

Package: emery (via r-universe)

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Title Accuracy Statistic Estimation for Imperfect Gold Standards

Version 0.5.1

Description Produce maximum likelihood estimates of common accuracy statistics for multiple measurement methods when a gold standard is not available. An R implementation of the expectation maximization algorithms described in Zhou et al. (2011) <[doi:10.1002/9780470906514](https://doi.org/10.1002/9780470906514)> with additional functions for creating simulated data and visualizing results. Supports binary, ordinal, and continuous measurement methods.

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RdMacros Rdpack

Suggests knitr, rmarkdown

VignetteBuilder knitr

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boot_ML	<i>Bootstrap ML accuracy statistic estimation for multi-method data</i>
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Description

boot_ML() is a function used to generate bootstrap estimates of results generated by estimate_ML() primarily for use in creating nonparametric confidence intervals.

Usage

```
boot_ML(
  type = c("binary", "ordinal", "continuous"),
  data,
  n_boot = 100,
  max_iter = 1000,
  tol = 1e-07,
  seed = NULL,
  ...
)
```

Arguments

type	A string specifying the data type of the methods under evaluation.
data	An n_obs by n_method <i>matrix</i> containing the observed values for each method. If the dimensions are named, row names will be used to name each observation (obs_names) and column names will be used to name each measurement method (method_names).
n_boot	number of bootstrap estimates to compute
max_iter	The maximum number of EM algorithm iterations to compute before reporting a result.
tol	The minimum change in statistic estimates needed to continue iterating the EM algorithm.
seed	optional seed for RNG
...	Additional arguments

Value

a list containing accuracy estimates, v , and the parameters used.

<code>v_0</code>	result from original data
<code>v_star</code>	list containing results from each bootstrap resampling
<code>params</code>	list containing the parameters used

Examples

```
# Set seed for this example
set.seed(11001101)

# Generate data for 4 binary methods
my_sim <- generate_multimethod_data(
  "binary",
  n_obs = 75,
  n_method = 4,
  se = c(0.87, 0.92, 0.79, 0.95),
  sp = c(0.85, 0.93, 0.94, 0.80),
  method_names = c("alpha", "beta", "gamma", "delta"))

# Bootstrap ML results
boot_ex <- boot_ML(
  "binary",
  data = my_sim$generated_data,
  n_boot = 20)
```

<code>censor_data</code>	<i>Censor data randomly rowwise</i>
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Description

Censor data randomly rowwise

Usage

```
censor_data(
  n_obs = dis$n_obs,
  first_reads_all = first_reads_all,
  n_method_subset = n_method_subset,
  n_method = n_method
)
```

Arguments

n_obs	An integer representing the number of observations to simulate.
first_reads_all	Used for binary methods. A logical which forces method 1 to have a result for every observation
n_method_subset	Used for binary methods. An integer defining how many methods to select at random to produce a result for each observation
n_method	An integer representing the number of methods to simulate.

define_disease_state *Define the True disease state of a simulated sample*

Description

Define the True disease state of a simulated sample

Usage

```
define_disease_state(D = NULL, n_obs = NULL, prev = NULL)
```

Arguments

D	Optional binary vector representing the true classification of each observation.
n_obs	An integer representing the number of observations to simulate.
prev	A value between 0-1 which represents the proportion of "positive" results in the target population.

Value

A list of features defining the true disease status of each observation

estimate_ML *Estimate maximum likelihood accuracy statistics by expectation maximization*

Description

estimate_ML() is a general function for estimating the maximum likelihood accuracy statistics for a set of methods with no known reference value, i.e. "truth", or "gold standard".

