

Package ‘MIC’

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Title Analysis of Antimicrobial Minimum Inhibitory Concentration Data

Version 2.0.0

Description Analyse, plot, and tabulate antimicrobial minimum inhibitory concentration (MIC) data. Validate the results of an MIC experiment by comparing observed MIC values to a gold standard assay, in line with standards from the International Organization for Standardization (2021) <<https://www.iso.org/standard/79377.html>>.

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as.sir_vectorised	<i>Convert MIC or Disk Diffusion to SIR, vectorised over antimicrobials</i>
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Description

The AMR::as.sir function is not vectorised over antimicrobials. This function provides vectorisation over antimicrobials. Due to the overhead of running AMR::as.sir, this function tries to be efficient by only running AMR::as.sir as little as necessary.

Usage

```
as.sir_vectorised(mic, mo, ab, accept_ecoff = FALSE, ...)
```

Arguments

mic	vector of MIC values
mo	vector of microorganism names
ab	vector of antibiotic names
accept_ecoff	if TRUE, ECOFFs will be used when no clinical breakpoints are available
...	additional arguments that are passed to AMR::as.sir

Value

S3 sir values

Examples

```
mic <- c("<0.25", "8", "64", ">64")
mo <- c("B_ESCHR_COLI", "B_ESCHR_COLI", "B_ESCHR_COLI", "B_ESCHR_COLI")
ab <- c("AMK", "AMK", "AMK", "AMK")
as.sir_vectorised(mic, mo, ab)
# using different microorganisms and antibiotics
mic <- c("<0.25", "8", "64", ">64")
mo <- c("B_ESCHR_COLI", "B_ESCHR_COLI", "B_PROTS_MRBL", "B_PROTS_MRBL")
ab <- c("AMK", "AMK", "CIP", "CIP")
as.sir_vectorised(mic, mo, ab)
```

bias	<i>Calculate MIC bias</i>
------	---------------------------

Description

Calculate the bias between two AMR::mic vectors. The bias is calculated as the percentage of test MICs that are above the gold standard MICs minus the percentage of test MICs that are below the gold standard MICs.

Usage

```
bias(gold_standard, test)
```

Arguments

```
gold_standard  AMR::mic vector
test           AMR::mic vector
```

Value

numeric value

References

International Organization for Standardization. ISO 20776-2:2021 Available from: <https://www.iso.org/standard/79377.html>

Examples

```
gold_standard <- c("<0.25", "8", "64", ">64")
test <- c("<0.25", "2", "16", "64")
bias(gold_standard, test)
```

clean_raw_mic	<i>Clean up raw MIC for use as a feature</i>
---------------	--

Description

Removes leading "=" which can sometimes be present in raw MIC results. Also converts co-trimoxazole to trimethprim component only.

Usage

```
clean_raw_mic(mic)
```

Arguments

mic	character containing MIC/s
-----	----------------------------

Value

character of clean MIC/s

Examples

```
clean_raw_mic(c("==>64", "0.25/8.0"))
```

compare_mic	<i>Compare and validate MIC values</i>
-------------	--

Description

This function compares an vector of MIC values to another. Generally, this is in the context of a validation experiment – an investigational assay or method (the "test") is compared to a gold standard. The rules used by this function are in line with "ISO 20776-2:2021 Part 2: Evaluation of performance of antimicrobial susceptibility test devices against reference broth micro-dilution."

There are two levels of detail that are provided. If only the MIC values are provided, the function will look for essential agreement between the two sets of MIC. If the organism and antibiotic arguments are provided, the function will also calculate the categorical agreement using EUCAST breakpoints (or, if breakpoint not available and accept_ecoff = TRUE, ECOFFs).

The function returns a special dataframe of results, which is also an mic_validation object. This object can be summarised using summary() for summary metrics, plotted using plot() for an essential agreement confusion matrix, and tabulated using table().

