# Package 'AMR'

January 5, 2021

```
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Description Functions to simplify the analysis and prediction of Antimicrobial
      Resistance (AMR) and to work with microbial and antimicrobial properties by
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     <doi:10.1111/j.1469-0691.2011.03703.x> and the Clinical and Laboratory
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# R topics documented:

o_trom_text	
p_property	
ge	
ge_groups	. 9
MR	. 11
ntibiotics	. 12
ntibiotic_class_selectors	. 15
ab	
disk	
mic	
.mo	
.rsi	
c_online_property	
e_onnie_property	
ranability	
utalogue_of_life	
utalogue_of_life_version	
ount	
ıcast_rules	
cample_isolates	
cample_isolates_unclean	
ter_ab_class	
rst_isolate	
test	
et_episode	
gplot_pca	
gplot_rsi	. 66
ness_ab_col	
trinsic_resistant	. 72
in	. 73
ey_antibiotics	. 74
ırtosis	. 78
fecycle	. 79
KE	. 80
dro	
icroorganisms	
icroorganisms.codes	
icroorganisms.old	
o matching score	
o_property	
o_source	
Ca	
roportion	
ndom	
sistance_predict	
i_translation	
rewness	
anslate	
WHONET	
HONET	118

ab\_from\_text 3

Index 121

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,
   ...
)
```

## **Arguments**

text text to analyse

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

see Examples

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

ter 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(). Defaults to FALSE. Using

TRUE is equal to using "name".

thorough\_search

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.

... 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 agents. 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.

4 ab\_from\_text

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

## Maturing lifecycle

The lifecycle of this function is **maturing**. The unlying code of a maturing function has been roughed out, but finer details might still change. Since this function needs wider usage and more extensive testing, you are very welcome to suggest changes at our repository or write us an email (see section 'Contact Us').

# Read more on our website!

On our website https://msberends.github.io/AMR/ you can find a comprehensive tutorial about how to conduct AMR analysis, the complete documentation of all functions and an example analysis using WHONET data. As we would like to better understand the backgrounds and needs of our users, please participate in our survey!

# **Examples**

```
# 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 tds")

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")
ab_group(abx[[1]])

if (require("dplyr")) {</pre>
```

ab\_property 5

```
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 = "|"))
}
```

ab\_property

Get properties of an antibiotic

## **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_locale(), tolower = FALSE, ...)
ab_atc(x, ...)
ab_cid(x, ...)
ab_synonyms(x, ...)
ab_tradenames(x, ...)
ab_group(x, language = get_locale(), ...)
ab_atc_group1(x, language = get_locale(), ...)
ab_atc_group2(x, language = get_locale(), ...)
ab_loinc(x, ...)
ab_loinc(x, ...)
ab_info(x, language = get_locale(), ...)
ab_info(x, language = get_locale(), ...)
ab_url(x, open = FALSE, ...)
ab_property(x, property = "name", language = get_locale(), ...)
```

6 ab\_property

# **Arguments**

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

language of the returned text, defaults to system language (see get\_locale())
and can also be set with getOption("AMR\_locale"). Use language = NULL or
language = "" to prevent translation.

tolower logical to indicate whether the first character of every output should be transformed to a lower case character. This will lead to e.g. "polymyxin B" and not
"polymyxin b".

other arguments passed on to as.ab()

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

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

units a logical to indicate whether the units instead of the DDDs itself must be re-

turned, see Examples

open browse the URL using utils::browseURL()

property one of the column names of one of the antibiotics data set

#### 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.

#### Value

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

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

## Source

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

WHONET 2019 software: http://www.whonet.org/software.html

 $European\ Commission\ Public\ Health\ PHARMACEUTICALS\ -\ COMMUNITY\ REGISTER\ :\ http://ec.europa.eu/health/documents/community-register/html/atc.htm$ 

ab\_property 7

#### Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

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## See Also

antibiotics

# **Examples**

```
# all properties:
ab_name("AMX")
                     # "Amoxicillin"
ab_atc("AMX")
                    # J01CA04 (ATC code from the WHO)
ab_cid("AMX")
                    # 33613 (Compound ID from PubChem)
ab_synonyms("AMX")  # a list with brand names of amoxicillin
ab_tradenames("AMX") # same
                   # "Beta-lactams/penicillins"
ab_group("AMX")
ab_atc_group1("AMX") # "Beta-lactam antibacterials, penicillins"
ab_atc_group2("AMX") # "Penicillins with extended spectrum"
ab_url("AMX")
                     # link to the official WHO page
# smart lowercase tranformation
ab_name(x = c("AMC", "PLB")) # "Amoxicillin/clavulanic acid" "Polymyxin B"
ab_name(x = c("AMC", "PLB"),
                              # "amoxicillin/clavulanic acid" "polymyxin B"
        tolower = TRUE)
# defined daily doses (DDD)
ab_ddd("AMX", "oral")
                                    # 1
ab_ddd("AMX", "oral", units = TRUE) # "g"
ab_ddd("AMX", "iv") # 1
ab_ddd("AMX", "iv", units = TRUE)
                                   # "g"
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")
                       # ATC code of AMP (ampicillin)
ab_group("J01CA01")
                        # Drug group of ampicillins ATC code
ab_loinc("ampicillin") # LOINC codes of ampicillin
ab_name("21066-6")
                        # "Ampicillin" (using LOINC)
                        # "Ampicillin" (using CID)
ab_name(6249)
ab_name("J01CA01")
                        # "Ampicillin" (using ATC)
# spelling from different languages and dyslexia are no problem
ab_atc("ceftriaxon")
ab_atc("cephtriaxone")
ab_atc("cephthriaxone")
```

8 age

```
ab_atc("seephthriaaksone")
```

age Age in years of individuals

## **Description**

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

## Usage

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

#### **Arguments**

X	date(s), will be coerced with as.POSIX1t()
reference	reference date(s) (defaults to today), will be coerced with as.POSIX1t()
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.

## Value

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

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

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## See Also

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

age\_groups 9

#### **Examples**

```
# 10 random birth dates
df <- data.frame(birth_date = Sys.Date() - runif(10) * 25000)
# add ages
df$age <- age(df$birth_date)
# add exact ages
df$age_exact <- age(df$birth_date, exact = TRUE)
df</pre>
```

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, defaults to 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
```

# 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

10 age\_groups

#### Stable lifecycle

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#### See Also

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

## **Examples**

```
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, 17))
age_groups(ages, "children")
# resistance of ciprofloxacin per age group
if (require("dplyr")) {
  example_isolates %>%
    filter_first_isolate() %>%
    filter(mo == as.mo("E. coli")) %>%
    group_by(age_group = age_groups(age)) %>%
    select(age_group, CIP) %>%
    ggplot_rsi(x = "age_group", minimum = 0)
}
```

AMR The AMR Package

## **Description**

Welcome to the AMR package.

#### **Details**

AMR is a free, open-source and independent R package to simplify the analysis and prediction of Antimicrobial Resistance (AMR) and to work with microbial and antimicrobial data and properties, by using evidence-based methods. Our aim is to provide a standard for clean and reproducible antimicrobial resistance data analysis, that can therefore empower epidemiological analyses to continuously enable surveillance and treatment evaluation in any setting.

After installing this package, R knows ~70,000 distinct microbial species and all ~550 antibiotic, antimycotic and antiviral drugs by name and code (including ATC, EARS-NET, LOINC and SNOMED CT), and knows all about valid R/SI and MIC values. It supports any data format, including WHONET/EARS-Net data.

This package is fully independent of any other R package and works on Windows, macOS and Linux with all versions of R since R-3.0.0 (April 2013). It was designed to work in any setting, including those with very limited resources. It was created for both routine data analysis and academic research at the Faculty of Medical Sciences of the University of Groningen, in collaboration with non-profit organisations Certe Medical Diagnostics and Advice and University Medical Center Groningen. This R package is actively maintained and free software; you can freely use and distribute it for both personal and commercial (but not patent) purposes under the terms of the GNU General Public License version 2.0 (GPL-2), as published by the Free Software Foundation.

This package can be used for:

- Reference for the taxonomy of microorganisms, since the package contains all microbial (sub)species from the Catalogue of Life and List of Prokaryotic names with Standing in Nomenclature
- Interpreting raw MIC and disk diffusion values, based on the latest CLSI or EUCAST guidelines
- Retrieving antimicrobial drug names, doses and forms of administration from clinical health care records
- Determining first isolates to be used for AMR analysis
- Calculating antimicrobial resistance
- Determining multi-drug resistance (MDR) / multi-drug resistant organisms (MDRO)
- Calculating (empirical) susceptibility of both mono therapy and combination therapies
- Predicting future antimicrobial resistance using regression models
- Getting properties for any microorganism (such as Gram stain, species, genus or family)
- Getting properties for any antibiotic (such as name, code of EARS-Net/ATC/LOINC/PubChem, defined daily dose or trade name)
- · Plotting antimicrobial resistance
- Applying EUCAST expert rules
- Getting SNOMED codes of a microorganism, or getting properties of a microorganism based on a SNOMED code

12 antibiotics

 Getting LOINC codes of an antibiotic, or getting properties of an antibiotic based on a LOINC code

- Machine reading the EUCAST and CLSI guidelines from 2011-2020 to translate MIC values and disk diffusion diameters to R/SI
- Principal component analysis for AMR

# Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

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#### **Contact Us**

For suggestions, comments or questions, please contact us at:

Matthijs S. Berends

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The Netherlands https://msberends.github.io/AMR/

If you have found a bug, please file a new issue at:

https://github.com/msberends/AMR/issues

antibiotics

Data sets with 557 antimicrobials

## **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.

# Usage

antibiotics

antivirals

antibiotics 13

#### **Format**

## For the antibiotics data set: a data.frame with 455 observations and 14 variables::

ah

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

ato

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

• cid

Compound ID as found in PubChem

• name

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

group

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

• atc\_group1

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

• 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

• oral\_units

Units of oral\_ddd

• iv\_ddd

Defined Daily Dose (DDD), parenteral treatment

• iv\_units

Units of iv\_ddd

• loinc

All LOINC codes (Logical Observation Identifiers Names and Codes) associated with the name of the antimicrobial agent. Use ab\_loinc() to retrieve them quickly, see ab\_property().

## For the antivirals data set: a data.frame with 102 observations and 9 variables::

• atc

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

cid

Compound ID as found in PubChem

name

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

• atc\_group

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

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

• oral\_units

Units of oral\_ddd

14 antibiotics

- iv\_ddd
   Defined Daily Dose (DDD), parenteral treatment
- iv\_units Units of iv\_ddd

An object of class data. frame with 102 rows and 9 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) are derived from the Compound ID (cid) and consequently only available where a CID is available.

#### Direct download:

These data sets are available as 'flat files' for use even without R - you can find the files here:

- https://github.com/msberends/AMR/raw/master/data-raw/antibiotics.txt
- https://github.com/msberends/AMR/raw/master/data-raw/antivirals.txt

Files in R format (with preserved data structure) can be found here:

- https://github.com/msberends/AMR/raw/master/data/antibiotics.rda
- https://github.com/msberends/AMR/raw/master/data/antivirals.rda

# Reference data publicly available

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#### **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 (http://ec.europa.eu/health/documents/community-register/html/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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#### **Source**

```
World Health Organization (WHO) Collaborating Centre for Drug Statistics Methodology (WHOCC): https://www.whocc.no/atc_ddd_index/
WHONET 2019 software: http://www.whonet.org/software.html
European Commission Public Health PHARMACEUTICALS - COMMUNITY REGISTER: http://ec.europa.eu/health/documents/community-register/html/atc.htm
```

#### See Also

microorganisms, intrinsic\_resistant

```
antibiotic_class_selectors

Antibiotic class selectors
```

# **Description**

These functions help to select the columns of antibiotics that are of a specific antibiotic class, without the need to define the columns or antibiotic abbreviations.

# Usage

```
ab_class(ab_class)
aminoglycosides()
carbapenems()
cephalosporins()
cephalosporins_1st()
cephalosporins_2nd()
cephalosporins_3rd()
cephalosporins_4th()
cephalosporins_5th()
fluoroquinolones()
glycopeptides()
macrolides()
penicillins()
tetracyclines()
```

#### **Arguments**

ab\_class

an antimicrobial class, like "carbapenems". The columns group, atc\_group1 and atc\_group2 of the antibiotics data set will be searched (case-insensitive) for this value.

#### **Details**

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

# Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

## Read more on our website!

On our website https://msberends.github.io/AMR/ you can find a comprehensive tutorial about how to conduct AMR analysis, the complete documentation of all functions and an example analysis using WHONET data. As we would like to better understand the backgrounds and needs of our users, please participate in our survey!

# See Also

```
filter_ab_class() for the filter() equivalent.
```

# Examples

```
# `example_isolates` is a dataset available in the AMR package.
# See ?example_isolates.
# this will select columns 'IPM' (imipenem) and 'MEM' (meropenem):
example_isolates[, c(carbapenems())]
\# this will select columns 'mo', 'AMK', 'GEN', 'KAN' and 'TOB':
example_isolates[, c("mo", aminoglycosides())]
if (require("dplyr")) {
  # this will select columns 'IPM' (imipenem) and 'MEM' (meropenem):
  example_isolates %>%
   select(carbapenems())
  # this will select columns 'mo', 'AMK', 'GEN', 'KAN' and 'TOB':
  example_isolates %>%
   select(mo, aminoglycosides())
  # this will select columns 'mo' and all antimycobacterial drugs ('RIF'):
  example_isolates %>%
    select(mo, ab_class("mycobact"))
```

as.ab 17

as.ab

Transform input to an antibiotic ID

# **Description**

Use this function to determine the antibiotic 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 = TRUE, ...)
is.ab(x)
```

## **Arguments**

x character vector to determine to antibiotic ID

flag\_multiple\_results
logical to indicate whether a note should be printed to the console that probably more than one antibiotic code or name can be retrieved from a single input value.

info logical to indicate whether a progress bar should be printed
... arguments passed on to internal functions

#### 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.

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

18 as.ab

• 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 agents. This may fail on some systems.

#### 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/

WHONET 2019 software: http://www.whonet.org/software.html

European Commission Public Health PHARMACEUTICALS - COMMUNITY REGISTER: http://ec.europa.eu/health/documents/community-register/html/atc.htm

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

#### 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, <a href="https://www.whocc.no">https://www.whocc.no</a>) and the Pharmaceuticals Community Register of the European Commission (<a href="https://ec.europa.eu/health/documents/community-register/html/atc.htm">http://ec.europa.eu/health/documents/community-register/html/atc.htm</a>).

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/.

as.disk 19

#### Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

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#### 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)

## **Examples**

```
# 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") # "Erythromycin"
ab_name("eryt")
                      # "Erythromycin"
```

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.

20 as.disk

#### **Usage**

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

#### **Arguments**

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

## **Details**

Interpret disk values as RSI values with as.rsi(). It supports guidelines from EUCAST and CLSI.

#### Value

An integer with additional class disk

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

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## See Also

```
as.rsi()
```

# **Examples**

as.mic 21

```
mo = "Strep pneu", # `mo` will be coerced with as.mo()
ab = "ampicillin", # and `ab` with as.ab()
guideline = "EUCAST")
as.rsi(df)
```

as.mic

Transform input to minimum inhibitory concentrations (MIC)

## **Description**

This transforms a vector to a new class mic, which is an ordered factor with valid minimum inhibitory concentrations (MIC) as levels. Invalid MIC values will be translated as NA with a warning.

# Usage

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

# **Arguments**

x vector

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

## **Details**

To interpret MIC values as RSI values, use as.rsi() on MIC values. It supports guidelines from EUCAST and CLSI.

# Value

Ordered factor with additional class mic

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

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#### See Also

```
as.rsi()
```

## **Examples**

as.mo

Transform input to a microorganism ID

# Description

Use this function to determine a valid microorganism ID (mo). Determination is done using intelligent rules and the complete taxonomic kingdoms Bacteria, Chromista, Protozoa, Archaea 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. Please see *Examples*.

# Usage

```
as.mo(
    x,
    Becker = FALSE,
    Lancefield = FALSE,
    allow_uncertain = TRUE,
    reference_df = get_mo_source(),
    ignore_pattern = getOption("AMR_ignore_pattern"),
    language = get_locale(),
    ...
)

is.mo(x)

mo_failures()
```

```
mo_uncertainties()
mo_renamed()
```

#### **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.* (1,2,3).

This excludes *Staphylococcus aureus* at default, use Becker = "all" to also cat-

egorise S. aureus as "CoPS".

Lancefield a logical to indicate whether beta-haemolytic *Streptococci* should be categorised

into Lancefield groups instead of their own species, according to Rebecca C. Lancefield (4). These *Streptococci* will be categorised in their first group, e.g. *Streptococcus dysgalactiae* will be group C, although officially it was also cate-

gorised into groups G and L.

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

"all" to also categorise all *Enterococci* as group D.

allow\_uncertain

a number between 0 (or "none") and 3 (or "all"), or TRUE (= 2) or FALSE (= 0) to indicate whether the input should be checked for less probable results, please

see Details

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

 ${\sf set\_mo\_source()}$  and  ${\sf get\_mo\_source()}$  to automate the usage of your own

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

ignore\_pattern a 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 option AMR\_ignore\_pattern, e.g. options(AMR\_ignore\_pattern =

"(not reported|contaminated flora)").

language language to translate text like "no growth", which defaults to the system lan-

guage (see get\_locale())

... other arguments passed on to functions

# Details

# General info:

A microorganism ID 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
   1
       П
   \---> subspecies, a 4-5 letter acronym
   \---> species, a 4-5 letter acronym
   \----> genus, a 5-7 letter acronym
\----> taxonomic kingdom: A (Archaea), AN (Animalia), B (Bacteria),
                      C (Chromista), F (Fungi), P (Protozoa)
```

Values that cannot be coerced will be considered 'unknown' and will get the MO code UNKNOWN. Use the mo\_\* functions to get properties based on the returned code, see Examples.