Usage

```

estimate_ML(
  type = c("binary", "ordinal", "continuous"),
  data,
  init = list(NULL),
  max_iter = 1000,
  tol = 1e-07,
  save_progress = TRUE,
  ...
)

estimate_ML_binary(
  data,
  init = list(prev_1 = NULL, se_1 = NULL, sp_1 = NULL),
  max_iter = 100,
  tol = 1e-07,
  save_progress = TRUE
)

estimate_ML_continuous(
  data,
  init = list(prev_1 = NULL, mu_i1_1 = NULL, sigma_i1_1 = NULL, mu_i0_1 = NULL,
    sigma_i0_1 = NULL),
  max_iter = 100,
  tol = 1e-07,
  save_progress = TRUE
)

estimate_ML_ordinal(
  data,
  init = list(pi_1_1 = NULL, phi_1ij_1 = NULL, phi_0ij_1 = NULL, n_level = NULL),
  level_names = NULL,
  max_iter = 1000,
  tol = 1e-07,
  save_progress = TRUE
)

```

Arguments

<code>type</code>	A string specifying the data type of the methods under evaluation.
<code>data</code>	An <code>n_obs</code> by <code>n_method</code> <code>matrix</code> containing the observed values for each method. If the dimensions are named, row names will be used to name each observation (<code>obs_names</code>) and column names will be used to name each measurement method (<code>method_names</code>).
<code>init</code>	An optional list of initial values used to seed the EM algorithm. If initial values are not provided, the <code>pollinate_ML()</code> function will be called on the data to estimate starting values. It is recommended to try several sets of starting param-

	eters to ensure that the algorithm converges to the same results. This is to verify that the result does not represent a local extrema.
max_iter	The maximum number of EM algorithm iterations to compute before reporting a result.
tol	The minimum change in statistic estimates needed to continue iterating the EM algorithm.
save_progress	A logical indication of whether to save interim calculations used in the EM algorithm.
...	Additional arguments
level_names	An optional, ordered, character vector of unique names corresponding to the levels of the methods.

Details

The lack of an infallible reference method is referred to as an imperfect gold standard (GS). Accuracy statistics which rely on a GS method, such as sensitivity, specificity, and AUC, can be estimated using imperfect gold standards by iteratively estimating the maximum likelihood values of these statistics while the conditional independence assumption holds. `estimate_ML()` relies on a collection of expectation maximization (EM) algorithms to achieve this. The EM algorithms used in this function are based on those presented in *Statistical Methods in Diagnostic Medicine, Second Edition* (Zhou et al. 2011) and have been validated on several examples therein. Additional details about these algorithms can be found for binary (Walter and Irwig 1988), ordinal (Zhou et al. 2005), and continuous (Hsieh et al. 2009) methods. Minor changes to the literal calculations have been made for efficiency, code readability, and the like, but the underlying steps remain functionally unchanged.

Value

`estimate_ML()` returns an S4 object of class "MultiMethodMLEstimate" containing the maximum likelihood accuracy statistics calculated by EM.

References

- Zhou X, Obuchowski NA, McClish DK (2011). *Statistical Methods in Diagnostic Medicine*. Wiley. doi:10.1002/9780470906514.
- Walter SD, Irwig LM (1988). "Estimation of test error rates, disease prevalence and relative risk from misclassified data: a review." *J. Clin. Epidemiol.*, **41**(9), 923–937. doi:10.1016/0895-4356(88)901102.
- Zhou X, Castelluccio P, Zhou C (2005). "Nonparametric estimation of ROC curves in the absence of a gold standard." *Biometrics*, **61**(2), 600–609. doi:10.1111/j.15410420.2005.00324.x.
- Hsieh H, Su H, Zhou X (2009). "Interval Estimation for the Difference in Paired Areas under the ROC Curves in the Absence of a Gold Standard Test." *Statistics in Medicine*. <https://doi.org/10.1002/sim.3661>.

