrapper around grepl() to match a pattern: x %like% pattern. It always returns a logical vector and is always case-insensitive (use  $x \%like\_case\%$  pattern for case-sensitive matching). Also, pattern can be as long as x to compare items of each index in both vectors, or they both can have the same length to iterate over all cases.

## Usage

```
like(x, pattern, ignore.case = TRUE)
x %like% pattern
x %unlike% pattern
x %like_case% pattern
x %unlike_case% pattern
```

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### **Arguments**

X	a character vector where matches are sought, or an object which can be coerced by as.character() to a character vector.
pattern	a character vector containing regular expressions (or a character string for fixed = TRUE) to be matched in the given character vector. Coerced by as.character() to a character string if possible.
ignore.case	if FALSE, the pattern matching is <i>case sensitive</i> and if TRUE, case is ignored during matching.

#### **Details**

These like() and %like%/%unlike% functions:

- Are case-insensitive (use %like\_case%/%unlike\_case% for case-sensitive matching)
- Support multiple patterns
- Check if pattern is a valid regular expression and sets fixed = TRUE if not, to greatly improve speed (vectorised over pattern)
- Always use compatibility with Perl unless fixed = TRUE, to greatly improve speed

Using RStudio? The %like%/%unlike% functions can also be directly inserted in your code from the Addins menu and can have its own keyboard shortcut like Shift+Ctrl+L or Shift+Cmd+L (see menu Tools > Modify Keyboard Shortcuts...). If you keep pressing your shortcut, the inserted text will be iterated over %like% -> %unlike% -> %like\_case% -> %unlike\_case%.

### Value

A logical vector

#### Source

Idea from the like function from the data. table package, although altered as explained in Details.

#### See Also

```
grepl()
```

# **Examples**

```
# data.table has a more limited version of %like%, so unload it:
try(detach("package:data.table", unload = TRUE), silent = TRUE)

a <- "This is a test"
b <- "TEST"
a %like% b
b %like% a

# also supports multiple patterns
a <- c("Test case", "Something different", "Yet another thing")
b <- c("case", "diff", "yet")</pre>
```

```
a %like% b
a %unlike% b
a %unlike% b
a %like% b[1]

# get isolates whose name start with 'Entero' (case-insensitive)
example_isolates[which(mo_name() %like% "^entero"), ]

if (require("dplyr")) {
   example_isolates %>%
     filter(mo_name() %like% "^ent")
}
```

mdro

Determine Multidrug-Resistant Organisms (MDRO)

### **Description**

Determine which isolates are multidrug-resistant organisms (MDRO) according to international, national and custom guidelines.

# Usage

```
mdro(
  x = NULL
  guideline = "CMI2012",
  col_mo = NULL,
  info = interactive(),
  pct_required_classes = 0.5,
  combine_SI = TRUE,
  verbose = FALSE,
  only_sir_columns = FALSE,
)
custom_mdro_guideline(..., as_factor = TRUE)
brmo(x = NULL, only_sir_columns = FALSE, ...)
mrgn(x = NULL, only_sir_columns = FALSE, ...)
mdr_tb(x = NULL, only_sir_columns = FALSE, ...)
mdr_cmi2012(x = NULL, only_sir_columns = FALSE, ...)
eucast_exceptional_phenotypes(x = NULL, only_sir_columns = FALSE, ...)
```

### **Arguments**

x a data.frame with antibiotics columns, like AMX or amox. Can be left blank for

automatic determination.

guideline a specific guideline to follow, see sections Supported international / national

guidelines and Using Custom Guidelines below. When left empty, the publica-

tion by Magiorakos et al. (see below) will be followed.

col\_mo column name of the names or codes of the microorganisms (see as.mo()) - the

default is the first column of class mo. Values will be coerced using as.mo().

info a logical to indicate whether progress should be printed to the console - the

default is only print while in interactive sessions

pct\_required\_classes

minimal required percentage of antimicrobial classes that must be available per isolate, rounded down. For example, with the default guideline, 17 antimicrobial classes must be available for *S. aureus*. Setting this pct\_required\_classes argument to 0.5 (default) means that for every *S. aureus* isolate at least 8 different classes must be available. Any lower number of available classes will return NA

for that isolate.

combine\_SI a logical to indicate whether all values of S and I must be merged into one, so

resistance is only considered when isolates are R, not I. As this is the default behaviour of the mdro() function, it follows the redefinition by EUCAST about the interpretation of I (increased exposure) in 2019, see section 'Interpretation of S, I and R' below. When using combine\_SI = FALSE, resistance is considered

when isolates are R or I.

verbose a logical to turn Verbose mode on and off (default is off). In Verbose mode,

the function does not return the MDRO results, but instead returns a data set in logbook form with extensive info about which isolates would be MDRO-

positive, or why they are not.

only\_sir\_columns

a logical to indicate whether only antibiotic columns must be detected that were

transformed to class sir (see as.sir()) on beforehand (default is FALSE)

in case of custom\_mdro\_guideline(): a set of rules, see section *Using Cus*-

tom Guidelines below. Otherwise: column name of an antibiotic, see section

Antibiotics below.

as\_factor a logical to indicate whether the returned value should be an ordered factor

(TRUE, default), or otherwise a character vector

### **Details**

These functions are context-aware. This means that the x argument can be left blank if used inside a data.frame call, see *Examples*.

For the pct\_required\_classes argument, values above 1 will be divided by 100. This is to support both fractions (0.75 or 3/4) and percentages (75).

**Note:** Every test that involves the Enterobacteriaceae family, will internally be performed using its newly named *order* Enterobacterales, since the Enterobacteriaceae family has been taxonomically reclassified by Adeolu *et al.* in 2016. Before that, Enterobacteriaceae was the only family under

the Enterobacteriales (with an i) order. All species under the old Enterobacteriaceae family are still under the new Enterobacterales (without an i) order, but divided into multiple families. The way tests are performed now by this mdro() function makes sure that results from before 2016 and after 2016 are identical.

# Value

- CMI 2012 paper function mdr\_cmi2012() or mdro():
  Ordered factor with levels Negative < Multi-drug-resistant (MDR) < Extensively drug-resistant (XDR) < Pandrug-resistant (PDR)
- TB guideline function mdr\_tb() or mdro(..., guideline = "TB"):

  Ordered factor with levels Negative < Mono-resistant < Poly-resistant < Multi-drug-resistant < Extensively drug-resistant
- German guideline function mrgn() or mdro(..., guideline = "MRGN"):
   Ordered factor with levels Negative < 3MRGN < 4MRGN</li>
- Everything else, except for custom guidelines:
  Ordered factor with levels Negative < Positive, unconfirmed < Positive. The value
  "Positive, unconfirmed" means that, according to the guideline, it is not entirely sure if the
  isolate is multi-drug resistant and this should be confirmed with additional (e.g. molecular)
  tests

# **Supported International / National Guidelines**

Currently supported guidelines are (case-insensitive):

- guideline = "CMI2012" (default)
  - Magiorakos AP, Srinivasan A *et al.* "Multidrug-resistant, extensively drug-resistant and pandrug-resistant bacteria: an international expert proposal for interim standard definitions for acquired resistance." Clinical Microbiology and Infection (2012) (doi:10.1111/j.14690691.2011.03570.x)
- guideline = "EUCAST3.3" (or simply guideline = "EUCAST")

  The European international guideline EUCAST Expert Rules Version 3.3 "Intrinsic Resistance and Unusual Phenotypes" (link)
- guideline = "EUCAST3.2"
   The European international guideline EUCAST Expert Rules Version 3.2 "Intrinsic Resistance and Unusual Phenotypes" (link)
- guideline = "EUCAST3.1"
   The European international guideline EUCAST Expert Rules Version 3.1 "Intrinsic Resistance and Exceptional Phenotypes Tables" (link)
- guideline = "TB"

  The international guideline for multi-drug resistant tuberculosis World Health Organization
  "Companion handbook to the WHO guidelines for the programmatic management of drugresistant tuberculosis" (link)
- guideline = "MRGN"

  The German national guideline Mueller et al. (2015) Antimicrobial Resistance and Infection
  Control 4:7; doi:10.1186/s1375601500476

• guideline = "BRMO"

The Dutch national guideline - Rijksinstituut voor Volksgezondheid en Milieu "WIP-richtlijn BRMO (Bijzonder Resistente Micro-Organismen) (ZKH)" (link)

Please suggest your own (country-specific) guidelines by letting us know: https://github.com/msberends/AMR/issues/new.

# **Using Custom Guidelines**

Custom guidelines can be set with the custom\_mdro\_guideline() function. This is of great importance if you have custom rules to determine MDROs in your hospital, e.g., rules that are dependent on ward, state of contact isolation or other variables in your data.

If you are familiar with the case\_when() function of the dplyr package, you will recognise the input method to set your own rules. Rules must be set using what R considers to be the 'formula notation'. The rule is written *before* the tilde (~) and the consequence of the rule is written *after* the tilde:

If a row/an isolate matches the first rule, the value after the first ~ (in this case '*Elderly Type A*') will be set as MDRO value. Otherwise, the second rule will be tried and so on. The number of rules is unlimited.

You can print the rules set in the console for an overview. Colours will help reading it if your console supports colours.

```
custom
#> A set of custom MDRO rules:
#> 1. CIP is "R" and age is higher than 60 -> Elderly Type A
#> 2. ERY is "R" and age is higher than 60 -> Elderly Type B
#> 3. Otherwise -> Negative
#>
```

The outcome of the function can be used for the guideline argument in the mdro() function:

#> Unmatched rows will return NA.

Rules can also be combined with other custom rules by using c():

The rules set (the custom object in this case) could be exported to a shared file location using saveRDS() if you collaborate with multiple users. The custom rules set could then be imported using readRDS().

#### **Antibiotics**

To define antibiotics column names, leave as it is to determine it automatically with guess\_ab\_col() or input a text (case-insensitive), or use NULL to skip a column (e.g. TIC = NULL to skip ticarcillin). Manually defined but non-existing columns will be skipped with a warning.

The following antibiotics are eligible for the functions eucast\_rules() and mdro(). These are shown below in the format 'name (antimicrobial ID, ATC code)', sorted alphabetically:

