# **Package**

June 21, 2024

**Title** Bayesian network meta-analyses in compliance with best practice and reporting guidelines **Version** 1.1.2

Description BUGSnet (Bayesian inference Using Gibbs Sampling to conduct NETwork meta-analysis) is a feature-rich R package to conduct Bayesian network meta-analyses in compliance with best practice and reporting guidelines. Bayesian analyses are conducted with JAGS. Outputs are highly customizable and include network plots, tables of network characteristics, league tables and league heat plots, SUCRA plots, rankograms, forest plots, leverage plots, traceplots, posterior mean deviance comparison plots.

```
License CC BY-NC-SA 4.0
License_is_FOSS yes
License_restricts_use yes
URL https://bugsnetsoftware.github.io
BugReports https://github.com/audrey-b/BUGSnet/issues
Depends R (>= 4.2)
Imports coda,
     dplyr,
     ggplot2,
     graphics,
     gridExtra,
     igraph,
     magrittr,
     meta,
     plyr,
     purrr,
     RColorBrewer,
     Rdpack,
     rjags,
     rlang,
     scales,
     stringr,
     tibble,
     tidyr,
     tidyselect,
     utils
```

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afib Atrial Fibrillation data

# **Description**

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The atrial fibrillation data was previously used to demonstrate the network meta-analysis capabilities of the R package 'gemtc'. This dataset was originally generated by Cooper et al. (2009) and was subsequently revised by van Valkenhoef et al (2013). This dataset provides information from 25 clinical trials examining the effects of various interventions (17) on the risk of stroke (binary outcome) among individuals with non-rheumatic atrial fibrillation. The stroke variable is a covariate indicating the proportion of patients in the study that had a prior stroke.

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#### Usage

data(afib)

#### **Format**

A dataset.

### References

Cooper NJ, Sutton AJ, Morris D, Ades AE, Welton NJ (2009). "Adressing between-study heterogeneity and inconsistency in mixed treatment comparisons: Application to stroke prevention treatments in individuals with non-rheumatic atrial fibrillation." *Statistics in Medicine*, **28**, 1861-1881.

van Valkenhoef, G, Lu, G, de Brock, B, Hillege, H, Ades, AE, Welton, NJ (2012). "Automating network meta-analysis." *Research Synthesis Methods*, **3**(4), 285-299.

# **Examples**

data(afib)

BUGSnet

BUGSnet: Bayesian network meta-analysis in compliance with best practice guidelines.

### **Description**

BUGSnet (Bayesian inference Using Gibbs Sampling to conduct NETwork meta-analysis) is a feature-rich R package to conduct Bayesian network meta-analyses in compliance with best practice and reporting guidelines. Bayesian analyses are conducted with JAGS and BUGS code is automatically generated by the package based on the user's inputs. Outputs are highly customizable and include network plots, tables of network characteristics, league tables and league heat plots, SUCRA plots, rankograms, forest plots, leverage plots, traceplots, posterior mean deviance comparison plots.

#### Details

To be able to use this package, you must have installed JAGS (Just Another Gibbs Sampler) on your computer.

How to cite this work? Type citation("BUGSnet").

#### References

Plummer M (2017). *JAGS Version 4.3.0 user manual*. http://people.stat.sc.edu/hansont/stat740/jags\_user\_manual.pdf, last accessed 2019-03-06.

Plummer M (2017). "JAGS download page." http://mcmc-jags.sourceforge.net/, last accessed 2019-03-06.

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data.plot

Patient Characteristic Plot

### **Description**

Plots a particular patient characteristic by study or by treatment. Useful for assessing differences in potential effect modifiers.

# Usage

```
data.plot(
  data,
  covariate,
  half.length = NULL,
  by = "study",
  avg.hline = TRUE,
  fill.str = NULL,
  text.size = 20,
  coord.flip = FALSE
)
```

# **Arguments**

data A BUGSnetData object produced by data.prep() covariate A string indicating the name of the patient characteristic to be plotted half.length A string indicating how to calculate the half-length of error bars (optional) If by="study" then data from arms will be grouped by study/trial. If by="treatment" by then bar graph is grouped by treatment. avg.hline If TRUE, adds overall average line to plot. Default is TRUE. An optional string indicating the variable to categorize measurements. For infill.str stance, some studies report the mean treatment effect and others may report the median treatment effect. If there is a variable in data called "type.measure" indicating whether the mean or median is reported, setting fill.str="type.measure" would colour all studies reporting the mean as red, and all the studies reporting the median as turquoise. text.size Font size of the text. Default is 20. If TRUE, coordinates will be flipped to display the plot in portrait mode. Default coord.flip is FALSE.

#### See Also

```
data.prep
```

```
data(diabetes.sim)
diabetes.slr <- data.prep(arm.data = diabetes.sim,
varname.t = "Treatment",
varname.s = "Study")</pre>
```

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data.prep

Data preparation

### **Description**

Puts data into appropriate form for subsequent analysis using BUGSnet.

# Usage

```
data.prep(arm.data, varname.t, varname.s)
```

### **Arguments**

arm.data
Data with 1 row for each study arm.

varname.t
A string indicating the name of the treatment variable.

varname.s
A string indicating the name of the study variable.