Examples

```
# Set seed for this example
set.seed(11001101)

# Generate data for 4 binary methods
my_sim <- generate_multimethod_data(
  "binary",
  n_obs = 75,
  n_method = 4,
  se = c(0.87, 0.92, 0.79, 0.95),
  sp = c(0.85, 0.93, 0.94, 0.80),
  method_names = c("alpha", "beta", "gamma", "delta"))

# View the data
my_sim$generated_data

# View the parameters used to generate the data
my_sim$params

# Estimate ML accuracy values by EM algorithm
my_result <- estimate_ML(
  "binary",
  data = my_sim$generated_data,
  save_progress = FALSE # this reduces the data stored in the resulting object
)

# View results of ML estimate
my_result@results
```

generate_multimethod_data

Create data sets which simulate paired measurements of multiple methods

Description

generate_multimethod_data() is a general function for creating a data set which simulates the results one might see when using several different methods to measure a set of objects.

Usage

```
generate_multimethod_data(
  type = c("binary", "ordinal", "continuous"),
  n_method = 3,
  n_obs = 100,
  prev = 0.5,
  D = NULL,
  method_names = NULL,
```

```
    obs_names = NULL,  
    ...  
  )  
  
generate_multimethod_binary(  
  n_method = 3,  
  n_obs = 100,  
  prev = 0.5,  
  D = NULL,  
  se = rep(0.9, n_method),  
  sp = rep(0.9, n_method),  
  method_names = NULL,  
  obs_names = NULL,  
  n_method_subset = n_method,  
  first_reads_all = FALSE  
)  
  
generate_multimethod_ordinal(  
  n_method = 3,  
  n_obs = 100,  
  prev = 0.5,  
  D = NULL,  
  n_level = 5,  
  pmf_pos = matrix(rep(1:n_level - 1, n_method), nrow = n_method, byrow = TRUE),  
  pmf_neg = matrix(rep(n_level:1 - 1, n_method), nrow = n_method, byrow = TRUE),  
  method_names = NULL,  
  level_names = NULL,  
  obs_names = NULL,  
  n_method_subset = n_method,  
  first_reads_all = FALSE  
)  
  
generate_multimethod_continuous(  
  n_method = 2,  
  n_obs = 100,  
  prev = 0.5,  
  D = NULL,  
  mu_i1 = rep(12, n_method),  
  sigma_i1 = diag(n_method),  
  mu_i0 = rep(10, n_method),  
  sigma_i0 = diag(n_method),  
  method_names = NULL,  
  obs_names = NULL,  
  n_method_subset = n_method,  
  first_reads_all = FALSE  
)
```


Arguments

type	A string specifying the data type of the methods being simulated.
n_method	An integer representing the number of methods to simulate.
n_obs	An integer representing the number of observations to simulate.
prev	A value between 0-1 which represents the proportion of "positive" results in the target population.
D	Optional binary vector representing the true classification of each observation.
method_names	Optional vector of names used to identify each method.
obs_names	Optional vector of names used to identify each observation.
...	Additional parameters
se, sp	Used for binary methods. A vector of length n_method of values between 0-1 representing the sensitivity and specificity of the methods.
n_method_subset	Used for binary methods. An integer defining how many methods to select at random to produce a result for each observation
first_reads_all	Used for binary methods. A logical which forces method 1 to have a result for every observation
n_level	Used for ordinal methods. An integer representing the number of ordinal levels each method has
pmf_pos, pmf_neg	Used for ordinal methods. A n_method by n_level matrix representing the probability mass functions for positive and negative results, respectively
level_names	Used for ordinal methods. Optional vector of names used to identify each level
mu_i1, mu_i0	Used for continuous methods. Vectors of length n_method of the method mean values for positive (negative) observations
sigma_i1, sigma_i0	Used for continuous methods. Covariance matrices of method positive (negative) observations

Details

The function supports binary measurement methods, e.g., Pass/Fail; ordinal measurement methods, e.g., the Likert scale; and continuous measurement methods, e.g., height. The data are generated under the assumption that the underlying population consists of a mixture of two groups. The primary application of this is to simulate a sample from a population which has some prevalence of disease.