Amikacin (AMK, J01GB06), amoxicillin (AMX, J01CA04), amoxicillin/clavulanic acid (AMC, J01CR02), ampicillin (AMP, J01CA01), ampicillin/sulbactam (SAM, J01CR01), arbekacin (ARB, J01GB12), aspoxicillin (APX, J01CA19), azidocillin (AZD, J01CE04), azithromycin (AZM, J01FA10), azlocillin (AZL, J01CA09), aztreonam (ATM, J01DF01), bacampicillin (BAM, J01CA06), bekanamycin (BEK, J01GB13), benzathine benzylpenicillin (BNB, J01CE08), benzathine phenoxymethylpenicillin (BNP, J01CE10), benzylpenicillin (PEN, J01CE01), besifloxacin (BES, S01AE08), biapenem (BIA, J01DH05), carbenicillin (CRB, J01CA03), carindacillin (CRN, J01CA05), cefacetrile (CAC, J01DB10), cefaclor (CEC, J01DC04), cefadroxil (CFR, J01DB05), cefalexin (LEX, J01DB01), cefaloridine (RID, J01DB02), cefalotin (CEP, J01DB03), cefamandole (MAN, J01DC03), cefapirin (HAP, J01DB08), cefatrizine (CTZ, J01DB07), cefazedone (CZD, J01DB06), cefazolin (CZO, J01DB04), cefcapene (CCP, J01DD17), cefdinir (CDR, J01DD15), cefditoren (DIT, J01DD16), cefepime (FEP, J01DE01), cefetamet (CAT, J01DD10), cefixime (CFM, J01DD08), cefmenoxime (CMX, J01DD05), cefmetazole (CMZ, J01DC09), cefodizime (DIZ, J01DD09), cefonicid (CID, J01DC06), cefoperazone (CFP, J01DD12), cefoperazone/sulbactam (CSL, J01DD62), ceforanide (CND, J01DC11), cefotaxime (CTX, J01DD01), cefotaxime/clavulanic acid (CTC, J01DD51), cefotetan (CTT, J01DC05), cefotiam (CTF, J01DC07), cefoxitin (F0X, J01DC01), cefozopran (Z0P, J01DE03), cefpiramide (CPM, J01DD11), cefpirome (CPO, J01DE02), cefpodoxime (CPD, J01DD13), cefprozil (CPR, J01DC10), cefroxadine (CRD, J01DB11), cefsulodin (CFS, J01DD03), ceftaroline (CPT, J01DI02), ceftazidime (CAZ, J01DD02), ceftazidime/clavulanic acid (CCV, J01DD52), cefteram (CEM, J01DD18), ceftezole (CTL, J01DB12), ceftibuten (CTB, J01DD14), ceftizoxime (CZX, J01DD07), ceftobiprole medocaril (CFM1, J01DI01), ceftolozane/tazobactam (CZT, J01DI54), ceftriaxone (CRO, J01DD04), ceftriaxone/betalactamase inhibitor (CEB, J01DD63), cefuroxime (CXM, J01DC02), cephradine (CED, J01DB09), chloramphenicol (CHL, J01BA01), ciprofloxacin (CIP, J01MA02), clarithromycin (CLR, J01FA09), clindamycin (CLI, J01FF01), clometocillin (CLM, J01CE07), cloxacillin (CLO, J01CF02), colistin (COL, J01XB01), cycloserine (CYC, J04AB01), dalbavancin (DAL, J01XA04), daptomycin (DAP, J01XX09), delafloxacin (DFX, J01MA23), dibekacin (DKB, J01GB09), dicloxacillin (DIC, J01CF01), dirithromycin (DIR, J01FA13), doripenem (DOR, J01DH04), doxycycline (DOX, J01AA02), enoxacin (ENX, J01MA04), epicillin (EPC, J01CA07), ertapenem (ETP, J01DH03), erythromycin (ERY, J01FA01), fleroxacin (FLE, J01MA08), flucloxacillin (FLC, J01CF05), flurithromycin (FLR1, J01FA14), fosfomycin (FOS, J01XX01), framycetin (FRM, D09AA01), fusidic acid (FUS, J01XC01), garenoxacin (GRN, J01MA19), gatifloxacin (GAT, J01MA16), gemifloxacin (GEM, J01MA15), gentamicin (GEN, J01GB03), grepafloxacin (GRX, J01MA11), hetacillin (HET, J01CA18), imipenem (IPM, J01DH51), imipenem/relebactam (IMR, J01DH56), isepamicin (ISE, J01GB11), josamycin (JOS, J01FA07), kanamycin (KAN, J01GB04), lascufloxacin (LSC, J01MA25), latamoxef (LTM, J01DD06), levofloxacin (LVX, J01MA12), levonadifloxacin (LND, J01MA24), lincomycin (LIN, J01FF02), linezolid (LNZ, J01XX08), lomefloxacin (LOM, J01MA07), loracarbef (LOR, J01DC08), mecillinam (MEC, J01CA11), meropenem (MEM, J01DH02), meropenem/vaborbactam (MEV, J01DH52), metampicillin (MTM, J01CA14),

meticillin (MET, J01CF03), mezlocillin (MEZ, J01CA10), micronomicin (MCR, S01AA22), midecamycin (MID, J01FA03), minocycline (MNO, J01AA08), miocamycin (MCM, J01FA11), moxifloxacin (MFX, J01MA14), nadifloxacin (NAD, D10AF05), nafcillin (NAF, J01CF06), nalidixic acid (NAL, J01MB02), neomycin (NEO, J01GB05), netilmicin (NET, J01GB07), nitrofurantoin (NIT, J01XE01), norfloxacin (NOR, J01MA06), ofloxacin (0FX, J01MA01), oleandomycin (0LE, J01FA05), oritavancin (ORI, J01XA05), oxacillin (OXA, J01CF04), panipenem (PAN, J01DH55), pazufloxacin (PAZ, J01MA18), pefloxacin (PEF, J01MA03), penamecillin (PNM, J01CE06), pheneticillin (PHE, J01CE05), phenoxymethylpenicillin (PHN, J01CE02), piperacillin (PIP, J01CA12), piperacillin/tazobactam (TZP, J01CR05), pivampicillin (PVM, J01CA02), pivmecillinam (PME, J01CA08), plazomicin (PLZ, J01GB14), polymyxin B (PLB, J01XB02), pristinamycin (PRI, J01FG01), procaine benzylpenicillin (PRB, J01CE09), propicillin (PRP, J01CE03), prulifloxacin (PRU, J01MA17), quinupristin/dalfopristin (QDA, J01FG02), ribostamycin (RST, J01GB10), rifampicin (RIF, J04AB02), rokitamycin (R0K, J01FA12), roxithromycin (RXT, J01FA06), rufloxacin (RFL, J01MA10), sisomicin (SIS, J01GB08), sitafloxacin (SIT, J01MA21), solithromycin (SOL, J01FA16), sparfloxacin (SPX, J01MA09), spiramycin (SPI, J01FA02), streptoduocin (STR, J01GA02), streptomycin (STR1, J01GA01), sulbactam (SUL, J01CG01), sulbenicillin (SBC, J01CA16), sulfadiazine (SDI, J01EC02), sulfadiazine/trimethoprim (SLT1, J01EE02), sulfadimethoxine (SUD, J01ED01), sulfadimidine (SDM, J01EB03), sulfadimidine/trimethoprim (SLT2, J01EE05), sulfafurazole (SLF, J01EB05), sulfaisodimidine (SLF1, J01EB01), sulfalene (SLF2, J01ED02), sulfamazone (SZO, J01ED09), sulfamerazine (SLF3, J01ED07), sulfamerazine/trimethoprim (SLT3, J01EE07), sulfamethizole (SLF4, J01EB02), sulfamethoxazole (SMX, J01EC01), sulfamethoxypyridazine (SLF5, J01ED05), sulfametomidine (SLF6, J01ED03), sulfametoxydiazine (SLF7, J01ED04), sulfametrole/trimethoprim (SLT4, J01EE03), sulfamoxole (SLF8, J01EC03), sulfamoxole/trimethoprim (SLT5, J01EE04), sulfanilamide (SLF9, J01EB06), sulfaperin (SLF10, J01ED06), sulfaphenazole (SLF11, J01ED08), sulfapyridine (SLF12, J01EB04), sulfathiazole (SUT, J01EB07), sulfathiourea (SLF13, J01EB08), sultamicillin (SLT6, J01CR04), talampicillin (TAL, J01CA15), tazobactam (TAZ, J01CG02), tebipenem (TBP, J01DH06), tedizolid (TZD, J01XX11), teicoplanin (TEC, J01XA02), telavancin (TLV, J01XA03), telithromycin (TLT, J01FA15), temafloxacin (TMX, J01MA05), temocillin (TEM, J01CA17), tetracycline (TCY, J01AA07), ticarcillin (TIC, J01CA13), ticarcillin/clavulanic acid (TCC, J01CR03), tigecycline (TGC, J01AA12), tilbroquinol (TBQ, P01AA05), tobramycin (TOB, J01GB01), tosufloxacin (TFX, J01MA22), trimethoprim (TMP, J01EA01), trimethoprim/sulfamethoxazole (SXT, J01EE01), troleandomycin (TRL, J01FA08), trovafloxacin (TVA, J01MA13), vancomycin (VAN, J01XA01)

#### **Interpretation of SIR**

In 2019, the European Committee on Antimicrobial Susceptibility Testing (EUCAST) has decided to change the definitions of susceptibility testing categories S, I, and R as shown below (https://www.eucast.org/newsiandr):

#### S - Susceptible, standard dosing regimen

A microorganism is categorised as "Susceptible, standard dosing regimen", when there is a high likelihood of therapeutic success using a standard dosing regimen of the agent.

# • I - Susceptible, increased exposure

A microorganism is categorised as "Susceptible, Increased exposure" when there is a high likelihood of therapeutic success because exposure to the agent is increased by adjusting the dosing regimen or by its concentration at the site of infection.

#### • R = Resistant

A microorganism is categorised as "Resistant" when there is a high likelihood of therapeutic failure even when there is increased exposure.

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Exposure is a function of how the mode of administration, dose, dosing interval, infusion time, as well as distribution and excretion of the antimicrobial agent will influence the infecting organism at the site of infection.

This AMR package honours this insight. Use susceptibility() (equal to proportion\_SI()) to determine antimicrobial susceptibility and count\_susceptible() (equal to count\_SI()) to count susceptible isolates.

#### **Source**

See the supported guidelines above for the list of publications used for this function.

### **Examples**

```
out <- mdro(example_isolates, guideline = "EUCAST")</pre>
table(out)
out <- mdro(example_isolates,</pre>
  guideline = custom_mdro_guideline(
    AMX == "R" ~ "Custom MDRO 1",
    VAN == "R" ~ "Custom MDRO 2"
table(out)
if (require("dplyr")) {
  example_isolates %>%
    mdro() %>%
    table()
  # no need to define `x` when used inside dplyr verbs:
  example_isolates %>%
    mutate(MDRO = mdro()) %>%
    pull(MDRO) %>%
    table()
}
```

mean\_amr\_distance

Calculate the Mean AMR Distance

# Description

Calculates a normalised mean for antimicrobial resistance between multiple observations, to help to identify similar isolates without comparing antibiograms by hand.

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## Usage

```
mean_amr_distance(x, ...)
## S3 method for class 'sir'
mean_amr_distance(x, ..., combine_SI = TRUE)
## S3 method for class 'data.frame'
mean_amr_distance(x, ..., combine_SI = TRUE)
amr_distance_from_row(amr_distance, row)
```

#### **Arguments**

X	a vector of class sir, mic or disk, or a data.frame containing columns of any of these classes
• • •	variables to select (supports tidyselect language such as column1:column4 and where(is.mic), and can thus also be antibiotic selectors
combine_SI	a logical to indicate whether all values of S and I must be merged into one, so the input only consists of S+I vs. R (susceptible vs. resistant) - the default is TRUE
amr_distance	the outcome of mean_amr_distance()
row	an index, such as a row number

# **Details**

The mean AMR distance is effectively the Z-score; a normalised numeric value to compare AMR test results which can help to identify similar isolates, without comparing antibiograms by hand.

MIC values (see as.mic()) are transformed with log2() first; their distance is thus calculated as (log2(x) - mean(log2(x))) / sd(log2(x)).

SIR values (see as.sir()) are transformed using "S" = 1, "I" = 2, and "R" = 3. If combine\_SI is TRUE (default), the "I" will be considered to be 1.

For data sets, the mean AMR distance will be calculated per column, after which the mean per row will be returned, see *Examples*.

Use amr\_distance\_from\_row() to subtract distances from the distance of one row, see Examples.

# Interpretation

Isolates with distances less than 0.01 difference from each other should be considered similar. Differences lower than 0.025 should be considered suspicious.