Usage

```
compare_mic(
  gold_standard,
  test,
  ab = NULL,
  mo = NULL,
  accept_ecoff = FALSE,
  simplify = TRUE,
  ea_mode = "categorical",
  tolerate_censoring = "gold_standard",
  tolerate_matched_censoring = "both",
  tolerate_leq = TRUE,
  tolerate_geq = TRUE,
  ...
)
```

Arguments

gold_standard	vector of MICs to compare against.
test	vector of MICs that are under investigation
ab	character vector (same length as MIC) of antibiotic names (optional)
mo	character vector (same length as MIC) of microorganism names (optional)
accept_ecoff	if TRUE, ECOFFs will be used when no clinical breakpoints are available
simplify	if TRUE, MIC values will be coerced into the closest halving dilution (e.g., 0.55 will be converted to 0.5)
ea_mode	"categorical" or "numeric", see essential_agreement
tolerate_censoring	"strict", "gold_standard", "test", or "both" - how to handle censored data (see essential_agreement for details). Generally, this should be left as "gold_standard" since this setting "tolerates" a test that has higher granularity (i.e., less censoring) than the gold standard. Setting to "test" or "both" should be used with caution but may be appropriate in some cases where the test also produces censored results.
tolerate_matched_censoring	"strict", "gold_standard", "test", or "both" - how to handle situations where one of the values is censored, but both values match (e.g., gold_standard = ">2", test = "2"). Generally, this should be left as "both", since these values are considered to be in essential agreement. For more details, see essential_agreement .
tolerate_leq	whether to tolerate <= in essential agreement, e.g., <=2 and 4 will be considered in essential agreement. See essential_agreement for details.
tolerate_geq	whether to tolerate >= in essential agreement, e.g., >=4 and 2 will be considered in essential agreement. See essential_agreement for details.
...	additional arguments to be passed to <code>AMR::as.sir</code>

Value

S3 mic_validation object

Examples

```
# Just using MIC values only
gold_standard <- c("<0.25", "8", "64", ">64")
test <- c("<0.25", "2", "16", "64")
val <- compare_mic(gold_standard, test)
summary(val)

# Using MIC values and antibiotic and organism names
gold_standard <- c("<0.25", "8", "64", ">64")
test <- c("<0.25", "2", "16", "64")
ab <- c("AMK", "AMK", "AMK", "AMK")
mo <- c("B_ESCHR_COLI", "B_ESCHR_COLI", "B_ESCHR_COLI", "B_ESCHR_COLI")
val <- compare_mic(gold_standard, test, ab, mo)
"error" %in% names(val) # val now has categorical agreement
```

compare_sir

Compare SIR results and generate categorical agreement

Description

Compare two AMR::sir vectors and generate a categorical agreement vector with the following levels: M (major error), vM (very major error), m (minor error). The error definitions are:

1. Major error (M): The test result is resistant (R) when the gold standard is susceptible (S).
2. vM (very major error): The test result is susceptible (S) when the gold standard is resistant (R).
3. Minor error (m): The test result is intermediate (I) when the gold standard is susceptible (S) or resistant (R), or vice versa.

Usage

```
compare_sir(gold_standard, test)
```

Arguments

```
gold_standard  Susceptibility results in AMR::sir format
test           Susceptibility results in AMR::sir format
```

Value

factor vector with the following levels: M, vM, m.

Examples

```
gold_standard <- c("S", "R", "I", "I")
gold_standard <- AMR::as.sir(gold_standard)
test <- c("S", "I", "R", "R")
test <- AMR::as.sir(test)
compare_sir(gold_standard, test)
```

droplevels.mic_validation

Droplevels for MIC validation object

Description

Quite often, MIC values are being compared across methods with different levels of granularity. For example, the true MIC may be measured across a higher range of values than the test method. This means that there may be MIC levels that don't provide much additional information (since they are only present in one of the methods). This function removes these unnecessary levels at both ranges of the MIC values.

This function ensure that the changes do not "change" the essential agreement interpretation. This can be suppressed using `safe = FALSE`, however this is probably not desired behaviour.

Usage

```
## S3 method for class 'mic_validation'
droplevels(x, safe = TRUE, ...)
```

Arguments

<code>x</code>	mic_validation object
<code>safe</code>	ensure that essential agreement is not changed after dropping levels
<code>...</code>	additional arguments

Value

mic_validation object

Examples

```
gold_standard <- c("<0.25", "0.25", "0.5", "1", "2", "1", "0.5")
test <- c("0.004", "0.08", "<0.25", "0.5", "1", "0.5", "0.5")
val <- compare_mic(gold_standard, test)
droplevels(val)
```

ecoffs

ECOFF data

Description

A dataset containing the epidemiological cut-off values (ECOFFs) for different antibiotics and microorganisms. Currently, only the ECOFF values for *Escherichia coli* are included.