# Value

data.prep returns an object of class BUGSnetData which is a list containing the following components:

arm. data - A tibble containing the arm level study data

treatments - A list of all treatments in the network

studies - A list of all studies in the network

n.arms - A tibble containing the number of arms for each study

varname.t - A string containing the name of the treatment variable

varname.s - A string containing the name of the study variable

# See Also

```
net.plot, net.tab, data.plot, nma.model
```

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# **Examples**

```
data(diabetes.sim)
diabetes.slr <- data.prep(arm.data = diabetes.sim,
varname.t = "Treatment",
varname.s = "Study")
diabetes.slr$arm.data
diabetes.slr$treatments
diabetes.slr$studies</pre>
```

diabetes

Diabetes data

# **Description**

The diabetes dataset has been used to illustrate network meta-analysis within the NICE-DSU TSD 2 (examples 3a and 3b). The dataset is comprised of information from 22 clinical trials examining the effect of antihypertensive drugs (there were six treatments including a placebo arm) on the risk of developing diabetes (a binary outcome). This dataset also contains information on the years of follow-up within each arm.

# Usage

```
data(diabetes)
```

#### **Format**

A dataset.

## References

Dias S, Welton NJ, Sutton AJ, Ades AE (2011). "NICE DSU technical support document 2: a generalised linear modelling framework for pairwise and network meta-analysis of randomised controlled trials." National Institute for Health and Clinical Excellence.

Elliott WJ, Meyer PM (2007). "Incident diabetes in clinical trials of antihypertensive drugs: a network meta-analysis." *The Lancet*, **369**(9557), 201–207.

```
data(diabetes)
```

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diabetes.sim

Diabetes data with simulated age covariate

# **Description**

This is the same as the Diabetes dataset but we have added a simulated age variable to this dataset for the purpose of demonstrating certain features within BUGSNET.

# Usage

```
data(diabetes.sim)
```

### **Format**

A dataset.

#### References

Dias S, Welton NJ, Sutton AJ, Ades AE (2011). "NICE DSU technical support document 2: a generalised linear modelling framework for pairwise and network meta-analysis of randomised controlled trials." National Institute for Health and Clinical Excellence.

Elliott WJ, Meyer PM (2007). "Incident diabetes in clinical trials of antihypertensive drugs: a network meta-analysis." *The Lancet*, **369**(9557), 201–207.

### **Examples**

```
data(diabetes.sim)
```

net.plot

Network Plot

# Description

Produces network plot where nodes represent treatments and edges represent direct evidence (e.g an RCT) comparing treatments.

# Usage

```
net.plot(
  data,
  node.scale = 5,
  edge.scale = 2,
  flag = NULL,
  study.counts = FALSE,
  label.offset1 = 0,
  label.offset2 = 1,
  graph.scale = TRUE,
  node.lab.cex = 1,
  edge.lab.cex = 1,
  node.colour = "#f69c54",
```

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```
edge.colour = "grey",
edge.lab.colour = "blue",
flag.edge.colour = "lightpink",
layout = "layout_in_circle",
layout.params = NULL
)
```

### **Arguments**

A BUGSnetData object produced by data.prep(). data node.scale Size of the nodes (default=5) Thickness of the edges (default=2). edge.scale flag Used to highlight direct comparisons to particular treatments (optional). Set this value to treatment(s) of interest and it will highlight, in red, all of the edges going into the specified treatment(s). study.counts If TRUE, prints the number of studies on each edge. label.offset1 Node label location (x-axis) relative to node. Default=0 label.offset2 Node label location (y-axis) relative to node. Default=1 graph.scale Whether to make edges and nodes proportionnaly larger with the number of studies/arms. Default is TRUE. Size of node labels node.lab.cex edge.lab.cex Size of edge labels node.colour Node colour (string) edge.colour Edge colour (string) edge.lab.colour Edge label colour (string) flag.edge.colour Color of flagged edges (string) Specifies how the nodes should be layed out in the graph. Default is "laylayout

out\_in\_circle". See igraph::layout\_for layout options

layout.params Additional parameters to be passed to the chosen 'layout' function. See igraph::layout\_

for more information

#### See Also

```
data.prep, igraph::layout_
```

```
data(diabetes.sim)
diabetes.slr <- data.prep(
  arm.data = diabetes.sim,
  varname.t = "Treatment",
  varname.s = "Study"
)
# use default settings
net.plot(diabetes.slr)</pre>
```

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```
# Highlight all direct comparisons with Placebo. Adjust node and edge size, centre node labels
net.plot(
   data = diabetes.slr,
   node.scale = 4,
   edge.scale = 1.5,
   flag="Placebo",
   label.offset1 = 0,
   label.offset2 = 0
)
```

net.tab

Generate Network Characteristics

# Description

Generates tables of network characteristics

#### Usage

```
net.tab(data, outcome, N, type.outcome, time = NULL)
```

### **Arguments**

data A BUGSnetData object produced by data.prep()
outcome A string indicating the name of the outcome variable

N A string indicating the name of the variable containing the number of partici-

pants in each arm

type.outcome A string. Options are: "binomial", "continuous", "rate" (e.g # of events and #

person-time reported), "rate2" (e.g # events and followup time reported)

time A string required when type.outcome = "rate" or "rate2". Name of variable

indicating person-time followup (e.g person years) or study followup time

# Value

```
network - Table of network characteristics
intervention - Summary statistics broken down by treatment
comparison - Summary statistics broken down by treatment comparison
```

# See Also

```
data.prep
```

```
data(diabetes.sim)
diabetes.slr <- data.prep(arm.data = diabetes.sim,
varname.t = "Treatment",
varname.s = "Study")
network.char <- net.tab(data = diabetes.slr,</pre>
```

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```
outcome = "diabetes",
N = "n",
type.outcome = "rate2",
time = "followup")
```

nma.compare

Consistency vs Inconsistency plot

# **Description**

Plots the posterior mean deviance of a consistency model vs an inconsistency model. This plot can help identify loops where inconsistency is present. Ideally, both models will contribute approximately 1 to the posterior mean deviance.

# Usage

```
nma.compare(consistency.model.fit, inconsistency.model.fit, ...)
```

# Arguments

```
consistency.model.fit
Results of nma.fit() of an consistency model.

inconsistency.model.fit
Results of nma.fit() of an inconsistency model.