The algorithm uses data from the Catalogue of Life (see below) and from one other source (see microorganisms).

The as.mo() function uses several coercion rules for fast and logical results. It assesses the input matching criteria in the following order:

- 1. Human pathogenic prevalence: the function starts with more prevalent microorganisms, followed by less prevalent ones;
- 2. Taxonomic kingdom: the function starts with determining Bacteria, then Fungi, then Protozoa, then others;
- 3. Breakdown of input values to identify possible matches.

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.

## **Coping with uncertain results:**

In addition, the as.mo() function can differentiate four levels of uncertainty to guess valid results:

- Uncertainty level 0: no additional rules are applied;
- Uncertainty level 1: allow previously accepted (but now invalid) taxonomic names and minor spelling errors;
- Uncertainty level 2: allow all of level 1, strip values between brackets, inverse the words of the input, strip off text elements from the end keeping at least two elements;
- Uncertainty level 3: allow all of level 1 and 2, strip off text elements from the end, allow any part of a taxonomic name.

The level of uncertainty can be set using the argument allow\_uncertain. The default is allow\_uncertain = TRUE, which is equal to uncertainty level 2. Using allow\_uncertain = FALSE is equal to uncertainty level 0 and will skip all rules. You can also use e.g. as. $mo(...,allow\_uncertain = 1)$  to only allow up to level 1 uncertainty.

With the default setting (allow\_uncertain = TRUE, level 2), below examples will lead to valid results:

- "Streptococcus group B (known as S. agalactiae)". The text between brackets will be removed and a warning will be thrown that the result *Streptococcus group B* (B\_STRPT\_GRPB) needs review.
- "S. aureus -please mind: MRSA". The last word will be stripped, after which the function will try to find a match. If it does not, the second last word will be stripped, etc. Again, a warning will be thrown that the result *Staphylococcus aureus* (B\_STPHY\_AURS) needs review.
- "Fluoroquinolone-resistant Neisseria gonorrhoeae". The first word will be stripped, after which the function will try to find a match. A warning will be thrown that the result *Neisseria gonorrhoeae* (B\_NESSR\_GNRR) needs review.

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 *Background on matching score*).
- 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 intelligent rules consider the prevalence of microorganisms in humans grouped into three groups, which is available as the prevalence columns in the microorganisms and microorganisms.old data sets. 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. Becker K *et al.* **Coagulase-Negative Staphylococci**. 2014. Clin Microbiol Rev. 27(4): 870–926; doi: 10.1128/CMR.0010913
- 2. Becker K *et al.* 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). 2019. Clin Microbiol Infect; doi: 10.1016/j.cmi.2019.02.028
- 3. Becker K *et al.* **Emergence of coagulase-negative staphylococci** 2020. Expert Rev Anti Infect Ther. 18(4):349-366; doi: 10.1080/14787210.2020.1730813
- 4. Lancefield RC A serological differentiation of human and other groups of hemolytic streptococci. 1933. J Exp Med. 57(4): 571–95; doi: 10.1084/jem.57.4.571
- 5. Catalogue of Life: Annual Checklist (public online taxonomic database), http://www.catalogueoflife.org (check included annual version with catalogue\_of\_life\_version()).

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

## 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:

$$m_{(x,n)} = \frac{l_n - 0.5 \cdot \min \left\{ l_n \operatorname{lev}(x, n) \atop l_n \cdot p_n \cdot k_n \right\}}{l_n \cdot p_n \cdot k_n}$$

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, which counts any insertion, deletion and substitution as 1 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 = 2, Protozoa = 3, Archaea = 4, others = 5.

The grouping into human pathogenic prevalence (p) is based on experience from several microbiological laboratories in the Netherlands in conjunction with international reports on pathogen prevalence. Group 1 (most prevalent microorganisms) consists of all microorganisms where the taxonomic class is Gammaproteobacteria or where the taxonomic genus is Enterococcus, Staphylococcus or Streptococcus. This group consequently contains all common Gram-negative bacteria, such as Pseudomonas and Legionella and all species within the order Enterobacterales. Group 2 consists of all microorganisms where the taxonomic phylum is Proteobacteria, Firmicutes, Actinobacteria or Sarcomastigophora, or where the taxonomic genus is Absidia, Acremonium, Actinotignum, Alternaria, Anaerosalibacter, Apophysomyces, Arachnia, Aspergillus, Aureobacterium, Aureobasidium, Bacteroides, Basidiobolus, Beauveria, Blastocystis, Branhamella, Calymmatobacterium, Candida, Capnocytophaga, Catabacter, Chaetomium, Chryseobacterium, Chryseomonas, Chrysonilia, Cladophialophora, Cladosporium, Conidiobolus, Cryptococcus, Curvularia, Exophiala, Exserohilum, Flavobacterium, Fonsecaea, Fusarium, Fusobacterium, Hendersonula, Hypomyces, Koserella, Lelliottia, Leptosphaeria, Leptotrichia, Malassezia, Malbranchea, Mortierella, Mucor, Mycocentrospora, Mycoplasma, Nectria, Ochroconis, Oidiodendron, Phoma, Piedraia, Pithomyces, Pityrosporum, Prevotella, Pseudallescheria, Rhizomucor, Rhizopus, Rhodotorula, Scolecobasidium, Scopulariopsis, Scytalidium, Sporobolomyces, Stachybotrys, Stomatococcus, Treponema, Trichoderma, Trichophyton, Trichosporon, Tritirachium or Ureaplasma. Group 3 consists of all other microorganisms.

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=0.079, a less prevalent microorganism in humans), although the latter would alphabetically come first.

# Catalogue of Life

This package contains the complete taxonomic tree of almost all microorganisms (~70,000 species) from the authoritative and comprehensive Catalogue of Life (CoL, http://www.catalogueoflife.org). The CoL is the most comprehensive and authoritative global index of species currently available. Nonetheless, we supplemented the CoL data with data from the List of Prokaryotic names with Standing in Nomenclature (LPSN, lpsn.dsmz.de). This supplementation is needed until the CoL+ project is finished, which we await.

Click here for more information about the included taxa. Check which versions of the CoL and LSPN were included in this package with catalogue\_of\_life\_version().

# Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

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sis using WHONET data. As we would like to better understand the backgrounds and needs of our users, please participate in our survey!

## 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.

## **Examples**

```
# These examples all return "B_STPHY_AURS", the ID of S. aureus:
as.mo("sau") # WHONET code
as.mo("stau")
as.mo("STAU")
as.mo("staaur")
as.mo("S. aureus")
as.mo("S aureus")
as.mo("Staphylococcus aureus")
as.mo("Staphylococcus aureus (MRSA)")
as.mo("Zthafilokkoockus oureuz") # handles incorrect spelling
as.mo("MRSA") # Methicillin Resistant S. aureus
as.mo("VISA")
                # Vancomycin Intermediate S. aureus
as.mo("VRSA")  # Vancomycin Resistant S. aureus
as.mo(115329001) # SNOMED CT code
# Dyslexia is no problem - these all work:
as.mo("Ureaplasma urealyticum")
as.mo("Ureaplasma urealyticus")
as.mo("Ureaplasmium urealytica")
as.mo("Ureaplazma urealitycium")
as.mo("Streptococcus group A")
as.mo("GAS") # Group A Streptococci
as.mo("GBS") # Group B Streptococci
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")
                                              # returns "Escherichia"
mo_gramstain("E. coli")
                                              # returns "Gram negative"
mo_is_intrinsic_resistant("E. coli", "vanco") # returns TRUE
```

# **Description**

Interpret minimum inhibitory concentration (MIC) values and disk diffusion diameters according to EUCAST or CLSI, or clean up existing R/SI values. This transforms the input to a new class rsi, which is an ordered factor with levels S < I < R. Values that cannot be interpreted will be returned as NA with a warning.

# Usage

```
as.rsi(x, ...)
is.rsi(x)
is.rsi.eligible(x, threshold = 0.05)
## S3 method for class 'mic'
as.rsi(
  Х,
  mo = NULL,
  ab = deparse(substitute(x)),
  guideline = "EUCAST",
  uti = FALSE,
  conserve_capped_values = FALSE,
  add_intrinsic_resistance = FALSE,
  reference_data = AMR::rsi_translation,
)
## S3 method for class 'disk'
as.rsi(
  х,
  mo = NULL,
  ab = deparse(substitute(x)),
  guideline = "EUCAST",
  uti = FALSE,
  add_intrinsic_resistance = FALSE,
  reference_data = AMR::rsi_translation,
## S3 method for class 'data.frame'
as.rsi(
  х,
  col_mo = NULL,
  guideline = "EUCAST",
  uti = NULL,
  conserve_capped_values = FALSE,
  add_intrinsic_resistance = FALSE,
  reference_data = rsi_translation
)
```

## **Arguments**

threshold

uti

vector of values (for class mic: an MIC value in mg/L, for class disk: a disk Χ diffusion radius in millimetres)

for using on a data.frame: names of columns to apply as.rsi() on (supports tidy selection like AMX: VAN). Otherwise: arguments passed on to methods.

maximum fraction of invalid antimicrobial interpretations of x, please see Ex-

amples

any (vector of) text that can be coerced to a valid microorganism code with mo

as.mo(), will be determined automatically if the dplyr package is installed

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

guideline defaults to the latest included EUCAST guideline, see Details for all options

> (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.rsi() on a data.frame, this can also be a column containing logicals or when left blank, the data set will be searched for a 'specimen' and rows containing 'urin' (such as 'urine', 'urina') in that column 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' v3.2 from 2020.

reference\_data a data.frame to be used for interpretation, which defaults to the rsi\_translation 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 rsi translation data set (same column names and column types). Please note that the guideline argument will be ignored when reference\_data is manually set.

col\_mo

column name of the IDs of the microorganisms (see as.mo()), defaults to the first column of class mo. Values will be coerced using as.mo().

# **Details**

# How it works:

The as.rsi() 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 R/SI 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, R/SI interpretation can be done very easily with either:
    your_data %>% mutate_if(is.mic, as.rsi) # until dplyr 1.0.0
    your_data %>% mutate(across(where(is.mic), as.rsi)) # since dplyr 1.0.0
```

- 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, R/SI interpretation can be done very easily with either:
     your\_data %>% mutate\_if(is.disk, as.rsi) # until dplyr 1.0.0
     your\_data %>% mutate(across(where(is.disk), as.rsi)) # since dplyr 1.0.0
- 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.rsi(data).

## Supported guidelines:

For interpreting MIC values as well as disk diffusion diameters, supported guidelines to be used as input for the guideline argument are: "CLSI 2010", "CLSI 2011", "CLSI 2012", "CLSI 2013", "CLSI 2014", "CLSI 2015", "CLSI 2016", "CLSI 2017", "CLSI 2018", "CLSI 2019", "EUCAST 2011", "EUCAST 2012", "EUCAST 2013", "EUCAST 2014", "EUCAST 2015", "EUCAST 2016", "EUCAST 2017", "EUCAST 2018", "EUCAST 2019", "EUCAST 2020".

Simply using "CLSI" or "EUCAST" as input will automatically select the latest version of that guideline. You can set your own data set using the reference\_data argument. The guideline argument will then be ignored.

# After interpretation:

After using as.rsi(), 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 interpretation guidelines:

The repository of this package contains a machine readable version of all guidelines. This is a CSV file consisting of 18,650 rows and 10 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 agent 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.rsi.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.

31

#### Value

Ordered factor with new class rsi

#### Interpretation of R and S/I

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

## • 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.

#### S = Susceptible

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 = Increased exposure, but still susceptible

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.

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

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

# Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

# Read more on our website!

On our website https://msberends.github.io/AMR/ you can find a comprehensive tutorial about how to conduct AMR analysis, the complete documentation of all functions and an example analysis using WHONET data. As we would like to better understand the backgrounds and needs of our users, please participate in our survey!

#### See Also

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

# **Examples**

```
summary(example_isolates) # see all R/SI results at a glance
if (require("skimr")) {
 # class <rsi> supported in skim() too:
 skim(example_isolates)
}
# For INTERPRETING disk diffusion and MIC values ------
# a whole data set, even with combined MIC values and disk zones
df <- data.frame(microorganism = "Escherichia coli",</pre>
                AMP = as.mic(8),
                CIP = as.mic(0.256),
                GEN = as.disk(18),
                TOB = as.disk(16),
                NIT = as.mic(32),
                ERY = "R")
as.rsi(df)
# for single values
as.rsi(x = as.mic(2),
       mo = as.mo("S. pneumoniae"),
       ab = "AMP",
       guideline = "EUCAST")
as.rsi(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.rsi)
  df %>% mutate_if(function(x) is.mic(x) | is.disk(x), as.rsi)
  df %>% mutate(across(where(is.mic), as.rsi))
  df %>% mutate_at(vars(AMP:TOB), as.rsi)
  df %>% mutate(across(AMP:TOB, as.rsi))
    mutate_at(vars(AMP:TOB), as.rsi, 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.rsi(uti = "from_the_bladder")
  data.frame(mo = "E. coli",
            NIT = c(" \le 2", 32),
             specimen = c("urine", "blood")) %>%
```

atc\_online\_property 33

```
as.rsi() # automatically determines urine isolates
 df %>%
   mutate_at(vars(AMP:NIT), as.rsi, mo = "E. coli", uti = TRUE)
}
# For CLEANING existing R/SI values -----
as.rsi(c("S", "I", "R", "A", "B", "C"))
as.rsi("<= 0.002; S") # will return "S"
rsi_data <- as.rsi(c(rep("S", 474), rep("I", 36), rep("R", 370)))
is.rsi(rsi_data)
plot(rsi_data)
                 # for percentages
barplot(rsi_data) # for frequencies
# the dplyr way
if (require("dplyr")) {
  example_isolates %>%
   mutate_at(vars(PEN:RIF), as.rsi)
  # same:
 example_isolates %>%
   as.rsi(PEN:RIF)
  # fastest way to transform all columns with already valid AMR results to class `rsi`:
  example_isolates %>%
   mutate_if(is.rsi.eligible, as.rsi)
  # note: from dplyr 1.0.0 on, this will be:
  # example_isolates %>%
     mutate(across(where(is.rsi.eligible), as.rsi))
}
```

atc\_online\_property Get ATC properties from WHOCC website

# **Description**

Gets data from the WHO to determine properties of an 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, ...)
```

34 atc\_online\_property

#### **Arguments**

#### **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
- "m1" = milliliter (e.g. eyedrops)

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

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

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availability 35

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## **Source**

```
https://www.whocc.no/atc_ddd_alterations_cumulative/ddd_alterations/abbrevations/
```

## **Examples**

```
# oral DDD (Defined Daily Dose) of amoxicillin
atc_online_property("J01CA04", "DDD", "O")

# 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, defaults to 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

#### Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

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# **Examples**

```
availability(example_isolates)

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

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 publicable/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_locale(),
    minimum = 30,
    combine_SI = TRUE,
    combine_IR = FALSE,
    add_ab_group = TRUE,
    remove_intrinsic_resistant = FALSE,
    decimal.mark = getOption("OutDec"),
```

```
big.mark = ifelse(decimal.mark == ",", ".", ","),
...
)
```

## **Arguments**

X	data with antibiotic columns, such as amox, AMX and AMC			
col_mo	column name of the IDs of the microorganisms (see as.mo()), defaults to the first column of class mo. Values will be coerced using as.mo().			
FUN	function to call on the mo column to transform the microorganism IDs, defaults to $mo\_shortname()$			
	arguments passed on to FUN			
translate_ab	character of length 1 containing column names of the antibiotics data set			
language	language of the returned text, defaults to system language (see <pre>get_locale())</pre> and can also be set with <pre>getOption("AMR_locale")</pre> . 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 Source.			
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). This used to be the argument combine_IR, but this now follows the redefinition by EUCAST about the interpretation of I (increased exposure) in 2019, see section 'Interpretation of S, I and R' below. Default is TRUE.			
combine_IR	logical to indicate whether values R and I should be summed			
add_ab_group	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. Use combine\_IR = FALSE (default) to test R vs. S+I and combine\_IR = TRUE to test R+I vs. S.

## Value

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

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

38 catalogue\_of\_life

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

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### Source

M39 Analysis and Presentation of Cumulative Antimicrobial Susceptibility Test Data, 4th Edition, 2014, Clinical and Laboratory Standards Institute (CLSI). https://clsi.org/standards/products/microbiology/documents/m39/.

### **Examples**

catalogue\_of\_life

The Catalogue of Life

## Description

This package contains the complete taxonomic tree of almost all microorganisms from the authoritative and comprehensive Catalogue of Life.

# Catalogue of Life

This package contains the complete taxonomic tree of almost all microorganisms (~70,000 species) from the authoritative and comprehensive Catalogue of Life (CoL, http://www.catalogueoflife.org). The CoL is the most comprehensive and authoritative global index of species currently available. Nonetheless, we supplemented the CoL data with data from the List of Prokaryotic names with Standing in Nomenclature (LPSN, lpsn.dsmz.de). This supplementation is needed until the CoL+ project is finished, which we await.

Click here for more information about the included taxa. Check which versions of the CoL and LSPN were included in this package with catalogue\_of\_life\_version().

catalogue\_of\_life 39

#### Included taxa

Included are:

• All ~55,000 (sub)species from the kingdoms of Archaea, Bacteria, Chromista and Protozoa

- All ~5,000 (sub)species from these orders of the kingdom of Fungi: Eurotiales, Microascales, Mucorales, Onygenales, Pneumocystales, Saccharomycetales, Schizosaccharomycetales and Tremellales, as well as ~4,600 other fungal (sub)species. 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 and including everything would tremendously slow down our algorithms too. By only including the aforementioned taxonomic orders, the most relevant fungi are covered (such as all species of Aspergillus, Candida, Cryptococcus, Histplasma, Pneumocystis, Saccharomyces and Trichophyton).
- All ~2,200 (sub)species from ~50 other relevant genera from the kingdom of Animalia (such as *Strongyloides* and *Taenia*)
- All ~13,000 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 responsible author(s) and year of scientific publication

The Catalogue of Life (http://www.catalogueoflife.org) is the most comprehensive and authoritative global index of species currently available. It holds essential information on the names, relationships and distributions of over 1.9 million species. The Catalogue of Life is used to support the major biodiversity and conservation information services such as the Global Biodiversity Information Facility (GBIF), Encyclopedia of Life (EoL) and the International Union for Conservation of Nature Red List. It is recognised by the Convention on Biological Diversity as a significant component of the Global Taxonomy Initiative and a contribution to Target 1 of the Global Strategy for Plant Conservation.

The syntax used to transform the original data to a cleansed R format, can be found here: https://github.com/msberends/AMR/blob/master/data-raw/reproduction\_of\_microorganisms.R.

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# See Also

Data set microorganisms for the actual data.

Function as.mo() to use the data for intelligent determination of microorganisms.