Usage

ecoffs

Format

ecoffs:

A data frame with 85 rows and 25 columns:

organism Microorganism code in AMR::mo format

antibiotic Antibiotic code in AMR::ab format

0.002:512 Counts of isolates in each concentration "bin"

Distributions see EUCAST documentation below

Observations Number of observations

(T)ECOFF see EUCAST documentation below

Confidence interval see EUCAST documentation below

Source

EUCAST <https://www.eucast.org/bacteria/mic-and-zone-distributions-ecoffs/>

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essential_agreement *Essential agreement for MIC validation*

Description

Essential agreement calculation for comparing two MIC vectors.

Usage

```
essential_agreement(
  x,
  y,
  coerce_mic = TRUE,
  tolerate_censoring = "strict",
  tolerate_matched_censoring = "both",
  tolerate_leq = TRUE,
  tolerate_geq = TRUE,
  mode = "categorical"
)
```


Arguments

x	AMR::mic or coercible
y	AMR::mic or coercible
coerce_mic	convert to AMR::mic
tolerate_censoring	"strict", "x", "y", or "both" - whether to tolerate censoring in x, y, or both. See details.
tolerate_matched_censoring	"strict", "x", "y", or "both" - how to handle situations where one of the values is censored, but both values match (e.g., x = ">2", y = "2"). For most situations, this is considered essential agreement. so should be left as "both".
tolerate_leq	whether to tolerate \leq in essential agreement, e.g., ≤ 2 and 4 will be considered in essential agreement (because ≤ 2 includes 2mg/L, which is within 1 dilution of 4mg/L). This argument respects the tolerate_censoring argument, so if tolerate_censoring is "strict", this will not be applied.
tolerate_geq	whether to tolerate \geq in essential agreement, e.g., ≥ 4 and 2 will be considered in essential agreement (because ≥ 4 includes 4mg/L, which is within 1 dilution of 2mg/L). This argument respects the tolerate_censoring argument, so if tolerate_censoring is strict, this will not be applied.
mode	"categorical" or "numeric", see details

Details

Essential agreement is a central concept in the comparison of two sets of MIC values. It is most often used when validating a new method against a gold standard. This function reliably performs essential agreement in line with ISO 20776-2:2021. The function can be used in two modes: categorical and numeric. In categorical mode, the function will use traditional MIC concentrations to determine the MIC (therefore it will use `force_mic()` to convert both x and y to a clean MIC – see [force_mic](#)). In numeric mode, the function will compare the ratio of the two MICs, after removing censoring (values that are ">" and "<" are multiplied and divided by 2, respectively — see [mic_uncensor](#)). In most cases, categorical mode provides more reliable results. Values within +/- 1 dilutions are considered to be in essential agreement.

The `tolerate_censoring` argument controls how the function handles censored data. If set to "strict", the function will return NA for any pair of values that are both censored (and not equal). If set to "x" or "y", the function will allow one of the values to be censored and will compare the uncensored value to the other value. When set to "both", the function will allow one of the values to be censored. If using "both" and both values are censored, the function will attempt to determine essential agreement based on the ratio of the two values, but a warning will be raised.

Value

logical vector

References

International Organization for Standardization. ISO 20776-2:2021 Available from: <https://www.iso.org/standard/79377.html>

Examples

```
x <- AMR::as.mic(c("<0.25", "8", "64", ">64"))
y <- AMR::as.mic(c("<0.25", "2", "16", "64"))
essential_agreement(x, y)
# TRUE FALSE FALSE TRUE

# examples using tolerate_censoring
x <- AMR::as.mic("<4")
y <- AMR::as.mic("0.25")

essential_agreement(x, y, tolerate_censoring = "x") # TRUE
essential_agreement(x, y, tolerate_censoring = "y") # FALSE
essential_agreement(x, y, tolerate_censoring = "both") # TRUE (same as "x")

# strict returns FALSE as it wants the censoring cut-offs to be close
essential_agreement(x, y, tolerate_censoring = "strict")
```

example_mics

Example MIC data

Description

Example minimum inhibitory concentration validation data for three antimicrobials on *Escherichia coli* strains. This data is synthetic and generated to give an example of different MIC distribution.