# **Examples**

```
sir <- random_sir(10)
sir
mean_amr_distance(sir)
mic <- random_mic(10)</pre>
```

```
mean_amr_distance(mic)
# equal to the Z-score of their log2:
(log2(mic) - mean(log2(mic))) / sd(log2(mic))
disk <- random_disk(10)</pre>
disk
mean_amr_distance(disk)
y <- data.frame(</pre>
  id = LETTERS[1:10],
  amox = random_sir(10, ab = "amox", mo = "Escherichia coli"),
  cipr = random_disk(10, ab = "cipr", mo = "Escherichia coli"),
  gent = random_mic(10, ab = "gent", mo = "Escherichia coli"),
  tobr = random_mic(10, ab = "tobr", mo = "Escherichia coli")
)
У
mean_amr_distance(y)
y$amr_distance <- mean_amr_distance(y, where(is.mic))</pre>
y[order(y$amr_distance), ]
if (require("dplyr")) {
  y %>%
    mutate(
      amr_distance = mean_amr_distance(y),
      check_id_C = amr_distance_from_row(amr_distance, id == "C")
    arrange(check_id_C)
}
if (require("dplyr")) {
  # support for groups
  example_isolates %>%
    filter(mo_genus() == "Enterococcus" & mo_species() != "") %>%
    select(mo, TCY, carbapenems()) %>%
    group_by(mo) %>%
   mutate(dist = mean_amr_distance(.)) %>%
    arrange(mo, dist)
}
```

microorganisms

Data Set with 52 171 Microorganisms

# **Description**

A data set containing the full microbial taxonomy (last updated: December 11th, 2022) of five kingdoms from the List of Prokaryotic names with Standing in Nomenclature (LPSN) and the Global Biodiversity Information Facility (GBIF). This data set is the backbone of this AMR package. MO codes can be looked up using as.mo().

#### Usage

microorganisms

#### **Format**

A tibble with 52 171 observations and 23 variables:

• mc

ID of microorganism as used by this package. This is a unique identifier.

• fullname

Full name, like "Escherichia coli". For the taxonomic ranks genus, species and subspecies, this is the 'pasted' text of genus, species, and subspecies. For all taxonomic ranks higher than genus, this is the name of the taxon. *This is a unique identifier.* 

• status

Status of the taxon, either "accepted" or "synonym"

- kingdom, phylum, class, order, family, genus, species, subspecies Taxonomic rank of the microorganism
- rank

Text of the taxonomic rank of the microorganism, such as "species" or "genus"

• ref

Author(s) and year of related scientific publication. This contains only the *first surname* and year of the *latest* authors, e.g. "Wallis *et al.* 2006 *emend*. Smith and Jones 2018" becomes "Smith *et al.*, 2018". This field is directly retrieved from the source specified in the column source. Moreover, accents were removed to comply with CRAN that only allows ASCII characters.

• lpsn

Identifier ('Record number') of the List of Prokaryotic names with Standing in Nomenclature (LPSN). This will be the first/highest LPSN identifier to keep one identifier per row. For example, *Acetobacter ascendens* has LPSN Record number 7864 and 11011. Only the first is available in the microorganisms data set.

• oxygen\_tolerance

Oxygen tolerance, either "aerobe", "anaerobe", "anaerobe/microaerophile", "facultative anaerobe", "likely facultative anaerobe", or "microaerophile". These data were retrieved from Bac-Dive (see *Source*). Items that contain "likely" are missing from Bac-Dive and were extrapolated from other species within the same genus to guess the oxygen tolerance. Currently 73.4% of all ~37 000 bacteria in the data set contain an oxygen tolerance.

- lpsn\_parent
  - LPSN identifier of the parent taxon
- lpsn\_renamed\_to

LPSN identifier of the currently valid taxon

• gbif

Identifier ('taxonID') of the Global Biodiversity Information Facility (GBIF)

• gbif\_parent

GBIF identifier of the parent taxon

- gbif\_renamed\_to
   GBIF identifier of the currently valid taxon
- source Either "GBIF", "LPSN", or "manually added" (see Source)
- prevalence
   Prevalence of the microorganism according to Bartlett *et al.* (2022, doi:10.1099/mic.0.001269),
   see mo\_matching\_score() for the full explanation
- snomed Systematized Nomenclature of Medicine (SNOMED) code of the microorganism, version of July 1st, 2021 (see *Source*). Use mo\_snomed() to retrieve it quickly, see mo\_property().

#### **Details**

Please note that entries are only based on the List of Prokaryotic names with Standing in Nomenclature (LPSN) and the Global Biodiversity Information Facility (GBIF) (see below). Since these sources incorporate entries based on (recent) publications in the International Journal of Systematic and Evolutionary Microbiology (IJSEM), it can happen that the year of publication is sometimes later than one might expect.

For example, *Staphylococcus pettenkoferi* was described for the first time in Diagnostic Microbiology and Infectious Disease in 2002 (doi:10.1016/s07328893(02)003991), but it was not before 2007 that a publication in IJSEM followed (doi:10.1099/ijs.0.643810). Consequently, the AMR package returns 2007 for mo\_year("S. pettenkoferi").

#### **Included Taxa**

Included taxonomic data are:

- All ~37 000 (sub)species from the kingdoms of Archaea and Bacteria
- ~7 900 (sub)species from the kingdom of Fungi. The kingdom of Fungi is a very large taxon with almost 300,000 different (sub)species, of which most are not microbial (but rather macroscopic, like mushrooms). Because of this, not all fungi fit the scope of this package. Only relevant fungi are covered (such as all species of Aspergillus, Candida, Cryptococcus, Histoplasma, Pneumocystis, Saccharomyces and Trichophyton).
- ~5 100 (sub)species from the kingdom of Protozoa
- ~1 400 (sub)species from 43 other relevant genera from the kingdom of Animalia (such as *Strongyloides* and *Taenia*)
- All ~9 800 previously accepted names of all included (sub)species (these were taxonomically renamed)
- The complete taxonomic tree of all included (sub)species: from kingdom to subspecies
- The identifier of the parent taxons
- The year and first author of the related scientific publication

#### **Manual additions:**

For convenience, some entries were added manually:

• ~1 500 entries of Salmonella, such as the city-like serovars and groups A to H

• 36 species groups (such as the beta-haemolytic *Streptococcus* groups A to K, coagulase-negative *Staphylococcus* (CoNS), *Mycobacterium tuberculosis* complex, etc.), of which the group compositions are stored in the microorganisms.groups data set

- 1 entry of *Blastocystis* (*B. hominis*), although it officially does not exist (Noel *et al.* 2005, PMID 15634993)
- 1 entry of *Moraxella* (*M. catarrhalis*), which was formally named *Branhamella catarrhalis* (Catlin, 1970) though this change was never accepted within the field of clinical microbiology
- 8 other 'undefined' entries (unknown, unknown Gram-negatives, unknown Gram-positives, unknown yeast, unknown fungus, and unknown anaerobic Gram-pos/Gram-neg bacteria)

The syntax used to transform the original data to a cleansed R format, can be found here: https://github.com/msberends/AMR/blob/main/data-raw/reproduction\_of\_microorganisms.R.

#### Direct download:

Like all data sets in this package, this data set is publicly available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

#### About the Records from LPSN (see Source)

LPSN is the main source for bacteriological taxonomy of this AMR package.

The List of Prokaryotic names with Standing in Nomenclature (LPSN) provides comprehensive information on the nomenclature of prokaryotes. LPSN is a free to use service founded by Jean P. Euzeby in 1997 and later on maintained by Aidan C. Parte.

### Source

- Parte, AC *et al.* (2020). **List of Prokaryotic names with Standing in Nomenclature (LPSN) moves to the DSMZ.** International Journal of Systematic and Evolutionary Microbiology, 70, 5607-5612; doi:10.1099/ijsem.0.004332. Accessed from https://lpsn.dsmz.de on December 11th, 2022.
- GBIF Secretariat (2022). GBIF Backbone Taxonomy. Checklist dataset doi:10.15468/39omei. Accessed from https://www.gbif.org on December 11th, 2022.
- Reimer, LC *et al.* (2022). *BacDive* in 2022: the knowledge base for standardized bacterial and archaeal data. Nucleic Acids Res., 50(D1):D741-D74; doi:10.1093/nar/gkab961. Accessed from https://bacdive.dsmz.de on May 12th, 2023.
- Public Health Information Network Vocabulary Access and Distribution System (PHIN VADS).
   US Edition of SNOMED CT from 1 September 2020. Value Set Name 'Microorganism', OID 2.16.840.1.114222.4.11.1009 (v12). URL: https://phinvads.cdc.gov
- Grimont *et al.* (2007). Antigenic Formulae of the Salmonella Serovars, 9th Edition. WHO Collaborating Centre for Reference and Research on *Salmonella* (WHOCC-SALM).
- Bartlett et al. (2022). A comprehensive list of bacterial pathogens infecting humans Microbiology 168:001269; doi:10.1099/mic.0.001269

#### See Also

130 microorganisms.codes

### **Examples**

microorganisms

microorganisms.codes Data Set with 4 957 Common Microorganism Codes

#### **Description**

A data set containing commonly used codes for microorganisms, from laboratory systems and WHONET. Define your own with set\_mo\_source(). They will all be searched when using as.mo() and consequently all the mo\_\* functions.

# Usage

microorganisms.codes

### **Format**

A tibble with 4 957 observations and 2 variables:

- code
   Commonly used code of a microorganism. This is a unique identifier.
- mo
   ID of the microorganism in the microorganisms data set

### **Details**

Like all data sets in this package, this data set is publicly available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

#### See Also

```
as.mo() microorganisms
```

### **Examples**

```
microorganisms.codes
# 'ECO' or 'eco' is the WHONET code for E. coli:
microorganisms.codes[microorganisms.codes$code == "ECO", ]
# and therefore, 'eco' will be understood as E. coli in this package:
mo_info("eco")
# works for all AMR functions:
mo_is_intrinsic_resistant("eco", ab = "vancomycin")
```

microorganisms.groups 131

microorganisms.groups Data Set with 521 Microorganisms In Species Groups

### **Description**

A data set containing species groups and microbiological complexes, which are used in the clinical breakpoints table.

# Usage

```
microorganisms.groups
```

### **Format**

A tibble with 521 observations and 4 variables:

- mo\_group
   ID of the species group / microbiological complex
- mo
   ID of the microorganism belonging in the species group / microbiological complex
- mo\_group\_name
   Name of the species group / microbiological complex, as retrieved with mo\_name()
- mo\_name
   Name of the microorganism belonging in the species group / microbiological complex, as retrieved with mo\_name()

# Details

Like all data sets in this package, this data set is publicly available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

# See Also

```
as.mo() microorganisms
```

### **Examples**

```
microorganisms.groups
# these are all species in the Bacteroides fragilis group, as per WHONET:
microorganisms.groups[microorganisms.groups$mo_group == "B_BCTRD_FRGL-C", ]
```

mo\_matching\_score

mo\_matching\_score

Calculate the Matching Score for Microorganisms

## **Description**

This algorithm is used by as.mo() and all the mo\_\* functions to determine the most probable match of taxonomic records based on user input.

#### **Usage**

```
mo_matching_score(x, n)
```

# **Arguments**

x Any user input value(s)

n A full taxonomic name, that exists in microorganisms\$fullname

#### **Matching Score for Microorganisms**

With ambiguous user input in as.mo() and all the mo\_\* functions, the returned results are chosen based on their matching score using mo\_matching\_score(). This matching score m, is calculated as:

$$m_{(x,n)} = \frac{l_n - 0.5 \cdot \min \begin{cases} l_n \\ \text{lev}(x,n) \end{cases}}{l_n \cdot p_n \cdot k_n}$$

where:

- x is the user input;
- n is a taxonomic name (genus, species, and subspecies);
- $l_n$  is the length of n;
- lev is the Levenshtein distance function (counting any insertion as 1, and any deletion or substitution as 2) that is needed to change x into n;
- $p_n$  is the human pathogenic prevalence group of n, as described below;
- $k_n$  is the taxonomic kingdom of n, set as Bacteria = 1, Fungi = 1.25, Protozoa = 1.5, Archaea = 2, others = 3.

The grouping into human pathogenic prevalence p is based on recent work from Bartlett  $et \, al.$  (2022, doi:10.1099/mic.0.001269) who extensively studied medical-scientific literature to categorise all bacterial species into these groups:

- **Established**, if a taxonomic species has infected at least three persons in three or more references. These records have prevalence = 1.0 in the microorganisms data set;
- **Putative**, if a taxonomic species has fewer than three known cases. These records have prevalence = 1.25 in the microorganisms data set.

