

Package ‘graphicalMCP’

November 8, 2024

Type Package

Title Graphical Multiple Comparison Procedures

Version 0.2.6

Description Multiple comparison procedures (MCPs) control the familywise error rate in clinical trials. Graphical MCPs include many commonly used procedures as special cases; see Bretz et al. (2011) <[doi:10.1002/bimj.201000239](https://doi.org/10.1002/bimj.201000239)>, Lu (2016) <[doi:10.1002/sim.6985](https://doi.org/10.1002/sim.6985)>, and Xi et al. (2017) <[doi:10.1002/bimj.201600233](https://doi.org/10.1002/bimj.201600233)>. This package is a low-dependency implementation of graphical MCPs which allow mixed types of tests. It also includes power simulations and visualization of graphical MCPs.

License Apache License (>= 2)

URL <https://github.com/openpharma/graphicalMCP>

BugReports <https://github.com/openpharma/graphicalMCP/issues>

Depends R (>= 4.1.0)

Imports matrixStats, mvtnorm

Suggests bench, dplyr, forcats, ggplot2, gMCP, gt, here, htmltools, igraph, knitr, lrrstat, prompt, rmarkdown, scales, testthat (>= 3.0.0), tibble, tictoc, tidyr, xfun

VignetteBuilder knitr

Config/testthat/edition 3

Encoding UTF-8

RoxygenNote 7.3.0

NeedsCompilation no

Author Dong Xi [aut, cre],
Ethan Brockmann [aut],
Gilead Sciences, Inc. [cph, fnd]

Maintainer Dong Xi <dong.xi1@gilead.com>

Repository CRAN

Date/Publication 2024-11-08 21:00:02 UTC

Contents

adjust_p_bonferroni	2
adjust_weights_parametric	4
as_initial_graph	6
bonferroni	8
graph_calculate_power	10
graph_create	13
graph_generate_weights	16
graph_rejection_orderings	17
graph_test_closure	19
graph_test_shortcut	22
graph_update	24
plot.initial_graph	26
plot.updated_graph	29
print.graph_report	30
print.initial_graph	32
print.power_report	33
print.updated_graph	34

Index	36
--------------	-----------

adjust_p_bonferroni	<i>Calculate adjusted p-values</i>
---------------------	------------------------------------

Description

For an intersection hypothesis, an adjusted p-value is the smallest significance level at which the intersection hypothesis can be rejected. The intersection hypothesis can be rejected if its adjusted p-value is less than or equal to α . Currently, there are three test types supported:

- Bonferroni tests for `adjust_p_bonferroni()`,
- Parametric tests for `adjust_p_parametric()`,
 - Note that one-sided tests are required for parametric tests.
- Simes tests for `adjust_p_simes()`.

Usage

```
adjust_p_bonferroni(p, hypotheses)
```

```
adjust_p_parametric(
  p,
  hypotheses,
  test_corr = NULL,
  maxpts = 25000,
  abseps = 1e-06,
  releps = 0
```

```
)
adjust_p_simes(p, hypotheses)
```

Arguments

p	A numeric vector of p-values (unadjusted, raw), whose values should be between 0 & 1. The length should match the length of hypotheses.
hypotheses	A numeric vector of hypothesis weights. Must be a vector of values between 0 & 1 (inclusive). The length should match the length of p. The sum of hypothesis weights should not exceed 1.
test_corr	(Optional) A numeric matrix of correlations between test statistics, which is needed to perform parametric tests using <code>adjust_p_parametric()</code> . The number of rows and columns of this correlation matrix should match the length of p.
maxpts	(Optional) An integer scalar for the maximum number of function values, which is needed to perform parametric tests using the <code>mvtnorm::GenzBretz</code> algorithm. The default is 25000.
abseps	(Optional) A numeric scalar for the absolute error tolerance, which is needed to perform parametric tests using the <code>mvtnorm::GenzBretz</code> algorithm. The default is 1e-6.
releps	(Optional) A numeric scalar for the relative error tolerance as double, which is needed to perform parametric tests using the <code>mvtnorm::GenzBretz</code> algorithm. The default is 0.

Value

A single adjusted p-value for the intersection hypothesis.

References

- Bretz, F., Maurer, W., Brannath, W., and Posch, M. (2009). A graphical approach to sequentially rejective multiple test procedures. *Statistics in Medicine*, 28(4), 586-604.
- Lu, K. (2016). Graphical approaches using a Bonferroni mixture of weighted Simes tests. *Statistics in Medicine*, 35(22), 4041-4055.
- Xi, D., Glimm, E., Maurer, W., and Bretz, F. (2017). A unified framework for weighted parametric multiple test procedures. *Biometrical Journal*, 59(5), 918-931.

See Also

`adjust_weights_parametric()` for adjusted hypothesis weights using parametric tests, `adjust_weights_simes()` for adjusted hypothesis weights using Simes tests.

Examples

```
hypotheses <- c(H1 = 0.5, H2 = 0.25, H3 = 0.25)
p <- c(0.019, 0.025, 0.05)
adjust_p_bonferroni(p, hypotheses)
```

```

set.seed(1234)
hypotheses <- c(H1 = 0.5, H2 = 0.25, H3 = 0.25)
p <- c(0.019, 0.025, 0.05)
# Using the `mvtnorm::GenzBretz` algorithm
corr <- matrix(0.5, nrow = 3, ncol = 3)
diag(corr) <- 1
adjust_p_parametric(p, hypotheses, corr)
hypotheses <- c(H1 = 0.5, H2 = 0.25, H3 = 0.25)
p <- c(0.019, 0.025, 0.05)
adjust_p_simes(p, hypotheses)

```

```
adjust_weights_parametric
```

Calculate adjusted hypothesis weights

Description

An intersection hypothesis can be rejected if its p-values are less than or equal to their adjusted significance levels, which are their adjusted hypothesis weights times α . For Bonferroni tests, their adjusted hypothesis weights are their hypothesis weights of the intersection hypothesis. Additional adjustment is needed for parametric and Simes tests:

- Parametric tests for [adjust_weights_parametric\(\)](#),
 - Note that one-sided tests are required for parametric tests.
- Simes tests for [adjust_weights_simes\(\)](#).

Usage

```

adjust_weights_parametric(
  matrix_weights,
  matrix_intersections,
  test_corr,
  alpha,
  test_groups,
  maxpts = 25000,
  abseps = 1e-06,
  releps = 0
)

adjust_weights_simes(matrix_weights, p, test_groups)

```

Arguments

`matrix_weights` (Optional) A matrix of hypothesis weights of all intersection hypotheses. This can be obtained as the second half of columns from the output of [graph_generate_weights\(\)](#).

`matrix_intersections` (Optional) A matrix of hypothesis indicators of all intersection hypotheses. This can be obtained as the first half of columns from the output of [graph_generate_weights\(\)](#).

test_corr	(Optional) A numeric matrix of correlations between test statistics, which is needed to perform parametric tests using <code>adjust_weights_parametric()</code> . The number of rows and columns of this correlation matrix should match the length of <code>p</code> .
alpha	(Optional) A numeric value of the overall significance level, which should be between 0 & 1. The default is 0.025 for one-sided hypothesis testing problems; another common choice is 0.05 for two-sided hypothesis testing problems. Note when parametric tests are used, only one-sided tests are supported.
test_groups	(Optional) A list of numeric vectors specifying hypotheses to test together. Grouping is needed to correctly perform Simes and parametric tests.
maxpts	(Optional) An integer scalar for the maximum number of function values, which is needed to perform parametric tests using the <code>mvtnorm::GenzBretz</code> algorithm. The default is 25000.
abseps	(Optional) A numeric scalar for the absolute error tolerance, which is needed to perform parametric tests using the <code>mvtnorm::GenzBretz</code> algorithm. The default is $1e-6$.
releps	(Optional) A numeric scalar for the relative error tolerance as double, which is needed to perform parametric tests using the <code>mvtnorm::GenzBretz</code> algorithm. The default is 0.
p	(Optional) A numeric vector of p-values (unadjusted, raw), whose values should be between 0 & 1. The length should match the number of columns of <code>matrix_weights</code> .