```
# Get version info of included data set
catalogue_of_life_version()

# Get a note when a species was renamed
mo_shortname("Chlamydophila psittaci")
# Note: 'Chlamydophila psittaci' (Everett et al., 1999) was renamed back to
```

```
'Chlamydia psittaci' (Page, 1968)
#> [1] "C. psittaci"
# Get any property from the entire taxonomic tree for all included species
mo_class("E. coli")
#> [1] "Gammaproteobacteria"
mo_family("E. coli")
#> [1] "Enterobacteriaceae"
mo_gramstain("E. coli") # based on kingdom and phylum, see ?mo_gramstain
#> [1] "Gram-negative"
mo_ref("E. coli")
#> [1] "Castellani et al., 1919"
\# Do not get mistaken - this package is about microorganisms
mo_kingdom("C. elegans")
#> [1] "Fungi"
                               # Fungi?!
mo_name("C. elegans")
#> [1] "Cladosporium elegans" # Because a microorganism was found
```

catalogue\_of\_life\_version

Version info of included Catalogue of Life

## **Description**

This function returns information about the included data from the Catalogue of Life.

## Usage

```
catalogue_of_life_version()
```

### Details

For DSMZ, see microorganisms.

# Value

a list, which prints in pretty format

### Catalogue of Life

This package contains the complete taxonomic tree of almost all microorganisms (~70,000 species) from the authoritative and comprehensive Catalogue of Life (CoL, http://www.catalogueoflife.org). The CoL is the most comprehensive and authoritative global index of species currently available. Nonetheless, we supplemented the CoL data with data from the List of Prokaryotic names with Standing in Nomenclature (LPSN, lpsn.dsmz.de). This supplementation is needed until the CoL+ project is finished, which we await.

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## See Also

microorganisms

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, please 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_rsi(..., only_all_tested = FALSE)
count_df(
  data,
  translate_ab = "name",
  language = get_locale(),
  combine_SI = TRUE,
  combine_IR = FALSE
)
```

#### **Arguments**

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

only\_all\_tested

(for combination therapies, i.e. using more than one variable for  $\dots$ ): 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 rsi (see as.rsi())

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, defaults to system language (see get\_locale())

and can also be set with getOption("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). This used to be the argument combine\_IR, but this now follows the redefinition by EUCAST about the interpretation of I (increased exposure) in 2019, see section 'Interpretation

of S. I and R' below. Default is TRUE.

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

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

outdated, see argument combine\_SI.

### **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  $n_rsi()$  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 rsi class (created with as.rsi()) and counts the number of S's, I's and R's. It also supports grouped variables. The function rsi\_df() works exactly like count\_df(), but adds the percentage of S, I and R.

## Value

An integer

# Stable lifecycle

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### Interpretation of R and S/I

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

#### • 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.

### • S = Susceptible

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 = Increased exposure, but still susceptible

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.

This AMR package honours this new 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 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>	-	-	-	-

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.

#### Read more on our website!

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#### See Also

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

```
# example_isolates is a data set available in the AMR package.
?example_isolates
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$AMX)
n_rsi(example_isolates$AMX)
# n_rsi() 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_rsi(example_isolates$AMX)
if (require("dplyr")) {
  example_isolates %>%
    group_by(hospital_id) %>%
    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_rsi(CIP), # same - analogous to n_distinct
              total = n())
                                   # NOT the number of tested isolates!
  # 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)
  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)
  # 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(hospital_id, AMX, CIP) %>%
   group_by(hospital_id) %>%
   count_df(translate = FALSE)
}
```

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://eucast.org), see *Source*.

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 = 10,
    version_expertrules = 3.2,
    ampc_cephalosporin_resistance = NA,
    ...
)
```

#### **Arguments**

```
x data with antibiotic columns, such as amox, AMX and AMC col_mo column name of the IDs of the microorganisms (see as.mo()), defaults to 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, defaults

to 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", "all", and defaults to c("breakpoints", "expert")

The default value can be set to another value, e.g. using options (AMR\_eucastrules

= "all").

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. Currently supported: 10.0.

version\_expertrules

the version number to use for the EUCAST Expert Rules and Intrinsic Resistance guideline. Currently supported: 3.1, 3.2.

ampc\_cephalosporin\_resistance

a character value that should be applied for AmpC de-repressed cephalosporin-resistant mutants, defaults to NA. Currently only works when version\_expertrules is 3.2; 'EUCAST Expert Rules v3.2 on Enterobacterales' states that susceptible (S) results of cefotaxime, ceftriaxone and ceftazidime should be reported with a note, or results should be suppressed (emptied) for these agents. A value of NA for this argument will remove results for these agents, while e.g. a value of "R" will make the results for these agents resistant. Use NULL to not alter the results for AmpC de-repressed cephalosporin-resistant mutants.

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

. . column name of an antibiotic, please see section *Antibiotics* below

### Details

Note: This function does not translate MIC values to RSI values. Use as .rsi() 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/master/data-raw/eucast\_rules.tsv.

#### '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 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 used 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), avoparcin (AVO, no ATC code), azithromycin (AZM, J01FA10), azlocillin (AZL, J01CA09), aztreonam (ATM, J01DF01), bacampicillin (BAM, J01CA06), benzylpenicillin (PEN, J01CE01), cadazolid (CDZ, J01DD09), carbenicillin (CRB, J01CA03), carindacillin (CRN, J01CA05), cefacetrile (CAC, J01DB10), cefaclor (CEC, J01DC04), cefadroxil (CFR, J01DB05), cefaloridine (RID, J01DB02), cefamandole (MAN, J01DC03), cefatrizine (CTZ, J01DB07), cefazedone (CZD, J01DB06), cefazolin (CZO, J01DB04), cefcapene (CCP, no ATC code), cefcapene pivoxil (CCX, no ATC code), cefdinir (CDR, J01DD15), cefditoren (DIT, J01DD16), cefditoren pivoxil (DIX, no ATC code), cefepime (FEP, J01DE01), cefetamet (CAT, J01DD10), cefetamet pivoxil (CPI, no ATC code), 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, no ATC code), cefotaxime/sulbactam (CTS, no ATC code), cefotetan (CTT, J01DC05), cefotiam (CTF, J01DC07), cefotiam hexetil (CHE, no ATC code), cefovecin (FOV, no ATC code), cefoxitin (FOX, J01DC01), cefoxitin screening (FOX1, no ATC code), cefpimizole (CFZ, no ATC code), cefpiramide (CPM, J01DD11), cefpirome (CPO, J01DE02), cefpodoxime (CPD, J01DD13), cefpodoxime proxetil (CPX, no ATC code), cefpodoxime/clavulanic acid (CDC, no ATC code), cefprozil (CPR, J01DC10), cefroxadine (CRD, J01DB11), cefsulodin (CFS, J01DD03), ceftaroline (CPT, J01DI02), ceftazidime (CAZ, J01DD02), ceftazidime/avibactam (CZA, no ATC code), ceftazidime/clavulanic acid (CCV, J01DD52), cefteram (CEM, no ATC code), cefteram pivoxil (CPL, no ATC code), ceftezole (CTL, J01DB12), ceftibuten (CTB, J01DD14), ceftiofur (TIO, no ATC code), ceftizoxime (CZX, J01DD07), ceftizoxime alapivoxil (CZP, no ATC code), ceftobiprole (BPR, J01DI01), ceftobiprole medocaril (CFM1, J01DI01), ceftolozane/enzyme inhibitor (CEI, J01DI54), ceftriaxone (CRO, J01DD04), cefuroxime (CXM, J01DC02), cephalexin (LEX, J01DB01), cephalothin (CEP, J01DB03), cephapirin (HAP, J01DB08), cephradine (CED, J01DB09), chloramphenicol (CHL, J01BA01), ciprofloxacin (CIP, J01MA02), clarithromycin (CLR, J01FA09), clindamycin (CLI, J01FF01), colistin (COL, J01XB01), cycloserine (CYC, J04AB01), dalbavancin (DAL, J01XA04), daptomycin (DAP, J01XX09), dibekacin (DKB, J01GB09), 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 (F0S, J01XX01), fusidic acid (FUS, J01XC01), gatifloxacin (GAT, J01MA16), gemifloxacin (GEM, J01MA15), gentamicin (GEN, J01GB03), grepafloxacin (GRX, J01MA11), hetacillin (HET, J01CA18), imipenem (IPM, J01DH51), isepamicin (ISE, J01GB11), josamycin (J0S, J01FA07), kanamycin (KAN, J01GB04), latamoxef (LTM, J01DD06), levofloxacin (LVX, J01MA12), lincomycin (LIN, J01FF02), linezolid (LNZ, J01XX08), lomefloxacin (LOM, J01MA07), loracarbef (LOR, J01DC08), mecillinam (Amdinocillin)

(MEC, J01CA11), meropenem (MEM, J01DH02), meropenem/vaborbactam (MEV, J01DH52), metampicillin (MTM, J01CA14), mezlocillin (MEZ, J01CA10), midecamycin (MID, J01FA03), minocycline (MNO, J01AA08), miocamycin (MCM, J01FA11), moxifloxacin (MFX, J01MA14), nalidixic acid (NAL, J01MB02), neomycin (NEO, J01GB05), netilmicin (NET, J01GB07), nitrofurantoin (NIT, J01XE01), norfloxacin (NOR, J01MA06), norvancomycin (NVA, no ATC code), novobiocin (NOV, QJ01XX95), ofloxacin (0FX, J01MA01), oleandomycin (0LE, J01FA05), oritavancin (0RI, J01XA05), oxacillin (OXA, J01CF04), pazufloxacin (PAZ, J01MA18), pefloxacin (PEF, J01MA03), phenoxymethylpenicillin (PHN, J01CE02), piperacillin (PIP, J01CA12), piperacillin/tazobactam (TZP, J01CR05), pirlimycin (PRL, no ATC code), pivampicillin (PVM, J01CA02), pivmecillinam (PME, J01CA08), polymyxin B (PLB, J01XB02), pristinamycin (PRI, J01FG01), prulifloxacin (PRU, J01MA17), quinupristin/dalfopristin (QDA, J01FG02), ramoplanin (RAM, no ATC code), ribostamycin (RST, J01GB10), rifampicin (RIF, J04AB02), rokitamycin (R0K, J01FA12), roxithromycin (RXT, J01FA06), rufloxacin (RFL, J01MA10), sisomicin (SIS, J01GB08), sparfloxacin (SPX, J01MA09), spiramycin (SPI, J01FA02), streptoduocin (STR, J01GA02), streptomycin (STR1, J01GA01), 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 (SZ0, 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), talampicillin (TAL, J01CA15), tedizolid (TZD, J01XX11), teicoplanin (TEC, J01XA02), teicoplaninmacromethod (TCM, no ATC code), telavancin (TLV, J01XA03), telithromycin (TLT, J01FA15), temafloxacin (TMX, J01MA05), temocillin (TEM, J01CA17), tetracycline (TCY, J01AA07), thiacetazone (THA, no ATC code), ticarcillin (TIC, J01CA13), ticarcillin/clavulanic acid (TCC, J01CR03), tigecycline (TGC, J01AA12), tobramycin (T0B, J01GB01), trimethoprim (TMP, J01EA01), trimethoprim/sulfamethoxazole (SXT, J01EE01), troleandomycin (TRL, J01FA08), trovafloxacin (TVA, J01MA13), vancomycin (VAN, J01XA01)

### Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

### Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

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sis using WHONET data. As we would like to better understand the backgrounds and needs of our users, please participate in our survey!

### **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 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)

```
a <- data.frame(mo = c("Staphylococcus aureus",</pre>
                      "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)
а
                       mo VAN AMX COL CAZ CXM PEN FOX
#
# 1 Staphylococcus aureus -
                                                     S
                                                          S
# 2 Enterococcus faecalis
                                                     S
                                                          S
# 3
         Escherichia coli
                                                     S
                                                          S
# 4 Klebsiella pneumoniae
                                                     S
                                                          S
# 5 Pseudomonas aeruginosa
# apply EUCAST rules: some results wil be changed
b <- eucast_rules(a)</pre>
b
#
                       mo VAN AMX COL
                                         CAZ
                                                   PEN
                                                        FOX
                                              CXM
# 1 Staphylococcus aureus
                                                     S
                                                          S
                                  S
                                      R
                                           R
                                                S
# 2 Enterococcus faecalis
                                       R
                                            R
                                                     S
                                                          R
                                                R
         Escherichia coli R -
                                                          S
# 3
                                                     R
# 4 Klebsiella pneumoniae
                           R
                               R
                                                     R
                                                          S
# 5 Pseudomonas aeruginosa
```

<sup>#</sup> do not apply EUCAST rules, but rather get a data.frame

50 example\_isolates

```
# containing all details about the transformations:
c <- eucast_rules(a, verbose = TRUE)</pre>
```

example\_isolates

Data set with 2,000 example isolates

## Description

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

#### Usage

```
example_isolates
```

#### **Format**

A data.frame with 2,000 observations and 49 variables:

- date date of receipt at the laboratory
- hospital\_id
   ID of the hospital, from A to D
- ward\_icu logical to determine if ward is an intensive care unit
- ward\_clinical logical to determine if ward is a regular clinical ward
- ward\_outpatient logical to determine if ward is an outpatient clinic
- age age of the patient
- gender gender of the patient
- patient\_idID of the patient
- mo
   ID of microorganism created with as.mo(), see also microorganisms
- PEN:RIF
   40 different antibiotics with class rsi (see as.rsi()); these column names occur in the antibiotics data set and can be translated with ab\_name()

# Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

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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 analysis. This data set can be used for practice.

# Usage

example\_isolates\_unclean

#### **Format**

A data.frame 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.rsi()

### Reference data publicly available

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52 filter\_ab\_class

filter\_ab\_class

Filter isolates on result in antimicrobial class

### **Description**

Filter isolates on results in specific antimicrobial classes. This makes it easy to filter on isolates that were tested for e.g. any aminoglycoside, or to filter on carbapenem-resistant isolates without the need to specify the drugs.

## Usage

```
filter_ab_class(x, ab_class, result = NULL, scope = "any", ...)
filter_aminoglycosides(x, result = NULL, scope = "any", ...)
filter_carbapenems(x, result = NULL, scope = "any", ...)
filter_cephalosporins(x, result = NULL, scope = "any", ...)
filter_1st_cephalosporins(x, result = NULL, scope = "any", ...)
filter_2nd_cephalosporins(x, result = NULL, scope = "any", ...)
filter_3rd_cephalosporins(x, result = NULL, scope = "any", ...)
filter_4th_cephalosporins(x, result = NULL, scope = "any", ...)
filter_5th_cephalosporins(x, result = NULL, scope = "any", ...)
filter_fluoroquinolones(x, result = NULL, scope = "any", ...)
filter_glycopeptides(x, result = NULL, scope = "any", ...)
filter_macrolides(x, result = NULL, scope = "any", ...)
filter_penicillins(x, result = NULL, scope = "any", ...)
filter_tetracyclines(x, result = NULL, scope = "any", ...)
```

#### Arguments

. ....

X	a data set
ab_class	an antimicrobial class, like "carbapenems". The columns group, atc_group1 and atc_group2 of the antibiotics data set will be searched (case-insensitive) for this value.
result	an antibiotic result: S, I or R (or a combination of more of them)
scope	the scope to check which variables to check, can be "any" (default) or "all"
• • •	previously used when this package still depended on the dplyr package, now ignored

filter\_ab\_class 53

#### **Details**

All columns of x will be searched for known antibiotic names, abbreviations, brand names and codes (ATC, EARS-Net, WHO, etc.). This means that a filter function like e.g. filter\_aminoglycosides() will include column names like 'gen', 'genta', 'J01GB03', 'tobra', 'Tobracin', etc.

## Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

### See Also

```
antibiotic_class_selectors() for the select() equivalent.
```

```
filter_aminoglycosides(example_isolates)
if (require("dplyr")) {
  # filter on isolates that have any result for any aminoglycoside
  example_isolates %>% filter_aminoglycosides()
  example_isolates %>% filter_ab_class("aminoglycoside")
  # this is essentially the same as (but without determination of column names):
  example_isolates %>%
    filter_at(.vars = vars(c("GEN", "TOB", "AMK", "KAN")),
              .vars_predicate = any_vars(. %in% c("S", "I", "R")))
  # filter on isolates that show resistance to ANY aminoglycoside
  example_isolates %>% filter_aminoglycosides("R", "any")
  # filter on isolates that show resistance to ALL aminoglycosides
  example_isolates %>% filter_aminoglycosides("R", "all")
  # filter on isolates that show resistance to
  # any aminoglycoside and any fluoroquinolone
  example_isolates %>%
   filter_aminoglycosides("R") %>%
   filter_fluoroquinolones("R")
  # filter on isolates that show resistance to
  # all aminoglycosides and all fluoroquinolones
  example_isolates %>%
    filter_aminoglycosides("R", "all") %>%
    filter_fluoroquinolones("R", "all")
  # with dplyr 1.0.0 and higher (that adds 'across()'), this is equal:
```

```
# (though the row names on the first are more correct)
example_isolates %>% filter_carbapenems("R", "all")
example_isolates %>% filter(across(carbapenems(), ~. == "R"))
}
```

first\_isolate

Determine first (weighted) isolates

## **Description**

Determine first (weighted) isolates of all microorganisms of every patient per episode and (if needed) per specimen type. To determine patient episodes not necessarily based on microorganisms, use is\_new\_episode() that also supports grouping with the dplyr package.

## Usage

```
first_isolate(
  х,
  col_date = NULL,
  col_patient_id = NULL,
  col_mo = NULL,
  col_testcode = NULL,
  col_specimen = NULL,
  col_icu = NULL,
  col_keyantibiotics = NULL,
  episode_days = 365,
  testcodes_exclude = NULL,
  icu_exclude = FALSE,
  specimen_group = NULL,
  type = "keyantibiotics",
  ignore_I = TRUE,
  points_threshold = 2,
  info = interactive(),
  include_unknown = FALSE,
)
filter_first_isolate(
  col_date = NULL,
  col_patient_id = NULL,
  col_mo = NULL,
)
filter_first_weighted_isolate(
  col_date = NULL,
  col_patient_id = NULL,
  col_mo = NULL,
```

```
col_keyantibiotics = NULL,
...
)
```

#### **Arguments**

x a data.frame containing isolates. Can be left blank when used inside dplyr

verbs, such as filter(), mutate() and summarise().

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

to the first column with a date class

col\_patient\_id column name of the unique IDs of the patients, defaults to the first column that

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

col\_mo column name of the IDs of the microorganisms (see as.mo()), defaults to 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)

col\_keyantibiotics

column name of the key antibiotics to determine first *weighted* isolates, see key\_antibiotics(). Defaults to the first column that starts with 'key' followed by 'ab' or 'antibiotics' (case insensitive). Use col\_keyantibiotics =

FALSE to prevent this.

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

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

icu\_exclude logical 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 "keyantibiotics" or "points",

see Details

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

when type = "keyantibiotics", see Details

points\_threshold

points until the comparison of key antibiotics will lead to inclusion of an isolate

when type = "points", see Details

info print progress

include\_unknown

logical to determine 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.

arguments passed on to first\_isolate() when using filter\_first\_isolate(),

or arguments passed on to key\_antibiotics() when using filter\_first\_weighted\_isolate()

#### **Details**

These functions are context-aware when used inside dplyr verbs, such as filter(), mutate() and summarise(). This means that then the x argument can be left blank, please 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.

## Why this is so important:

To conduct an analysis of antimicrobial resistance, you should only include the first isolate of every patient per episode (Hindler *et al.* 2007). If you would not do this, you could easily get an overestimate or underestimate of the resistance of an antibiotic. Imagine that a patient was admitted with an MRSA and that it was found in 5 different blood cultures the following week. The resistance percentage of oxacillin of all *S. aureus* isolates would be overestimated, because you included this MRSA more than once. It would be selection bias.

```
filter_*() shortcuts:
```

The functions filter\_first\_isolate() and filter\_first\_weighted\_isolate() are helper functions to quickly filter on first isolates.

The function filter\_first\_isolate() is essentially equal to either:

filter(only\_weighted\_firsts == TRUE) %>%
select(-only\_weighted\_firsts, -keyab)

```
x[first_isolate(x, ...), ]
x %>% filter(first_isolate(...))
The function filter_first_weighted_isolate() is essentially equal to:
x %>%
    mutate(keyab = key_antibiotics(.)) %>%
    mutate(only_weighted_firsts = first_isolate(x, col_keyantibiotics = "keyab", ...)) %>%
```

#### Value

A logical vector

#### **Key antibiotics**

There are two ways to determine whether isolates can be included as first *weighted* isolates which will give generally the same results:

- Using type = "keyantibiotics" and argument ignore\_I
   Any difference from S to R (or vice versa) will (re)select an isolate as a first weighted isolate. With ignore\_I = FALSE, also differences from I to SIR (or vice versa) will lead to this. This is a reliable method and 30-35 times faster than method 2. Read more about this in the key\_antibiotics() function.
- 2. Using type = "points" and argument points\_threshold
  A difference from I to SIR (or vice versa) means 0.5 points, a difference from S to R (or vice versa) means 1 point. When the sum of points exceeds points\_threshold, which default to 2, an isolate will be (re)selected as a first weighted isolate.

#### Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

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#### Source

Methodology of this function is strictly based on:

M39 Analysis and Presentation of Cumulative Antimicrobial Susceptibility Test Data, 4th Edition, 2014, Clinical and Laboratory Standards Institute (CLSI). https://clsi.org/standards/products/microbiology/documents/m39/.

### See Also

```
key_antibiotics()
```

```
# `example_isolates` is a dataset available in the AMR package.
# See ?example_isolates.
# basic filtering on first isolates
example_isolates[first_isolate(example_isolates), ]
# filtering based on isolates ------
if (require("dplyr")) {
  # filter on first isolates:
  example_isolates %>%
   mutate(first_isolate = first_isolate(.)) %>%
   filter(first_isolate == TRUE)
  # short-hand versions:
  example_isolates %>%
   filter(first_isolate())
  example_isolates %>%
   filter_first_isolate()
  example_isolates %>%
   filter_first_weighted_isolate()
  # now let's see if first isolates matter:
  A <- example_isolates %>%
```

58 g.test

```
group_by(hospital_id) %>%
   summarise(count = n_rsi(GEN),
                                             # gentamicin availability
              resistance = resistance(GEN)) # gentamicin resistance
  B <- example_isolates %>%
   filter_first_weighted_isolate() %>%
                                             # the 1st isolate filter
   group_by(hospital_id) %>%
   summarise(count = n_rsi(GEN),
                                             # gentamicin availability
             resistance = resistance(GEN)) # gentamicin resistance
  # Have a look at A and B.
  # B is more reliable because every isolate is counted only once.
  # Gentamicin resistance in hospital D appears to be 3.7% higher than
  # when you (erroneously) would have used all isolates for analysis.
}
```

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 of $x$ . 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.

## 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

g.test 59

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 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>
```

60 g.test

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:

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

statistic the value the chi-squared test statistic. the degrees of freedom of the approximate chi-squared distribution of the test parameter 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. the Pearson residuals, (observed -expected) / sqrt(expected). residuals 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

### **Questioning lifecycle**

The lifecycle of this function is **questioning**. This function might be no longer be optimal approach, or is it questionable whether this function should be in this AMR package at all.

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## Source

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

- 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()
```

get\_episode 61

#### **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 \leftarrow g.test(x, p = c(1, 2, 1) / 4)
# G$p.value = 0.12574.
# 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)
# p = 0.01787343
# There is a significant difference from a 1:1 ratio.
# Meaning: there are significantly more left-billed birds.
```

get\_episode

Determine (new) episodes for patients

# Description

These functions determine which items in a vector can be considered (the start of) a new episode, based on the argument episode\_days. This can be used to determine clinical episodes for any epidemiological analysis. The <code>get\_episode()</code> function returns the index number of the episode per group, while the <code>is\_new\_episode()</code> function returns values TRUE/FALSE to indicate whether an item in a vector is the start of a new episode.

### Usage

```
get_episode(x, episode_days, ...)
is_new_episode(x, episode_days, ...)
```

62 get\_episode

#### **Arguments**

```
x vector of dates (class Date or POSIXt)
episode_days length of the required episode in days, please see Details
... arguments passed on to as.Date()
```

#### Details

Dates are first sorted from old to new. The oldest date will mark the start of the first episode. After this date, the next date will be marked that is at least episode\_days days later than the start of the first episode. From that second marked date on, the next date will be marked that is at least episode\_days days later than the start of the second episode which will be the start of the third episode, and so on. Before the vector is being returned, the original order will be restored.

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.

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

### Value

```
get_episode(): a double vectoris_new_episode(): a logical vector
```

### Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

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## See Also

```
first_isolate()
```

```
# `example_isolates` is a dataset available in the AMR package.
# See ?example_isolates.

get_episode(example_isolates$date, episode_days = 60)
is_new_episode(example_isolates$date, episode_days = 60)
# filter on results from the third 60-day episode only, using base R
```

ggplot\_pca 63

```
example_isolates[which(get_episode(example_isolates$date, 60) == 3), ]
if (require("dplyr")) {
  # is_new_episode() can also be used in dplyr verbs to determine patient
  # episodes based on any (combination of) grouping variables:
  example_isolates %>%
   mutate(condition = sample(x = c("A", "B", "C"),
                              size = 2000.
                              replace = TRUE)) %>%
   group_by(condition) %>%
   mutate(new_episode = is_new_episode(date, 365))
  example_isolates %>%
   group_by(hospital_id, patient_id) %>%
   transmute(date,
              patient_id,
              new_index = get_episode(date, 60),
              new_logical = is_new_episode(date, 60))
  example_isolates %>%
   group_by(hospital_id) %>%
    summarise(patients = n_distinct(patient_id),
              n_episodes_365 = sum(is_new_episode(date, episode_days = 365)),
              n_episodes_60 = sum(is_new_episode(date, episode_days = 60)),
              n_episodes_30 = sum(is_new_episode(date, episode_days = 30)))
  # grouping on patients and microorganisms leads to the same results
  # as first_isolate():
  x <- example_isolates %>%
   filter(first_isolate(., include_unknown = TRUE))
  y <- example_isolates %>%
   group_by(patient_id, mo) %>%
    filter(is_new_episode(date, 365))
  identical(x$patient_id, y$patient_id)
  # but is_new_episode() has a lot more flexibility than first_isolate(),
  # since you can now group on anything that seems relevant:
  example_isolates %>%
   group_by(patient_id, mo, hospital_id, ward_icu) %>%
   mutate(flag_episode = is_new_episode(date, 365))
}
```

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.

64 ggplot\_pca

#### 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 <= scale <= 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

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

ggplot\_pca 65

using the pca() function as input for x, this will be determined automatically based on the attribute non\_numeric\_cols, see pca(). 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 the size of the points points\_alpha the alpha (transparency) of the points 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

ellipse

points\_size

arrows\_size

arrows

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, like scale\_colour\_viridis\_d() or scale\_colour\_brewer().

### Maturing lifecycle

The lifecycle of this function is maturing. The unlying code of a maturing function has been roughed out, but finer details might still change. Since this function needs wider usage and more extensive testing, you are very welcome to suggest changes at our repository or write us an email (see section 'Contact Us').

## 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
- 5. Cleaned all syntax based on the lintr package, fixed grammatical errors and added integrity
- 6. Updated documentation

#### **Examples**

```
# `example_isolates` is a dataset available in the AMR package.
# See ?example_isolates.
# See ?pca for more info about Principal Component Analysis (PCA).
if (require("dplyr")) {
 pca_model <- example_isolates %>%
   filter(mo_genus(mo) == "Staphylococcus") %>%
   group_by(species = mo_shortname(mo)) %>%
   summarise_if (is.rsi, resistance) %>%
   pca(FLC, AMC, CXM, GEN, TOB, TMP, SXT, CIP, TEC, TCY, ERY)
  # old (base R)
  biplot(pca_model)
  # new
  ggplot_pca(pca_model)
  if (require("ggplot2")) {
   ggplot_pca(pca_model) +
     scale_colour_viridis_d() +
     labs(title = "Title here")
 }
}
```

ggplot\_rsi

AMR plots with ggplot2

### **Description**

Use these functions to create bar plots for antimicrobial resistance analysis. All functions rely on ggplot2 functions.

## Usage

```
ggplot_rsi(
 data,
 position = NULL,
 x = "antibiotic",
 fill = "interpretation",
 facet = NULL,
 breaks = seq(0, 1, 0.1),
 limits = NULL,
  translate_ab = "name",
  combine_SI = TRUE,
  combine_IR = FALSE,
 minimum = 30,
 language = get_locale(),
 nrow = NULL,
  colours = c(S = "#61a8ff", SI = "#61a8ff", I = "#61f7ff", IR = "#ff6961", R =
    "#ff6961"),
  datalabels = TRUE,
```

```
datalabels.size = 2.5,
  datalabels.colour = "grey15",
  title = NULL,
  subtitle = NULL,
  caption = NULL,
  x.title = "Antimicrobial",
  y.title = "Proportion",
geom_rsi(
  position = NULL,
  x = c("antibiotic", "interpretation"),
  fill = "interpretation",
  translate_ab = "name",
  minimum = 30,
  language = get_locale(),
  combine_SI = TRUE,
  combine_IR = FALSE,
)
facet_rsi(facet = c("interpretation", "antibiotic"), nrow = NULL)
scale_y_percent(breaks = seq(0, 1, 0.1), limits = NULL)
scale_rsi_colours(
  colours = c(S = "#61a8ff", SI = "#61a8ff", I = "#61f7ff", IR = "#ff6961", R =
    "#ff6961")
)
theme_rsi()
labels_rsi_count(
  position = NULL,
  x = "antibiotic",
  translate_ab = "name",
  minimum = 30,
  language = get_locale(),
  combine_SI = TRUE,
  combine_IR = FALSE,
  datalabels.size = 3,
  datalabels.colour = "grey15"
)
```

## **Arguments**

```
data a data.frame with column(s) of class rsi (see as.rsi())

position position adjustment of bars, either "fill", "stack" or "dodge"

x variable to show on x axis, either "antibiotic" (default) or "interpretation" or a grouping variable
```

fill variable to categorise using the plots legend, either "antibiotic" (default) or "interpretation" or a grouping variable variable to split plots by, either "interpretation" (default) or "antibiotic" facet or a grouping variable breaks numeric vector of positions limits numeric vector of length two providing limits of the scale, use NA to refer to the existing minimum or maximum a column name of the antibiotics data set to translate the antibiotic abbreviations translate\_ab 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). This used to be the argument combine\_IR, but this now follows the redefinition by EUCAST about the interpretation of I (increased exposure) in 2019, see section 'Interpretation of S, I and R' below. Default is TRUE. a logical to indicate whether all values of I and R must be merged into one, so combine\_IR the output only consists of S vs. I+R (susceptible vs. non-susceptible). This is outdated, see argument combine\_SI. 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, defaults to system language (see get\_locale()) and can also be set with getOption("AMR\_locale"). Use language = NULL or language = "" to prevent translation. (when using facet) number of rows nrow colours a named vector with colours for the bars. The names must be one or more of: S, SI, I, IR, R or be FALSE to use default ggplot2 colours. datalabels show datalabels using labels\_rsi\_count() datalabels.size size of the datalabels datalabels.colour colour of the datalabels title text to show as title of the plot subtitle text to show as subtitle of the plot caption text to show as caption of the plot x.title text to show as x axis description y.title text to show as y axis description other arguments passed on to geom\_rsi() . . .

### **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\_rsi() will take any variable from the data that has an rsi class (created with as.rsi()) using rsi\_df() and will plot bars with the percentage R, I and S. The default behaviour is to have the bars stacked and to have the different antibiotics on the x axis.