Graphical arguments such as main=, ylab=, and xlab= may be passed in plot().
```

# See Also

nma.run

```
data(diabetes.sim)
diabetes.slr <- data.prep(</pre>
  arm.data = diabetes.sim,
 varname.t = "Treatment",
  varname.s = "Study"
)
#Random effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.re.c <- nma.model(</pre>
  data = diabetes.slr,
  outcome = "diabetes",
 N = "n"
  reference = "Placebo",
  family = "binomial",
  link = "cloglog",
  effects = "random",
  type = "consistency",
  time = "followup"
```

nma.diag

```
diabetes.re.c.res <- nma.run(</pre>
  model = diabetes.re.c,
  n.adapt = 100,
 n.burnin = 0,
 n.iter = 100
)
#Random effects, inconsistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.re.i <- nma.model(</pre>
  data = diabetes.slr,
  outcome = "diabetes",
 N = "n"
  reference = "Placebo",
  family = "binomial",
  link = "cloglog",
  effects = "random",
  type = "inconsistency",
  time = "followup"
diabetes.re.i.res <- nma.run(</pre>
  model = diabetes.re.i,
  n.adapt = 100,
 n.burnin = 0,
  n.iter = 100
)
# Assess model fit for a both an inconsistency model and consistency model using nma.fit()
assess.consistency <- nma.fit(diabetes.re.c.res)</pre>
assess.inconsistency <- nma.fit(diabetes.re.i.res)</pre>
#Plot the results against each other to assess inconsistency
nma.compare(assess.consistency, assess.inconsistency)
```

nma.diag

Trace plots and convergence diagnostics for MCMC chains

# **Description**

Produces trace plots and Gelman-Rubin and Geweke convergence diagnostics for the MCMC chains obtained from nma.run(). The Gelman-Rubin and Geweke diagnostics are implemented using functions from the coda package.

# Usage

```
nma.diag(
  nma,
  trace = TRUE,
  gelman.rubin = TRUE,
  geweke = TRUE,
  params = "all",
  thin = 1,
  ncol = 1,
```

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```
nrow = 3,
plot_prompt = TRUE,
geweke_frac1 = 0.1,
geweke_frac2 = 0.5
)
```

### **Arguments**

nma A BUGSnetRun object produced by nma.run() trace If TRUE, outputs trace plots. Default is TRUE.

gelman.rubin If TRUE, runs Gelman-Rubin diagnostic. Default is TRUE.

geweke If TRUE, runs Geweke diagnostic. Default is TRUE.

params Integer or character vector which specifies which parameters to produce trace

plots for when trace is set to TRUE. Default is "all" which plots every monitored

parameter.

thin Thinning factor for the meme chains when producing trace plots. Default is 1.

ncol Number of columns in each batch of trace plots

nrow Number rows in each batch of trace plots

plot\_prompt If TRUE, prompts the user to hit enter before plotting each additional batch of

trace plots. Default is TRUE.

geweke\_frac1 Fraction to use from beginning of chain. Default is 0.1.

geweke\_frac2 Fraction to use from end of chain. Default is 0.5.

### Value

gelman.rubin An object of class gelman.rubin.results containing the Gelman-Rubin diagnostic results. A formatted table with custom PSRF threshold can be printed using print(x, gelman.rubin.threshold = 1.2).

geweke An object of class geweke.results containing the Geweke diagnostic results. A formatted table with custom significance level can be printed using print(x, alpha = 0.05).

# See Also

```
nma.run
```

nma.fit

|--|

### **Description**

Computes the Deviance Information Criteria (DIC) and produces a leverage plot (as in the NICE Technical Support Document 2) for a given model. These can be used to assess and compare the fit of different models (i.e fixed vs random effects, consistency vs inconsistency). nma.fit also produces a plot comparing the leverage of each data point against their contribution to the total posterior deviance. Points lying outside the purple dotted line are generally identified as contributing to the model's poor fit. Points with high leverage are influencial i.e. they have a stong influence on the estimates.

# Usage

```
nma.fit(nma, plot.pD = TRUE, plot.DIC = TRUE, plot.Dres = TRUE, ...)
```

#### **Arguments**

nma	A BUGSnetRun object produced by running nma.run().
plot.pD	Whether to include pD on the plot. Default is TRUE.
plot.DIC	Whether to include DIC on the plot. Default is TRUE.
plot.Dres	Whether to include Dres on the plot. Default is TRUE.
	Graphical arguments such as main=, ylab=, and xlab= may be passed as in plot(). These arguments will only effect the leverage plot.