Value

- `adjust_weights_parametric()` returns a matrix with the same dimensions as `matrix_weights`, whose hypothesis weights have been adjusted according to parametric tests.
- `adjust_weights_simes()` returns a matrix with the same dimensions as `matrix_weights`, whose hypothesis weights have been adjusted according to Simes tests.

References

Lu, K. (2016). Graphical approaches using a Bonferroni mixture of weighted Simes tests. *Statistics in Medicine*, 35(22), 4041-4055.

Xi, D., Glimm, E., Maurer, W., and Bretz, F. (2017). A unified framework for weighted parametric multiple test procedures. *Biometrical Journal*, 59(5), 918-931.

See Also

`adjust_p_parametric()` for adjusted p-values using parametric tests, `adjust_p_simes()` for adjusted p-values using Simes tests.

Examples

```
alpha <- 0.025
p <- c(0.018, 0.01, 0.105, 0.006)
num_hyps <- length(p)
g <- bonferroni_holm(rep(1 / 4, 4))
```

```

weighting_strategy <- graph_generate_weights(g)
matrix_intersections <- weighting_strategy[, seq_len(num_hyps)]
matrix_weights <- weighting_strategy[, -seq_len(num_hyps)]

set.seed(1234)
adjust_weights_parametric(
  matrix_weights = matrix_weights,
  matrix_intersections = matrix_intersections,
  test_corr = diag(4),
  alpha = alpha,
  test_groups = list(1:4)
)
alpha <- 0.025
p <- c(0.018, 0.01, 0.105, 0.006)
num_hyps <- length(p)
g <- bonferroni_holm(rep(1 / 4, 4))
weighting_strategy <- graph_generate_weights(g)
matrix_intersections <- weighting_strategy[, seq_len(num_hyps)]
matrix_weights <- weighting_strategy[, -seq_len(num_hyps)]

adjust_weights_simes(
  matrix_weights = matrix_weights,
  p = p,
  test_groups = list(1:4)
)

```

as_initial_graph

Convert between graphicalMCP, gMCP, and igraph graph classes

Description

Graph objects have different structures and attributes in graphicalMCP, gMCP, and igraph R packages. These functions convert between different classes to increase compatibility.

Note that igraph and gMCP have additional attributes for vertices, edges, or a graph itself. These conversion functions only handle attributes related to hypothesis names, hypothesis weights and transition weights. Other attributes will be dropped when converting.

Usage

```

as_initial_graph(graph)

## S3 method for class 'graphMCP'
as_initial_graph(graph)

## S3 method for class 'igraph'
as_initial_graph(graph)

as_graphMCP(graph)

```

```
## S3 method for class 'initial_graph'  
as_graphMCP(graph)  
  
as_igraph(graph)  
  
## S3 method for class 'initial_graph'  
as_igraph(graph)
```

Arguments

graph An `initial_graph` object from the `graphicalMCP` package, a `graphMCP` object from the `gMCP` package, or an `igraph` object from the `igraph` package, depending on the conversion type.

Value

- `as_graphMCP()` returns a `graphMCP` object for the `gMCP` package.
- `as_igraph()` returns an `igraph` object for the `igraph` package.
- `as_initial_graph()` returns an `initial_graph` object for the `graphicalMCP` package.

References

Csardi, G., Nepusz, T., Traag, V., Horvat, S., Zanini, F., Noom, D., and Mueller, K. (2024). *igraph*: Network analysis and visualization in R. R package version 2.0.3. <https://CRAN.R-project.org/package=igraph>.

Rohmeyer, K., and Klinglmueller, K. (2024). *gMCP*: Graph based multiple test procedures. R package version 0.8-17. <https://cran.r-project.org/package=gMCP>.

See Also

[graph_create\(\)](#) for the initial graph used in the `graphicalMCP` package.

Examples

```
g_graphicalMCP <- random_graph(5)  
  
if (requireNamespace("gMCP", quietly = TRUE)) {  
  g_gMCP <- as_graphMCP(g_graphicalMCP)  
  
  all.equal(g_graphicalMCP, as_initial_graph(g_gMCP))  
}  
  
if (requireNamespace("igraph", quietly = TRUE)) {  
  g_igraph <- as_igraph(g_graphicalMCP)  
  
  all.equal(g_graphicalMCP, as_initial_graph(g_igraph))  
}
```

 bonferroni

Example graphs of commonly used multiple comparison procedures

Description

Built-in functions to quickly generate select graphical multiple comparison procedures.

Usage

```
bonferroni(hypotheses, hyp_names = NULL)
```

```
bonferroni_holm(hypotheses, hyp_names = NULL)
```

```
huque_etal(hyp_names = NULL)
```

```
fallback(hypotheses, hyp_names = NULL)
```

```
fallback_improved_1(hypotheses, hyp_names = NULL)
```

```
fallback_improved_2(hypotheses, epsilon = 1e-04, hyp_names = NULL)
```

```
fixed_sequence(num_hyps, hyp_names = NULL)
```

```
simple_successive_1(hyp_names = NULL)
```

```
simple_successive_2(hyp_names = NULL)
```

```
random_graph(num_hyps, hyp_names = NULL)
```

```
two_doses_two_primary_two_secondary(hyp_names = NULL)
```

```
three_doses_two_primary_two_secondary(hyp_names = NULL)
```

Arguments

hypotheses	(Optional) A numeric vector of hypothesis weights in a graphical multiple comparison procedure. Must be a vector of values between 0 & 1 (inclusive). The length should match num_hyps and the length of hyp_names. The sum of hypothesis weights should not exceed 1.
hyp_names	(Optional) A character vector of hypothesis names. The length should match num_hyps and the length of hypotheses. If hyp_names are not specified, hypotheses will be named sequentially as H1, H2,
epsilon	(Optional) A numeric scalar indicating the value of the ϵ edge. This should be a much smaller value than hypothesis and transition weights. The default is 1e-4.
num_hyps	(Optional) Number of hypotheses in a graphical multiple comparison procedure.

Value

An S3 object as returned by `graph_create()`.

References

- Bretz, F., Maurer, W., Brannath, W., and Posch, M. (2009). A graphical approach to sequentially rejective multiple test procedures. *Statistics in Medicine*, 28(4), 586-604.
- Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.
- Huque, M. F., Alosch, M., and Bhole, R. (2011). Addressing multiplicity issues of a composite endpoint and its components in clinical trials. *Journal of Biopharmaceutical Statistics*, 21(4), 610-634.
- Maurer, W., Hothorn, L., and Lehman, W. (1995). Multiple comparisons in drug clinical trials and preclinical assays: a-priori ordered hypotheses. *Biometrie in der chemisch-pharmazeutischen Industrie*, 6, 3-18.
- Westfall, P. H., and Krishen, A. (2001). Optimally weighted, fixed sequence and gatekeeper multiple testing procedures. *Journal of Statistical Planning and Inference*, 99(1), 25-40.
- Wiens, B. L. (2003). A fixed sequence Bonferroni procedure for testing multiple endpoints. *Pharmaceutical Statistics*, 2(3), 211-215.
- Wiens, B. L., and Dmitrienko, A. (2005). The fallback procedure for evaluating a single family of hypotheses. *Journal of Biopharmaceutical Statistics*, 15(6), 929-942.
- Xi, D., and Bretz, F. (2019). Symmetric graphs for equally weighted tests, with application to the Hochberg procedure. *Statistics in Medicine*, 38(27), 5268-5282.