```
facet_rsi() 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_rsi_colours() sets colours to the bars: pastel blue for S, pastel turquoise for I and pastel red for R, using ggplot2::scale_fill_manual().

theme_rsi() is a [ggplot2 theme][ggplot2::theme() with minimal distraction.

labels_rsi_count() print datalabels on the bars with percentage and amount of isolates using ggplot2::geom_text().

ggplot_rsi() 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.
```

### Maturing lifecycle

The lifecycle of this function is **maturing**. The unlying code of a maturing function has been roughed out, but finer details might still change. Since this function needs wider usage and more extensive testing, you are very welcome to suggest changes at our repository or write us an email (see section 'Contact Us').

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```
if (require("ggplot2") & require("dplyr")) {
  # get antimicrobial results for drugs against a UTI:
  ggplot(example_isolates %>% select(AMX, NIT, FOS, TMP, CIP)) +
   geom_rsi()
  # prettify the plot using some additional functions:
  df <- example_isolates %>% select(AMX, NIT, FOS, TMP, CIP)
  ggplot(df) +
   geom_rsi() +
   scale_y_percent() +
   scale_rsi_colours() +
   labels_rsi_count() +
   theme_rsi()
  # or better yet, simplify this using the wrapper function - a single command:
  example_isolates %>%
    select(AMX, NIT, FOS, TMP, CIP) %>%
   ggplot_rsi()
  # get only proportions and no counts:
  example_isolates %>%
   select(AMX, NIT, FOS, TMP, CIP) %>%
   ggplot_rsi(datalabels = FALSE)
  # add other ggplot2 arguments as you like:
  example_isolates %>%
    select(AMX, NIT, FOS, TMP, CIP) %>%
   ggplot_rsi(width = 0.5,
```

70 guess\_ab\_col

```
colour = "black",
               size = 1,
               linetype = 2,
               alpha = 0.25)
  example_isolates %>%
    select(AMX) %>%
    ggplot_rsi(colours = c(SI = "yellow"))
}
# resistance of ciprofloxacine per age group
example_isolates %>%
  mutate(first_isolate = first_isolate(.)) %>%
  filter(first_isolate == TRUE,
         mo == as.mo("E. coli")) %>%
  # age_groups() is also a function in this AMR package:
  group_by(age_group = age_groups(age)) %>%
  select(age_group,
         CIP) %>%
  ggplot_rsi(x = "age_group")
# for colourblind mode, use divergent colours from the viridis package:
example_isolates %>%
  select(AMX, NIT, FOS, TMP, CIP) %>%
  ggplot_rsi() +
  scale_fill_viridis_d()
# a shorter version which also adjusts data label colours:
example_isolates %>%
  select(AMX, NIT, FOS, TMP, CIP) %>%
  ggplot_rsi(colours = FALSE)
# it also supports groups (don't forget to use the group var on `x` or `facet`):
example_isolates %>%
  select(hospital_id, AMX, NIT, FOS, TMP, CIP) %>%
  group_by(hospital_id) %>%
  ggplot_rsi(x = "hospital_id",
             facet = "antibiotic",
             nrow = 1,
             title = "AMR of Anti-UTI Drugs Per Hospital",
             x.title = "Hospital",
             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.

guess\_ab\_col 71

#### Usage

```
guess_ab_col(x = NULL, search_string = NULL, verbose = 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
```

#### **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. **Longer columns names take precedence over shorter column names.** 

### Value

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

#### Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

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72 intrinsic\_resistant

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 data.frame with 93,892 observations and 2 variables:

- microorganism

  Name of the microorganism
- antibiotic Name of the antibiotic drug

#### **Details**

The repository of this AMR package contains a file comprising this exact data set: https://github.com/msberends/AMR/blob/master/data-raw/intrinsic\_resistant.txt. This file allows for machine reading EUCAST guidelines about intrinsic resistance, which is almost impossible with the Excel and PDF files distributed by EUCAST. The file is updated automatically.

This data set is based on 'EUCAST Expert Rules' and 'EUCAST Intrinsic Resistance and Unusual Phenotypes' v3.2 from 2020.

## Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

join 73

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# **Examples**

```
if (require("dplyr")) {
  intrinsic_resistant %>%
    filter(antibiotic == "Vancomycin", microorganism %like% "Enterococcus") %>%
    pull(microorganism)
# [1] "Enterococcus casseliflavus" "Enterococcus gallinarum"
}
```

join

Join microorganisms to a data set

### **Description**

Join the data set microorganisms easily to an existing table or 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 table to join, or character vector
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

#### **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() function from base R will be used.

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

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### **Examples**

```
left_join_microorganisms(as.mo("K. pneumoniae"))
left_join_microorganisms("B_KLBSL_PNE")
if (require("dplyr")) {
  example_isolates %>%
    left_join_microorganisms() %>%
    colnames()
  df <- data.frame(date = seq(from = as.Date("2018-01-01"),</pre>
                               to = as.Date("2018-01-07"),
                               by = 1),
                   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)
}
```

#### **Description**

These function can be used to determine first isolates (see first\_isolate()). Using key antibiotics to determine first isolates is more reliable than without key antibiotics. These selected isolates can then be called first *weighted* isolates.

### Usage

```
key_antibiotics(
  col_mo = NULL,
  universal_1 = guess_ab_col(x, "amoxicillin"),
  universal_2 = guess_ab_col(x, "amoxicillin/clavulanic acid"),
  universal_3 = guess_ab_col(x, "cefuroxime"),
  universal_4 = guess_ab_col(x, "piperacillin/tazobactam"),
  universal_5 = guess_ab_col(x, "ciprofloxacin"),
  universal_6 = guess_ab_col(x, "trimethoprim/sulfamethoxazole"),
  GramPos_1 = guess_ab_col(x, "vancomycin"),
  GramPos_2 = guess_ab_col(x, "teicoplanin"),
GramPos_3 = guess_ab_col(x, "tetracycline"),
  GramPos_4 = guess_ab_col(x, "erythromycin"),
  GramPos_5 = guess_ab_col(x, "oxacillin"),
  GramPos_6 = guess_ab_col(x, "rifampin"),
  GramNeg_1 = guess_ab_col(x, "gentamicin"),
  GramNeg_2 = guess_ab_col(x, "tobramycin"),
  GramNeg_3 = guess_ab_col(x, "colistin"),
  GramNeg_4 = guess_ab_col(x, "cefotaxime"),
GramNeg_5 = guess_ab_col(x, "ceftazidime"),
  GramNeg_6 = guess_ab_col(x, "meropenem"),
  warnings = TRUE,
)
key_antibiotics_equal(
  у,
  z,
  type = c("keyantibiotics", "points"),
  ignore_I = TRUE,
  points_threshold = 2,
  info = FALSE
)
```

# **Arguments**

```
a data.frame with antibiotics columns, like AMX or amox. Can be left blank when used inside dplyr verbs, such as filter(), mutate() and summarise().

col_mo

column name of the IDs of the microorganisms (see as.mo()), defaults to the first column of class mo. Values will be coerced using as.mo().

universal_1, universal_2, universal_3, universal_4, universal_5, universal_6

column names of broad-spectrum antibiotics, case-insensitive. See details for which antibiotics will be used at default (which are guessed with guess_ab_col()).
```

GramPos\_1, GramPos\_2, GramPos\_3, GramPos\_4, GramPos\_5, GramPos\_6 column names of antibiotics for Gram-positives, case-insensitive. See details for which antibiotics will be used at default (which are guessed with guess\_ab\_col()). GramNeg\_1, GramNeg\_2, GramNeg\_3, GramNeg\_4, GramNeg\_5, GramNeg\_6 column names of antibiotics for Gram-negatives, case-insensitive. See details for which antibiotics will be used at default (which are guessed with guess\_ab\_col()). give a warning about missing antibiotic columns (they will be ignored) warnings other arguments passed on to functions y, z character vectors to compare type to determine weighed isolates; can be "keyantibiotics" or "points", type see Details logical to determine whether antibiotic interpretations with "I" will be ignored ignore\_I when type = "keyantibiotics", see Details points\_threshold points until the comparison of key antibiotics will lead to inclusion of an isolate

when type = "points", see Details

info print progress

#### **Details**

The key\_antibiotics() function is context-aware when used inside dplyr verbs, such as filter(), mutate() and summarise(). This means that then the x argument can be left blank, please see *Examples*.

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

The first\_isolate() function only uses this function on the same microbial species from the same patient. Using this, e.g. an MRSA will be included after a susceptible *S. aureus* (MSSA) is found within the same patient episode. Without key antibiotic comparison it would not. See first\_isolate() for more info.

At default, the antibiotics that are used for **Gram-positive bacteria** are:

- Amoxicillin
- · Amoxicillin/clavulanic acid
- Cefuroxime
- · Piperacillin/tazobactam
- · Ciprofloxacin
- Trimethoprim/sulfamethoxazole
- Vancomycin
- Teicoplanin
- Tetracycline
- Erythromycin
- Oxacillin
- Rifampin

At default the antibiotics that are used for Gram-negative bacteria are:

- Amoxicillin
- · Amoxicillin/clavulanic acid
- Cefuroxime
- Piperacillin/tazobactam
- · Ciprofloxacin
- Trimethoprim/sulfamethoxazole
- Gentamicin
- · Tobramycin
- Colistin
- · Cefotaxime
- · Ceftazidime
- Meropenem

The function key\_antibiotics\_equal() checks the characters returned by key\_antibiotics() for equality, and returns a logical vector.

#### Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

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# **Key antibiotics**

There are two ways to determine whether isolates can be included as first *weighted* isolates which will give generally the same results:

- Using type = "keyantibiotics" and argument ignore\_I
   Any difference from S to R (or vice versa) will (re)select an isolate as a first weighted isolate. With ignore\_I = FALSE, also differences from I to SIR (or vice versa) will lead to this. This is a reliable method and 30-35 times faster than method 2. Read more about this in the key\_antibiotics() function.
- 2. Using type = "points" and argument points\_threshold
  A difference from I to SIR (or vice versa) means 0.5 points, a difference from S to R (or vice versa) means 1 point. When the sum of points exceeds points\_threshold, which default to 2, an isolate will be (re)selected as a first weighted isolate.

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## See Also

```
first_isolate()
```

78 kurtosis

#### **Examples**

```
\mbox{\# `example\_isolates` is a dataset available in the AMR package.}
# See ?example_isolates.
# output of the `key_antibiotics()` function could be like this:
strainA <- "SSSRR.S.R..S"</pre>
strainB <- "SSSIRSSSRSSS"</pre>
# those strings can be compared with:
key_antibiotics_equal(strainA, strainB)
# TRUE, because I is ignored (as well as missing values)
key_antibiotics_equal(strainA, strainB, 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_antibiotics()) %>% # no need to define `x`
    mutate(
      # now calculate first isolates
      first_regular = first_isolate(col_keyantibiotics = FALSE),
      # and first WEIGHTED isolates
      first_weighted = first_isolate(col_keyantibiotics = "keyab")
  # Check the difference, in this data set it results in a lot 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)
## 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)
```

lifecycle 79

## **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 <i>excess kurtosis</i> should be returned, defined as the kurtosis minus 3.

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

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### See Also

skewness()

lifecycle	Lifecycles of functions in the AMR package

# Description

Functions in this AMR package are categorised using the lifecycle circle of the Tidyverse as found on www.tidyverse.org/lifecycle.

This page contains a section for every lifecycle (with text borrowed from the aforementioned Tidyverse website), so they can be used in the manual pages of the functions.

# **Experimental lifecycle**

The lifecycle of this function is **experimental**. An experimental function is in early stages of development. The unlying code might be changing frequently. Experimental functions might be removed without deprecation, so you are generally best off waiting until a function is more mature before you use it in production code. Experimental functions are only available in development versions of this AMR package and will thus not be included in releases that are submitted to CRAN, since such functions have not yet matured enough.

80 like

#### Maturing lifecycle

The lifecycle of this function is **maturing**. The unlying code of a maturing function has been roughed out, but finer details might still change. Since this function needs wider usage and more extensive testing, you are very welcome to suggest changes at our repository or write us an email (see section 'Contact Us').

# Stable lifecycle

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### Retired lifecycle

The lifecycle of this function is **retired**. A retired function is no longer under active development, and (if appropiate) a better alternative is available. No new arguments will be added, and only the most critical bugs will be fixed. In a future version, this function will be removed.

# Questioning lifecycle

The lifecycle of this function is **questioning**. This function might be no longer be optimal approach, or is it questionable whether this function should be in this AMR package at all.

like

Pattern matching with keyboard shortcut

# **Description**

Convenient wrapper around grep() 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 %like_case% pattern
```

like 81

## **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 string containing a regular expression (or character string for fixed =

TRUE) to be matched in the given character vector. Coerced by as.character() to a character string if possible. If a character vector of length 2 or more is

supplied, the first element is used with a warning.

ignore.case if FALSE, the pattern matching is case sensitive and if TRUE, case is ignored

during matching.

#### **Details**

The %like% function:

• Is case-insensitive (use %like\_case% for case-sensitive matching)

• Supports multiple patterns

 Checks if pattern is a regular expression and sets fixed = TRUE if not, to greatly improve speed

• Tries again with perl = TRUE if regex fails

Using RStudio? The text %like% can also be directly inserted in your code from the Addins menu and can have its own Keyboard Shortcut like Ctrl+Shift+L or Cmd+Shift+L (see Tools > Modify Keyboard Shortcuts...).

#### Value

A logical vector

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

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#### **Source**

Idea from the like function from the data. table package

## See Also

grep()

# **Examples**

```
# simple test
a <- "This is a test"
b <- "TEST"
a %like% b
#> TRUE
b %like% a
#> FALSE
\mbox{\tt\#} also supports multiple patterns, length must be equal to x
a <- c("Test case", "Something different", "Yet another thing")
                                           "yet")
b <- c( "case",
                              "diff",
a %like% b
#> TRUE TRUE TRUE
# get isolates whose name start with 'Ent' or 'ent'
if (require("dplyr")) {
  example_isolates %>%
    filter(mo_name(mo) %like% "^ent")
}
```

mdro

Determine multidrug-resistant organisms (MDRO)

# **Description**

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

# Usage

```
mdro(
    x,
    guideline = "CMI2012",
    col_mo = NULL,
    info = interactive(),
    pct_required_classes = 0.5,
    combine_SI = TRUE,
    verbose = FALSE,
    ...
)

brmo(x, guideline = "BRMO", ...)

mrgn(x, guideline = "TB", ...)

mdr_tb(x, guideline = "TB", ...)

mdr_cmi2012(x, guideline = "CMI2012", ...)

eucast_exceptional_phenotypes(x, guideline = "EUCAST", ...)
```

## **Arguments**

x a data.frame with antibiotics columns, like AMX or amox. Can be left blank when used inside dplyr verbs, such as filter(), mutate() and summarise().

guideline a specific guideline to follow. When left empty, the publication by Magiorakos

et al. (2012, Clinical Microbiology and Infection) will be followed, please see

Details.

col\_mo column name of the IDs of the microorganisms (see as.mo()), defaults to 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, defaults

to 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.

... column name of an antibiotic, please see section *Antibiotics* below

### **Details**

These functions are context-aware when used inside dplyr verbs, such as filter(), mutate() and summarise(). This means that then the x argument can be left blank, please 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).

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) (link)

guideline = "EUCAST3.2" (or simply guideline = "EUCAST")
 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.