Usage

```
example_mics
```

Format

example_mics:

A data frame with 300 rows and 4 columns:

gs Gold standard MICs

test Test MICs

mo Microorganism code in AMR::mo format

ab Antibiotic code in AMR::ab format

Source

Synthetic data

fill_dilution_levels	<i>Fill MIC dilution levels</i>
----------------------	---------------------------------

Description

Fill MIC dilution levels

Usage

```
fill_dilution_levels(x, cap_upper = TRUE, cap_lower = TRUE, as.mic = TRUE)
```

Arguments

x	MIC vector
cap_upper	If True, will the top level will be the highest MIC dilution in x
cap_lower	If True, will the bottom level will be the lowest MIC dilution in x
as.mic	By default, returns an ordered factor. Set as.mic = TRUE to return as AMR::mic

Value

ordered factor (or AMR::mic if as.mic = TRUE)

Examples

```
# use in combination with droplevels to clean up levels:
x <- AMR::as.mic(c("<0.25", "8", "64", ">64"))
x <- droplevels(x)
fill_dilution_levels(x)
```

force_mic	<i>Force MIC-like into MIC-compatible format</i>
-----------	--

Description

Convert a value that is "almost" an MIC into a valid MIC value.

Usage

```
force_mic(
  value,
  levels_from_AMR = FALSE,
  max_conc = 512,
  min_conc = 0.002,
  method = "closest",
  prefer = "max",
  leq = TRUE,
  geq = NULL
)
```

Arguments

value	vector of MIC-like values (numeric or character)
levels_from_AMR	conform to AMR::as.mic levels
max_conc	maximum concentration to force to
min_conc	minimum concentration to force to
method	method to use when forcing MICs (closest or round_up)
prefer	where value is in between MIC (e.g., 24mg/L) chose the higher MIC ("max") or lower MIC ("min"); only applies to method = "closest"
leq	whether to force \leq for lower censored values (i.e., $<$). If TRUE, then all values below the limit of detection are converted to \leq . If FALSE, then they are converted to $<$. If NULL, they are not changed.
geq	whether to force \geq for higher censored values (i.e., $>$). If TRUE, then all values above the limit of detection are converted to \geq . If FALSE, then they are converted to $>$. If NULL, they are not changed.

Details

Some experimental or analytical conditions measure MIC (or surrogate) in a way that does not fully conform to traditional MIC levels (i.e., concentrations). This function allows these values to be coerced into an MIC value that is compatible with the AMR::mic class. When using method = "closest", the function will choose the closest MIC value to the input value (e.g., 2.45 will be coerced to 2). When using method = "round up", the function will round up to the next highest MIC value (e.g., 2.45 will be coerced to 4). "Round up" is technically the correct approach if the input value was generated from an experiment that censored between concentrations (e.g., broth or agar dilution). However, "closest" may be more appropriate in some cases.

Please note that this function will not make any changes to censored values (beyond some simple cleaning, e.g., ≤ 2 is converted to ≤ 2). This is because it is not possible to make assumptions about censored data.

The leq and geq arguments convert censored values to \leq or \geq . When MIC is measured using an inhibitory dilution method, the lower limit should be reported as \leq (since the lowest dilution could be inhibitory itself), and the upper limit should be reported as $>$ (growth in the highest dilution means that it is not an inhibitory concentration). The default values for leq and geq enforce this.

Value

AMR::as.mic compatible character

Examples

```
force_mic(c("2.32", "<4.12", ">1.01"))
```

get_mic

*Get MIC meta-data from feature database***Description**

This function helps extract MICs from a database of results. It is compatible with the PATRIC meta data format when used on a tidy_patric_db object, created using tidy_patric_db().

If more than one MIC is present for a particular observation, the function can return the higher MIC by setting prefer_high_mic = TRUE. If prefer_high_mic = FALSE, the lower MIC will be returned.