Value

A list containing a simulated data set and the parameters used to create it

Examples

```

# Set seed for this example
set.seed(11001101)

# Generate data for 4 binary methods
my_sim <- generate_multimethod_data(
  "binary",
  n_obs = 75,
  n_method = 4,
  se = c(0.87, 0.92, 0.79, 0.95),
  sp = c(0.85, 0.93, 0.94, 0.80),
  method_names = c("alpha", "beta", "gamma", "delta"))

# View the data
my_sim$generated_data

# View the parameters used to generate the data
my_sim$params

# Estimate ML accuracy values by EM algorithm
my_result <- estimate_ML(
  "binary",
  data = my_sim$generated_data,
  save_progress = FALSE # this reduces the data stored in the resulting object
)

# View results of ML estimate
my_result@results

```

MultiMethodMLEstimate-class

S4 object containing the results of multi-method ML accuracy estimates

Description

S4 object containing the results of multi-method ML accuracy estimates

Slots

`results` a list of estimated accuracy statistics
`names` a list containing vectors of names of various dimensions
`data` a copy of the data used to generate the estimated values
`iter` an integer number of iterations needed for the EM algorithm to converge
`prog` a list containing the values calculated during each iteration of the EM algorithm
`type` a string describing the data type

name_thing	<i>Create unique names for a set of things</i>
------------	--

Description

Create unique names for a set of things

Usage

```
name_thing(thing = "", n = 1)
```

Arguments

thing	a string that describes the set of items to name
n	an integer number of unique names to create

Value

a vector of unique names

plot, MultiMethodMLEstimate-method	<i>Create plots from a MultiMethodMLEstimate object</i>
------------------------------------	---

Description

Create a list of plots visualizing the expectation maximization process and resulting accuracy statistics stored in a MultiMethodMLEstimate object.

Usage

```
## S4 method for signature 'MultiMethodMLEstimate'
plot(x, y, ...)
```

Arguments

x	a MultiMethodMLEstimate S4 object
y	not used
...	Arguments passed on to plot_ML
params	A list of population parameters. This is primarily used to evaluate results from a simulation where the target parameters are known, but can be used to visualize results with respect to some True value.

Value

A list of ggplot2 plots

plot_ML

Create plots visualizing the ML estimation process and results.

Description

plot_ML() is a general function for visualizing results generated by estimate_ML().

Usage

```
plot_ML(ML_est, params = NULL)
```

```
plot_ML_binary(
  ML_est,
  params = list(prev = NULL, se = NULL, sp = NULL, D = NULL)
)
```

```
plot_ML_ordinal(
  ML_est,
  params = list(pi_1_1 = NULL, phi_1ij_1 = NULL, phi_0ij_1 = NULL, D = NULL)
)
```

```
plot_ML_continuous(
  ML_est,
  params = list(prev_1 = NULL, mu_i1_1 = NULL, sigma_i1_1 = NULL, mu_i0_1 = NULL,
    sigma_i0_1 = NULL, D = NULL)
)
```

Arguments

ML_est	A MultiMethodMLEstimate class object
params	A list of population parameters. This is primarily used to evaluate results from a simulation where the target parameters are known, but can be used to visualize results with respect to some True value.

Value

A list of ggplot2 plots.

Binary:

prev	A plot showing how the prevalence estimate changes with each iteration of the EM algorithm
se	A plot showing how the sensitivity estimates of each method change with each iteration of the EM algorithm
sp	A plot showing how the specificity estimates of each method change with each iteration of the EM algorithm