**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)</li>
- 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:

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

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

# 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 used 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), avoparcin (AV0, no ATC code), azithromycin (AZM, J01FA10), azlocillin (AZL, J01CA09), aztreonam (ATM, J01DF01), bacampicillin (BAM, J01CA06), benzylpenicillin (PEN, J01CE01), cadazolid (CDZ, J01DD09), carbenicillin (CRB, J01CA03), carindacillin (CRN, J01CA05), cefacetrile (CAC, J01DB10), cefaclor (CEC, J01DC04), cefadroxil (CFR, J01DB05), cefaloridine (RID, J01DB02), cefamandole (MAN, J01DC03), cefatrizine (CTZ, J01DB07), cefazedone (CZD, J01DB06), cefazolin (CZO, J01DB04), cefcapene (CCP, no ATC code), cefcapene pivoxil (CCX, no ATC code), cefdinir (CDR, J01DD15), cefditoren (DIT, J01DD16), cefditoren pivoxil (DIX, no ATC code), cefepime (FEP, J01DE01), cefetamet (CAT, J01DD10), cefetamet pivoxil (CPI, no ATC code), 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, no ATC code), cefotaxime/sulbactam (CTS, no ATC code), cefotetan (CTT, J01DC05), cefotiam (CTF, J01DC07), cefotiam hexetil (CHE, no ATC code), cefovecin (FOV, no ATC code), cefoxitin (FOX, J01DC01), cefoxitin screening (FOX1, no ATC code), cefpimizole (CFZ, no ATC code), cefpiramide (CPM, J01DD11), cefpirome (CPO, J01DE02), cefpodoxime (CPD, J01DD13), cefpodoxime proxetil (CPX, no ATC code), cefpodoxime/clavulanic acid (CDC, no ATC code), cefprozil (CPR, J01DC10), cefroxadine (CRD, J01DB11), cefsulodin (CFS, J01DD03), ceftaroline (CPT, J01DI02), ceftazidime (CAZ, J01DD02), ceftazidime/avibactam (CZA, no ATC code), ceftazidime/clavulanic acid (CCV, J01DD52), cefteram (CEM, no ATC code), cefteram pivoxil (CPL, no ATC code), ceftezole (CTL, J01DB12), ceftibuten (CTB, J01DD14), ceftiofur (TIO, no ATC code), ceftizoxime (CZX, J01DD07), ceftizoxime alapivoxil (CZP, no ATC code), ceftobiprole (BPR, J01DI01), ceftobiprole medocaril (CFM1, J01DI01), ceftolozane/enzyme inhibitor (CEI, J01DI54), ceftriaxone (CRO, J01DD04), cefuroxime (CXM, J01DC02), cephalexin (LEX, J01DB01), cephalothin (CEP, J01DB03), cephapirin (HAP, J01DB08), cephradine (CED, J01DB09), chloramphenicol (CHL, J01BA01), ciprofloxacin (CIP, J01MA02), clarithromycin (CLR, J01FA09), clindamycin (CLI, J01FF01), colistin (COL, J01XB01), cycloserine (CYC, J04AB01), dalbavancin (DAL, J01XA04), daptomycin (DAP, J01XX09), dibekacin (DKB, J01GB09), 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 (F0S, J01XX01), fusidic acid (FUS, J01XC01), gatifloxacin (GAT, J01MA16), gemifloxacin (GEM, J01MA15), gentamicin (GEN, J01GB03), grepafloxacin (GRX, J01MA11), hetacillin (HET, J01CA18), imipenem (IPM, J01DH51), isepamicin (ISE, J01GB11), josamycin (J0S, J01FA07), kanamycin (KAN, J01GB04), latamoxef (LTM, J01DD06), levofloxacin (LVX, J01MA12), lincomycin (LIN, J01FF02), linezolid (LNZ, J01XX08), lomefloxacin (LOM, J01MA07), loracarbef (LOR, J01DC08), mecillinam (Amdinocillin) (MEC, J01CA11), meropenem (MEM, J01DH02), meropenem/vaborbactam (MEV, J01DH52), metampicillin (MTM, J01CA14), mezlocillin (MEZ, J01CA10), midecamycin (MID, J01FA03), minocycline (MNO, J01AA08), miocamycin (MCM, J01FA11), moxifloxacin (MFX, J01MA14), nalidixic acid (NAL, J01MB02), neomycin (NEO, J01GB05), netilmicin (NET, J01GB07), nitrofurantoin (NIT, J01XE01), norfloxacin (NOR, J01MA06), norvancomycin (NVA, no ATC code), novobiocin (NOV, QJ01XX95), ofloxacin (OFX, J01MA01), oleandomycin (OLE, J01FA05), oritavancin (ORI, J01XA05), oxacillin (OXA, J01CF04), pazufloxacin (PAZ, J01MA18), pefloxacin (PEF, J01MA03), phenoxymethylpenicillin (PHN, J01CE02), piperacillin (PIP, J01CA12), piperacillin/tazobactam (TZP, J01CR05), pirlimycin (PRL, no ATC code), pivampicillin (PVM, J01CA02), pivmecillinam (PME, J01CA08), polymyxin B (PLB, J01XB02), pristinamycin (PRI, J01FG01), prulifloxacin (PRU, J01MA17), quinupristin/dalfopristin (QDA, J01FG02), ramoplanin (RAM, no ATC code), ribostamycin (RST, J01GB10), rifampicin (RIF, J04AB02), rokitamycin (ROK, J01FA12), roxithromycin (RXT, J01FA06), rufloxacin (RFL, J01MA10), sisomicin (SIS, J01GB08), sparfloxacin (SPX, J01MA09), spiramycin (SPI, J01FA02), streptoduocin (STR, J01GA02), streptomycin (STR1, J01GA01), 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 (SZ0, 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), sulfamilamide (SLF9, J01EB06), sulfaperin (SLF10, J01ED06), sulfaphenazole (SLF11, J01ED08), sulfapyridine (SLF12, J01EB04), sulfathiazole (SUT, J01EB07), sulfathiourea (SLF13, J01EB08), talampicillin (TAL, J01CA15), tedizolid (TZD, J01XX11), teicoplanin (TEC, J01XA02), teicoplaninmacromethod (TCM, no ATC code), telavancin (TLV, J01XA03), telithromycin (TLT, J01FA15), temafloxacin (TMX, J01MA05), temocillin (TEM, J01CA17), tetracycline (TCY, J01AA07), thiacetazone (THA, no ATC code), ticarcillin (TIC, J01CA13), ticarcillin/clavulanic acid (TCC, J01CR03), tigecycline (TGC, J01AA12), tobramycin (T0B, J01GB01), trimethoprim (TMP, J01EA01), trimethoprim/sulfamethoxazole (SXT, J01EE01), troleandomycin (TRL, J01FA08), trovafloxacin (TVA, J01MA13), vancomycin (VAN, J01XA01)

#### Interpretation of R and S/I

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

#### • 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.

### • S = Susceptible

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 = Increased exposure, but still susceptible

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.

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

#### Read more on our website!

On our website https://msberends.github.io/AMR/ you can find a comprehensive tutorial about how to conduct AMR analysis, the complete documentation of all functions and an example analysis using WHONET data. As we would like to better understand the backgrounds and needs of our users, please participate in our survey!

#### **Source**

Please see *Details* for the list of publications used for this function.

microorganisms 87

#### **Examples**

microorganisms

Data set with 67,151 microorganisms

# **Description**

A data set containing the microbial taxonomy of six kingdoms from the Catalogue of Life. MO codes can be looked up using as.mo().

# Usage

microorganisms

# **Format**

A data.frame with 67,151 observations and 16 variables:

- mc
  - ID of microorganism as used by this package
- fullname

Full name, like "Escherichia coli"

- kingdom, phylum, class, order, family, genus, species, subspecies Taxonomic rank of the microorganism
- rank

Text of the taxonomic rank of the microorganism, like "species" or "genus"

- ref
- Author(s) and year of concerning scientific publication
- species\_id

ID of the species as used by the Catalogue of Life

source

Either "CoL", "DSMZ" (see Source) or "manually added"

• prevalence

Prevalence of the microorganism, see as.mo()

• snomed

SNOMED code of the microorganism. Use mo\_snomed() to retrieve it quickly, see mo\_property().

88 microorganisms

#### **Details**

Manually added were:

• 11 entries of *Streptococcus* (beta-haemolytic: groups A, B, C, D, F, G, H, K and unspecified; other: viridans, milleri)

- 2 entries of Staphylococcus (coagulase-negative (CoNS) and coagulase-positive (CoPS))
- 3 entries of *Trichomonas* (*Trichomonas vaginalis*, and its family and genus)
- 1 entry of Candida (Candida krusei), that is not (yet) in the Catalogue of Life
- 1 entry of *Blastocystis (Blastocystis hominis*), although it officially does not exist (Noel *et al.* 2005, PMID 15634993)
- 5 other 'undefined' entries (unknown, unknown Gram negatives, unknown Gram positives, unknown yeast and unknown fungus)
- 6 families under the Enterobacterales order, according to Adeolu *et al.* (2016, PMID 27620848), that are not (yet) in the Catalogue of Life
- 7,411 species from the DSMZ (Deutsche Sammlung von Mikroorganismen und Zellkulturen) since the DSMZ contain the latest taxonomic information based on recent publications

#### Direct download:

This data set is available as 'flat file' for use even without R - you can find the file here:

https://github.com/msberends/AMR/raw/master/data-raw/microorganisms.txt

The file in R format (with preserved data structure) can be found here:

https://github.com/msberends/AMR/raw/master/data/microorganisms.rda

#### About the records from DSMZ (see source)

Names of prokaryotes are defined as being validly published by the International Code of Nomenclature of Bacteria. Validly published are all names which are included in the Approved Lists of Bacterial Names and the names subsequently published in the International Journal of Systematic Bacteriology (IJSB) and, from January 2000, in the International Journal of Systematic and Evolutionary Microbiology (IJSEM) as original articles or in the validation lists. (from https://www.dsmz.de/services/online-tools/prokaryotic-nomenclature-up-to-date)

In February 2020, the DSMZ records were merged with the List of Prokaryotic names with Standing in Nomenclature (LPSN).

## Catalogue of Life

This package contains the complete taxonomic tree of almost all microorganisms (~70,000 species) from the authoritative and comprehensive Catalogue of Life (CoL, http://www.catalogueoflife.org). The CoL is the most comprehensive and authoritative global index of species currently available. Nonetheless, we supplemented the CoL data with data from the List of Prokaryotic names with Standing in Nomenclature (LPSN, lpsn.dsmz.de). This supplementation is needed until the CoL+ project is finished, which we await.

Click here for more information about the included taxa. Check which versions of the CoL and LSPN were included in this package with catalogue\_of\_life\_version().

microorganisms.codes 89

#### Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

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## Source

Catalogue of Life: Annual Checklist (public online taxonomic database), http://www.catalogueoflife.org (check included annual version with catalogue\_of\_life\_version()).

Parte, A.C. (2018). LPSN — List of Prokaryotic names with Standing in Nomenclature (bacterio.net), 20 years on. International Journal of Systematic and Evolutionary Microbiology, 68, 1825-1829; doi: 10.1099/ijsem.0.002786

Leibniz Institute DSMZ-German Collection of Microorganisms and Cell Cultures, Germany, Prokary-otic Nomenclature Up-to-Date, https://www.dsmz.de/services/online-tools/prokaryotic-nomenclature-up-and https://lpsn.dsmz.de (check included version with catalogue\_of\_life\_version()).

### See Also

as.mo(), mo\_property(), microorganisms.codes, intrinsic\_resistant

microorganisms.codes Data set with 5,583 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 data frame with 5.583 observations and 2 variables:

- code
   Commonly used code of a microorganism
- mo
   ID of the microorganism in the microorganisms data set

90 microorganisms.old

#### Reference data publicly available

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Click here for more information about the included taxa. Check which versions of the CoL and LSPN were included in this package with catalogue\_of\_life\_version().

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# See Also

as.mo() microorganisms

microorganisms.old

Data set with previously accepted taxonomic names

# **Description**

A data set containing old (previously valid or accepted) taxonomic names according to the Catalogue of Life. This data set is used internally by as.mo().

## Usage

microorganisms.old

#### **Format**

A data.frame with 12,708 observations and 4 variables:

- fullname
  - Old full taxonomic name of the microorganism
- fullname\_new
   New full taxonomic name of the microorganism
- ref
   Author(s) and year of concerning scientific publication
- prevalence Prevalence of the microorganism, see as.mo()

mo\_matching\_score 91

#### Catalogue of Life

This package contains the complete taxonomic tree of almost all microorganisms (~70,000 species) from the authoritative and comprehensive Catalogue of Life (CoL, http://www.catalogueoflife.org). The CoL is the most comprehensive and authoritative global index of species currently available. Nonetheless, we supplemented the CoL data with data from the List of Prokaryotic names with Standing in Nomenclature (LPSN, lpsn.dsmz.de). This supplementation is needed until the CoL+ project is finished, which we await.

Click here for more information about the included taxa. Check which versions of the CoL and LSPN were included in this package with catalogue\_of\_life\_version().

### Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

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# **Source**

Catalogue of Life: Annual Checklist (public online taxonomic database), http://www.catalogueoflife.org (check included annual version with catalogue\_of\_life\_version()).

Parte, A.C. (2018). LPSN — List of Prokaryotic names with Standing in Nomenclature (bacterio.net), 20 years on. International Journal of Systematic and Evolutionary Microbiology, 68, 1825-1829; doi: 10.1099/ijsem.0.002786

# See Also

```
as.mo() mo_property() microorganisms
```

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)
```

92 mo\_matching\_score

#### **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:

$$m_{(x,n)} = \frac{l_n - 0.5 \cdot \min \left\{ l_n \operatorname{lev}(x, n) \atop l_n \cdot p_n \cdot k_n \right\}}$$

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, which counts any insertion, deletion and substitution as 1 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 = 2, Protozoa = 3, Archaea = 4, others = 5.

The grouping into human pathogenic prevalence (p) is based on experience from several microbiological laboratories in the Netherlands in conjunction with international reports on pathogen prevalence. Group 1 (most prevalent microorganisms) consists of all microorganisms where the taxonomic class is Gammaproteobacteria or where the taxonomic genus is Enterococcus, Staphylococcus or Streptococcus. This group consequently contains all common Gram-negative bacteria, such as Pseudomonas and Legionella and all species within the order Enterobacterales. Group 2 consists of all microorganisms where the taxonomic phylum is Proteobacteria, Firmicutes, Actinobacteria or Sarcomastigophora, or where the taxonomic genus is Absidia, Acremonium, Actinotignum, Alternaria, Anaerosalibacter, Apophysomyces, Arachnia, Aspergillus, Aureobacterium, Aureobasidium, Bacteroides, Basidiobolus, Beauveria, Blastocystis, Branhamella, Calymmatobacterium, Candida, Capnocytophaga, Catabacter, Chaetomium, Chryseobacterium, Chryseomonas, Chrysonilia, Cladophialophora, Cladosporium, Conidiobolus, Cryptococcus, Curvularia, Exophiala, Exserohilum, Flavobacterium, Fonsecaea, Fusarium, Fusobacterium, Hendersonula, Hypomyces, Koserella, Lelliottia, Leptosphaeria, Leptotrichia, Malassezia, Malbranchea, Mortierella, Mucor, Mycocentrospora, Mycoplasma, Nectria, Ochroconis, Oidiodendron, Phoma, Piedraia, Pithomyces, Pityrosporum, Prevotella,\Pseudallescheria, Rhizomucor, Rhizopus, Rhodotorula, Scolecobasidium, Scopulariopsis, Scytalidium, Sporobolomyces, Stachybotrys, Stomatococcus, Treponema, Trichoderma, Trichophyton, Trichosporon, Tritirachium or Ureaplasma. Group 3 consists of all other microorganisms.

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=0.079, a less prevalent microorganism in humans), although the latter would alphabetically come first.

#### Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

#### Author(s)

Matthijs S. Berends

# **Examples**

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. Please see *Examples*.

# Usage

```
mo_name(x, language = get_locale(), ...)
mo_fullname(x, language = get_locale(), ...)
mo_shortname(x, language = get_locale(), ...)
mo_subspecies(x, language = get_locale(), ...)
mo_species(x, language = get_locale(), ...)
mo_genus(x, language = get_locale(), ...)
mo_family(x, language = get_locale(), ...)
mo_order(x, language = get_locale(), ...)
mo_class(x, language = get_locale(), ...)
mo_phylum(x, language = get_locale(), ...)
```

```
mo_kingdom(x, language = get_locale(), ...)
   mo_domain(x, language = get_locale(), ...)
   mo_type(x, language = get_locale(), ...)
   mo_gramstain(x, language = get_locale(), ...)
   mo_is_gram_negative(x, language = get_locale(), ...)
   mo_is_gram_positive(x, language = get_locale(), ...)
   mo_is_intrinsic_resistant(x, ab, language = get_locale(), ...)
   mo_snomed(x, language = get_locale(), ...)
   mo_ref(x, language = get_locale(), ...)
   mo_authors(x, language = get_locale(), ...)
   mo_year(x, language = get_locale(), ...)
   mo_rank(x, language = get_locale(), ...)
   mo_taxonomy(x, language = get_locale(), ...)
   mo_synonyms(x, language = get_locale(), ...)
   mo_info(x, language = get_locale(), ...)
   mo_url(x, open = FALSE, language = get_locale(), ...)
   mo_property(x, property = "fullname", language = get_locale(), ...)
Arguments
                    any character (vector) that can be coerced to a valid microorganism code with
    х
                    as.mo(). Can be left blank for auto-guessing the column containing microor-
                    ganism codes when used inside dplyr verbs, such as filter(), mutate() and
                     summarise(), please see Examples.
   language
                    language of the returned text, defaults to system language (see get_locale())
                    and can be overwritten by setting the option AMR_locale, e.g. options(AMR_locale
                    = "de"), see translate. Also used to translate text like "no growth". Use language
                    = NULL or language = "" to prevent translation.
                    other arguments passed on to as.mo(), such as 'allow_uncertain' and 'ignore_pattern'
                    any (vector of) text that can be coerced to a valid antibiotic code with as.ab()
   ab
                    browse the URL using browseURL()
   open
                    one of the column names of the microorganisms data set: "mo", "fullname",
   property
                     "kingdom", "phylum", "class", "order", "family", "genus", "species", "subspecies",
                     "rank", "ref", "species_id", "source", "prevalence", "snomed", or must be
```

"shortname"

#### **Details**

All functions will return the most recently known taxonomic property according to the Catalogue of Life, except for mo\_ref(), mo\_authors() and mo\_year(). Please refer to this example, knowing that *Escherichia blattae* was renamed to *Shimwellia blattae* in 2010:

- mo\_name("Escherichia blattae") will return "Shimwellia blattae" (with a message about the renaming)
- mo\_ref("Escherichia blattae") will return "Burgess et al., 1973" (with a message about the renaming)
- mo\_ref("Shimwellia blattae") will return "Priest et al., 2010" (without a message)

The short name - mo\_shortname() - almost always returns the first character of the genus and the full species, like "E. coli". 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*. In other words, 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.

The Gram stain - mo\_gramstain() - will be determined based on the taxonomic kingdom and phylum. According to Cavalier-Smith (2002, PMID 11837318), who defined subkingdoms Negibacteria and Posibacteria, only these phyla are Posibacteria: Actinobacteria, Chloroflexi, Firmicutes and Tenericutes. These bacteria are considered Gram-positive - 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 (except when the input is NA or the MO code is UNKNOWN), thus always return FALSE for species outside the taxonomic kingdom of Bacteria.

Intrinsic resistance - mo\_is\_intrinsic\_resistant() - will be determined based on the intrinsic\_resistant data set, which is based on 'EUCAST Expert Rules' and 'EUCAST Intrinsic Resistance and Unusual Phenotypes' v3.2 from 2020. The mo\_is\_intrinsic\_resistant() can be vectorised over arguments x (input for microorganisms) and over ab (input for antibiotics).

All output will be translated where possible.

The function mo\_url() will return the direct URL to the online database entry, which also shows the scientific reference of the concerned species.

# Value

- An integer in case of mo\_year()
- A list in case of mo\_taxonomy() and mo\_info()
- A named character in case of mo\_url()
- A double in case of mo\_snomed()
- · A character in all other cases

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

#### 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:

$$m_{(x,n)} = \frac{l_n - 0.5 \cdot \min \left\{ l_n \operatorname{lev}(x, n) \atop l_n \cdot p_n \cdot k_n \right\}}{l_n \cdot p_n \cdot k_n}$$

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, which counts any insertion, deletion and substitution as 1 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 = 2, Protozoa = 3, Archaea = 4, others = 5.

The grouping into human pathogenic prevalence (p) is based on experience from several microbiological laboratories in the Netherlands in conjunction with international reports on pathogen prevalence. Group 1 (most prevalent microorganisms) consists of all microorganisms where the taxonomic class is Gammaproteobacteria or where the taxonomic genus is Enterococcus, Staphylococcus or Streptococcus. This group consequently contains all common Gram-negative bacteria, such as Pseudomonas and Legionella and all species within the order Enterobacterales. Group 2 consists of all microorganisms where the taxonomic phylum is Proteobacteria, Firmicutes, Actinobacteria or Sarcomastigophora, or where the taxonomic genus is Absidia, Acremonium, Actinotignum, Alternaria, Anaerosalibacter, Apophysomyces, Arachnia, Aspergillus, Aureobacterium, Aureobasidium, Bacteroides, Basidiobolus, Beauveria, Blastocystis, Branhamella, Calymmatobacterium, Candida, Capnocytophaga, Catabacter, Chaetomium, Chryseobacterium, Chryseomonas, Chrysonilia, Cladophialophora, Cladosporium, Conidiobolus, Cryptococcus, Curvularia, Exophiala, Exserohilum, Flavobacterium, Fonsecaea, Fusarium, Fusobacterium, Hendersonula, Hypomyces, Koserella, Lelliottia, Leptosphaeria, Leptotrichia, Malassezia, Malbranchea, Mortierella, Mucor, Mycocentrospora, Mycoplasma, Nectria, Ochroconis, Oidiodendron, Phoma, Piedraia, Pithomyces, Pityrosporum, Prevotella,\Pseudallescheria, Rhizomucor, Rhizopus, Rhodotorula, Scolecobasidium, Scopulariopsis, Scytalidium, Sporobolomyces, Stachybotrys, Stomatococcus, Treponema, Trichoderma, Trichophyton, Trichosporon, Tritirachium or Ureaplasma. Group 3 consists of all other microorganisms.

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=0.079, a less prevalent microorganism in humans), although the latter would alphabetically come first.

# Catalogue of Life

This package contains the complete taxonomic tree of almost all microorganisms (~70,000 species) from the authoritative and comprehensive Catalogue of Life (CoL, http://www.catalogueoflife.org). The CoL is the most comprehensive and authoritative global index of species currently available. Nonetheless, we supplemented the CoL data with data from the List of Prokaryotic names with Standing in Nomenclature (LPSN, lpsn.dsmz.de). This supplementation is needed until the CoL+ project is finished, which we await.

Click here for more information about the included taxa. Check which versions of the CoL and LSPN were included in this package with catalogue\_of\_life\_version().

#### Source

- Becker K *et al.* Coagulase-Negative Staphylococci. 2014. Clin Microbiol Rev. 27(4): 870–926; doi: 10.1128/CMR.0010913
- 2. Becker K *et al.* 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). 2019. Clin Microbiol Infect; doi: 10.1016/j.cmi.2019.02.028
- 3. Becker K *et al.* **Emergence of coagulase-negative staphylococci** 2020. Expert Rev Anti Infect Ther. 18(4):349-366; doi: 10.1080/14787210.2020.1730813
- 4. Lancefield RC A serological differentiation of human and other groups of hemolytic streptococci. 1933. J Exp Med. 57(4): 571–95; doi: 10.1084/jem.57.4.571
- 5. Catalogue of Life: Annual Checklist (public online taxonomic database), http://www.catalogueoflife.org (check included annual version with catalogue\_of\_life\_version()).

# Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

### Read more on our website!

On our website https://msberends.github.io/AMR/ you can find a comprehensive tutorial about how to conduct AMR analysis, the complete documentation of all functions and an example analysis using WHONET data. As we would like to better understand the backgrounds and needs of our users, please participate in our survey!

# See Also

microorganisms

# **Examples**