Usage

```
get_mic(
  x,
  ids,
  ab_col,
  id_col = NULL,
  as_mic = TRUE,
  prefer_high_mic = TRUE,
  simplify = TRUE
)
```

Arguments

x	dataframe containing meta-data
ids	vector of IDs to get meta-data for
ab_col	column name containing MIC results
id_col	column name containing IDs
as_mic	return as AMR::as.mic
prefer_high_mic	where multiple MIC results per ID, prefer the higher MIC
simplify	return as vector of MICs (vs dataframe)

Value

vector containing MICs, or dataframe of IDs and MICs

Examples

```
df <- data.frame(genome_id = c("a_12", "b_42", "x_21", "x_21", "r_75"),
  gentamicin = c(0.25, 0.125, 32.0, 16.0, "<0.0125"))
get_mic(df,
  ids = c("b_42", "x_21"),
  ab_col = "gentamicin",
  id_col = "genome_id",
  as_mic = FALSE,
```

```
prefer_high_mic = TRUE,
simplify = TRUE)
```

mic_censor

Censor MIC values

Description

MIC datasets often arise from different laboratories or experimental conditions. In practice, this means that there can be different levels of censoring (\leq and $>$) within the data. This function can be used to harmonise the dataset to a single level of censoring. The function requires a set of rules that specify the censoring levels (see example).

Usage

```
mic_censor(mic, ab = NULL, mo = NULL, rules = NULL, max = Inf, min = -Inf)
```

Arguments

mic	MIC (coercible to AMR::as.mic)
ab	antibiotic name (coercible to AMR::as.ab)
mo	microorganism name (coercible to AMR::as.mo)
rules	censor rules - named list of pathogen (in AMR::as.mo code) to antibiotic (in AMR::as.ab code) to censoring rules. The censoring rules should provide a min or max value to censor MICs to. See example for more.
max	maximum concentration to censor to (default = Inf), will override any rules provided
min	minimum concentration to censor to (default = -Inf), will override any rules provided

Value

censored MIC values (S3 mic class)

Examples

```
example_rules <- list("B_ESCHR_COLI" = list(
  "AMK" = list(min = 2, max = 32),
  "CHL" = list(min = 4, max = 64),
  "GEN" = list(min = 1, max = 16),
  "CIP" = list(min = 0.015, max = 4),
  "MEM" = list(min = 0.016, max = 16),
  "AMX" = list(min = 2, max = 64),
  "AMC" = list(min = 2, max = 64),
  "FEP" = list(min = 0.5, max = 64),
  "CAZ" = list(min = 1, max = 128),
  "TGC" = list(min = 0.25, max = 1)
```

```

    ))

    mic_censor(AMR::as.mic(512),
               "AMK",
               "B_ESCHR_COLI",
               example_rules) == AMR::as.mic(">32")

```

mic_range	<i>Generate dilution series</i>
-----------	---------------------------------

Description

Generate dilution series

Usage

```
mic_range(start = 512, dilutions = Inf, min = 0.002, precise = FALSE)
```

Arguments

start	starting (highest) concentration
dilutions	number of dilutions
min	minimum (lowest) concentration
precise	force range to be high precision (not usually desired behaviour)

Value

Vector of numeric concentrations

Examples

```

mic_range(128)
mic_range(128, dilutions = 21) # same results

```

mic_r_breakpoint	<i>R breakpoint for MIC</i>
------------------	-----------------------------

Description

R breakpoint for MIC

Usage

```
mic_r_breakpoint(mo, ab, accept_ecoff = FALSE, ...)
```

Arguments

mo	mo name (coerced using AMR::as.mo)
ab	ab name (coerced using AMR::as.ab)
accept_ecoff	if TRUE, ECOFFs will be used when no clinical breakpoints are available
...	additional arguments to pass to AMR::as.sir, which is used to calculate the R breakpoint

Value

MIC value

Examples

```
mic_r_breakpoint("B_ESCHR_COLI", "AMK")
mic_r_breakpoint("B_ESCHR_COLI", "CHL", accept_ecoff = TRUE)
```

mic_s_breakpoint	<i>S breakpoint for MIC</i>
------------------	-----------------------------

Description

S breakpoint for MIC

Usage

```
mic_s_breakpoint(mo, ab, accept_ecoff = FALSE, ...)
```

Arguments

mo	mo name (coerced using AMR::as.mo)
ab	ab name (coerced using AMR::as.ab)
accept_ecoff	if TRUE, ECOFFs will be used when no clinical breakpoints are available
...	additional arguments to pass to AMR::as.sir, which is used to calculate the S breakpoint