### Value

DIC - A number indicating the Deviance Information Criteria. The DIC is calculated as the sum of Dres and pD. A larger DIC is indicative of a worse model fit.

leverage - A vector with one value per study arm indicating the leverage of each data point (study arm). Leverage is defined as pmdev minus the deviance at the posterior mean of the fitted values.

w - A vector with one value per study arm. The magnitude of w represents the data point's contribution to the posterior mean deviance of the model and is simply the square root of pmdev. The sign indicates whether the data point is being over (negative sign) or under (positive sign) estimated by the model and is calculated as the sign of the difference of the observed outcome minus the predicted outcome.

pmdev - A vector with one value per study arm representing the posterior mean residual deviance for each data point (study arm).

Dres - The posterior mean of the residual deviance.

pD - The effective number of parameters, calculated as the sum of the leverages.

# See Also

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```
data(diabetes.sim)
diabetes.slr <- data.prep(</pre>
  arm.data = diabetes.sim,
  varname.t = "Treatment",
  varname.s = "Study"
#Random effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.re.c <- nma.model(</pre>
  data = diabetes.slr,
 outcome = "diabetes",
 N = "n"
 reference = "Placebo",
  family = "binomial",
 link = "cloglog",
 effects = "random",
  type = "consistency",
 time = "followup"
)
diabetes.re.c.res <- nma.run(</pre>
 model = diabetes.re.c,
 n.adapt = 100,
 n.burnin = 0,
 n.iter = 100
#Fixed effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.fe.c <- nma.model(</pre>
 data = diabetes.slr,
 outcome = "diabetes",
 N = "n"
 reference = "Placebo",
  family = "binomial",
 link = "cloglog",
 effects = "fixed",
  type="consistency",
  time="followup"
diabetes.fe.c.res <- nma.run(</pre>
 model = diabetes.fe.c,
 n.adapt = 100,
 n.burnin = 0,
 n.iter = 100
#Compare fixed vs random effects via leverage plots and DIC
par(mfrow=c(1,2))
nma.fit(diabetes.fe.c.res, main = "Fixed Effects Model")
nma.fit(diabetes.re.c.res, main= "Random Effects Model")
```

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nma.forest	Forest Plot

# Description

Produces a forest plot of point estimates and 95% credible intervals obtained with the quantile method.

# Usage

```
nma.forest(
  nma,
  comparator,
  central.tdcy = "median",
  order = NULL,
  log.scale = FALSE,
  lwd = 1,
  x.trans = NULL,
  cov.value = NULL
)
```

# Arguments

nma	A BUGSnetRun object produced by running nma.run().
comparator	The treatment to use as a comparator
central.tdcy	The posterior statistic used in order to measure relative effectiveness. The options are "mean" and "median". Default is median.
order	Optional. A vector of strings representing the order in which to display the treatments.
log.scale	If TRUE, odds ratios, relative risk or hazard ratios are reported on the log scale. Default is FALSE.
lwd	Line width relative to the default (default=1).
x.trans	Optional. A string indicating a transformation to apply to the x-axis. Setting this parameter to "log" is useful when there are extreme values or to allow an easier interpretation of odds ratios and relative ratios (if e.g. treatment B is twice as far from the line y=1 then treatment A then it's OR/RR is twice that of treatment A.)
cov.value	Must be specified for meta-regression. This is the value of the covariate for which to report the results.

# Value

forestplot - A forest plot.

# See Also

```
nma.run, nma.league, nma.rank
```

nma.league

### **Examples**

```
data(diabetes.sim)
diabetes.slr <- data.prep(arm.data = diabetes.sim,</pre>
varname.t = "Treatment",
varname.s = "Study")
#Random effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.re.c <- nma.model(</pre>
  data = diabetes.slr,
  outcome = "diabetes",
 N = "n"
  reference = "Placebo",
  family = "binomial",
  link = "cloglog",
  effects = "random",
  type = "consistency",
  time = "followup"
)
diabetes.re.c.res <- nma.run(</pre>
  model = diabetes.re.c,
  n.adapt = 100,
 n.burnin = 0,
  n.iter = 100
#make forest plot
nma.forest(nma = diabetes.re.c.res, comparator="Placebo")
```

nma.league

League Table and Heat Plot

# Description

Produces a league table and a league heat plot that contain point estimates of relative effectiveness for all possible pairs of treatments point estimates along with 95% credible intervals obtained with the quantile method.

# Usage

```
nma.league(
  nma,
  central.tdcy = "median",
  log.scale = FALSE,
  order = NULL,
  low.colour = "darkgoldenrod1",
  mid.colour = "white",
  high.colour = "cornflowerblue",
  cov.value = NULL,
  digits = 2
)
```

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# **Arguments**

nma	A BUGSnetRun object produced by running nma.run().
central.tdcy	The statistic that you want to use in order to measure relative effectiveness. The options are "mean" and "median".
log.scale	If TRUE, odds ratios, relative risk or hazard ratios are reported on the log scale. Default is FALSE.
order	A vector of strings representing the order in which to display the treatments.
low.colour	A string indicating the colour of low relative treatment effects for the heat plot (e.g relative risk of $0.5$ ).
mid.colour	A string indicating the colour of null relative treatment effects for the heat plot (e.g relative risk of $\sim$ 1.0).
high.colour	A string indicating the colour of high relative treatment effects for the heat plot (e.g relative risk of $\sim$ 2.0).
cov.value	Must be specified for meta-regression. This is the value of the covariate for which to report the results.
digits	Number of digits to display after the decimal point

### Value

table - A league table. Row names indicate comparator treatments.

longtable - League table in the long format.

heatplot - League heat plot, where a color scale is used to represent relative treatment effects and \*\* are used to highlight statistically significant differences.

# See Also