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#### Furthermore,

 Any genus present in the established list also has prevalence = 1.0 in the microorganisms data set;

- Any other genus present in the **putative** list has prevalence = 1.25 in the microorganisms data set:
- Any other species or subspecies of which the genus is present in the two aforementioned groups, has prevalence = 1.5 in the microorganisms data set;
- Any non-bacterial genus, species or subspecies of which the genus is present in the following list, has prevalence = 1.25 in the microorganisms data set: Absidia, Acanthamoeba, Acremonium, Aedes, Alternaria, Amoeba, Ancylostoma, Angiostrongylus, Anisakis, Anopheles, Apophysomyces, Aspergillus, Aureobasidium, Basidiobolus, Beauveria, Blastocystis, Blastomyces, Candida, Capillaria, Chaetomium, Chrysonilia, Cladophialophora, Cladosporium, Conidiobolus, Contracaecum, Cordylobia, Cryptococcus, Curvularia, Demodex, Dermatobia, Dientamoeba, Diphyllobothrium, Dirofilaria, Echinostoma, Entamoeba, Enterobius, Exophiala, Exserohilum, Fasciola, Fonsecaea, Fusarium, Giardia, Haloarcula, Halobacterium, Halococcus, Hendersonula, Heterophyes, Histomonas, Histoplasma, Hymenolepis, Hypomyces, Hysterothylacium, Leishmania, Malassezia, Malbranchea, Metagonimus, Meyerozyma, Microsporidium, Microsporum, Mortierella, Mucor, Mycocentrospora, Necator, Nectria, Ochroconis, Oesophagostomum, Oidiodendron, Opisthorchis, Pediculus, Penicillium, Phlebotomus, Phoma, Pichia, Piedraia, Pithomyces, Pityrosporum, Pneumocystis, Pseudallescheria, Pseudoterranova, Pulex, Rhizomucor, Rhizopus, Rhodotorula, Saccharomyces, Sarcoptes, Scolecobasidium, Scopulariopsis, Scytalidium, Spirometra, Sporobolomyces, Stachybotrys, Strongyloides, Syngamus, Taenia, Talaromyces, Toxocara, Trichinella, Trichobilharzia, Trichoderma, Trichomonas, Trichophyton, Trichosporon, Trichostrongylus, Trichuris, Tritirachium, Trombicula, Trypanosoma, Tunga, or Wuchereria;
- All other records have prevalence = 2.0 in the microorganisms data set.

When calculating the matching score, all characters in x and n are ignored that are other than A-Z, a-z, 0-9, spaces and parentheses.

All matches are sorted descending on their matching score and for all user input values, the top match will be returned. This will lead to the effect that e.g., "E. coli" will return the microbial ID of *Escherichia coli* (m=0.688, a highly prevalent microorganism found in humans) and not *Entamoeba coli* (m=0.381, a less prevalent microorganism in humans), although the latter would alphabetically come first.

#### Reference Data Publicly Available

All data sets in this AMR package (about microorganisms, antibiotics, SIR interpretation, EUCAST rules, etc.) are publicly and freely available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. We also provide tab-separated plain text files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

#### Note

This algorithm was originally described in: Berends MS et al. (2022). AMR: An R Package for Working with Antimicrobial Resistance Data. Journal of Statistical Software, 104(3), 1-31;

### doi:10.18637/jss.v104.i03.

Later, the work of Bartlett A *et al.* about bacterial pathogens infecting humans (2022, doi:10.1099/mic.0.001269) was incorporated.

# Author(s)

Dr. Matthijs Berends, 2018

# **Examples**

```
mo_reset_session()
as.mo("E. coli")
mo_uncertainties()

mo_matching_score(
    x = "E. coli",
    n = c("Escherichia coli", "Entamoeba coli")
)
```

mo\_property

Get Properties of a Microorganism

# **Description**

Use these functions to return a specific property of a microorganism based on the latest accepted taxonomy. All input values will be evaluated internally with as.mo(), which makes it possible to use microbial abbreviations, codes and names as input. See *Examples*.

# Usage

```
mo_name(
    x,
    language = get_AMR_locale(),
    keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
    ...
)

mo_fullname(
    x,
    language = get_AMR_locale(),
    keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
    ...
)

mo_shortname(
    x,
    language = get_AMR_locale(),
```

```
keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_subspecies(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_species(
 х,
  language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_genus(
  Χ,
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_family(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_order(
  language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_class(
 language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_phylum(
 х,
```

```
language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_kingdom(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_domain(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_type(
 Х,
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_status(
 х,
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_pathogenicity(
 Х,
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_gramstain(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_is_gram_negative(
```

```
Χ,
  language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_is_gram_positive(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_is_yeast(
  language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_is_intrinsic_resistant(
 Χ,
  ab,
  language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_oxygen_tolerance(
  language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_is_anaerobic(
 language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo\_snomed(
 language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
```

```
mo_ref(
  х,
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_authors(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_year(
 Х,
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_lpsn(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_gbif(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_rank(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_taxonomy(
 language = get_AMR_locale(),
 keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
```

```
)
mo_synonyms(
  language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_current(x, language = get_AMR_locale(), ...)
mo_info(
  х,
  language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_url(
  Х,
  open = FALSE,
  language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
mo_property(
  х,
  property = "fullname",
  language = get_AMR_locale(),
  keep_synonyms = getOption("AMR_keep_synonyms", FALSE),
)
```

# Arguments

keep\_synonyms

any character (vector) that can be coerced to a valid microorganism code with as.mo(). Can be left blank for auto-guessing the column containing microorganism codes if used in a data set, see *Examples*.

language language to translate text like "no growth", which defaults to the system language (see get\_AMR\_locale())

a logical to indicate if old, previously valid taxonomic names must be preserved and not be corrected to currently accepted names. The default is FALSE, which will return a note if old taxonomic names were processed. The default can be set with the package option AMR\_keep\_synonyms, i.e. options(AMR\_keep\_synonyms = TRUE) or options(AMR\_keep\_synonyms = FALSE).

... other arguments passed on to as.mo(), such as 'minimum\_matching\_score', 'ignore\_pattern', and 'remove\_from\_input'

ab any (vector of) text that can be coerced to a valid antibiotic drug code with as.ab()

open browse the URL using browseURL()

property one of the column names of the microorganisms data set: "mo", "fullname",

"status", "kingdom", "phylum", "class", "order", "family", "genus", "species", "subspecies", "rank", "ref", "oxygen\_tolerance", "source", "lpsn", "lpsn\_parent", "lpsn\_renamed\_to", "gbif", "gbif\_parent", "gbif\_renamed\_to", "prevalence", or

"snomed", or must be "shortname"

#### **Details**

All functions will, at default, **not** keep old taxonomic properties, as synonyms are automatically replaced with the current taxonomy. Take for example *Enterobacter aerogenes*, which was initially named in 1960 but renamed to *Klebsiella aerogenes* in 2017:

- mo\_genus("Enterobacter aerogenes") will return "Klebsiella" (with a note about the renaming)
- mo\_genus("Enterobacter aerogenes", keep\_synonyms = TRUE) will return "Enterobacter" (with a once-per-session warning that the name is outdated)
- mo\_ref("Enterobacter aerogenes") will return "Tindall et al., 2017" (with a note)
- mo\_ref("Enterobacter aerogenes", keep\_synonyms = TRUE) will return "Hormaeche et al., 1960" (with a warning)

The short name (mo\_shortname()) returns the first character of the genus and the full species, such as "E. coli", for species and subspecies. Exceptions are abbreviations of staphylococci (such as "CoNS", Coagulase-Negative Staphylococci) and beta-haemolytic streptococci (such as "GBS", Group B Streptococci). Please bear in mind that e.g. E. coli could mean Escherichia coli (kingdom of Bacteria) as well as Entamoeba coli (kingdom of Protozoa). Returning to the full name will be done using as.mo() internally, giving priority to bacteria and human pathogens, i.e. "E. coli" will be considered Escherichia coli. As a result, mo\_fullname(mo\_shortname("Entamoeba coli")) returns "Escherichia coli".

Since the top-level of the taxonomy is sometimes referred to as 'kingdom' and sometimes as 'domain', the functions mo\_kingdom() and mo\_domain() return the exact same results.

Determination of human pathogenicity (mo\_pathogenicity()) is strongly based on Bartlett *et al.* (2022, doi:10.1099/mic.0.001269). This function returns a factor with the levels *Pathogenic*, *Potentially pathogenic*, *Non-pathogenic*, and *Unknown*.

Determination of the Gram stain (mo\_gramstain()) will be based on the taxonomic kingdom and phylum. Originally, Cavalier-Smith defined the so-called subkingdoms Negibacteria and Posibacteria (2002, PMID 11837318), and only considered these phyla as Posibacteria: Actinobacteria, Chloroflexi, Firmicutes, and Tenericutes. These phyla were later renamed to Actinomycetota, Chloroflexota, Bacillota, and Mycoplasmatota (2021, PMID 34694987). Bacteria in these phyla are considered Gram-positive in this AMR package, except for members of the class Negativicutes (within phylum Bacillota) which are Gram-negative. All other bacteria are considered Gram-negative. Species outside the kingdom of Bacteria will return a value NA. Functions mo\_is\_gram\_negative() and mo\_is\_gram\_positive() always return TRUE or FALSE (or NA when the input is NA or the MO code is UNKNOWN), thus always return FALSE for species outside the taxonomic kingdom of Bacteria.

Determination of yeasts (mo\_is\_yeast()) will be based on the taxonomic kingdom and class. *Budding yeasts* are fungi of the phylum Ascomycota, class Saccharomycetes (also called Hemiascomycetes). *True yeasts* are aggregated into the underlying order Saccharomycetales. Thus, for all microorganisms that are member of the taxonomic class Saccharomycetes, the function will return TRUE. It returns FALSE otherwise (or NA when the input is NA or the MO code is UNKNOWN).

Determination of intrinsic resistance (mo\_is\_intrinsic\_resistant()) will be based on the intrinsic\_resistant data set, which is based on 'EUCAST Expert Rules' and 'EUCAST Intrinsic Resistance and Unusual Phenotypes' v3.3 (2021). The mo\_is\_intrinsic\_resistant() function can be vectorised over both argument x (input for microorganisms) and ab (input for antibiotics).

Determination of bacterial oxygen tolerance (mo\_oxygen\_tolerance()) will be based on Bac-Dive, see *Source*. The function mo\_is\_anaerobic() only returns TRUE if the oxygen tolerance is "anaerobe", indicting an obligate anaerobic species or genus. It always returns FALSE for species outside the taxonomic kingdom of Bacteria.

The function mo\_url() will return the direct URL to the online database entry, which also shows the scientific reference of the concerned species.

SNOMED codes (mo\_snomed()) are from the version of July 1st, 2021. See *Source* and the microorganisms data set for more info.

Old taxonomic names (so-called 'synonyms') can be retrieved with mo\_synonyms() (which will have the scientific reference as name), the current taxonomic name can be retrieved with mo\_current(). Both functions return full names.

All output will be translated where possible.

#### Value

- An integer in case of mo\_year()
- An ordered factor in case of mo\_pathogenicity()
- A list in case of mo\_taxonomy(), mo\_synonyms(), mo\_snomed() and mo\_info()
- A named character in case of mo\_url()
- A character in all other cases

### **Matching Score for Microorganisms**

This function uses as.mo() internally, which uses an advanced algorithm to translate arbitrary user input to valid taxonomy using a so-called matching score. You can read about this public algorithm on the MO matching score page.