Value

MIC value

Examples

```
mic_s_breakpoint("B_ESCHR_COLI", "AMK")
mic_s_breakpoint("B_ESCHR_COLI", "CHL", accept_ecoff = TRUE)
```


mic_uncensor

*Uncensor MICs***Description**

Uncensor MICs

Usage

```
mic_uncensor(
  mic,
  method = "scale",
  scale = 2,
  ab = NULL,
  mo = NULL,
  distros = NULL
)
```

Arguments

mic	vector of MICs to uncensor; will be coerced to MIC using <code>AMR::as.mic</code>
method	method to uncensor MICs (scale, simple, or bootstrap)
scale	scalar to multiply or divide MIC by (for method = scale)
ab	antibiotic name (for method = bootstrap)
mo	microorganism name (for method = bootstrap)
distros	dataframe of epidemiological distributions (only used, optionally, for method = bootstrap)

Details

Censored MIC data is generally unsuitable for modelling without some conversion of censored data. The default behaviour (method = scale) is to halve MICs under the limit of detection (\leq) and double MICs above the limit of detection ($>$). When used with method = simple, this function effectively just removes the censoring symbols, e.g., ≤ 2 becomes 2, and > 64 becomes 64.

The bootstrap method is the more complex of the three available methods. It attempts to use a second (uncensored) MIC distribution to sample values in the censored range. These values are then used to populate and uncensor the MIC data provided as input (mic). The second (uncensored) MIC distribution is ideally provided from similar experimental conditions. Alternatively, epidemiological distributions can be used. These distributions should be provided as a dataframe to the distros argument. The format for this dataframe is inspired by the EUCAST epidemiological distributions, see: <https://www.eucast.org/bacteria/mic-and-zone-distributions-ecoffs/>. The dataframe should contain columns for antimicrobial (converted using `AMR::as.ab`), organism (converted using `AMR::as.mo`), and MIC concentrations. An example is provided in the 'ecoffs' dataset available with this package. Currently, only *Escherichia coli* is available in this dataset. Each observation (row) consists of the frequency a particular MIC concentration is observed in the distribution. If such a dataframe is not provided to distros, the function will attempt to use 'ecoffs', but remains limited to *E. coli*.

Value

vector of MICs in AMR::mic format

References

<https://www.eucast.org/bacteria/mic-and-zone-distributions-ecoffs/>

Examples

```
mic_uncensor(c(">64.0", "<0.25", "8.0"), method = "scale", scale = 2)
```

plot.mic_validation	<i>Plot MIC validation results</i>
---------------------	------------------------------------

Description

Plot MIC validation results

Usage

```
## S3 method for class 'mic_validation'
plot(
  x,
  match_axes = TRUE,
  add_missing_dilutions = TRUE,
  facet_wrap_ncol = NULL,
  facet_wrap_nrow = NULL,
  ...
)
```

Arguments

x	object generated using compare_mic
match_axes	Same x and y axis
add_missing_dilutions	Axes will include dilutions that are not
facet_wrap_ncol	Facet wrap into n columns by antimicrobial (optional, only available when more than one antimicrobial in validation)
facet_wrap_nrow	Facet wrap into n rows by antimicrobial (optional, only available when more than one antimicrobial in validation) represented in the data, based on a series of dilutions generated using mic_range().
...	additional arguments

Value

ggplot object

Examples

```
gold_standard <- c("<0.25", "8", "64", ">64")
test <- c("<0.25", "2", "16", "64")
val <- compare_mic(gold_standard, test)
plot(val)

# if the validation contains multiple antibiotics, i.e.,
ab <- c("CIP", "CIP", "AMK", "AMK")
val <- compare_mic(gold_standard, test, ab)
# the following will plot all antibiotics in a single plot (pooled results)
plot(val)
# use the faceting arguments to split the plot by antibiotic
plot(val, facet_wrap(ncol = 2))
```

```
print.mic_validation  Print MIC validation object
```

Description

Print MIC validation object

Usage

```
## S3 method for class 'mic_validation'
print(x, ...)
```

Arguments

x	mic_validation object
...	additional arguments

Value

character

Examples

```
gold_standard <- c("<0.25", "8", "64", ">64")
test <- c("<0.25", "2", "16", "64")
val <- compare_mic(gold_standard, test)
print(val)
```

```
print.mic_validation_summary
      Print MIC validation summary
```