```
nma.run, nma.rank, nma.forest
```

```
data(diabetes.sim)
diabetes.slr <- data.prep(</pre>
 arm.data = diabetes.sim,
  varname.t = "Treatment",
  varname.s = "Study"
#Random effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.re.c <- nma.model(</pre>
 data = diabetes.slr,
 outcome = "diabetes",
 N = "n"
 reference = "Placebo",
  family = "binomial",
 link = "cloglog",
  effects = "random",
  type = "consistency",
  time = "followup"
```

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```
diabetes.re.c.res <- nma.run(
  model = diabetes.re.c,
  n.adapt = 100,
  n.burnin = 0,
  n.iter = 100)

league_table <- nma.league(
  nma = diabetes.re.c.res,
  central.tdcy = "median"
)

league_table$heatplot
league_table$table
league_table$longtable</pre>
```

nma.model

Create Bugs Model for Arm-Level Data

# Description

Creates BUGS code which can be ran through nma.run().

# Usage

```
nma.model(
  data = NULL,
  outcome,
  N = NULL,
  sd = NULL,
  reference,
  type = "consistency",
  time = NULL,
  family = NULL,
  link = NULL,
  effects,
  prior.mu = "DEFAULT",
  prior.d = "DEFAULT",
  prior.sigma = "DEFAULT",
  prior.beta = NULL,
  covariate = NULL
)
```

# Arguments

Ν

data A BUGSnetData object containing the data from arm-based trials produced by

data.prep()

outcome A string indicating the name of your outcome variable.

A string indicating the name of the variable containing the number of partici-

pants in each arm.

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A string (only required for continuous outcomes) indicating variable name of

the standard deviation of the outcome. Standard errors should be converted to standard deviation by multiplying by the square root of the sample size prior to

using this function.

reference A string for the treatment that will be seen as the 'referent' comparator and

labeled as treatment 1 in the BUGS code. This is often a placebo or control drug

of some kind.

type If type="inconsistency", an inconsistency model will be built. By default, type="consistency"

and a consistency model is built. will be built.

time A string (only required for binomial-cloglog or poisson-log models) indicating

the name of variable indicating person-time followup (e.g person years) or study

followup time.

family A string indicating the family of the distribution of the outcome. Options are:

"binomial", "normal", "poisson".

link The link function for the nma model. Options are "logit" (binomial family),

"log" (binomial or poisson family), "cloglog" (binomial family), "identity" (nor-

mal family).

effects A string indicating the type of treatment effect relative to baseline. Options are

"fixed" or "random".

prior.mu A string of BUGS code that defines priors on the baseline treatment effects. By

default, independent normal priors are used with mean 0 and standard deviation 15u, where u is the largest maximum likelihood estimator in single trials (see

van Valkenhoef, G et al. 2012).

prior.d A string of BUGS code that defines define priors on relative treatment effects.

By default, independent normal priors are used with mean 0 and standard deviation 15u, where u is the largest maximum likelihood estimator in single trials

(see van Valkenhoef, G et al. 2012).

prior.sigma A string of BUGS code that defines the prior on the standard deviation of relative

treatment effects. By default, a uniform distribution with range 0 to u is used, where u is the largest maximum likelihood estimator in single trials (see van

Valkenhoef, G et al. 2012).

prior.beta Optional string that defines the prior on the meta-regression coefficients. Op-

tions are "UNRELATED", "EXCHANGEABLE", "EQUAL" (TSD3) or a string

of BUGS code.

covariate Optional string indicating the name of the variable in your data set that you

would like to adjust for via meta regression. By default, covariate=NULL and no covariate adjustment is applied. If the specified covariate is numeric then it will be centered for the analysis. If it is a character or factor then it will be treated as categorical. Currently only categorical variables with fewer than 3

levels are supported.

## **Details**

For meta-regression, the prespecified prior choices for the regression coefficients  $\beta_{(1,2)}, \dots, \beta_{(1,K)}$  are:

**Unrelated:** 

 $iidt(0, u^2, 1)$ 

**Exchangeable:** 

 $iidN(b, \gamma^2), b t(0, u^2, 1), \gamma U(0, u)$ 

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#### **Equal:**

$$\beta_2 = \dots = \beta_T = B, B \ t(0, u^2, 1)$$

where u is the largest maximum likelihood estimator in single trials (see van Valkenhoef, G et al. 2012).

#### Value

nma.model returns an object of class BUGSnetModel which is a list containing the following components:

bugs - A long character string containing BUGS code that will be run in jags.

data - The data used in the BUGS code.

scale - The scale of the outcome, based on the chosen family and link function examples are "Risk Ratio" (relative risk), "Odds Ratio", "Mean Difference", "Hazard Ratio"

trt.key - Treatments mapped to integer numbers, used to run BUGS code.

•••

#### References

van Valkenhoef, G, Lu, G, de Brock, B, Hillege, H, Ades, AE, Welton, NJ (2012). "Automating network meta-analysis." *Research Synthesis Methods*, **3**(4), 285-299.