#### Source

- 1. Berends MS *et al.* (2022). **AMR: An R Package for Working with Antimicrobial Resistance Data**. *Journal of Statistical Software*, 104(3), 1-31; doi:10.18637/jss.v104.i03
- 2. Becker K *et al.* (2014). **Coagulase-Negative Staphylococci.** *Clin Microbiol Rev.* 27(4): 870-926; doi:10.1128/CMR.0010913
- 3. Becker K et al. (2019). Implications of identifying the recently defined members of the S. aureus complex, S. argenteus and S. schweitzeri: A position paper of members of the ESCMID Study Group for staphylococci and Staphylococcal Diseases (ESGS). Clin Microbiol Infect; doi:10.1016/j.cmi.2019.02.028

4. Becker K *et al.* (2020). **Emergence of coagulase-negative staphylococci.** *Expert Rev Anti Infect Ther.* 18(4):349-366; doi:10.1080/14787210.2020.1730813

- 5. Lancefield RC (1933). A serological differentiation of human and other groups of hemolytic streptococci. *J Exp Med.* 57(4): 571-95; doi:10.1084/jem.57.4.571
- Berends MS et al. (2022). Trends in Occurrence and Phenotypic Resistance of Coagulase-Negative Staphylococci (CoNS) Found in Human Blood in the Northern Netherlands between 2013 and 2019/ Micro.rganisms 10(9), 1801; doi:10.3390/microorganisms10091801
- Parte, AC et al. (2020). List of Prokaryotic names with Standing in Nomenclature (LPSN) moves to the DSMZ. International Journal of Systematic and Evolutionary Microbiology, 70, 5607-5612; doi:10.1099/ijsem.0.004332. Accessed from https://lpsn.dsmz.de on December 11th, 2022.
- 8. GBIF Secretariat (2022). GBIF Backbone Taxonomy. Checklist dataset doi:10.15468/39omei. Accessed from https://www.gbif.org on December 11th, 2022.
- Reimer, LC et al. (2022). BacDive in 2022: the knowledge base for standardized bacterial and archaeal data. Nucleic Acids Res., 50(D1):D741-D74; doi:10.1093/nar/gkab961. Accessed from https://bacdive.dsmz.de on May 12th, 2023.
- 10. Public Health Information Network Vocabulary Access and Distribution System (PHIN VADS). US Edition of SNOMED CT from 1 September 2020. Value Set Name 'Microorganism', OID 2.16.840.1.114222.4.11.1009 (v12). URL: https://phinvads.cdc.gov
- 11. Bartlett A *et al.* (2022). **A comprehensive list of bacterial pathogens infecting humans** *Microbiology* 168:001269; doi:10.1099/mic.0.001269

### Reference Data Publicly Available

All data sets in this AMR package (about microorganisms, antibiotics, SIR interpretation, EUCAST rules, etc.) are publicly and freely available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. We also provide tab-separated plain text files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

#### See Also

Data set microorganisms

# **Examples**

```
# taxonomic tree -----
mo_kingdom("Klebsiella pneumoniae")
mo_phylum("Klebsiella pneumoniae")
mo_class("Klebsiella pneumoniae")
mo_order("Klebsiella pneumoniae")
mo_family("Klebsiella pneumoniae")
mo_genus("Klebsiella pneumoniae")
mo_species("Klebsiella pneumoniae")
mo_subspecies("Klebsiella pneumoniae")
```

```
# full names and short names ------
mo_name("Klebsiella pneumoniae")
mo_fullname("Klebsiella pneumoniae")
mo_shortname("Klebsiella pneumoniae")
# other properties ------
mo_pathogenicity("Klebsiella pneumoniae")
mo_gramstain("Klebsiella pneumoniae")
mo_snomed("Klebsiella pneumoniae")
mo_type("Klebsiella pneumoniae")
mo_rank("Klebsiella pneumoniae")
mo_url("Klebsiella pneumoniae")
mo_is_yeast(c("Candida", "Trichophyton", "Klebsiella"))
# scientific reference -------
mo_ref("Klebsiella aerogenes")
mo_authors("Klebsiella aerogenes")
mo_year("Klebsiella aerogenes")
mo_lpsn("Klebsiella aerogenes")
mo_gbif("Klebsiella aerogenes")
mo_synonyms("Klebsiella aerogenes")
# abbreviations known in the field ------
mo_genus("MRSA")
mo_species("MRSA")
mo_shortname("VISA")
mo_gramstain("VISA")
mo_genus("EHEC")
mo_species("EIEC")
mo_name("UPEC")
# known subspecies ------
mo_fullname("K. pneu rh")
mo_shortname("K. pneu rh")
# Becker classification, see ?as.mo ------
mo_fullname("Staph epidermidis")
mo_fullname("Staph epidermidis", Becker = TRUE)
mo_shortname("Staph epidermidis")
mo_shortname("Staph epidermidis", Becker = TRUE)
```

mo\_source

```
# Lancefield classification, see ?as.mo ------
mo_fullname("Strep agalactiae")
mo_fullname("Strep agalactiae", Lancefield = TRUE)
mo_shortname("Strep agalactiae")
mo_shortname("Strep agalactiae", Lancefield = TRUE)
# language support ------
mo_gramstain("Klebsiella pneumoniae", language = "de") # German
mo_gramstain("Klebsiella pneumoniae", language = "nl") # Dutch
mo_gramstain("Klebsiella pneumoniae", language = "es") # Spanish
mo_gramstain("Klebsiella pneumoniae", language = "el") # Greek
mo_gramstain("Klebsiella pneumoniae", language = "uk") # Ukrainian
# mo_type is equal to mo_kingdom, but mo_kingdom will remain untranslated
mo_kingdom("Klebsiella pneumoniae")
mo_type("Klebsiella pneumoniae")
mo_kingdom("Klebsiella pneumoniae", language = "zh") # Chinese, no effect
mo_type("Klebsiella pneumoniae", language = "zh") # Chinese, translated
mo_fullname("S. pyogenes", Lancefield = TRUE, language = "de")
mo_fullname("S. pyogenes", Lancefield = TRUE, language = "uk")
# gram stains and intrinsic resistance can be used as a filter in dplyr verbs
if (require("dplyr")) {
 example_isolates %>%
    filter(mo_is_gram_positive()) %>%
    count(mo_genus(), sort = TRUE)
}
if (require("dplyr")) {
 example_isolates %>%
    filter(mo_is_intrinsic_resistant(ab = "vanco")) %>%
    count(mo_genus(), sort = TRUE)
}
# get a list with the complete taxonomy (from kingdom to subspecies)
mo_taxonomy("Klebsiella pneumoniae")
# get a list with the taxonomy, the authors, Gram-stain,
# SNOMED codes, and URL to the online database
mo_info("Klebsiella pneumoniae")
```

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#### **Description**

These functions can be used to predefine your own reference to be used in as.mo() and consequently all mo\_\* functions (such as mo\_genus() and mo\_gramstain()).

This is **the fastest way** to have your organisation (or analysis) specific codes picked up and translated by this package, since you don't have to bother about it again after setting it up once.

#### Usage

```
set_mo_source(
  path,
  destination = getOption("AMR_mo_source", "~/mo_source.rds")
)
get_mo_source(destination = getOption("AMR_mo_source", "~/mo_source.rds"))
```

## **Arguments**

path location of your reference file, this can be any text file (comma-, tab- or pipe-

separated) or an Excel file (see Details). Can also be "", NULL or FALSE to delete

the reference file.

destination destination of the compressed data file - the default is the user's home directory.

#### **Details**

The reference file can be a text file separated with commas (CSV) or tabs or pipes, an Excel file (either 'xls' or 'xlsx' format) or an R object file (extension '.rds'). To use an Excel file, you will need to have the readxl package installed.

set\_mo\_source() will check the file for validity: it must be a data.frame, must have a column named "mo" which contains values from microorganisms\$mo or microorganisms\$fullname and must have a reference column with your own defined values. If all tests pass, set\_mo\_source() will read the file into R and will ask to export it to "~/mo\_source.rds". The CRAN policy disallows packages to write to the file system, although 'exceptions may be allowed in interactive sessions if the package obtains confirmation from the user'. For this reason, this function only works in interactive sessions so that the user can specifically confirm and allow that this file will be created. The destination of this file can be set with the destination argument and defaults to the user's home directory. It can also be set with the package option AMR\_mo\_source, e.g. options(AMR\_mo\_source = "my/location/file.rds").

The created compressed data file "mo\_source.rds" will be used at default for MO determination (function as.mo() and consequently all mo\_\* functions like mo\_genus() and mo\_gramstain()). The location and timestamp of the original file will be saved as an attribute to the compressed data file.

The function get\_mo\_source() will return the data set by reading "mo\_source.rds" with readRDS(). If the original file has changed (by checking the location and timestamp of the original file), it will call set\_mo\_source() to update the data file automatically if used in an interactive session.

Reading an Excel file (.xlsx) with only one row has a size of 8-9 kB. The compressed file created with set\_mo\_source() will then have a size of 0.1 kB and can be read by get\_mo\_source() in only a couple of microseconds (millionths of a second).

mo\_source

# **How to Setup**

Imagine this data on a sheet of an Excel file. The first column contains the organisation specific codes, the second column contains valid taxonomic names:

We save it as "home/me/ourcodes.xlsx". Now we have to set it as a source:

```
set_mo_source("home/me/ourcodes.xlsx")
#> NOTE: Created mo_source file '/Users/me/mo_source.rds' (0.3 kB) from
#> '/Users/me/Documents/ourcodes.xlsx' (9 kB), columns
#> "Organisation XYZ" and "mo"
```

It has now created a file "~/mo\_source.rds" with the contents of our Excel file. Only the first column with foreign values and the 'mo' column will be kept when creating the RDS file.

And now we can use it in our functions:

```
as.mo("lab_mo_ecoli")
#> Class 'mo'
#> [1] B_ESCHR_COLI

mo_genus("lab_mo_kpneumoniae")
#> [1] "Klebsiella"

# other input values still work too
as.mo(c("Escherichia coli", "E. coli", "lab_mo_ecoli"))
#> NOTE: Translation to one microorganism was guessed with uncertainty.
#> Use mo_uncertainties() to review it.
#> Class 'mo'
#> [1] B_ESCHR_COLI B_ESCHR_COLI B_ESCHR_COLI
```

If we edit the Excel file by, let's say, adding row 4 like this:

	1	Α	1	В	1
	- -		- -		-
1		Organisation XYZ		mo	1
2	1	lab_mo_ecoli		Escherichia coli	1
3	I	lab_mo_kpneumoniae	I	Klebsiella pneumoniae	1
4	I	lab_Staph_aureus	I	Staphylococcus aureus	1
5	١				1

...any new usage of an MO function in this package will update your data file:

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```
as.mo("lab_mo_ecoli")
#> NOTE: Updated mo_source file '/Users/me/mo_source.rds' (0.3 kB) from
#> '/Users/me/Documents/ourcodes.xlsx' (9 kB), columns
#> "Organisation XYZ" and "mo"
#> Class 'mo'
#> [1] B_ESCHR_COLI

mo_genus("lab_Staph_aureus")
#> [1] "Staphylococcus"

To delete the reference data file, just use "", NULL or FALSE as input for set_mo_source():
set_mo_source(NULL)
#> Removed mo_source file '/Users/me/mo_source.rds'
```

If the original file (in the previous case an Excel file) is moved or deleted, the mo\_source.rds file will be removed upon the next use of as.mo() or any mo\_\* function.

рса

Principal Component Analysis (for AMR)

# **Description**

Performs a principal component analysis (PCA) based on a data set with automatic determination for afterwards plotting the groups and labels, and automatic filtering on only suitable (i.e. non-empty and numeric) variables.

## Usage