See Also

`graph_create()` for a general way to create the initial graph.

Examples

```
# Bretz et al. (2009)
bonferroni(hypotheses = rep(1 / 3, 3))
# Bretz et al. (2009)
bonferroni_holm(hypotheses = rep(1 / 3, 3))
# Huque et al. (2011)
huque_etal()
# Wiens (2003)
fallback(hypotheses = rep(1 / 3, 3))
# Wiens and Dmitrienko (2005)
fallback_improved_1(hypotheses = rep(1 / 3, 3))
# Bretz et al. (2009)
fallback_improved_2(hypotheses = rep(1 / 3, 3))
# Maurer et al. (1995); Westfall and Krishen (2001)
fixed_sequence(num_hyps = 3)
# Figure 1 in Bretz et al. (2011)
simple_successive_1()
# Figure 4 in Bretz et al. (2011)
```

```

simple_successive_2()
# Create a random graph with three hypotheses
random_graph(num_hyps = 3)
# Figure 6 in Xi and Bretz et al. (2019)
two_doses_two_primary_two_secondary()
# Add another dose to Figure 6 in Xi and Bretz et al. (2019)
three_doses_two_primary_two_secondary()

```

graph_calculate_power *Calculate power values for a graphical multiple comparison procedure*

Description

Under the alternative hypotheses, the distribution of test statistics is assumed to be a multivariate normal distribution. Given this distribution, this function calculates power values for a graphical multiple comparison procedure. By default, it calculate the local power, which is the probability to reject an individual hypothesis, the probability to reject at least one hypothesis, the probability to reject all hypotheses, the expected number of rejections, and the probability of user-defined success criteria. See vignette("shortcut-testing") and vignette("closed-testing") for more illustration of power calculation.

Usage

```

graph_calculate_power(
  graph,
  alpha = 0.025,
  power_marginal = rep(alpha, length(graph$hypotheses)),
  test_groups = list(seq_along(graph$hypotheses)),
  test_types = c("bonferroni"),
  test_corr = rep(list(NA), length(test_types)),
  sim_n = 1e+05,
  sim_corr = diag(length(graph$hypotheses)),
  sim_success = NULL,
  verbose = FALSE
)

```

Arguments

graph	An initial graph as returned by <code>graph_create()</code> .
alpha	A numeric value of the one-sided overall significance level, which should be between 0 & 1. The default is 0.025 for one-sided hypothesis testing. Note that only one-sided tests are supported.
power_marginal	A numeric vector of marginal power values to use when simulating p-values. See Details for more on the simulation process.
test_groups	A list of numeric vectors specifying hypotheses to test together. Grouping is needed to correctly perform Simes and parametric tests.

test_types	A character vector of test types to apply to each test group. This is needed to correctly perform Simes and parametric tests. The length should match the number of elements in test_groups.
test_corr	(Optional) A list of numeric correlation matrices. Each entry in the list should correspond to each test group. For a test group using Bonferroni or Simes tests, its corresponding entry in test_corr should be NA. For a test group using parametric tests, its corresponding entry in test_corr should be a numeric correlation matrix specifying the correlation between test statistics for hypotheses in this test group. The length should match the number of elements in test_groups.
sim_n	An integer scalar specifying the number of simulations. The default is 1e5.
sim_corr	A numeric matrix of correlations between test statistics for all hypotheses. The dimensions should match the number of hypotheses in graph.
sim_success	A list of user-defined functions to specify the success criteria. Functions must take one simulation's logical vector of results as an input, and return a length-one logical vector. For instance, if "success" means rejecting hypotheses 1 and 2, use <code>sim_success = list("1 and 2" = function(x) x[1] && x[2])</code> . If the list is not named, the function body will be used as the name. Lambda functions also work starting with R 4.1, e.g. <code>sim_success = list(\(x) x[3] x[4])</code> .
verbose	A logical scalar specifying whether the details of power simulations should be included in results. The default is <code>verbose = FALSE</code> .

Value

A `power_report` object with a list of 3 elements:

- `inputs` - Input parameters, which is a list of:
 - `graph` - Initial graph,
 - `alpha` - Overall significance level,
 - `test_groups` - Groups of hypotheses for different types of tests,
 - `test_types` - Different types of tests,
 - `test_corr` - Correlation matrices for parametric tests,
 - `sim_n` - Number of simulations,
 - `power_marginal` - Marginal power of all hypotheses
 - `sim_corr` - Correlation matrices for simulations,
 - `sim_success` - User-defined success criteria.
- `power` - A list of power values
 - `power_local` - Local power of all hypotheses, which is the proportion of simulations in which each hypothesis is rejected,
 - `rejection_expected` - Expected (average) number of rejected hypotheses,
 - `power_at_least_1` - Power to reject at least one hypothesis,
 - `power_all` - Power to reject all hypotheses,
 - `power_success` - Power of user-defined success, which is the proportion of simulations in which the user-defined success criterion
 - `sim_success` is met.
- `details` - An optional list of datasets showing simulated p-values and results for each simulation.

Simulation details

The power calculation is based on simulations. The distribution to simulate from is determined as a multivariate normal distribution by `power_marginal` and `sim_corr`. In particular, `power_marginal` is a vector of marginal power values for all hypotheses. The marginal power is the power to reject the null hypothesis at the significance level α *without multiplicity adjustment*. This value could be readily available from standard software and other R packages. Then we can determine the mean of the multivariate normal distribution as

$$\Phi^{-1}(1 - \alpha) - \Phi^{-1}(1 - d_i)$$

, which is often called the non-centrality parameter or the drift parameter. Here d_i is the marginal power `power_marginal` of hypothesis i . Given the correlation matrix `sim_corr`, we can simulate from this multivariate normal distribution using the `mvtnorm` R package (Genz and Bretz, 2009).

Each set simulated values can be used to calculate the corresponding one-sided p-values. Then this set of p-values are plugged into the graphical multiple comparison procedure to determine which hypotheses are rejected. This process is repeated `n_sim` times to produce the power values as the proportion of simulations in which a particular success criterion is met.