Dias S, Sutton AJ, Welton NJ, Ades AE (2011). "NICE DSU technical support document 3. Heterogeneity: subgroups, meta-regression, bias and bias-adjustment." National Institute for Health and Clinical Excellence. Last updated April 2012.

### See Also

```
data.prep, nma.run
```

```
data(diabetes.sim)
diabetes.slr <- data.prep(arm.data = diabetes.sim,</pre>
varname.t = "Treatment",
varname.s = "Study")
#Random effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.re.c <- nma.model(data = diabetes.slr,</pre>
       outcome = "diabetes",
       N = "n"
       reference = "Placebo",
       family = "binomial",
       link = "cloglog",
       effects = "random"
       type="consistency",
       time="followup"
#Fixed effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.fe.c <- nma.model(data = diabetes.slr,</pre>
       outcome = "diabetes",
```

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```
N = "n",
reference = "Placebo",
family = "binomial",
link = "cloglog",
effects = "fixed",
type="consistency",
time="followup"
)
```

nma.model.contrast

Create Bugs Model for contrast-Level data

# Description

Creates BUGS code which can be ran through nma.run().

# Usage

```
nma.model.contrast(
  data_contrast = NULL,
  differences,
  se.diffs,
  var.arm1 = NULL,
  reference,
  type = "consistency",
  effects,
  scale,
  prior.mu = "DEFAULT",
  prior.d = "DEFAULT",
  prior.sigma = "DEFAULT")
```

# **Arguments**

data_contrast	A BUGSnetData object containing the data from contrast-based trials produced by data.prep()
differences	A string indicating the name of the differences for contrast-based studies
se.diffs	A string indicating the variable name of the standard errors of the differences.
var.arm1	A string (only required for networks with multi-arm trials) indicating the variable name of the variance of the treatment in arm 1 of each study
reference	A string for the treatment that will be seen as the 'referent' comparator and labeled as treatment 1 in the BUGS code. This is often a placebo or control drug of some kind.
type	If type="inconsistency", an inconsistency model will be built. By default, type="consistency" and a consistency model is built. will be built.
effects	A string indicating the type of treatment effect relative to baseline. Options are "fixed" or "random".
scale	A string indicating the scale of the data, such as "Mean Difference" or "Log-Odds Ratio".

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prior.mu	A string of BUGS code that defines priors on the baseline treatment effects. By default, independent normal priors are used with mean 0 and standard deviation 15u, where u is the largest maximum likelihood estimator in single trials (see van Valkenhoef, G et al. 2012).
prior.d	A string of BUGS code that defines define priors on relative treatment effects. By default, independent normal priors are used with mean 0 and standard deviation 15u, where u is the largest maximum likelihood estimator in single trials (see van Valkenhoef, G et al. 2012).

prior.sigma A string of BU

A string of BUGS code that defines the prior on the variance of relative treatment effects. By default, a uniform distribution with range 0 to u is used, where u is the largest maximum likelihood estimator in single trials (see van Valkenhoef, G et al. 2012).

### Value

nma.model returns an object of class BUGSnetModel which is a list containing the following components:

bugs - A long character string containing BUGS code that will be run in jags.

data - The data used in the BUGS code.

scale - The scale of the outcome, based on the chosen family and link function examples are "Risk Ratio" (relative risk), "Odds Ratio", "Mean Difference", "Hazard Ratio"

trt.key - Treatments mapped to integer numbers, used to run BUGS code.

...

#### References

van Valkenhoef, G, Lu, G, de Brock, B, Hillege, H, Ades, AE, Welton, NJ (2012). "Automating network meta-analysis." *Research Synthesis Methods*, **3**(4), 285-299.

Dias S, Sutton AJ, Welton NJ, Ades AE (2011). "NICE DSU technical support document 3. Heterogeneity: subgroups, meta-regression, bias and bias-adjustment." National Institute for Health and Clinical Excellence. Last updated April 2012.

# See Also

```
data.prep, nma.run
```

```
data(diabetes.sim)

diabetes.slr <- data.prep(
    arm.data = diabetes.sim,
    varname.t = "Treatment",
    varname.s = "Study"
)

#Random effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.re.c <- nma.model(
    data = diabetes.slr,
    outcome = "diabetes",
    N = "n",</pre>
```

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```
reference = "Placebo",
  family = "binomial",
  link = "cloglog",
  effects = "random"
  type = "consistency",
  time = "followup"
#Fixed effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.fe.c <- nma.model(</pre>
  data = diabetes.slr,
  outcome = "diabetes",
 N = "n"
  reference = "Placebo",
  family = "binomial",
  link = "cloglog",
  effects = "fixed",
  type = "consistency",
  time = "followup"
```

nma.model.shared

Create Bugs Model with Shared Parameters

# Description

Creates BUGS code which can be run through nma.run(). Handles a combination of arm-level and contrast-level data.