```
pca(
    x,
    ...,
    retx = TRUE,
    center = TRUE,
    scale. = TRUE,
    tol = NULL,
    rank. = NULL
)
```

## **Arguments**

x a data.frame containing numeric columns

... columns of x to be selected for PCA, can be unquoted since it supports quasiquotation.

retx a logical value indicating whether the rotated variables should be returned.

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center a logical value indicating whether the variables should be shifted to be zero

centered. Alternately, a vector of length equal the number of columns of  $\boldsymbol{x}$  can

be supplied. The value is passed to scale.

scale. a logical value indicating whether the variables should be scaled to have unit

variance before the analysis takes place. The default is FALSE for consistency with S, but in general scaling is advisable. Alternatively, a vector of length equal the number of columns of y can be supplied. The value is passed to cool or

the number of columns of x can be supplied. The value is passed to scale.

a value indicating the magnitude below which components should be omitted. (Components are omitted if their standard deviations are less than or equal to tol times the standard deviation of the first component.) With the default null setting, no components are omitted (unless rank. is specified less than

min(dim(x)).). Other settings for tol could be tol = 0 or tol = sqrt(.Machine\$double.eps),

which would omit essentially constant components.

rank. optionally, a number specifying the maximal rank, i.e., maximal number of prin-

cipal components to be used. Can be set as alternative or in addition to to1, useful notably when the desired rank is considerably smaller than the dimensions

of the matrix.

#### **Details**

tol

The pca() function takes a data.frame as input and performs the actual PCA with the R function prcomp().

The result of the pca() function is a prcomp object, with an additional attribute non\_numeric\_cols which is a vector with the column names of all columns that do not contain numeric values. These are probably the groups and labels, and will be used by ggplot\_pca().

#### Value

An object of classes pca and prcomp

# **Examples**

```
# `example_isolates` is a data set available in the AMR package.
# See ?example_isolates.

if (require("dplyr")) {
    # calculate the resistance per group first
    resistance_data <- example_isolates %>%
        group_by(
        order = mo_order(mo), # group on anything, like order
        genus = mo_genus(mo)
    ) %>% # and genus as we do here;
    filter(n() >= 30) %>% # filter on only 30 results per group
    summarise_if(is.sir, resistance) # then get resistance of all drugs

# now conduct PCA for certain antimicrobial drugs
pca_result <- resistance_data %>%
    pca(AMC, CXM, CTX, CAZ, GEN, TOB, TMP, SXT)
```

```
pca_result
summary(pca_result)

# old base R plotting method:
biplot(pca_result)
# new ggplot2 plotting method using this package:
if (require("ggplot2")) {
   ggplot_pca(pca_result)

   ggplot_pca(pca_result) +
        scale_colour_viridis_d() +
        labs(title = "Title here")
}
```

plot

Plotting for Classes sir, mic and disk

# Description

Functions to plot classes sir, mic and disk, with support for base R and ggplot2.

# Usage

```
## S3 method for class 'mic'
plot(
  Х,
 mo = NULL,
 ab = NULL,
  guideline = "EUCAST",
 main = deparse(substitute(x)),
 ylab = translate_AMR("Frequency", language = language),
 xlab = translate_AMR("Minimum Inhibitory Concentration (mg/L)", language = language),
  colours_SIR = c("#3CAEA3", "#F6D55C", "#ED553B"),
  language = get_AMR_locale(),
  expand = TRUE,
  include_PKPD = getOption("AMR_include_PKPD", TRUE),
  breakpoint_type = getOption("AMR_breakpoint_type", "human"),
)
## S3 method for class 'mic'
autoplot(
  object,
 mo = NULL,
  ab = NULL,
```

```
guideline = "EUCAST",
  title = deparse(substitute(object)),
 ylab = translate_AMR("Frequency", language = language),
 xlab = translate_AMR("Minimum Inhibitory Concentration (mg/L)", language = language),
  colours_SIR = c("#3CAEA3", "#F6D55C", "#ED553B"),
 language = get_AMR_locale(),
  expand = TRUE,
  include_PKPD = getOption("AMR_include_PKPD", TRUE),
 breakpoint_type = getOption("AMR_breakpoint_type", "human"),
)
## S3 method for class 'mic'
fortify(object, ...)
## S3 method for class 'disk'
plot(
  Х,
 main = deparse(substitute(x)),
 ylab = translate_AMR("Frequency", language = language),
 xlab = translate_AMR("Disk diffusion diameter (mm)", language = language),
 mo = NULL,
 ab = NULL,
  guideline = "EUCAST",
  colours_SIR = c("#3CAEA3", "#F6D55C", "#ED553B"),
  language = get_AMR_locale(),
  expand = TRUE,
  include_PKPD = getOption("AMR_include_PKPD", TRUE),
 breakpoint_type = getOption("AMR_breakpoint_type", "human"),
)
## S3 method for class 'disk'
autoplot(
 object,
 mo = NULL,
  ab = NULL,
  title = deparse(substitute(object)),
  ylab = translate_AMR("Frequency", language = language),
 xlab = translate_AMR("Disk diffusion diameter (mm)", language = language),
  guideline = "EUCAST",
  colours_SIR = c("#3CAEA3", "#F6D55C", "#ED553B"),
 language = get_AMR_locale(),
  expand = TRUE,
  include_PKPD = getOption("AMR_include_PKPD", TRUE),
 breakpoint_type = getOption("AMR_breakpoint_type", "human"),
)
```

```
## S3 method for class 'disk'
fortify(object, ...)
## S3 method for class 'sir'
plot(
 х,
 ylab = translate_AMR("Percentage", language = language),
 xlab = translate_AMR("Antimicrobial Interpretation", language = language),
 main = deparse(substitute(x)),
 language = get_AMR_locale(),
)
## S3 method for class 'sir'
autoplot(
 object,
  title = deparse(substitute(object)),
 xlab = translate_AMR("Antimicrobial Interpretation", language = language),
 ylab = translate_AMR("Frequency", language = language),
 colours_SIR = c("#3CAEA3", "#F6D55C", "#ED553B"),
 language = get_AMR_locale(),
)
## S3 method for class 'sir'
fortify(object, ...)
```

## **Arguments**

x, object	<pre>values created with as.mic(), as.disk() or as.sir() (or their random_* vari- ants, such as random_mic())</pre>
mo	any (vector of) text that can be coerced to a valid microorganism code with as.mo()
ab	any (vector of) text that can be coerced to a valid antimicrobial drug code with as.ab()
guideline	interpretation guideline to use - the default is the latest included EUCAST guideline, see <i>Details</i>
main, title	title of the plot
xlab, ylab	axis title
colours_SIR	colours to use for filling in the bars, must be a vector of three values (in the order S, I and R). The default colours are colour-blind friendly.
language	language to be used to translate 'Susceptible', 'Increased exposure'/'Intermediate' and 'Resistant' - the default is system language (see <pre>get_AMR_locale()</pre> ) and can be overwritten by setting the <pre>package</pre> option AMR_locale, e.g. options(AMR_locale = "de"), see <pre>translate</pre> . Use language = NULL or language = "" to prevent translation.

expand a logical to indicate whether the range on the x axis should be expanded between

the lowest and highest value. For MIC values, intermediate values will be factors of 2 starting from the highest MIC value. For disk diameters, the whole diameter

range will be filled.

include\_PKPD a logical to indicate that PK/PD clinical breakpoints must be applied as a last re-

sort - the default is TRUE. Can also be set with the package option AMR\_include\_PKPD.

breakpoint\_type

the type of breakpoints to use, either "ECOFF", "animal", or "human". ECOFF stands for Epidemiological Cut-Off values. The default is "human", which can

also be set with the package option AMR\_breakpoint\_type.

... arguments passed on to methods

#### **Details**

The interpretation of "I" will be named "Increased exposure" for all EUCAST guidelines since 2019, and will be named "Intermediate" in all other cases.

For interpreting MIC values as well as disk diffusion diameters, supported guidelines to be used as input for the guideline argument are: "EUCAST 2023", "EUCAST 2022", "EUCAST 2021", "EUCAST 2020", "EUCAST 2019", "EUCAST 2018", "EUCAST 2017", "EUCAST 2016", "EUCAST 2015", "EUCAST 2014", "EUCAST 2013", "EUCAST 2012", "EUCAST 2011", "CLSI 2023", "CLSI 2022", "CLSI 2021", "CLSI 2020", "CLSI 2019", "CLSI 2018", "CLSI 2017", "CLSI 2016", "CLSI 2015", "CLSI 2014", "CLSI 2013", "CLSI 2012", and "CLSI 2011".

Simply using "CLSI" or "EUCAST" as input will automatically select the latest version of that guideline.

#### Value

The autoplot() functions return a ggplot model that is extendible with any ggplot2 function.

The fortify() functions return a data.frame as an extension for usage in the ggplot2::ggplot() function.

## **Examples**

```
some_mic_values <- random_mic(size = 100)
some_disk_values <- random_disk(size = 100, mo = "Escherichia coli", ab = "cipro")
some_sir_values <- random_sir(50, prob_SIR = c(0.55, 0.05, 0.30))

plot(some_mic_values)
plot(some_disk_values)
plot(some_sir_values)

# when providing the microorganism and antibiotic, colours will show interpretations:
plot(some_mic_values, mo = "S. aureus", ab = "ampicillin")
plot(some_disk_values, mo = "Escherichia coli", ab = "cipro")
plot(some_disk_values, mo = "Escherichia coli", ab = "cipro", language = "nl")

if (require("ggplot2")) {
   autoplot(some_mic_values)</pre>
```

```
}
if (require("ggplot2")) {
   autoplot(some_disk_values, mo = "Escherichia coli", ab = "cipro")
}
if (require("ggplot2")) {
   autoplot(some_sir_values)
}
```

proportion

Calculate Antimicrobial Resistance

# **Description**

These functions can be used to calculate the (co-)resistance or susceptibility of microbial isolates (i.e. percentage of S, SI, I, IR or R). All functions support quasiquotation with pipes, can be used in summarise() from the dplyr package and also support grouped variables, see *Examples*.

resistance() should be used to calculate resistance, susceptibility() should be used to calculate susceptibility.

# Usage

```
resistance(..., minimum = 30, as_percent = FALSE, only_all_tested = FALSE)
susceptibility(..., minimum = 30, as_percent = FALSE, only_all_tested = FALSE)
sir_confidence_interval(
 ab_result = "R",
 minimum = 30,
 as_percent = FALSE,
 only_all_tested = FALSE,
 confidence_level = 0.95,
 side = "both",
 collapse = FALSE
)
proportion_R(..., minimum = 30, as_percent = FALSE, only_all_tested = FALSE)
proportion_IR(..., minimum = 30, as_percent = FALSE, only_all_tested = FALSE)
proportion_I(..., minimum = 30, as_percent = FALSE, only_all_tested = FALSE)
proportion_SI(..., minimum = 30, as_percent = FALSE, only_all_tested = FALSE)
proportion_S(..., minimum = 30, as_percent = FALSE, only_all_tested = FALSE)
```