References

- Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011a). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.
- Bretz, F., Maurer, W., and Hommel, G. (2011b). Test and power considerations for multiple endpoint analyses using sequentially rejective graphical procedures. *Statistics in Medicine*, 30(13), 1489-1501.
- Genz, A., and Bretz, F. (2009). *Computation of Multivariate Normal and t Probabilities*, series Lecture Notes in Statistics. Springer-Verlag, Heidelberg.
- Lu, K. (2016). Graphical approaches using a Bonferroni mixture of weighted Simes tests. *Statistics in Medicine*, 35(22), 4041-4055.
- Xi, D., Glimm, E., Maurer, W., and Bretz, F. (2017). A unified framework for weighted parametric multiple test procedures. *Biometrical Journal*, 59(5), 918-931.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 4 in Bretz et al. (2011a).
alpha <- 0.025
hypotheses <- c(0.5, 0.5, 0, 0)
delta <- 0.5
transitions <- rbind(
  c(0, delta, 1 - delta, 0),
  c(delta, 0, 0, 1 - delta),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)
```

```

marginal_power <- c(0.8, 0.8, 0.7, 0.9)
corr1 <- matrix(0.5, nrow = 2, ncol = 2)
diag(corr1) <- 1
corr <- rbind(
  cbind(corr1, 0.5 * corr1),
  cbind(0.5 * corr1, corr1)
)
success_fns <- list(
  # Probability to reject both H1 and H2
  `H1andH2` = function(x) x[1] & x[2],
  # Probability to reject both (H1 and H3) or (H2 and H4)
  `(H1andH3)or(H2andH4)` = function(x) (x[1] & x[3]) | (x[2] & x[4])
)
set.seed(1234)
# Bonferroni tests
# Reduce the number of simulations to save time for package compilation
power_output <- graph_calculate_power(
  g,
  alpha,
  sim_corr = corr,
  sim_n = 1e2,
  power_marginal = marginal_power,
  sim_success = success_fns
)

# Parametric tests for H1 and H2; Simes tests for H3 and H4
# User-defined success: to reject H1 or H2; to reject H1 and H2
# Reduce the number of simulations to save time for package compilation
graph_calculate_power(
  g,
  alpha,
  test_groups = list(1:2, 3:4),
  test_types = c("parametric", "simes"),
  test_corr = list(corr1, NA),
  sim_n = 1e2,
  sim_success = list(
    function(.) .[1] || .[2],
    function(.) .[1] && .[2]
  )
)
)

```

graph_create

Create the initial graph for a multiple comparison procedure

Description

A graphical multiple comparison procedure is represented by 1) a vector of initial hypothesis weights hypotheses, and 2) a matrix of initial transition weights transitions. This function creates the initial graph object using hypothesis weights and transition weights.

Usage

```
graph_create(hypotheses, transitions, hyp_names = NULL)
```

Arguments

hypotheses	A numeric vector of hypothesis weights in a graphical multiple comparison procedure. Must be a vector of values between 0 & 1 (inclusive). The length should match the row and column lengths of transitions. The sum of hypothesis weights should not exceed 1.
transitions	A numeric matrix of transition weights between hypotheses in a graphical multiple comparison procedure. Must be a square matrix of values between 0 & 1 (inclusive). The row and column lengths should match the length of hypotheses. Each row (Transition weights leaving a hypothesis) can sum to no more than 1. The diagonal entries (Transition weights from a hypothesis to itself) must be all 0s.
hyp_names	(Optional) A character vector of hypothesis names. If not provided, names from hypotheses and transitions will be used. If names are not specified, hypotheses will be named sequentially as H1, H2,

Value

An S3 object of class `initial_graph` with a list of 2 elements:

- Hypothesis weights hypotheses.
- Transition weights transitions.

Validation of inputs

Inputs are also validated to make sure of the validity of the graph:

- Hypothesis weights hypotheses are numeric.
- Transition weights transitions are numeric.
- Length of hypotheses and dimensions of transitions match.
- Hypothesis weights hypotheses must be non-negative and sum to no more than 1.
- Transition weights transitions:
 - Values must be non-negative.
 - Rows must sum to no more than 1.
 - Diagonal entries must be all 0.
- Hypothesis names hyp_names override names in hypotheses or transitions.

References

- Bretz, F., Maurer, W., Brannath, W., and Posch, M. (2009). A graphical approach to sequentially rejective multiple test procedures. *Statistics in Medicine*, 28(4), 586-604.
- Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

See Also

[graph_update\(\)](#) for the updated graph after hypotheses being deleted from the initial graph.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 1 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
transitions <- rbind(
  c(0, 0, 1, 0),
  c(0, 0, 0, 1),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
hyp_names <- c("H11", "H12", "H21", "H22")
g <- graph_create(hypotheses, transitions, hyp_names)
g

# Explicit names override names in `hypotheses` (with a warning)
hypotheses <- c(h1 = 0.5, h2 = 0.5, h3 = 0, h4 = 0)
transitions <- rbind(
  c(0, 0, 1, 0),
  c(0, 0, 0, 1),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions, hyp_names)
g

# Use names in `transitions`
hypotheses <- c(0.5, 0.5, 0, 0)
transitions <- rbind(
  H1 = c(0, 0, 1, 0),
  H2 = c(0, 0, 0, 1),
  H3 = c(0, 1, 0, 0),
  H4 = c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)
g

# Unmatched names in `hypotheses` and `transitions` (with an error)
hypotheses <- c(h1 = 0.5, h2 = 0.5, h3 = 0, h4 = 0)
transitions <- rbind(
  H1 = c(0, 0, 1, 0),
  H2 = c(0, 0, 0, 1),
  H3 = c(0, 1, 0, 0),
  H4 = c(1, 0, 0, 0)
)
try(
g <- graph_create(hypotheses, transitions)
)
```

```

# When names are not specified, hypotheses are numbered sequentially as
# H1, H2, ...
hypotheses <- c(0.5, 0.5, 0, 0)
transitions <- rbind(
  c(0, 0, 1, 0),
  c(0, 0, 0, 1),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)
g

```

graph_generate_weights

Generate the weighting strategy based on a graphical multiple comparison procedure

Description

A graphical multiple comparison procedure defines a closed test procedure, which tests each intersection hypothesis and reject an individual hypothesis if all intersection hypotheses involving it have been rejected. An intersection hypothesis represents the parameter space where individual null hypotheses involved are true simultaneously.

The closure based on a graph consists of all updated graphs (corresponding to intersection hypotheses) after all combinations of hypotheses are deleted. For a graphical multiple comparison procedure with m hypotheses, there are $2^m - 1$ updated graphs (intersection hypotheses), including the initial graph (the overall intersection hypothesis). The weighting strategy of this graph consists of hypothesis weights from all $2^m - 1$ updated graphs (intersection hypotheses). The algorithm to derive the weighting strategy is based on Algorithm 1 in Bretz et al. (2011).

Usage

```
graph_generate_weights(graph)
```

Arguments

graph An initial graph as returned by `graph_create()`.

Value

A numeric matrix of all intersection hypotheses and their hypothesis weights. For a graphical multiple comparison procedure with m hypotheses, the number of rows is $2^m - 1$, each of which corresponds to an intersection hypothesis. The number of columns is $2 \cdot m$. The first m columns indicate which individual hypotheses are included in a given intersection hypothesis and the second half of columns provide hypothesis weights for each individual hypothesis for a given intersection hypothesis.

Performance

Generation of intersection hypotheses is closely related to the power set of a given set of indices. As the number of hypotheses increases, the memory and time usage can grow quickly (e.g., at a rate of $O(2^n)$). There are also multiple ways to implement Algorithm 1 in Bretz et al. (2011). See vignette("generate-closure") for more information about generating intersection hypotheses and comparisons of different approaches to calculate weighting strategies.