### Usage

```
nma.model.shared(
  data_arm = NULL,
  data_contrast = NULL,
  outcome,
  differences,
  N = NULL,
  sd.a = NULL,
  se.diffs = NULL,
  var.arm1 = NULL,
  reference,
  type = "consistency",
  time = NULL,
  family = NULL,
  link = NULL,
  effects,
  prior.mu = "DEFAULT",
  prior.d = "DEFAULT",
  prior.sigma = "DEFAULT"
```

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### **Arguments**

A BUGSnetData object containing the data from arm-based trials produced by data\_arm data.prep() A BUGSnetData object containing the data from contrast-based trials produced data\_contrast by data.prep() A string indicating the name of your outcome variable for arm-based studies. outcome differences A string indicating the name of the differences (outcome) for contrast-based studies Ν A string indicating the name of the variable containing the number of participants in each arm for arm-based data A string (only required for continuous outcomes with arm-level data) indicatsd.a ing variable name of the standard deviation of the outcome. Standard errors should be converted to standard deviation by multiplying by the square root of the sample size prior to using this function. se.diffs A string indicating the variable name of the standard errors of the differences in data\_contrast. A string (only required for contrast-based continuous data in networks with var.arm1 multi-arm trials) indicating the variable name of the variance of the treatment in arm 1 in each study reference A string for the treatment that will be seen as the 'referent' comparator and labeled as treatment 1 in the BUGS code. This is often a placebo or control drug of some kind. If type="inconsistency", an inconsistency model will be built. By default, type="consistency" type and a consistency model is built. will be built. A string (only required for binomial-cloglog or poisson-log models) indicating time the name of variable indicating person-time followup (e.g person years) or study followup time. A string indicating the family of the distribution of the outcome for arm-based family trials. Options are: "binomial", "normal", "poisson" link The link function for the nma model for arm-based models. Options are "logit" (binomial family), "log" (binomial family), "cloglog" (poisson family), "identity" (normal family). effects A string indicating the type of treatment effect relative to baseline. Options are "fixed" or "random". A string of BUGS code that defines priors on the baseline treatment effects. By prior.mu default, independent normal priors are used with mean 0 and standard deviation 15u, where u is the largest maximum likelihood estimator in single trials (see van Valkenhoef, G et al. 2012). prior.d A string of BUGS code that defines define priors on relative treatment effects. By default, independent normal priors are used with mean 0 and standard deviation 15u, where u is the largest maximum likelihood estimator in single trials (see van Valkenhoef, G et al. 2012). A string of BUGS code that defines the prior on the variance of relative treatment prior.sigma effects. By default, a uniform distribution with range 0 to u is used, where u is the largest maximum likelihood estimator in single trials (see van Valkenhoef,

G et al. 2012).

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#### Value

nma.model returns an object of class BUGSnetModel which is a list containing the following components:

bugs - A long character string containing BUGS code that will be run in jags.

data - The data used in the BUGS code.

scale - The scale of the outcome, based on the chosen family and link function examples are "Risk Ratio" (relative risk), "Odds Ratio", "Mean Difference", "Hazard Ratio"

trt.key - Treatments mapped to integer numbers, used to run BUGS code.

...

### References

van Valkenhoef, G, Lu, G, de Brock, B, Hillege, H, Ades, AE, Welton, NJ (2012). "Automating network meta-analysis." *Research Synthesis Methods*, **3**(4), 285-299.

Dias S, Sutton AJ, Welton NJ, Ades AE (2011). "NICE DSU technical support document 3. Heterogeneity: subgroups, meta-regression, bias and bias-adjustment." National Institute for Health and Clinical Excellence. Last updated April 2012.

### See Also

```
data.prep, nma.run
```

```
data(diabetes.sim)
diabetes.slr <- data.prep(arm.data = diabetes.sim,</pre>
varname.t = "Treatment",
varname.s = "Study")
#Random effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.re.c <- nma.model(</pre>
  data = diabetes.slr,
  outcome = "diabetes",
 N = "n"
  reference = "Placebo",
  family = "binomial",
  link = "cloglog",
  effects = "random"
  type = "consistency",
  time = "followup"
#Fixed effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.fe.c <- nma.model(</pre>
  data = diabetes.slr,
  outcome = "diabetes",
 N = "n"
  reference = "Placebo",
  family = "binomial",
  link = "cloglog",
```

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```
effects = "fixed",
  type = "consistency",
  time = "followup"
)
```

nma.rank

Table and Plots of Treatment Rankings

### **Description**

Produces a SUCRA (Surface Under the Cumulative Ranking Curve) plot and table. A SUCRA table summarizes the probabilities that each treatment is the best, second best...worst treatment in the network.

# Usage

```
nma.rank(
  nma,
  largerbetter,
  sucra.lwd = 1,
  sucra.palette = "Set1",
  ranko.palette = "Blues",
  cov.value = NULL
)
```

# **Arguments**

nma A BUGSnetRun object produced by running nma.run().

largerbetter A boolean variable indicating whether a larger probability should indicate a

more effective treatment (TRUE) or if a smaller probability should indicate a

more effective treatment (FALSE).

sucra.lwd Line width relative to the default (default=1) in the SUCRA plot.

sucra.palette A string indicating the colour set from RcolorBrewer for the SUCRA plot.

"Set1" is used by default and is automatically extended if there are many treat-

ments in your network.

ranko.palette A string indicating the colour set from RcolorBrewer for the rankogram. "Blues"

is used by default and is automatically extended if there are many treatments in

your network.

cov.value Must be specified for meta-regression. This is the value of the covariate for

which to report the results.

### Value

ranktable - A rank table showing the probability of each treatment being the nth best treatment.

sucratable - A table showing the probability of each treatment being the nth best treatment or better and an overall SUCRA value for each treatment.

order - A vector containing the order of efficacy of treatments (from best to worst) based on their SUCRA value. This vector is useful for ordering treatments when creating the league heat plot with nma.league().

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longtable - A long form table of ranking probabilities and SUCRA value.

sucraplot - A SUCRA plot showing the probability of each treatment being the nth best treatment or better.

rankogram - A rankogram showing the probability of each treatment being the nth best treatment.

#### See Also