```
proportion_df(
  data,
  translate_ab = "name",
  language = get_AMR_locale(),
 minimum = 30,
  as_percent = FALSE,
  combine_SI = TRUE,
  confidence_level = 0.95
)
sir_df(
  data,
  translate_ab = "name",
  language = get_AMR_locale(),
 minimum = 30,
  as_percent = FALSE,
  combine_SI = TRUE,
  confidence_level = 0.95
)
```

#### **Arguments**

. . .

one or more vectors (or columns) with antibiotic interpretations. They will be transformed internally with as.sir() if needed. Use multiple columns to calculate (the lack of) co-resistance: the probability where one of two drugs have a resistant or susceptible result. See *Examples*.

minimum

the minimum allowed number of available (tested) isolates. Any isolate count lower than minimum will return NA with a warning. The default number of 30 isolates is advised by the Clinical and Laboratory Standards Institute (CLSI) as best practice, see *Source*.

as\_percent

a logical to indicate whether the output must be returned as a hundred fold with % sign (a character). A value of 0.123456 will then be returned as "12.3%".

only\_all\_tested

(for combination therapies, i.e. using more than one variable for . . . ): a logical to indicate that isolates must be tested for all antibiotics, see section *Combination Therapy* below

ab\_result

antibiotic results to test against, must be one or more values of "S", "I", or "R"

confidence\_level

the confidence level for the returned confidence interval. For the calculation, the number of S or SI isolates, and R isolates are compared with the total number of available isolates with R, S, or I by using binom.test(), i.e., the Clopper-Pearson method.

side

the side of the confidence interval to return. The default is "both" for a length 2 vector, but can also be (abbreviated as) "min"/"left"/"lower"/"less" or "max"/"right"/"higher"/"greater".

collapse	a logical to indicate whether the output values should be 'collapsed', i.e. be merged together into one value, or a character value to use for collapsing
data	a data.frame containing columns with class sir (see as.sir())
translate_ab	a column name of the antibiotics data set to translate the antibiotic abbreviations to, using ab_property()
language	language of the returned text - the default is the current system language (see <pre>get_AMR_locale()</pre> ) and can also be set with the <pre>package</pre> option AMR_locale. Use language = NULL or language = "" to prevent translation.
combine_SI	a logical to indicate whether all values of S and I must be merged into one, so the output only consists of S+I vs. R (susceptible vs. resistant) - the default is TRUE

#### **Details**

Remember that you should filter your data to let it contain only first isolates! This is needed to exclude duplicates and to reduce selection bias. Use first\_isolate() to determine them in your data set with one of the four available algorithms.

The function resistance() is equal to the function proportion\_R(). The function susceptibility() is equal to the function proportion\_SI().

Use sir\_confidence\_interval() to calculate the confidence interval, which relies on binom.test(), i.e., the Clopper-Pearson method. This function returns a vector of length 2 at default for antimicrobial resistance. Change the side argument to "left"/"min" or "right"/"max" to return a single value, and change the ab\_result argument to e.g. c("S", "I") to test for antimicrobial susceptibility, see Examples.

These functions are not meant to count isolates, but to calculate the proportion of resistance/susceptibility. Use the count\_\*() functions to count isolates. The function susceptibility() is essentially equal to count\_susceptible()/count\_all(). Low counts can influence the outcome - the proportion\_\*() functions may camouflage this, since they only return the proportion (albeit dependent on the minimum argument).

The function proportion\_df() takes any variable from data that has an sir class (created with as.sir()) and calculates the proportions S, I, and R. It also supports grouped variables. The function sir\_df() works exactly like proportion\_df(), but adds the number of isolates.

# Value

A double or, when as\_percent = TRUE, a character.

# **Combination Therapy**

When using more than one variable for . . . (= combination therapy), use only\_all\_tested to only count isolates that are tested for all antibiotics/variables that you test them for. See this example for two antibiotics, Drug A and Drug B, about how susceptibility() works to calculate the %SI:

```
only_all_tested = FALSE only_all_tested = TRUE

Drug A Drug B include as include as include as
```

proportion proportion

		numerator	denominator	numerator	denominator
S or I	S or I	Χ	Χ	Χ	Χ
R	S or I	Χ	Χ	Χ	Χ
<na></na>	S or I	Χ	Χ	-	-
S or I	R	Χ	Χ	Χ	Χ
R	R	-	Χ	-	Χ
<na></na>	R	-	-	-	-
S or I	<na></na>	Χ	Χ	-	-
R	<na></na>	-	-	-	-
<na></na>	<na></na>	<b>-</b>	<b>-</b>	<b>-</b>	<b>-</b>

Please note that, in combination therapies, for only\_all\_tested = TRUE applies that:

```
count_S() + count_I() + count_R() = count_all()
proportion_S() + proportion_I() + proportion_R() = 1
```

and that, in combination therapies, for only\_all\_tested = FALSE applies that:

```
count_S() + count_I() + count_R() >= count_all()
proportion_S() + proportion_I() + proportion_R() >= 1
```

Using only\_all\_tested has no impact when only using one antibiotic as input.

## **Interpretation of SIR**

In 2019, the European Committee on Antimicrobial Susceptibility Testing (EUCAST) has decided to change the definitions of susceptibility testing categories S, I, and R as shown below (https://www.eucast.org/newsiandr):

## • S - Susceptible, standard dosing regimen

A microorganism is categorised as "Susceptible, standard dosing regimen", when there is a high likelihood of therapeutic success using a standard dosing regimen of the agent.

# • I - Susceptible, increased exposure

A microorganism is categorised as "Susceptible, Increased exposure" when there is a high likelihood of therapeutic success because exposure to the agent is increased by adjusting the dosing regimen or by its concentration at the site of infection.

#### • R = Resistant

A microorganism is categorised as "Resistant" when there is a high likelihood of therapeutic failure even when there is increased exposure.

- *Exposure* is a function of how the mode of administration, dose, dosing interval, infusion time, as well as distribution and excretion of the antimicrobial agent will influence the infecting organism at the site of infection.

This AMR package honours this insight. Use susceptibility() (equal to proportion\_SI()) to determine antimicrobial susceptibility and count\_susceptible() (equal to count\_SI()) to count susceptible isolates.

## Source

M39 Analysis and Presentation of Cumulative Antimicrobial Susceptibility Test Data, 5th Edition, 2022, Clinical and Laboratory Standards Institute (CLSI). https://clsi.org/standards/products/microbiology/documents/m39/.

#### See Also

count() to count resistant and susceptible isolates.

## **Examples**

```
# example_isolates is a data set available in the AMR package.
# run ?example_isolates for more info.
example_isolates
# base R ------
# determines %R
resistance(example_isolates$AMX)
sir_confidence_interval(example_isolates$AMX)
sir_confidence_interval(example_isolates$AMX,
 confidence_level = 0.975
sir_confidence_interval(example_isolates$AMX,
 confidence_level = 0.975,
 collapse = ", "
)
# determines %S+I:
susceptibility(example_isolates$AMX)
sir_confidence_interval(example_isolates$AMX,
 ab_result = c("S", "I")
# be more specific
proportion_S(example_isolates$AMX)
proportion_SI(example_isolates$AMX)
proportion_I(example_isolates$AMX)
proportion_IR(example_isolates$AMX)
proportion_R(example_isolates$AMX)
# dplyr -----
if (require("dplyr")) {
 example_isolates %>%
   group_by(ward) %>%
   summarise(
     r = resistance(CIP),
     n = n_sir(CIP)
   ) # n_sir works like n_distinct in dplyr, see ?n_sir
if (require("dplyr")) {
```

proportion proportion

```
example_isolates %>%
   group_by(ward) %>%
   summarise(
     cipro_R = resistance(CIP),
     ci_min = sir_confidence_interval(CIP, side = "min"),
     ci_max = sir_confidence_interval(CIP, side = "max"),
   )
}
if (require("dplyr")) {
 # scoped dplyr verbs with antibiotic selectors
 # (you could also use across() of course)
 example_isolates %>%
   group_by(ward) %>%
    summarise_at(
     c(aminoglycosides(), carbapenems()),
     resistance
   )
}
if (require("dplyr")) {
 example_isolates %>%
   group_by(ward) %>%
   summarise(
     R = resistance(CIP, as_percent = TRUE),
     SI = susceptibility(CIP, as_percent = TRUE),
     n1 = count_all(CIP), # the actual total; sum of all three
     n2 = n_sir(CIP), # same - analogous to n_distinct
      total = n()
   ) # NOT the number of tested isolates!
 # Calculate co-resistance between amoxicillin/clav acid and gentamicin,
 # so we can see that combination therapy does a lot more than mono therapy:
 example_isolates %>% susceptibility(AMC) # %SI = 76.3%
 example_isolates %>% count_all(AMC) #  n = 1879
 example_isolates %>% susceptibility(GEN) # %SI = 75.4%
 example_isolates %>% count_all(GEN) # n = 1855
 example_isolates %>% susceptibility(AMC, GEN) # %SI = 94.1%
 example_isolates %>% count_all(AMC, GEN) # n = 1939
 # See Details on how `only_all_tested` works. Example:
 example_isolates %>%
    summarise(
     numerator = count_susceptible(AMC, GEN),
      denominator = count_all(AMC, GEN),
     proportion = susceptibility(AMC, GEN)
   )
 example_isolates %>%
    summarise(
     numerator = count_susceptible(AMC, GEN, only_all_tested = TRUE),
      denominator = count_all(AMC, GEN, only_all_tested = TRUE),
```

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```
proportion = susceptibility(AMC, GEN, only_all_tested = TRUE)
   )
 example_isolates %>%
   group_by(ward) %>%
   summarise(
     cipro_p = susceptibility(CIP, as_percent = TRUE),
     cipro_n = count_all(CIP),
     genta_p = susceptibility(GEN, as_percent = TRUE),
     genta_n = count_all(GEN),
     combination_p = susceptibility(CIP, GEN, as_percent = TRUE),
      combination_n = count_all(CIP, GEN)
 # Get proportions S/I/R immediately of all sir columns
 example_isolates %>%
    select(AMX, CIP) %>%
   proportion_df(translate = FALSE)
 # It also supports grouping variables
 # (use sir_df to also include the count)
 example_isolates %>%
    select(ward, AMX, CIP) %>%
   group_by(ward) %>%
   sir_df(translate = FALSE)
}
```

random

Random MIC Values/Disk Zones/SIR Generation

# **Description**

These functions can be used for generating random MIC values and disk diffusion diameters, for AMR data analysis practice. By providing a microorganism and antimicrobial drug, the generated results will reflect reality as much as possible.

## Usage

```
random_mic(size = NULL, mo = NULL, ab = NULL, ...)
random_disk(size = NULL, mo = NULL, ab = NULL, ...)
random_sir(size = NULL, prob_SIR = c(0.33, 0.33, 0.33), ...)
```

#### **Arguments**

size

desired size of the returned vector. If used in a data.frame call or dplyr verb, will get the current (group) size if left blank.

mo	any character that can be coerced to a valid microorganism code with as.mo()
ab	any character that can be coerced to a valid antimicrobial drug code with as.ab()
	ignored, only in place to allow future extensions
prob_SIR	a vector of length 3: the probabilities for "S" (1st value), "I" (2nd value) and "R" (3rd value)

#### **Details**

The base R function sample() is used for generating values.

Generated values are based on the EUCAST 2023 guideline as implemented in the clinical\_breakpoints data set. To create specific generated values per bug or drug, set the mo and/or ab argument.

## Value

```
class mic for random_mic() (see as.mic()) and class disk for random_disk() (see as.disk())
```

# **Examples**

```
random_mic(25)
random_disk(25)
random_sir(25)

# make the random generation more realistic by setting a bug and/or drug:
random_mic(25, "Klebsiella pneumoniae") # range 0.0625-64
random_mic(25, "Klebsiella pneumoniae", "meropenem") # range 0.0625-16
random_mic(25, "Streptococcus pneumoniae", "meropenem") # range 0.0625-4

random_disk(25, "Klebsiella pneumoniae") # range 8-50
random_disk(25, "Klebsiella pneumoniae", "ampicillin") # range 11-17
random_disk(25, "Streptococcus pneumoniae", "ampicillin") # range 12-27
```

resistance\_predict

Predict Antimicrobial Resistance

# **Description**

Create a prediction model to predict antimicrobial resistance for the next years on statistical solid ground. Standard errors (SE) will be returned as columns se\_min and se\_max. See *Examples* for a real live example.