References

Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

See Also

[graph_test_closure\(\)](#) for graphical multiple comparison procedures using the closed test.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 1 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
transitions <- rbind(
  c(0, 0, 1, 0),
  c(0, 0, 0, 1),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)

graph_generate_weights(g)
```

graph_rejection_orderings

Find alternate rejection orderings (sequences) for shortcut tests

Description

When multiple hypotheses are rejected by using [graph_test_shortcut\(\)](#), there may be multiple orderings or sequences in which hypotheses are rejected one by one. The default order in [graph_test_shortcut\(\)](#) is based on the adjusted p-values, from the smallest to the largest. This function [graph_rejection_orderings\(\)](#) provides all possible and valid orders (or sequences) of rejections. Although the order of rejection does not affect the final rejection decisions Bretz et al. (2009), different sequences could offer different ways to explain the step-by-step process of shortcut graphical multiple comparison procedures.

Usage

```
graph_rejection_orderings(shortcut_test_result)
```

Arguments

```
shortcut_test_result
```

A graph_report object as returned by `graph_test_shortcut()`.

Value

A modified graph_report object containing all valid orderings of rejections of hypotheses

References

Bretz, F., Maurer, W., Brannath, W., and Posch, M. (2009). A graphical approach to sequentially rejective multiple test procedures. *Statistics in Medicine*, 28(4), 586-604.

Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

See Also

`graph_test_shortcut()` for shortcut graphical multiple comparison procedures.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 4 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
delta <- 0.5
transitions <- rbind(
  c(0, delta, 1 - delta, 0),
  c(delta, 0, 0, 1 - delta),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)

p <- c(0.018, 0.01, 0.105, 0.006)
alpha <- 0.025

shortcut_testing <- graph_test_shortcut(g, p, alpha, verbose = TRUE)

# Reject H1, H2, and H4
shortcut_testing$outputs$rejected

# Default order of rejections: H2, H1, H4
shortcut_testing$details$del_seq

# There is another valid sequence of rejection: H2, H4, H1
```

```

graph_rejection_orderings(shortcut_testing)$valid_orderings

# Finally, intermediate updated graphs can be obtained by providing the order
# of rejections into `[graph_update()]`
graph_update(g, delete = c(2, 4, 1))

```

graph_test_closure *Perform closed graphical multiple comparison procedures*

Description

Closed graphical multiple comparison procedures, or graphical multiple comparison procedures based on the closure, generate the closure based on a graph consisting of all intersection hypotheses. It tests each intersection hypothesis and rejects an individual hypothesis if all intersection hypotheses involving it have been rejected. An intersection hypothesis represents the parameter space where individual null hypotheses involved are true simultaneously.

For a graphical multiple comparison procedure with m hypotheses, there are $2^m - 1$ intersection hypotheses. For each intersection hypothesis, a test type could be chosen to determine how to reject the intersection hypothesis. Current choices of test types include Bonferroni, Simes and parametric. This implementation offers a more general framework covering Bretz et al. (2011), Lu (2016), and Xi et al. (2017). See vignette("closed-testing") for more illustration of closed test procedures and interpretation of their outputs.

Usage

```

graph_test_closure(
  graph,
  p,
  alpha = 0.025,
  test_groups = list(seq_along(graph$hypotheses)),
  test_types = c("bonferroni"),
  test_corr = rep(list(NA), length(test_types)),
  verbose = FALSE,
  test_values = FALSE
)

```

Arguments

graph	An initial graph as returned by <code>graph_create()</code> .
p	A numeric vector of p-values (unadjusted, raw), whose values should be between 0 & 1. The length should match the number of hypotheses in graph.
alpha	A numeric value of the overall significance level, which should be between 0 & 1. The default is 0.025 for one-sided hypothesis testing problems; another common choice is 0.05 for two-sided hypothesis testing problems. Note when parametric tests are used, only one-sided tests are supported.
test_groups	A list of numeric vectors specifying hypotheses to test together. Grouping is needed to correctly perform Simes and parametric tests.

test_types	A character vector of test types to apply to each test group. This is needed to correctly perform Simes and parametric tests. The length should match the number of elements in test_groups.
test_corr	(Optional) A list of numeric correlation matrices. Each entry in the list should correspond to each test group. For a test group using Bonferroni or Simes tests, its corresponding entry in test_corr should be NA. For a test group using parametric tests, its corresponding entry in test_corr should be a numeric correlation matrix specifying the correlation between test statistics for hypotheses in this test group. The length should match the number of elements in test_groups.
verbose	A logical scalar specifying whether the details of the adjusted p-value calculations should be included in results. When verbose = TRUE, adjusted p-values are provided for each intersection hypothesis. The default is verbose = FALSE.
test_values	A logical scalar specifying whether adjusted significance levels should be provided for each hypothesis. When test_values = TRUE, it provides an equivalent way of performing graphical multiple comparison procedures by comparing each p-value with its significance level. If the p-value of a hypothesis is less than or equal to its significance level, the hypothesis is rejected. The default is test_values = FALSE.

Value

A graph_report object with a list of 4 elements:

- inputs - Input parameters, which is a list of:
 - graph - Initial graph,
 - p - (Unadjusted or raw) p-values,
 - alpha - Overall significance level,
 - test_groups - Groups of hypotheses for different types of tests,
 - test_types - Different types of tests,
 - test_corr - Correlation matrices for parametric tests.
- outputs - Output parameters, which is a list of:
 - adjusted_p - Adjusted p-values,
 - rejected - Rejected hypotheses,
 - graph - Updated graph after deleting all rejected hypotheses.
- details - Verbose outputs with adjusted p-values for intersection hypotheses, if verbose = TRUE.
- test_values - Adjusted significance levels, if test_values = TRUE.

Details for test specification

Test specification includes three components: test_groups, test_types, and test_corr. Alignment among entries in these components is important for correct implementation. There are two ways to provide test specification. The first approach is the "unnamed" approach, which assumes that all 3 components are ordered the same way, i.e., the n -th element of test_types and test_corr should apply to the n -th group in test_groups. The second "named" approach uses

the name of each element of each component to connect the element of `test_types` and `test_corr` with the correct element of `test_groups`. Consistency should be ensured for correct implementation.

References

Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

Lu, K. (2016). Graphical approaches using a Bonferroni mixture of weighted Simes tests. *Statistics in Medicine*, 35(22), 4041-4055.

Xi, D., Glimm, E., Maurer, W., and Bretz, F. (2017). A unified framework for weighted parametric multiple test procedures. *Biometrical Journal*, 59(5), 918-931.

See Also

[graph_test_shortcut\(\)](#) for shortcut graphical multiple comparison procedures.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses
# (H1 and H2) and two secondary hypotheses (H3 and H4)
# See Figure 4 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
delta <- 0.5
transitions <- rbind(
  c(0, delta, 1 - delta, 0),
  c(delta, 0, 0, 1 - delta),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)

p <- c(0.018, 0.01, 0.105, 0.006)
alpha <- 0.025

# Closed graphical multiple comparison procedure using Bonferroni tests
# Same results as `graph_test_shortcut(g, p, alpha)`
graph_test_closure(g, p, alpha)

# Closed graphical multiple comparison procedure using parametric tests for
# H1 and H2, and Bonferroni tests for H3 and H4
set.seed(1234)
corr_list <- list(matrix(c(1, 0.5, 0.5, 1), nrow = 2), NA)
graph_test_closure(
  graph = g,
  p = p,
  alpha = alpha,
  test_groups = list(1:2, 3:4),
  test_types = c("parametric", "bonferroni"),
  test_corr = corr_list
```

```

)
# The "named" approach to obtain the same results
# Note that "group2" appears before "group1" in `test_groups`
set.seed(1234)
corr_list <- list(group1 = matrix(c(1, 0.5, 0.5, 1), nrow = 2), group2 = NA)
graph_test_closure(
  graph = g,
  p = p,
  alpha = alpha,
  test_groups = list(group1 = 1:2, group2 = 3:4),
  test_types = c(group2 = "bonferroni", group1 = "parametric"),
  test_corr = corr_list
)