qk	A plot showing how the q values for each observation k change over each iteration of the EM algorithm
qk_hist	A histogram of q values. Observations, k, can be colored by True state if it is passed by params\$D.
se_sp	A plot showing the path the sensitivity and specificity estimates for each method follows during the EM algorithm. True sensitivity and specificity values can be passed by params\$se and params\$sp, respectively. This is useful for comparing algorithm results when applied to simulation data where True parameter values are known.
Ordinal:	
ROC	The Receiver Operator Characteristic (ROC) curves estimated for each method
q_k1	A plot showing how the q values for each observation, k, change when d=1 over each iteration of the EM algorithm. Observations can be colored by True state if it is passed (params\$D).
q_k0	A plot showing how the q values for each observation, k, change when d=0 over each iteration of the EM algorithm. Observations can be colored by True state if it is passed by params\$D.
q_k1_hist	A histogram of q_1 values. Observations, k, can be colored by True state if it is passed by params\$D.
phi_d	A stacked bar graph representing the estimated CMFs of each method when d=0 and d=1.
Continuous:	
ROC	The Receiver Operator Characteristic (ROC) curves estimated for each method
z_k1	A plot showing how the z_k1 values for each observation change over each iteration of the EM algorithm. Observations can be colored by True state if it is passed (params\$D).
z_k0	A plot showing how the z_k0 values for each observation change over each iteration of the EM algorithm. Observations can be colored by True state if it is passed (params\$D).
z_k1_hist	A histogram of z_k1 values. Observations can be colored by True state if it is passed (params\$D).

Examples

```
# Set seed for this example
set.seed(11001101)

# Generate data for 4 binary methods
my_sim <- generate_multimethod_data(
  "binary",
  n_obs = 75,
  n_method = 4,
  se = c(0.87, 0.92, 0.79, 0.95),
  sp = c(0.85, 0.93, 0.94, 0.80),
```

```

method_names = c("alpha", "beta", "gamma", "delta"))

# View the data
my_sim$generated_data

# View the parameters used to generate the data
my_sim$params

# Estimate ML accuracy values by EM algorithm
my_result <- estimate_ML(
  "binary",
  data = my_sim$generated_data,
  save_progress = FALSE # this reduces the data stored in the resulting object
)

# View results of ML estimate
my_result@results

```

pollinate_ML

Generate seed values for EM algorithm

Description

pollinate_ML() is a general helper function which can be used to generate starting values, i.e. seeds, for the estimate_ML() function from a multi-method data set.

Usage

```
pollinate_ML(type = c("binary", "ordinal", "continuous"), data, ...)
```

```
pollinate_ML_binary(data, ...)
```

```
pollinate_ML_ordinal(
  data,
  n_level = NULL,
  threshold_level = ceiling(n_level/2),
  level_names = NULL
)
```

```
pollinate_ML_continuous(
  data,
  prev = 0.5,
  q_seeds = c((1 - prev)/2, 1 - (prev/2)),
  high_pos = TRUE
)
```

Arguments

type	A string specifying the data type of the methods under evaluation.
data	An n_obs by n_method <i>matrix</i> containing the observed values for each method. If the dimensions are named, row names will be used to name each observation (obs_names) and column names will be used to name each measurement method (method_names).
...	Additional arguments
n_level	Used for ordinal methods. Integer number of levels each method contains
threshold_level	Used for ordinal methods. A value from 1 to n_level which indicates the initial threshold used to define positive and negative disease states.
level_names	Used for ordinal methods. Optional vector of length n_level containing names for each level.
prev	A double between 0-1 representing the proportion of positives in the population
q_seeds	Used for continuous methods. A vector of length 2 representing the quantiles at which the two groups are assumed to be centered
high_pos	Used for continuous methods. A logical indicating whether larger values are considered "positive"

Value

a list of EM algorithm initialization values

```
show,MultiMethodMLEstimate-method
```

Show a MultiMethodMLEstimate S4 object

Description

Print the accuracy statistic estimates stored in a MultiMethodMLEstimate object.

Usage

```
## S4 method for signature 'MultiMethodMLEstimate'
show(object)
```

Arguments

object An object of class MultiMethodMLEstimate.

Value

A list containing relevant accuracy statistic estimates. This is a subset of the list stored in results slot of the MultiMethodMLEstimate object.

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