## Usage

```
resistance_predict(
  х,
  col_ab,
  col_date = NULL,
 year_min = NULL,
 year_max = NULL,
 year_every = 1,
 minimum = 30,
 model = NULL,
 I_as_S = TRUE,
 preserve_measurements = TRUE,
  info = interactive(),
)
sir_predict(
 Х,
  col_ab,
  col_date = NULL,
 year_min = NULL,
 year_max = NULL,
 year_every = 1,
 minimum = 30,
 model = NULL,
  I_as_S = TRUE,
  preserve_measurements = TRUE,
  info = interactive(),
)
## S3 method for class 'resistance_predict'
plot(x, main = paste("Resistance Prediction of", x_name), ...)
ggplot_sir_predict(
 х,
 main = paste("Resistance Prediction of", x_name),
 ribbon = TRUE,
  . . .
)
## S3 method for class 'resistance_predict'
autoplot(
 object,
 main = paste("Resistance Prediction of", x_name),
 ribbon = TRUE,
)
```

# Arguments

Х	a data.frame containing isolates. Can be left blank for automatic determination, see <i>Examples</i> .
col_ab	column name of $x$ containing antimicrobial interpretations ("R", "I" and "S")
col_date	column name of the date, will be used to calculate years if this column doesn't consist of years already - the default is the first column of with a date class
year_min	lowest year to use in the prediction model, dafaults to the lowest year in col_date
year_max	highest year to use in the prediction model - the default is 10 years after today
year_every	unit of sequence between lowest year found in the data and year_max
minimum	minimal amount of available isolates per year to include. Years containing less observations will be estimated by the model.
model	the statistical model of choice. This could be a generalised linear regression model with binomial distribution (i.e. using glm(, family = binomial), assuming that a period of zero resistance was followed by a period of increasing resistance leading slowly to more and more resistance. See <i>Details</i> for all valid options.
I_as_S	a logical to indicate whether values "I" should be treated as "S" (will otherwise be treated as "R"). The default, TRUE, follows the redefinition by EUCAST about the interpretation of I (increased exposure) in 2019, see section <i>Interpretation of S, I and R</i> below.
preserve_measu	
	a <u>logical</u> to indicate whether predictions of years that are actually available in the data should be overwritten by the original data. The standard errors of those years will be NA.
info	a logical to indicate whether textual analysis should be printed with the name and summary() of the statistical model.
	arguments passed on to functions
main	title of the plot
ribbon	a logical to indicate whether a ribbon should be shown (default) or error bars
object	model data to be plotted

# **Details**

Valid options for the statistical model (argument model) are:

- "binomial" or "binom" or "logit": a generalised linear regression model with binomial distribution
- "loglin" or "poisson": a generalised log-linear regression model with poisson distribution
- "lin" or "linear": a linear regression model

#### Value

A data.frame with extra class resistance\_predict with columns:

- year
- value, the same as estimated when preserve\_measurements = FALSE, and a combination of observed and estimated otherwise
- se\_min, the lower bound of the standard error with a minimum of 0 (so the standard error will never go below 0%)
- se\_max the upper bound of the standard error with a maximum of 1 (so the standard error will never go above 100%)
- observations, the total number of available observations in that year, i.e. S + I + R
- observed, the original observed resistant percentages
- estimated, the estimated resistant percentages, calculated by the model

Furthermore, the model itself is available as an attribute: attributes(x)\$model, see Examples.

# **Interpretation of SIR**

In 2019, the European Committee on Antimicrobial Susceptibility Testing (EUCAST) has decided to change the definitions of susceptibility testing categories S, I, and R as shown below (https://www.eucast.org/newsiandr):

## • S - Susceptible, standard dosing regimen

A microorganism is categorised as "Susceptible, standard dosing regimen", when there is a high likelihood of therapeutic success using a standard dosing regimen of the agent.

## • I - Susceptible, increased exposure

A microorganism is categorised as "Susceptible, Increased exposure" when there is a high likelihood of therapeutic success because exposure to the agent is increased by adjusting the dosing regimen or by its concentration at the site of infection.

#### • R = Resistant

A microorganism is categorised as "Resistant" when there is a high likelihood of therapeutic failure even when there is increased exposure.

Exposure is a function of how the mode of administration, dose, dosing interval, infusion time, as well as distribution and excretion of the antimicrobial agent will influence the infecting organism at the site of infection.

This AMR package honours this insight. Use susceptibility() (equal to proportion\_SI()) to determine antimicrobial susceptibility and count\_susceptible() (equal to count\_SI()) to count susceptible isolates.

## See Also

The proportion() functions to calculate resistance

Models: lm() glm()

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## **Examples**

```
x <- resistance_predict(example_isolates,</pre>
 col_ab = "AMX",
  year_min = 2010,
  model = "binomial"
)
plot(x)
if (require("ggplot2")) {
  ggplot_sir_predict(x)
}
# using dplyr:
if (require("dplyr")) {
  x <- example_isolates %>%
    filter_first_isolate() %>%
    filter(mo_genus(mo) == "Staphylococcus") %>%
    resistance_predict("PEN", model = "binomial")
  print(plot(x))
  # get the model from the object
  mymodel <- attributes(x)$model</pre>
  summary(mymodel)
}
# create nice plots with ggplot2 yourself
if (require("dplyr") && require("ggplot2")) {
  data <- example_isolates %>%
    filter(mo == as.mo("E. coli")) %>%
    resistance_predict(
      col_ab = "AMX",
      col_date = "date"
      model = "binomial",
      info = FALSE,
      minimum = 15
   )
  head(data)
  autoplot(data)
}
```

skewness

Skewness of the Sample

# **Description**

Skewness is a measure of the asymmetry of the probability distribution of a real-valued random variable about its mean.

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When negative ('left-skewed'): the left tail is longer; the mass of the distribution is concentrated on the right of a histogram. When positive ('right-skewed'): the right tail is longer; the mass of the distribution is concentrated on the left of a histogram. A normal distribution has a skewness of 0.

# Usage

```
skewness(x, na.rm = FALSE)

## Default S3 method:
skewness(x, na.rm = FALSE)

## S3 method for class 'matrix'
skewness(x, na.rm = FALSE)

## S3 method for class 'data.frame'
skewness(x, na.rm = FALSE)
```

## **Arguments**

x a vector of values, a matrix or a data.frame

na.rm a logical value indicating whether NA values should be stripped before the com-

putation proceeds

## See Also

kurtosis()

# **Examples**

```
skewness(runif(1000))
```

translate

Translate Strings from the AMR Package

# **Description**

For language-dependent output of AMR functions, such as mo\_name(), mo\_gramstain(), mo\_type() and ab\_name().

# Usage

```
get_AMR_locale()
set_AMR_locale(language)
reset_AMR_locale()
translate_AMR(x, language = get_AMR_locale())
```

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## Arguments

language language to choose. Use one of these supported language names or ISO-639-1

codes: English (en), Chinese (zh), Czech (cs), Danish (da), Dutch (nl), Finnish (fi), French (fr), German (de), Greek (el), Italian (it), Japanese (ja), Norwegian (no), Polish (pl), Portuguese (pt), Romanian (ro), Russian (ru), Spanish (es),

Swedish (sv), Turkish (tr), or Ukrainian (uk).

x text to translate

#### **Details**

The currently 20 supported languages are English (en), Chinese (zh), Czech (cs), Danish (da), Dutch (nl), Finnish (fi), French (fr), German (de), Greek (el), Italian (it), Japanese (ja), Norwegian (no), Polish (pl), Portuguese (pt), Romanian (ro), Russian (ru), Spanish (es), Swedish (sv), Turkish (tr), and Ukrainian (uk). All these languages have translations available for all antimicrobial drugs and colloquial microorganism names.

To permanently silence the once-per-session language note on a non-English operating system, you can set the package option AMR\_locale in your .Rprofile file like this:

```
# Open .Rprofile file
utils::file.edit("~/.Rprofile")

# Then add e.g. Italian support to that file using:
options(AMR_locale = "Italian")
```

And then save the file.

Please read about adding or updating a language in our Wiki.

## **Changing the Default Language:**

The system language will be used at default (as returned by Sys.getenv("LANG") or, if LANG is not set, Sys.getlocale("LC\_COLLATE")), if that language is supported. But the language to be used can be overwritten in two ways and will be checked in this order:

- Setting the package option AMR\_locale, either by using e.g. set\_AMR\_locale("German")
  or by running e.g. options(AMR\_locale = "German").
  Note that setting an R option only works in the same session. Save the command options(AMR\_locale = "(your language)") to your .Rprofile file to apply it for every session. Run utils::file.edit("~/.Rprofile" to edit your .Rprofile file.
- 2. Setting the system variable LANGUAGE or LANG, e.g. by adding LANGUAGE="de\_DE.utf8" to your .Renviron file in your home directory.

Thus, if the package option AMR\_locale is set, the system variables LANGUAGE and LANG will be ignored.

# **Examples**

```
# Current settings (based on system language)
ab_name("Ciprofloxacin")
mo_name("Coagulase-negative Staphylococcus (CoNS)")
```

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```
# setting another language
set_AMR_locale("Dutch")
ab_name("Ciprofloxacin")
mo_name("Coagulase-negative Staphylococcus (CoNS)")
# setting yet another language
set_AMR_locale("German")
ab_name("Ciprofloxacin")
mo_name("Coagulase-negative Staphylococcus (CoNS)")
# set_AMR_locale() understands endonyms, English exonyms, and ISO-639-1:
set_AMR_locale("Deutsch")
set_AMR_locale("German")
set_AMR_locale("de")
ab_name("amox/clav")
# reset to system default
reset_AMR_locale()
ab_name("amox/clav")
```

WHOCC

WHOCC: WHO Collaborating Centre for Drug Statistics Methodology

## Description

All antimicrobial drugs and their official names, ATC codes, ATC groups and defined daily dose (DDD) are included in this package, using the WHO Collaborating Centre for Drug Statistics Methodology.

## WHOCC

This package contains **all ~550 antibiotic**, **antimycotic and antiviral drugs** and their Anatomical Therapeutic Chemical (ATC) codes, ATC groups and Defined Daily Dose (DDD) from the World Health Organization Collaborating Centre for Drug Statistics Methodology (WHOCC, https://www.whocc.no) and the Pharmaceuticals Community Register of the European Commission (https://ec.europa.eu/health/documents/community-register/html/reg\_hum\_atc.htm).

These have become the gold standard for international drug utilisation monitoring and research.

The WHOCC is located in Oslo at the Norwegian Institute of Public Health and funded by the Norwegian government. The European Commission is the executive of the European Union and promotes its general interest.

NOTE: The WHOCC copyright does not allow use for commercial purposes, unlike any other info from this package. See https://www.whocc.no/copyright\_disclaimer/.

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# **Examples**

```
as.ab("meropenem")
ab_name("J01DH02")
ab_tradenames("flucloxacillin")
```

WHONET

Data Set with 500 Isolates - WHONET Example

## **Description**

This example data set has the exact same structure as an export file from WHONET. Such files can be used with this package, as this example data set shows. The antibiotic results are from our example\_isolates data set. All patient names were created using online surname generators and are only in place for practice purposes.

## Usage

WHONET

#### **Format**

A tibble with 500 observations and 53 variables:

- Identification number ID of the sample
- Specimen number ID of the specimen
- Organism

Name of the microorganism. Before analysis, you should transform this to a valid microbial class, using as.mo().

- Country Country of origin
- Laboratory
  Name of laboratory
- Last name

Fictitious last name of patient

• First name

Fictitious initial of patient

• Sex

Fictitious gender of patient

• Age

Fictitious age of patient

• Age category

Age group, can also be looked up using age\_groups()

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- Date of admission

  Date of hospital admission
- Specimen date

  Date when specimen was received at laboratory
- Specimen type Specimen type or group
- Specimen type (Numeric)
  Translation of "Specimen type"
- Reason

Reason of request with Differential Diagnosis

- Isolate number ID of isolate
- Organism type

Type of microorganism, can also be looked up using mo\_type()

- Serotype Serotype of microorganism
- Beta-lactamase
   Microorganism produces beta-lactamase?
- Microorganism produces beta-lactamase?
- Microorganism produces extended spectrum beta-lactamase?
- Carbapenemase Microorganism produces carbapenemase?
- MRSA screening test Microorganism is possible MRSA?
- Inducible clindamycin resistance Clindamycin can be induced?
- Comment

Other comments

- Date of data entry

  Date this data was entered in WHONET
- AMP\_ND10:CIP\_EE

28 different antibiotics. You can lookup the abbreviations in the antibiotics data set, or use e.g. ab\_name("AMP") to get the official name immediately. Before analysis, you should transform this to a valid antibiotic class, using as.sir().

#### **Details**

Like all data sets in this package, this data set is publicly available for download in the following formats: R, MS Excel, Apache Feather, Apache Parquet, SPSS, SAS, and Stata. Please visit our website for the download links. The actual files are of course available on our GitHub repository.

# **Examples**

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# **Index**

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