# Closed graphical multiple comparison procedure using parametric tests for
# H1 and H2, and Simes tests for H3 and H4
set.seed(1234)
graph_test_closure(
  graph = g,
  p = p,
  alpha = alpha,
  test_groups = list(group1 = 1:2, group2 = 3:4),
  test_types = c(group1 = "parametric", group2 = "simes"),
  test_corr = corr_list
)

```

graph_test_shortcut	<i>Perform shortcut (sequentially rejective) graphical multiple comparison procedures</i>
---------------------	---

Description

Shortcut graphical multiple comparison procedures are sequentially rejective procedure based on Bretz et al. (2009). With m hypotheses, there are at most m steps to obtain all rejection decisions. These procedure are equivalent to closed graphical multiple comparison procedures using Bonferroni tests for intersection hypotheses, but shortcut procedures are faster to perform. See vignette("shortcut-testing") for more illustration of shortcut procedures and interpretation of their outputs.

Usage

```

graph_test_shortcut(
  graph,
  p,
  alpha = 0.025,
  verbose = FALSE,
  test_values = FALSE
)

```

Arguments

graph	An initial graph as returned by <code>graph_create()</code> .
p	A numeric vector of p-values (unadjusted, raw), whose values should be between 0 & 1. The length should match the number of hypotheses in graph.
alpha	A numeric scalar of the overall significance level, which should be between 0 & 1. The default is 0.025 for one-sided hypothesis testing problems; another common choice is 0.05 for two-sided hypothesis testing problems.
verbose	A logical scalar specifying whether the details of intermediate update graphs should be included in results. When <code>verbose = TRUE</code> , intermediate update graphs are provided after deleting each hypothesis, which has been rejected. The default is <code>verbose = FALSE</code> .
test_values	A logical scalar specifying whether adjusted significance levels should be provided for each hypothesis. When <code>test_values = TRUE</code> , it provides an equivalent way of performing graphical multiple comparison procedures by comparing each p-value with its significance level. If the p-value of a hypothesis is less than or equal to its significance level, the hypothesis is rejected. The order of rejection is based on the order of adjusted p-values from the smallest to the largest. The default is <code>test_values = FALSE</code> .

Value

An S3 object of class `graph_report` with a list of 4 elements:

- `inputs` - Input parameters, which is a list of:
 - `graph` - Initial graph, `*p` - (Unadjusted or raw) p-values,
 - `alpha` - Overall significance level,
 - `test_groups` - Groups of hypotheses for different types of tests, which are the list of all hypotheses for `graph_test_shortcut()`,
 - `test_types` - Different types of tests, which are "bonferroni" for `graph_test_shortcut()`.
- `Output parameters outputs`, which is a list of:
 - `adjusted_p` - Adjusted p-values,
 - `rejected` - Rejected hypotheses,
 - `graph` - Updated graph after deleting all rejected hypotheses.
- `details` - Verbose outputs with intermediate updated graphs, if `verbose = TRUE`.
- `test_values` - Adjusted significance levels, if `test_values = TRUE`.

References

- Bretz, F., Maurer, W., Brannath, W., and Posch, M. (2009). A graphical approach to sequentially rejective multiple test procedures. *Statistics in Medicine*, 28(4), 586-604.
- Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

See Also

- [graph_test_closure\(\)](#) for graphical multiple comparison procedures using the closed test,
- [graph_rejection_orderings\(\)](#) for all possible rejection orderings.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 1 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
transitions <- rbind(
  c(0, 0, 1, 0),
  c(0, 0, 0, 1),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)

p <- c(0.018, 0.01, 0.105, 0.006)
alpha <- 0.025
graph_test_shortcut(g, p, alpha)
```

graph_update	<i>Obtain an updated graph by updating an initial graphical after deleting hypotheses</i>
--------------	---

Description

After a hypothesis is deleted, an initial graph will be updated. The deleted hypothesis will have the hypothesis weight of 0 and the transition weight of 0. Remaining hypotheses will have updated hypothesis weights and transition weights according to Algorithm 1 of Bretz et al. (2009).

Usage

```
graph_update(graph, delete)
```

Arguments

graph	An initial graph as returned by graph_create() .
delete	A logical or integer vector, denoting which hypotheses to delete. A logical vector results in the "unordered mode", which means that hypotheses corresponding to TRUE in delete will be deleted. The sequence of deletion will follow the sequence of TRUE's in delete. In this case, the length of the logical vector must match the number of hypotheses in graph. An integer vector results in the "ordered mode", which means that delete specifies the sequence in which hypotheses should be deleted by indicating the location of deleted hypotheses, e.g., 1st, 2nd, etc. In this case, the integer vector can have any length, but must only contain valid hypothesis numbers (greater than 0, and less than or equal to the number of hypotheses in graph).

Value

An S3 object of class `updated_graph` with a list of 4 elements:

- `initial_graph`: The initial graph object.
- `updated_graph`: The updated graph object with specified hypotheses deleted.
- `deleted`: A numeric vector indicating which hypotheses were deleted.
- `intermediate_graphs`: When using the ordered mode, a list of intermediate updated graphs after each hypothesis is deleted according to the sequence specified by `delete`.

Sequence of deletion

When there are multiple hypotheses to be deleted from a graph, there are many sequences of deletion in which an initial graph is updated to an updated graph. If the interest is in the updated graph after all hypotheses specified by `delete` are deleted, this updated graph is the same no matter which sequence of deletion is used. This property has been proved by Bretz et al. (2009). If the interest is in the intermediate updated graph after each hypothesis is deleted according to the sequence specified by `delete`, an integer vector of `delete` should be specified and these detailed outputs will be provided.

References

Bretz, F., Maurer, W., Brannath, W., and Posch, M. (2009). A graphical approach to sequentially rejective multiple test procedures. *Statistics in Medicine*, 28(4), 586-604.

Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

See Also

- [graph_create\(\)](#) for the initial graph.
- [graph_rejection_orderings\(\)](#) for possible sequences of rejections for a graphical multiple comparison procedure using shortcut testing.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 1 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
transitions <- rbind(
  c(0, 0, 1, 0),
  c(0, 0, 0, 1),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)

# Delete the second and third hypotheses in the "unordered mode"
graph_update(g, delete = c(FALSE, TRUE, TRUE, FALSE))
```

```
# Equivalent way in the "ordered mode" to obtain the updated graph after
# deleting the second and third hypotheses
# Additional intermediate updated graphs are also provided
graph_update(g, delete = 2:3)
```

plot.initial_graph *S3 plot method for class initial_graph*

Description

The plot of an `initial_graph` translates the hypotheses into vertices and transitions into edges to create a network plot. Vertices are labeled with hypothesis names and hypothesis weights, and edges are labeled with transition weights. See `vignette("graph-examples")` for more illustration of commonly used multiple comparison procedure using graphs.

Usage