```
# taxonomic tree ------
mo_kingdom("E. coli")  # "Bacteria"
mo_phylum("E. coli")  # "Proteobacteria"
mo_class("E. coli")  # "Gammaproteobacteria"
mo_order("E. coli")  # "Enterobacterales"
mo_family("E. coli")  # "Enterobacteriaceae"
```

```
mo_genus("E. coli") # "Escherichia"
mo_species("E. coli") # "coli"
mo_subspecies("E. coli") # ""
# colloquial properties ------
mo_name("E. coli")  # "Escherichia coli"  mo_fullname("E. coli")  # "Escherichia coli" - same as mo_name()  mo_shortname("E. coli")  # "E. coli"
# other properties ------
mo_gramstain("E. coli")  # "Gram-negative"
mo_snomed("E. coli")  # 112283007, 116395006, ... (SNOMED codes)
mo_type("E. coli")  # "Bacteria" (equal to kingdom, but may be translated)
mo_rank("E. coli")  # "species"
mo_url("E. coli")  # get the direct url to the online database entry
mo_synonyms("E. coli")  # get previously accepted taxonomic names
# scientific reference ------
mo_ref("E. coli")  # "Castellani et al., 1919"
mo_authors("E. coli")  # "Castellani et al."
mo_year("E. coli")  # 1919
mo_year("E. coli")
                               # 1919
# abbreviations known in the field ------
mo_genus("MRSA") # "Staphylococcus"
mo_species("MRSA") # "aureus"
mo_shortname("VISA") # "S. aureus"
mo_gramstain("VISA") # "Gram-positive"
mo_genus("EHEC")
                               # "Escherichia"
                                # "coli"
mo_species("EHEC")
# known subspecies ------
mo_name("doylei")  # "Campylobacter jejuni doylei"
mo_genus("doylei")  # "Campylobacter"
mo_species("doylei")  # "jejuni"
mo_subspecies("doylei")  # "doylei"
mo_fullname("K. pneu rh")  # "Klebsiella pneumoniae rhinoscleromatis"
mo_shortname("K. pneu rh") # "K. pneumoniae"
# Becker classification, see ?as.mo -------
mo_fullname("S. epi")
                                           # "Staphylococcus epidermidis"
mo_shortname("S. epi", Becker = TRUE) # "CoNS"
# Lancefield classification, see ?as.mo -----
mo_fullname("S. pyo") # "Streptococcus pyogenes"
mo_fullname("S. pyo", Lancefield = TRUE) # "Streptococcus group A"
mo_shortname("S. pyo") # "S. pyogenes"
mo_shortname("S. pyo", Lancefield = TRUE) # "GAS" (='Group A Streptococci')
# language support ------
mo_gramstain("E. coli", language = "de") # "Gramnegativ"
mo_gramstain("E. coli", language = "nl") # "Gram-negatief"
```

mo\_source 99

```
mo_gramstain("E. coli", language = "es") # "Gram negativo"
# mo_type is equal to mo_kingdom, but mo_kingdom will remain official
                                       # "Bacteria" on a German system
mo_kingdom("E. coli")
                                       # "Bakterien" on a German system
mo_type("E. coli")
mo_type("E. coli")
                                       # "Bacteria" on an English system
mo_fullname("S. pyogenes",
           Lancefield = TRUE.
           language = "de")
                                     # "Streptococcus Gruppe A"
mo_fullname("S. pyogenes",
           Lancefield = TRUE,
           language = "nl")
                                       # "Streptococcus groep A"
# other ------
# gram stains and intrinsic resistance can also be used as a filter in dplyr verbs
if (require("dplyr")) {
 example_isolates %>%
   filter(mo_is_gram_positive())
 example_isolates %>%
   filter(mo_is_intrinsic_resistant(ab = "vanco"))
# get a list with the complete taxonomy (from kingdom to subspecies)
mo_taxonomy("E. coli")
# get a list with the taxonomy, the authors, Gram-stain and URL to the online database
mo_info("E. coli")
```

mo\_source

User-defined reference data set for microorganisms

# 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"))
```

mo\_source

## **Arguments**

path location of your reference file, see Details. Can be "", NULL or FALSE to delete the reference file.

destination destination of the compressed data file, default to 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 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 as an R option, using 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).

# How to setup

Imagine this data on a sheet of an Excel file (mo codes were looked up in the microorganisms data set). The first column contains the organisation specific codes, the second column contains an MO code from this package:

	I	Α	I	В	١
	٠   -		٠ [ -		
1		Organisation XYZ		mo	١
2		lab_mo_ecoli		B_ESCHR_COLI	
3		lab_mo_kpneumoniae		B_KLBSL_PNMN	
4	Ι		Ι		I

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"
```

mo\_source 101

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:

```
| A | B |

--|------|------|

1 | Organisation XYZ | mo |

2 | lab_mo_ecoli | B_ESCHR_COLI |

3 | lab_mo_kpneumoniae | B_KLBSL_PNMN |

4 | lab_Staph_aureus | B_STPHY_AURS |

5 | |
```

...any new usage of an MO function in this package will update your data file:

```
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.

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

102 pca

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

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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 $\boldsymbol{x}$ to be selected for PCA, can be unquoted since it supports quasiquotation.
retx	a logical value indicating whether the rotated variables should be returned.
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 $x$ 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 x can be supplied. The value is passed to scale.
tol	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

pca 103

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 principal components to be used. Can be set as alternative or in addition to tol, useful notably when the desired rank is considerably smaller than the dimensions of the matrix.

### **Details**

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

# **Maturing lifecycle**

The lifecycle of this function is **maturing**. The unlying code of a maturing function has been roughed out, but finer details might still change. Since this function needs wider usage and more extensive testing, you are very welcome to suggest changes at our repository or write us an email (see section 'Contact Us').

# **Examples**

```
# `example_isolates` is a dataset 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
                                        # then get resistance of all drugs
   summarise_if(is.rsi, resistance)
  # now conduct PCA for certain antimicrobial agents
  pca_result <- resistance_data %>%
   pca(AMC, CXM, CTX, CAZ, GEN, TOB, TMP, SXT)
  pca_result
  summary(pca_result)
  biplot(pca_result)
  ggplot_pca(pca_result) # a new and convenient plot function
```

proportion

Calculate microbial 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, please 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)
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_locale(),
 minimum = 30,
 as_percent = FALSE,
 combine_SI = TRUE,
  combine_IR = FALSE
)
rsi_df(
 data,
  translate_ab = "name",
 language = get_locale(),
 minimum = 30,
 as_percent = FALSE,
 combine_SI = TRUE,
  combine_IR = FALSE
)
```

## **Arguments**

one or more vectors (or columns) with antibiotic interpretations. They will be transformed internally with as.rsi() if needed. Use multiple columns to calculate (the lack of) co-resistance: the probability where one of two drugs have a resistant or supportible result. See Examples

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 *Combina*-

tion therapy below

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

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, defaults to system language (see get\_locale())

and can also be set with getOption("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). This used to be the argument combine\_IR, but this now follows the redefinition by EUCAST about the interpretation of I (increased exposure) in 2019, see section 'Interpretation

of S, I and R' below. Default is TRUE.

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

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

outdated, see argument combine\_SI.

# Details

The function resistance() is equal to the function proportion\_R(). The function susceptibility() is equal to the function proportion\_SI().

Remember that you should filter your table 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.

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 being dependent on the minimum argument).

The function proportion\_df() takes any variable from data that has an rsi class (created with as.rsi()) and calculates the proportions R, I and S. It also supports grouped variables. The function rsi\_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 denominator		include as denominator	
S or I	S or I	Χ	Х	X	Х	
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.

# Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

# Interpretation of R and S/I

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

#### • 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.

#### • S = Susceptible

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 = Increased exposure, but still susceptible

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.

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

## Read more on our website!

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#### Source

M39 Analysis and Presentation of Cumulative Antimicrobial Susceptibility Test Data, 4th Edition, 2014, 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.
?example_isolates
resistance(example_isolates$AMX)
                                     # determines %R
susceptibility(example_isolates$AMX) # determines %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)
if (require("dplyr")) {
  example_isolates %>%
    group_by(hospital_id) %>%
    summarise(r = resistance(CIP),
              n = n_rsi(CIP)) # n_rsi works like n_distinct in dplyr, see ?n_rsi
  example_isolates %>%
    group_by(hospital_id) %>%
    summarise(R = resistance(CIP, as_percent = TRUE),
              SI = susceptibility(CIP, as_percent = TRUE),
              n1 = count_all(CIP), # the actual total; sum of all three
```

108 random

```
n2 = n_rsi(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)
                                         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)
# 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),
           proportion = susceptibility(AMC, GEN, only_all_tested = TRUE))
example_isolates %>%
 group_by(hospital_id) %>%
 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 rsi columns
example_isolates %>%
  select(AMX, CIP) %>%
 proportion_df(translate = FALSE)
# It also supports grouping variables
example_isolates %>%
  select(hospital_id, AMX, CIP) %>%
 group_by(hospital_id) %>%
 proportion_df(translate = FALSE)
```

random

}

Random MIC values/disk zones/RSI generation

# **Description**

These functions can be used for generating random MIC values and disk diffusion diameters, for AMR analysis practice.

random 109

#### Usage

```
random_mic(size, mo = NULL, ab = NULL, ...)
random_disk(size, mo = NULL, ab = NULL, ...)
random_rsi(size, prob_RSI = c(0.33, 0.33, 0.33), ...)
```

#### **Arguments**

size	desired size of the returned vector
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 agent code with as . ab()
	extension for future versions, not used at the moment
prob_RSI	a vector of length 3: the probabilities for R (1st value), S (2nd value) and I (3rd value)

#### **Details**

The base R function sample() is used for generating values.

Generated values are based on the latest EUCAST guideline implemented in the rsi\_translation 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())
```

## **Maturing lifecycle**

The lifecycle of this function is **maturing**. The unlying code of a maturing function has been roughed out, but finer details might still change. Since this function needs wider usage and more extensive testing, you are very welcome to suggest changes at our repository or write us an email (see section 'Contact Us').

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## **Examples**

```
random_mic(100)
random_disk(100)
random_rsi(100)

# make the random generation more realistic by setting a bug and/or drug:
random_mic(100, "Klebsiella pneumoniae")  # range 0.0625-64
random_mic(100, "Klebsiella pneumoniae", "meropenem")  # range 0.0625-16
random_mic(100, "Streptococcus pneumoniae", "meropenem")  # range 0.0625-4

random_disk(100, "Klebsiella pneumoniae")  # range 11-50
```

```
random_disk(100, "Klebsiella pneumoniae", "ampicillin")  # range 6-14
random_disk(100, "Streptococcus pneumoniae", "ampicillin") # range 16-22
```

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(),
)
rsi_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_rsi_predict(
  main = paste("Resistance Prediction of", x_name),
```

```
ribbon = TRUE,
...
```

## **Arguments**

X	a data.frame containing isolates. Can be left blank when used inside dplyr verbs, such as filter(), mutate() and summarise().	
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, defaults to 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, defaults to 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 Details 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_measurements		
	a logical 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	

## **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, please see *Examples*.

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## Interpretation of R and S/I

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

## • 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.

# • S = Susceptible

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 = Increased exposure, but still susceptible

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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#### See Also

The proportion() functions to calculate resistance Models: lm() glm()

## **Examples**

```
x <- resistance_predict(example_isolates,</pre>
                        col_ab = "AMX",
                        year_min = 2010,
                        model = "binomial")
plot(x)
if (require("ggplot2")) {
  ggplot_rsi_predict(x)
# using dplyr:
if (require("dplyr")) {
  x <- example_isolates %>%
    filter_first_isolate() %>%
    filter(mo_genus(mo) == "Staphylococcus") %>%
    resistance_predict("PEN", model = "binomial")
  plot(x)
  # get the model from the object
  mymodel \leftarrow attributes(x) \mod el
  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)
  ggplot(data,
         aes(x = year)) +
    geom\_col(aes(y = value),
             fill = "grey75") +
    geom_errorbar(aes(ymin = se_min,
                      ymax = se_max),
                  colour = "grey50") +
    scale_y_continuous(limits = c(0, 1),
                       breaks = seq(0, 1, 0.1),
                       labels = paste0(seq(0, 100, 10), "%")) +
    labs(title = expression(paste("Forecast of Amoxicillin Resistance in ",
                                   italic("E. coli"))),
         y = "%R",
         x = "Year") +
    theme_minimal(base_size = 13)
}
```

114 rsi\_translation

rsi\_translation

Data set for R/SI interpretation

## **Description**

Data set to interpret MIC and disk diffusion to R/SI values. Included guidelines are CLSI (2010-2019) and EUCAST (2011-2020). Use as.rsi() to transform MICs or disks measurements to R/SI values.

# Usage

```
rsi_translation
```

#### **Format**

A data.frame with 18,650 observations and 10 variables:

- guideline Name of the guideline
- method Either "MIC" or "DISK"
- site Body site, e.g. "Oral" or "Respiratory"
- mo Microbial ID, see as.mo()
- ab
   Antibiotic ID, 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**

The repository of this AMR package contains a file comprising this exact data set: https://github.com/msberends/AMR/blob/master/data-raw/rsi\_translation.txt. This file allows for machine reading EUCAST and CLSI guidelines, which is almost impossible with the Excel and PDF files distributed by EUCAST and CLSI. The file is updated automatically.

skewness 115

## Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

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#### See Also

 $intrinsic\_resistant$ 

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.

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 computation proceeds

116 translate

#### Stable lifecycle

The lifecycle of this function is **stable**. In a stable function, major changes are unlikely. This means that the unlying code will generally evolve by adding new arguments; removing arguments or changing the meaning of existing arguments will be avoided.

If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

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#### See Also

kurtosis()

translate

Translate strings from AMR package

#### **Description**

For language-dependent output of AMR functions, like mo\_name(), mo\_gramstain(), mo\_type() and ab\_name().