Description

Print MIC validation summary

Usage

```
## S3 method for class 'mic_validation_summary'
print(x, ...)
```

Arguments

x	mic_validation_summary object
...	additional arguments

Value

character

Examples

```
gold_standard <- c("<0.25", "8", "64", ">64")
test <- c("<0.25", "2", "16", "64")
val <- compare_mic(gold_standard, test)
print(summary(val))
```

```
qc_in_range      Check that MIC is within QC range
```

Description

Check whether MIC values are within acceptable range for quality control (QC). Every MIC experiment should include a control strain with a known MIC. The results of the experiment are only valid if the control strain MIC falls within the acceptable range. This function checks whether an MIC result is within the acceptable range given: 1) a control strain (usually identified as an ATCC or NCTC number), 2) an antibiotic name, and 3) a guideline (EUCAST or CLSI). The acceptable range is defined by 'QC_table', which is a dataset which is loaded with this package.

The source of the QC values is the WHONET QC Ranges and Targets available from the 'Antimicrobial Resistance Test Interpretation Engine' (AMRIE) repository: <https://github.com/AClark-WHONET/AMRIE>

Usage

```
qc_in_range(
  measurement,
  strain,
  ab,
  ignore_na = TRUE,
  guideline = "EUCAST",
  year = "2023"
)
```

Arguments

measurement	measured QC MIC
strain	control strain identifier (usually ATCC)
ab	antibiotic name (will be coerced to AMR::as.ab)
ignore_na	ignores NA (returns TRUE)
guideline	Guideline to use (EUCAST or CLSI)
year	Guideline year (version)

Value

logical vector

References

O'Brien TF, Stelling JM. WHONET: An Information System for Monitoring Antimicrobial Resistance. *Emerg Infect Dis.* 1995 Jun;1(2):66–66.

Examples

```
qc_in_range(AMR::as.mic(0.5), 25922, "GEN") == TRUE
qc_in_range(AMR::as.mic(8.0), 25922, "GEN") == FALSE
```

qc_on_target	<i>Check that QC measurement is at the required target</i> [Experimental]
--------------	--

Description

MIC experiments should include a control strain with a known MIC. The MIC result for the control strain should be a particular target MIC. This function checks whether the target MIC was achieved given: 1) a control strain (usually identified as an ATCC or NCTC number), 2) an antibiotic name, and 3) a guideline (EUCAST or CLSI).

Since QC target values are currently not publicly available in an easy to use format, this function takes a pragmatic approach – for most antibiotics and QC strains, the target is assumed to be the midpoint of the acceptable range. This approximation is not necessarily equal to the QC target

reported by guideline setting bodies such as EUCAST. Therefore, this function is considered experimental and should be used with caution.

This function can be used alongside `qc_in_range()`, which checks whether the MIC is within the acceptable range.

The source of the QC values is the WHONET QC Ranges and Targets available from the 'Antimicrobial Resistance Test Interpretation Engine' (AMRIE) repository: <https://github.com/AClark-WHONET/AMRIE>

Usage

```
qc_on_target(
  measurement,
  strain,
  ab,
  ignore_na = TRUE,
  guideline = "EUCAST",
  year = "2023"
)
```

Arguments

<code>measurement</code>	measured QC MIC
<code>strain</code>	control strain identifier (usually ATCC)
<code>ab</code>	antibiotic name (will be coerced to AMR::as.ab)
<code>ignore_na</code>	ignores NA (returns TRUE)
<code>guideline</code>	Guideline to use (EUCAST or CLSI)
<code>year</code>	Guideline year (version)

Value

logical vector

References

O'Brien TF, Stelling JM. WHONET: An Information System for Monitoring Antimicrobial Resistance. *Emerg Infect Dis.* 1995 Jun;1(2):66–66.

Examples

```
qc_on_target(AMR::as.mic(0.5), 25922, "GEN") == TRUE
```

standardise_mic

Standardise MIC to control strain [Experimental]**Description**

MIC experiments are generally quality-controlled by including a control strain with a known MIC. The MIC result for the control strain should be a particular target MIC, or at least within an acceptable range. This function standardises a measured MIC to the target MIC given: 1) a control strain (usually identified as an ATCC or NCTC number), 2) an antibiotic name, and 3) a guideline (EUCAST or CLSI). The definition of standardisation in this context is to adjust the measured MIC based on the QC MIC. This is based on the following principles and assumption:

1. A measured MIC is composed of two components: the true MIC and a measurement error. The measurement error is considered to be inevitable when measuring MICs, and is likely to be further composed of variability in laboratory conditions and operator interpretation.
2. It is assumed that the MIC of the control strain in the experiment has also been affected by this error.