```
## S3 method for class 'initial_graph'
plot(
  x,
  ...,
  v_palette = c("#6baed6", "#cccccc"),
  layout = "grid",
  nrow = NULL,
  ncol = NULL,
  edge_curves = NULL,
  precision = 4,
  eps = NULL,
  background_color = "white",
  margins = c(0, 0, 0, 0)
)
```

Arguments

<code>x</code>	An object of class <code>initial_graph</code> to plot.
<code>...</code>	Other arguments passed on to <code>igraph::plot.igraph()</code> .
<code>v_palette</code>	A character vector of length two specifying the colors for retained and deleted hypotheses. More extensive color customization must be done with <code>vertex_color</code> .
<code>layout</code>	An <code>igraph</code> layout specification (See <code>?igraph.plotting</code>), or <code>"grid"</code> , which lays out hypotheses left-to-right and top-to-bottom. <code>nrow</code> and <code>ncol</code> control the grid shape.
<code>nrow</code>	An integer scalar specifying the number of rows in the vertex grid. If row and column counts are not specified, vertices will be laid out as close to a square as possible.

ncol	An integer scalar specifying the number of columns in the vertex grid. If row and column counts are not specified, vertices will be laid out as close to a square as possible.
edge_curves	A named numeric vector specifying the curvature of specific edges. Edge pairs (Where two vertices share an edge in each possible direction) are detected automatically and get 0.25 curvature. Adjust edges by adding an entry with name "vertex1 vertex2, and adjust default edge pairs curvature by adding an entry with name "pairs" - <code>edge_curves = c("pairs" = 0.5, "H1 H3" = 0.25, "H3 H4" = 0.75)</code> .
precision	An integer scalar indicating the number of decimal places to to display.
eps	A numeric scalar. The transition weight of eps will be displayed as ϵ , which indicates edges with infinitesimally small weights. See Bretz et al. (2009) for more details.
background_color	A character scalar specifying a background color for the whole plotting area. Passed directly to <code>graphics::par()</code> (bg).
margins	A length 4 numeric vector specifying the margins for the plot. Defaults to all 0, since igraph plots tend to have large margins. It is passed directly to <code>graphics::par()</code> (mar).

Value

An object `x` of class `initial_graph`, after plotting the initial graph.

Customization of graphs

There are a few values for `igraph::plot.igraph()` that get their defaults changed for graphicalMCP. These values can still be changed by passing them as arguments to `plot.initial_graph()`. Here are the new defaults:

- `vertex.color = "#6baed6"`,
- `vertex.label.color = "black"`,
- `vertex.size = 20`,
- `edge.arrow.size = 1`,
- `edge.arrow.width = 1`,
- `edge.label.color = "black"`
- `asp = 0`.

Neither graphicalMCP nor igraph does anything about overlapping edge labels. If you run into this problem, and vertices can't practically be moved enough to avoid collisions of edge labels, using edge curves can help. igraph puts edge labels closer to the tail of an edge when an edge is straight, and closer to the head of an edge when it's curved. By setting an edge's curve to some very small value, an effectively straight edge can be shifted to a new position.

References

Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

Xi, D., and Bretz, F. (2019). Symmetric graphs for equally weighted tests, with application to the Hochberg procedure. *Statistics in Medicine*, 38(27), 5268-5282.

See Also

[plot.updated_graph\(\)](#) for the plot method for the updated graph after hypotheses being deleted from the initial graph.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 4 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
delta <- 0.5
transitions <- rbind(
  c(0, delta, 1 - delta, 0),
  c(delta, 0, 0, 1 - delta),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)
plot(g)

# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and four secondary hypotheses (H31, H32, H41, and H42)
# See Figure 6 in Xi and Bretz (2019).
hypotheses <- c(0.5, 0.5, 0, 0, 0, 0)
epsilon <- 1e-5
transitions <- rbind(
  c(0, 0.5, 0.25, 0, 0.25, 0),
  c(0.5, 0, 0, 0.25, 0, 0.25),
  c(0, 0, 0, 0, 1, 0),
  c(epsilon, 0, 0, 0, 0, 1 - epsilon),
  c(0, epsilon, 1 - epsilon, 0, 0, 0),
  c(0, 0, 0, 1, 0, 0)
)
hyp_names <- c("H1", "H2", "H31", "H32", "H41", "H42")
g <- graph_create(hypotheses, transitions, hyp_names)

plot_layout <- rbind(
  c(0.15, 0.5),
  c(0.65, 0.5),
  c(0, 0),
  c(0.5, 0),
  c(0.3, 0),
  c(0.8, 0)
```

```
)
plot(g, layout = plot_layout, eps = epsilon, edge_curves = c(pairs = .5))
```

plot.updated_graph *S3 plot method for the class updated_graph*

Description

Plotting an updated graph is a *very* light wrapper around `plot.initial_graph()`, only changing the default vertex color to use gray for deleted hypotheses.

Usage

```
## S3 method for class 'updated_graph'
plot(x, ...)
```

Arguments

`x` An object of class `updated_graph` to plot.

`...` Arguments passed on to `plot.initial_graph`

`v_palette` A character vector of length two specifying the colors for retained and deleted hypotheses. More extensive color customization must be done with `vertex.color`.

`layout` An igraph layout specification (See `?igraph.plotting`), or "grid", which lays out hypotheses left-to-right and top-to-bottom. `nrow` and `ncol` control the grid shape.

`nrow` An integer scalar specifying the number of rows in the vertex grid. If row and column counts are not specified, vertices will be laid out as close to a square as possible.

`ncol` An integer scalar specifying the number of columns in the vertex grid. If row and column counts are not specified, vertices will be laid out as close to a square as possible.

`edge_curves` A named numeric vector specifying the curvature of specific edges. Edge pairs (Where two vertices share an edge in each possible direction) are detected automatically and get 0.25 curvature. Adjust edges by adding an entry with name "vertex1|vertex2", and adjust default edge pairs curvature by adding an entry with name "pairs" - `edge_curves = c("pairs" = 0.5, "H1|H3" = 0.25, "H3|H4" = 0.75)`.

`precision` An integer scalar indicating the number of decimal places to to display.

`eps` A numeric scalar. The transition weight of `eps` will be displayed as ϵ , which indicates edges with infinitesimally small weights. See Bretz et al. (2009) for more details.

`background_color` A character scalar specifying a background color for the whole plotting area. Passed directly to `graphics::par()` (`bg`).

margins A length 4 numeric vector specifying the margins for the plot. Defaults to all 0, since igraph plots tend to have large margins. It is passed directly to `graphics::par()` (`mar`).

Value

An object `x` of class `updated_graph`, after plotting the updated graph.

References

Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

See Also

`plot.initial_graph()` for the plot method for the initial graph.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 1 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
transitions <- rbind(
  c(0, 0, 1, 0),
  c(0, 0, 0, 1),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)

# Delete the second and third hypotheses in the "unordered mode"
plot(
  graph_update(
    g,
    c(FALSE, TRUE, TRUE, FALSE)
  ),
  layout = "grid"
)
```

print.graph_report *S3 print method for the class graph_report*

Description

A printed `graph_report` displays the initial graph, p-values and significance levels, rejection decisions, and optional detailed test results.

Usage

```
## S3 method for class 'graph_report'
print(x, ..., precision = 4, indent = 2, rows = 10)
```

Arguments

x	An object of class graph_report to print.
...	Other values passed on to other methods (currently unused)
precision	An integer scalar indicating the number of decimal places to to display.
indent	An integer scalar indicating how many spaces to indent results.
rows	An integer scalar indicating how many rows of detailed test results to print.

Value

An object x of class graph_report, after printing the report of conducting a graphical multiple comparison procedure.

References

Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 1 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
transitions <- rbind(
  c(0, 0, 1, 0),
  c(0, 0, 0, 1),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)

p <- c(0.018, 0.01, 0.105, 0.006)
alpha <- 0.025
graph_test_shortcut(g, p, alpha)
```

```
print.initial_graph    S3 print method for the class initial_graph
```

Description

A printed `initial_graph` displays a header stating "Initial graph", hypothesis weights, and transition weights.