# Usage

get\_locale()

## **Details**

Strings will be translated to foreign languages if they are defined in a local translation file. Additions to this file can be suggested at our repository. The file can be found here: https://github.com/msberends/AMR/blob/master/data-raw/translations.tsv. This file will be read by all functions where a translated output can be desired, like all mo\_\* functions (such as mo\_name(), mo\_gramstain(), mo\_type(), etc.) and ab\_\* functions (such as ab\_name(), ab\_group(), etc.).

Currently supported languages are: Dutch, English, French, German, Italian, Portuguese, Spanish. Please note that currently not all these languages have translations available for all antimicrobial agents and colloquial microorganism names.

Please suggest your own translations by creating a new issue on our repository.

## 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()), if that language is supported. But the language to be used can be overwritten in two ways and will be checked in this order:

- 1. Setting the R option AMR\_locale, e.g. by running options(AMR\_locale = "de")
- 2. Setting the system variable LANGUAGE or LANG, e.g. by adding LANGUAGE="de\_DE.utf8" to your .Renviron file in your home directory

So if the R option AMR\_locale is set, the system variables LANGUAGE and LANG will be ignored.

WHOCC 117

#### Stable lifecycle

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If the unlying code needs breaking changes, they will occur gradually. For example, a argument will be deprecated and first continue to work, but will emit an message informing you of the change. Next, typically after at least one newly released version on CRAN, the message will be transformed to an error.

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# **Examples**

```
# The 'language' argument of below functions
# will be set automatically to your system language
# with get_locale()
# English
mo_name("CoNS", language = "en")
#> "Coagulase-negative Staphylococcus (CoNS)"
# German
mo_name("CoNS", language = "de")
#> "Koagulase-negative Staphylococcus (KNS)"
# Dutch
mo_name("CoNS", language = "nl")
#> "Coagulase-negatieve Staphylococcus (CNS)"
# Spanish
mo_name("CoNS", language = "es")
#> "Staphylococcus coagulasa negativo (SCN)"
# Italian
mo_name("CoNS", language = "it")
#> "Staphylococcus negativo coagulasi (CoNS)"
# Portuguese
mo_name("CoNS", language = "pt")
#> "Staphylococcus coagulase negativo (CoNS)"
```

WHOCC: WHO Collaborating Centre for Drug Statistics Methodology

118 WHONET

#### **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 (http://ec.europa.eu/health/documents/community-register/html/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/.

#### Read more on our website!

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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 are created using online surname generators and are only in place for practice purposes.

# Usage

WHONET

WHONET 119

#### **Format**

A data.frame 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()

• 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?

ESBL

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.rsi().

## Reference data publicly available

All reference data sets (about microorganisms, antibiotics, R/SI interpretation, EUCAST rules, etc.) in this AMR package are publicly and freely available. We continually export our data sets to formats for use in R, SPSS, SAS, Stata and Excel. We also supply flat files that are machine-readable and suitable for input in any software program, such as laboratory information systems. Please find all download links on our website, which is automatically updated with every code change.

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# Index

* Becker	ab_info(), 6
as.mo, 22	ab_loinc(ab_property), 5
* Lancefield	ab_loinc(), <i>13</i>
as.mo, 22	ab_name (ab_property), 5
* becker	
	ab_name(), 4, 50, 68, 116
as.mo, 22 * datasets	ab_name(AMP), 120
	ab_property, 5
antibiotics, 12	ab_property(), 3, 13, 42, 68, 105
example_isolates, 50	ab_synonyms (ab_property), 5
example_isolates_unclean, 51	$ab\_synonyms(), 6$
intrinsic_resistant, 72	ab_tradenames (ab_property), 5
microorganisms, 87	ab_tradenames(), 6
microorganisms.codes, 89	ab_url (ab_property), 5
microorganisms.old, 90	ab_url(), <u>6</u>
rsi_translation, 114	age, 8
WHONET, 118	age(), 9, 10
* guess	age_groups, 9
as.mo, 22	age_groups(), 8, 119
* lancefield	aminoglycosides
as.mo, 22	(antibiotic_class_selectors),
* <b>mo</b>	15
as.mo, 22	aminoglycosides(), 16
%like% (like), 80	AMR, 11
%like_case% (like), 80	anti_join_microorganisms(join), 73
3MRGN (mdro), 82	antibiotic_class_selectors, 15
4MRGN (mdro), 82	antibiotic_class_selectors(), 53
-l- 10	antibiotics, 3, 5–7, 12, 12, 13, 16, 17, 19,
ab, 18	37, 42, 50, 52, 68, 70, 71, 105, 120
ab (as.ab), 17	antivirals, <i>13</i>
ab_*, 4, 12, 18, 116	antivirals (antibiotics), 12
ab_atc (ab_property), 5	as.ab, 17
ab_atc_group1 (ab_property), 5	
ab_atc_group2 (ab_property), 5	as.ab(), 3, 5, 6, 12, 17, 18, 29, 71, 94, 109,
ab_cid(ab_property), 5	
$ab\_cid(), 6$	as.character(), 81
ab_class(antibiotic_class_selectors),	as.Date(), 62
15	as.disk,19
ab_ddd (ab_property), 5	as.disk(), 30, 32, 109
$ab\_ddd(), 6$	as.mic, 21
ab_from_text, 3	as.mic(), $30$ , $32$ , $109$
ab_from_text(), 19	as.mo, 22
ab_group(ab_property), 5	as.mo(), 24, 25, 29, 32, 37, 39, 45, 50, 51, 55,
ab_group(), 4, 116	73, 75, 83, 87, 89–96, 99–101, 109,
ab_info(ab_property),5	114, 119

as.POSIXlt(),8	count_IR (count), 41
as.rsi,27	count_R (count), 41
as.rsi(), 20-22, 29, 30, 42, 46, 50, 51, 67,	count_R(), <i>42</i>
68, 105, 114, 120	<pre>count_resistant (count), 41</pre>
ATC (ab_property), 5	<pre>count_resistant(), 41, 42</pre>
atc_online_ddd(atc_online_property), 33	count_S (count), 41
atc_online_groups	count_SI (count), 41
(atc_online_property), 33	count_SI(), 31, 42, 43, 86, 107, 112
atc_online_property, 33	count_susceptible (count), 41
availability, 35	count_susceptible(), 31, 41-43, 86, 107,
biplot(), <i>63</i>	
BRMO (mdro), 82	data.frame, 13, 19, 23, 24, 27, 29, 35, 37, 42,
brmo (mdro), 82	47, 50, 51, 55, 67, 71, 72, 75, 79, 83,
browseURL(), 94	87, 89, 90, 100, 102, 103, 105, 111,
bug_drug_combinations, 36	114, 115, 119
bug_drug_combinations(),37	disk, 19, 20, 29, 30
	disk (as.disk), 19
carbapenems	double, 6, 8, 62, 95, 105
<pre>(antibiotic_class_selectors),</pre>	, , , , ,
15	EUCAST (eucast_rules), 45
catalogue_of_life,38	eucast_exceptional_phenotypes (mdro), 82
catalogue_of_life_version,40	eucast_rules, 45
catalogue_of_life_version(), 25, 26, 38,	eucast_rules(), 30, 47, 85
40, 88–91, 97	example_isolates, 50, 118
cephalosporins	example_isolates_unclean, 51
(antibiotic_class_selectors),	/
15	<pre>facet_rsi (ggplot_rsi), 66</pre>
cephalosporins_1st	facet_rsi(), 69
(antibiotic_class_selectors),	factor, 9, 21, 31, 84
15	filter(), 55, 62, 83, 94, 111
cephalosporins_2nd	filter_1st_cephalosporins
(antibiotic_class_selectors),	(filter_ab_class), 52
15	filter_2nd_cephalosporins
cephalosporins_3rd	(filter_ab_class), 52
(antibiotic_class_selectors),	filter_3rd_cephalosporins
15	(filter_ab_class), 52
cephalosporins_4th	filter_4th_cephalosporins
(antibiotic_class_selectors),	(filter_ab_class), 52
15	filter_5th_cephalosporins
cephalosporins_5th	(filter_ab_class), 52
(antibiotic_class_selectors),	filter_ab_class, 52
15	filter_ab_class(), 16
character, 4, 6, 18, 24, 25, 74, 81, 95, 105	filter_aminoglycosides
chisq.test(), $58-60$	(filter_ab_class), 52
Click here, 26, 38, 40, 88, 90, 91, 97	filter_aminoglycosides(), 53
	· · · · · · · · · · · · · · · · · ·
count, 41	filter_carbapenems (filter_ab_class), 52
count(), 105, 107	filter_cephalosporins  (filter_ch_close) 52
count_all(count), 41	(filter_ab_class), 52
count_all(), 42	filter_first_isolate (first_isolate), 54
count_df (count), 41	filter_first_isolate(), 55, 56
count_df(), 42, 68	filter_first_weighted_isolate
count_I (count), 41	(first_isolate), 54

filter_first_weighted_isolate(), 55, 56	intrinsic_resistant, 15, 29, 72, 89, 95, 115
filter_fluoroquinolones	is.ab (as.ab), 17
(filter_ab_class), 52	is.disk (as.disk), 19
filter_glycopeptides (filter_ab_class),	is.mic (as.mic), 21
52	is.mo (as.mo), 22
filter_macrolides (filter_ab_class), 52	is.rsi (as.rsi), 27
filter_penicillins (filter_ab_class), 52	is.rsi.eligible(), 30
filter_tetracyclines (filter_ab_class),	is_new_episode (get_episode), 61
52	is_new_episode(), <i>54</i> , <i>56</i> , <i>61</i> , <i>62</i>
first_isolate, 54	iain 72
first_isolate(), 55, 56, 62, 75–77, 105	join, 73
fisher.test(), 59	key_antibiotics, 74
fluoroquinolones	key_antibiotics(), 55–57, 76, 77
<pre>(antibiotic_class_selectors),</pre>	key_antibiotics_equal
15	(key_antibiotics), 74
format(), 36, 37	key_antibiotics_equal(), 76, 77
format.bug_drug_combinations	kurtosis, 78
(bug_drug_combinations), 36	kurtosis(), <i>116</i>
full_join_microorganisms(join), 73	Kui tosis(), 110
g.test, 58	<pre>labels_rsi_count (ggplot_rsi), 66</pre>
g.test(), 58	labels_rsi_count(), 68, 69
geom_rsi (ggplot_rsi), 66	<pre>left_join_microorganisms(join), 73</pre>
geom_rsi(), 68	lifecycle, 4, 6, 8, 10, 18, 20, 21, 25, 31, 34,
get_episode, 61	36, 37, 42, 48, 53, 57, 60, 62, 65, 69,
get_episode(), <i>61</i> , <i>62</i>	71, 74, 77, 79, 79, 80, 81, 84, 93, 95,
get_locale (translate), 116	101, 103, 106, 109, 112, 116, 117
get_locale(), 6, 23, 37, 42, 68, 94, 105	like, 80
get_mo_source (mo_source), 99	list, 3, 4, 6, 35, 40, 95
<pre>get_mo_source(), 23, 100</pre>	lm(), 113
ggplot2, 66, 68	logical, 9, 29, 46, 56, 62, 77, 80, 81, 83
<pre>ggplot2::facet_wrap(), 69</pre>	
<pre>ggplot2::geom_text(), 69</pre>	macrolides
<pre>ggplot2::scale_fill_manual(), 69</pre>	<pre>(antibiotic_class_selectors),</pre>
<pre>ggplot2::scale_y_continuous(), 69</pre>	15
ggplot2::theme(), 69	matrix, <i>79</i> , <i>115</i>
ggplot_pca, 63	MDR (mdro), 82
ggplot_pca(), 65, 103	mdr_cmi2012 (mdro), 82
ggplot_rsi,66	mdr_cmi2012(), 84
ggplot_rsi(), 69	mdr_tb (mdro), 82
ggplot_rsi_predict	mdr_tb(), <i>84</i>
(resistance_predict), 110	mdro, 82
glm(), 113	mdro(), 47, 83-85
glycopeptides	merge(), 74
(antibiotic_class_selectors),	mic, 21, 29, 30
15	mic(as.mic), 21
grep(), 80, 81	microorganisms, 15, 24, 25, 27, 39-41, 50,
guess_ab_col, 70	51, 73, 87, 89–91, 94, 97, 100
guess_ab_col(), 47, 75, 76, 84	microorganisms.codes, 89,89
	microorganisms.old, $25$ , $90$
inner_join(join), 73	microorganisms\$fullname, 92
<pre>inner_join_microorganisms(join), 73</pre>	microorganisms\$mo, 100
integer, 6, 8, 20, 42, 95	mo, 22, 23, 25, 29, 37, 45, 55, 73, 75, 83

mo (as.mo), 22	<pre>mo_uncertainties(), 24</pre>
mo_*, 24, 25, 27, 89, 91, 92, 96, 99, 101, 116	<pre>mo_url (mo_property), 93</pre>
<pre>mo_authors (mo_property), 93</pre>	$mo\_url(), 95$
mo_authors(), 95	<pre>mo_year (mo_property), 93</pre>
mo_class (mo_property), 93	mo_year(), 95
<pre>mo_domain (mo_property), 93</pre>	mrgn (mdro), 82
$mo\_domain(), 95$	mrgn(), 84
mo_failures (as.mo), 22	mutate(), 55, 62, 83, 94, 111
mo_failures(), 24	
mo_family(mo_property), 93	n_rsi(count),41
mo_fullname (mo_property), 93	n_rsi(), <i>4</i> 2
mo_genus (mo_property), 93	
mo_genus(), 27, 99, 100	pca, 102, <i>103</i>
mo_gramstain (mo_property), 93	pca(), <i>64</i> , <i>65</i> , <i>103</i>
mo_gramstain(), 27, 95, 99, 100, 116	PDR ( $mdro$ ), $82$
mo_info (mo_property), 93	penicillins
mo_info(), 95	<pre>(antibiotic_class_selectors),</pre>
mo_is_gram_negative (mo_property), 93	15
mo_is_gram_negative(), 95	<pre>plot.resistance_predict</pre>
mo_is_gram_positive (mo_property), 93	(resistance_predict), 110
mo_is_gram_positive(), 95	portion (proportion), 104
	prcomp, <i>103</i>
mo_is_intrinsic_resistant	prcomp(), 64, 103
(mo_property), 93	princomp, 64
mo_is_intrinsic_resistant(), 95	princomp(), 64
mo_kingdom (mo_property), 93	proportion, 104
mo_kingdom(), 95	<pre>proportion(), 113</pre>
mo_matching_score, 91	proportion_*,44
mo_matching_score(), 25, 92, 96	proportion_df (proportion), 104
mo_name (mo_property), 93	proportion_df(), 105
mo_name(), 116	proportion_I (proportion), 104
mo_order (mo_property), 93	proportion_IR (proportion), 104
<pre>mo_phylum (mo_property), 93</pre>	proportion_R (proportion), 104
mo_property, 93	proportion_R(), $105$
mo_property(), 87, 89, 91	proportion_S (proportion), 104
mo_rank (mo_property), 93	proportion_SI (proportion), 104
<pre>mo_ref (mo_property), 93</pre>	proportion_SI(), 31, 43, 86, 105, 107, 112
$mo\_ref(), 95$	F
mo_renamed(as.mo), 22	random, 108
mo_renamed(), 24	random_disk(random), 108
mo_shortname (mo_property), 93	random_disk(), 109
mo_shortname(), <i>37</i> , <i>95</i>	random_mic(random), 108
<pre>mo_snomed (mo_property), 93</pre>	random_mic(), <i>109</i>
$mo\_snomed(), 87, 95$	random_rsi (random), 108
mo_source, 99	readRDS(), 100
<pre>mo_species (mo_property), 93</pre>	resistance (proportion), 104
<pre>mo_subspecies (mo_property), 93</pre>	resistance(), 35, 42, 104, 105
mo_synonyms (mo_property), 93	resistance_predict, 110, 111
mo_taxonomy (mo_property), 93	right_join_microorganisms(join), 73
mo_taxonomy(), 95	rsi, 28, 31, 42, 50, 67, 68, 105
mo_type (mo_property), 93	rsi (as.rsi), 27
mo_type(), 116, 119	rsi_df (proportion), 104
mo_uncertainties (as.mo), 22	rsi_df(), 42, 68, 105
• • • • • • • • • • • • • • • • • • • •	TT I I I

```
rsi_predict(resistance_predict), 110
rsi_translation, 29, 109, 114
sample(), 109
scale, 102
scale_rsi_colours (ggplot_rsi), 66
scale_rsi_colours(), 69
scale_y_percent (ggplot_rsi), 66
scale_y_percent(),69
semi_join_microorganisms(join), 73
set_mo_source (mo_source), 99
set_mo_source(), 23, 89, 100, 101
skewness, 115
skewness(), 79
summarise(), 55, 62, 83, 94, 111
summary(), 111
susceptibility (proportion), 104
susceptibility(), 31, 35, 42, 43, 86,
         104–107, 112
Sys.getlocale(), 116
tetracyclines
        (antibiotic\_class\_selectors),
        15
theme_rsi (ggplot_rsi), 66
theme_rsi(), 69
translate, 6, 94, 95, 116
utils::browseURL(), 6
variable grouping, 62
vector, 18, 24, 25
WHOCC, 117
WHONET, 118
write us an email (see section
         'Contact Us'), 4, 65, 69, 80, 103,
        109, 112
XDR (mdro), 82
```