The standardisation applied by this function uses the measured QC strain MIC as a reference point, and scales the rest of the MICs to this reference. In general, this means that the MICs are doubled or halved, depending on the result of the QC MIC. A worked example is provided below and illustrates the transformation that this function applies.

There is no current evidence base for this approach, therefore, this function is considered experimental and should be used with caution.

Usage

```
standardise_mic(
  test_measurement,
  qc_measurement,
  strain,
  ab,
  prefer_upper = FALSE,
  ignore_na = TRUE,
  guideline = "EUCAST",
  year = "2023",
  force = TRUE
)
```

Arguments

test_measurement	Measured MIC to standardise
qc_measurement	Measured QC MIC to standardise to
strain	control strain identifier (usually ATCC)
ab	antibiotic name (will be coerced to AMR::as.ab)

prefer_upper	Where the target MIC is a range, prefer the upper value in the range
ignore_na	Ignore NA (returns AMR::NA_mic_)
guideline	Guideline to use (EUCAST or CLSI)
year	Guideline year (version)
force	Force into MIC-compatible format after standardisation

Value

AMR::mic vector

Examples

```
# Ref strain QC MIC for GEN is 0.5
standardise_mic(
  test_measurement = c(AMR::as.mic(">8.0"), # QC = 1, censored MIC remains censored
                      AMR::as.mic(4.0),   # QC = 0.5 which is on target, so stays same
                      AMR::as.mic(2),     # QC = 1, so scaled down to 1
                      AMR::as.mic(2)),    # QC = 0.25, so scaled up to 8
  qc_measurement = c(AMR::as.mic(1),
                    AMR::as.mic(0.5),
                    AMR::as.mic(1),
                    AMR::as.mic(0.25)),
  strain = 25922,
  ab = AMR::as.ab("GEN"))
```

subset.mic_validation *Subset MIC validation object*

Description

Subset MIC validation object

Usage

```
## S3 method for class 'mic_validation'
subset(x, subset, ...)
```

Arguments

x	mic_validation object
subset	logical expression to subset by
...	additional arguments

Value

mic_validation object

Examples

```
gold_standard <- c("<0.25", "8", "64", ">64")
test <- c("<0.25", "2", "16", "64")
ab <- AMR::as.ab(c("AMK", "AMK", "CIP", "CIP"))
mo <- AMR::as.mo(c("E. coli", "E. coli", "P. mirabilis", "P. mirabilis"))
val <- compare_mic(gold_standard, test, ab, mo)
subset(val, ab == AMR::as.ab("AMX"))
subset(val, mo == AMR::as.mo("E. coli"))
```

summary.mic_validation

Summary of MIC validation results

Description

Summarise the results of an MIC validation generated using compare_mic().

Usage

```
## S3 method for class 'mic_validation'
summary(object, ...)
```

Arguments

object	S3 mic_validation object
...	further optional parameters

Value

S3 mic_validation_summary object

Examples

```
gold_standard <- c("<0.25", "8", "64", ">64")
test <- c("<0.25", "2", "16", "64")
val <- compare_mic(gold_standard, test)
summary(val)
# or, for more detailed results
as.data.frame(summary(val))
```

table

Table

Description

Table

Usage

```
table(x, ...)

## Default S3 method:
table(x, ...)

## S3 method for class 'mic_validation'
table(
  x,
  format = "flextable",
  fill_dilutions = TRUE,
  bold = TRUE,
  ea_color = NULL,
  gold_standard_name = "Gold Standard",
  test_name = "Test",
  ...
)
```

Arguments

x	mic_validation S3 object
...	further arguments
format	simple or flextable
fill_dilutions	Fill dilutions that are not present in the data in order to match the y- and x- axes
bold	Bold cells where essential agreement is TRUE
ea_color	Background color for essential agreement cells
gold_standard_name	Name of the gold standard to display in output
test_name	Name of the test to display in output

Value

table or flextable object

Examples

```
gold_standard <- c("<0.25", "8", "64", ">64")
test <- c("<0.25", "2", "16", "64")
val <- compare_mic(gold_standard, test)
table(val)
```

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