Usage

```
## S3 method for class 'initial_graph'
print(x, ..., precision = 4, indent = 0)
```

Arguments

<code>x</code>	An object of class <code>initial_graph</code> to print.
<code>...</code>	Other values passed on to other methods (currently unused).
<code>precision</code>	An integer scalar indicating the number of decimal places to display.
<code>indent</code>	An integer scalar indicating how many spaces to indent results.

Value

An object `x` of class `initial_graph`, after printing the initial graph.

References

Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

See Also

[print.updated_graph\(\)](#) for the print method for the updated graph after hypotheses being deleted from the initial graph.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 1 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
transitions <- rbind(
  c(0, 0, 1, 0),
  c(0, 0, 0, 1),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
hyp_names <- c("H11", "H12", "H21", "H22")
```



```
g <- graph_create(hypotheses, transitions, hyp_names)
g
```

```
print.power_report      S3 print method for the class power_report
```

Description

A printed `power_report` displays the initial graph, testing and simulation options, power outputs, and optional detailed simulations and test results.

Usage

```
## S3 method for class 'power_report'
print(x, ..., precision = 4, indent = 2, rows = 10)
```

Arguments

<code>x</code>	An object of the class <code>power_report</code> to print
<code>...</code>	Other values passed on to other methods (currently unused)
<code>precision</code>	An integer scalar indicating the number of decimal places to display.
<code>indent</code>	An integer scalar indicating how many spaces to indent results.
<code>rows</code>	An integer scalar indicating how many rows of detailed test results to print.

Value

An object `x` of the class `power_report`, after printing the report of conducting power simulations based on a graphical multiple comparison procedure.

References

Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011a). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

Bretz, F., Maurer, W., and Hommel, G. (2011b). Test and power considerations for multiple end-point analyses using sequentially rejective graphical procedures. *Statistics in Medicine*, 30(13), 1489-1501.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 4 in Bretz et al. (2011).
alpha <- 0.025
hypotheses <- c(0.5, 0.5, 0, 0)
delta <- 0.5
transitions <- rbind(
```

```

    c(0, delta, 1 - delta, 0),
    c(delta, 0, 0, 1 - delta),
    c(0, 1, 0, 0),
    c(1, 0, 0, 0)
  )
  g <- graph_create(hypotheses, transitions)

  marginal_power <- c(0.8, 0.8, 0.7, 0.9)
  corr1 <- matrix(0.5, nrow = 2, ncol = 2)
  diag(corr1) <- 1
  corr <- rbind(
    cbind(corr1, 0.5 * corr1),
    cbind(0.5 * corr1, corr1)
  )
  success_fns <- list(
    # Probability to reject both H1 and H2
    `H1andH2` = function(x) x[1] & x[2],
    # Probability to reject both (H1 and H3) or (H2 and H4)
    `(H1andH3)or(H2andH4)` = function(x) (x[1] & x[3]) | (x[2] & x[4])
  )
  set.seed(1234)
  # Bonferroni tests
  power_output <- graph_calculate_power(
    g,
    alpha,
    sim_corr = corr,
    sim_n = 1e5,
    power_marginal = marginal_power,
    sim_success = success_fns
  )

```

```
print.updated_graph    S3 print method for the class updated_graph
```

Description

A printed `updated_graph` displays the initial graph, the (final) updated graph, and the sequence of intermediate updated graphs after hypotheses are deleted (if available).

Usage

```
## S3 method for class 'updated_graph'
print(x, ..., precision = 6, indent = 2)
```

Arguments

<code>x</code>	An object of the class <code>updated_graph</code> to print.
<code>...</code>	Other values passed on to other methods (currently unused).
<code>precision</code>	An integer scalar indicating the number of decimal places to to display.
<code>indent</code>	An integer scalar indicating how many spaces to indent results.

Value

An object `x` of the class `updated_graph`, after printing the updated graph.

References

Bretz, F., Posch, M., Glimm, E., Klinglmueller, F., Maurer, W., and Rohmeyer, K. (2011a). Graphical approaches for multiple comparison procedures using weighted Bonferroni, Simes, or parametric tests. *Biometrical Journal*, 53(6), 894-913.

See Also

[print.initial_graph\(\)](#) for the print method for the initial graph.

Examples

```
# A graphical multiple comparison procedure with two primary hypotheses (H1
# and H2) and two secondary hypotheses (H3 and H4)
# See Figure 1 in Bretz et al. (2011).
hypotheses <- c(0.5, 0.5, 0, 0)
transitions <- rbind(
  c(0, 0, 1, 0),
  c(0, 0, 0, 1),
  c(0, 1, 0, 0),
  c(1, 0, 0, 0)
)
g <- graph_create(hypotheses, transitions)

# Delete the second and third hypotheses in the "unordered mode"
graph_update(g, delete = c(FALSE, TRUE, TRUE, FALSE))

# Equivalent way in the "ordered mode" to obtain the updated graph after
# deleting the second and third hypotheses
# Additional intermediate updated graphs are also provided
graph_update(g, delete = 2:3)
```

Index

`adjust_p_bonferroni`, 2
`adjust_p_bonferroni()`, 2
`adjust_p_parametric`
 (`adjust_p_bonferroni`), 2
`adjust_p_parametric()`, 2, 3, 5
`adjust_p_simes` (`adjust_p_bonferroni`), 2
`adjust_p_simes()`, 2, 5
`adjust_weights_parametric`, 4
`adjust_weights_parametric()`, 3–5
`adjust_weights_simes`
 (`adjust_weights_parametric`), 4
`adjust_weights_simes()`, 3–5
`as_graphMCP` (`as_initial_graph`), 6
`as_igraph` (`as_initial_graph`), 6
`as_initial_graph`, 6

`bonferroni`, 8
`bonferroni_holm` (`bonferroni`), 8

`fallback` (`bonferroni`), 8
`fallback_improved_1` (`bonferroni`), 8
`fallback_improved_2` (`bonferroni`), 8
`fixed_sequence` (`bonferroni`), 8

`graph_calculate_power`, 10
`graph_create`, 13
`graph_create()`, 7, 9, 10, 16, 19, 23–25
`graph_generate_weights`, 16
`graph_generate_weights()`, 4
`graph_rejection_orderings`, 17
`graph_rejection_orderings()`, 17, 24, 25
`graph_test_closure`, 19
`graph_test_closure()`, 17, 24
`graph_test_shortcut`, 22
`graph_test_shortcut()`, 17, 18, 21, 23
`graph_update`, 24
`graph_update()`, 15
`graphics::par()`, 27, 29, 30

`huque_etal` (`bonferroni`), 8

`igraph::plot.igraph()`, 27

`plot.initial_graph`, 26, 29
`plot.initial_graph()`, 29, 30
`plot.updated_graph`, 29
`plot.updated_graph()`, 28
`print.graph_report`, 30
`print.initial_graph`, 32
`print.initial_graph()`, 35
`print.power_report`, 33
`print.updated_graph`, 34
`print.updated_graph()`, 32

`random_graph` (`bonferroni`), 8

`simple_successive_1` (`bonferroni`), 8
`simple_successive_2` (`bonferroni`), 8

`three_doses_two_primary_two_secondary`
 (`bonferroni`), 8
`two_doses_two_primary_two_secondary`
 (`bonferroni`), 8