```
nma.run, nma.league, nma.forest
```

# **Examples**

```
data(diabetes.sim)
diabetes.slr <- data.prep(</pre>
  arm.data = diabetes.sim,
  varname.t = "Treatment",
  varname.s = "Study"
#Random effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.re.c <- nma.model(</pre>
  data = diabetes.slr,
  outcome = "diabetes",
 N = "n"
  reference = "Placebo",
  family = "binomial",
  link = "cloglog",
  effects = "random",
  type = "consistency",
  time = "followup"
diabetes.re.c.res <- nma.run(</pre>
  model = diabetes.re.c,
  n.adapt = 100,
 n.burnin = 0,
 n.iter = 100
#get sucra results
sucra_results <- nma.rank(nma = diabetes.re.c.res, largerbetter = FALSE)</pre>
#plot sucra results
sucra_results$sucraplot
```

 ${\tt nma.regplot}$ 

Plot of relative treatment effects vs covariate values for metaregression.

# **Description**

Produces a plot of relative treatment effects on the linear scale vs covariate values for meta-regression.

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#### Usage

```
nma.regplot(nma, x.range = NULL, lwd = 1, palette = "Set1")
```

### Arguments

nma A BUGSnetRun object produced by running nma.run().

x.range The range of the x axis (covariate values). By default, the range will be the same

as in the data.

lwd Line width relative to the default (default=1).

palette A string indicating the colour set from RcolorBrewer for the plot. "set1" is

great, but you may need a different colour set if there are many treatments in

your network.

#### Value

regplot - A plot of the relative treatment effects vs covariate values for meta-regression.

### See Also

```
nma.run
```

```
data(afib)
afib.slr <- data.prep(arm.data = afib,</pre>
varname.t = "treatment",
varname.s = "study")
#Random effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
afib.re.c <- nma.model(</pre>
  data = afib.slr,
 outcome = "events"
 N = "sampleSize",
  reference = "02",
  family = "binomial",
  link = "logit",
  effects = "random",
  covariate = "stroke"
  prior.beta = "EXCHANGEABLE"
afib.re.c.res <- nma.run(
 model = afib.re.c,
  n.adapt = 100,
 n.burnin = 0,
  n.iter = 100)
nma.regplot(afib.re.c.res)
```

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nma.run Run NMA model

# Description

Takes bugs code from an object produced by nma.model and runs model using jags.

# Usage

```
nma.run(
  model,
  monitor = "DEFAULT",
  DIC = TRUE,
  n.adapt = 1000,
  n.burnin = floor(n.iter/2),
  n.iter,
  thin = 1,
  n.chains = 3,
  inits = "DEFAULT"
)
```

# **Arguments**

model	A BUGSnetModel object produced by running nma.model.
monitor	A vector of all variables that you would like to monitor. Default is "DEFAULT" which will monitor the relative treatment effects d as well as sigma when a random effects model is used and the regression coefficients beta when meta-regression is used.
DIC	Default is TRUE and nodes required to calculate the DIC and other fit statistics are monitored. Otherwise you may set it to FALSE.
n.adapt	Number of adaptations for the mcmc chains.
n.burnin	Number of burnin iterations for the mcmc chains.
n.iter	Number of iterations for the mcmc chains.
thin	Thinning factor for the mcmc chains. Default is 1.
n.chains	Number of mcmc chains. Default is 3.
inits	Specifies initial values and random number generator options for each chain. The "DEFAULT" option uses the R random seed to set the JAGS random seeds. Non-default options are passed directly to jags.model. In order to use the JAGS default initialization, set inits to NULL. See jags.model for more info.

# Value

```
nma.run returns an object of class BUGSnetRun which is a list containing the following components: samples - The MCMC samples produced by running the BUGS model.

model - The BUGSnetModel object obtained from nma.model which was used to run jags.

scale - The scale of the outcome, based on the chosen family and link function.

trt.key - Treatments mapped to numbers, used to run BUGS code.

family - Family that was used for the NMA model (e.g normal, binomial, poisson)

link - Link function that was used for the NMA model (e.g normal, binomial, poisson)
```

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### See Also

nma.model, nma.fit, nma.league, nma.rank, nma.forest, nma.regplot, nma.trace, jags.model

### **Examples**

```
data(diabetes.sim)
diabetes.slr <- data.prep(</pre>
  arm.data = diabetes.sim,
  varname.t = "Treatment",
  varname.s = "Study"
#Random effects, consistency model.
#Binomial family, cloglog link. This implies that the scale will be the Hazard Ratio.
diabetes.re.c <- nma.model(</pre>
  data = diabetes.slr,
 outcome = "diabetes",
 N = "n"
 reference = "Placebo",
  family = "binomial",
 link = "cloglog",
 effects = "random",
  type = "consistency",
  time = "followup"
)
diabetes.re.c.res <- nma.run(</pre>
 model = diabetes.re.c,
 n.adapt = 100,
 n.burnin = 0,
 n.iter = 100
```

nma.trace

Deprecated. Please use nma.diag instead.

### **Description**

Deprecated. Produces traceplots of the MCMC chains obtained from nma.run()

# Usage

```
nma.trace(nma, n = "all", thin = 1, colours = "DEFAULT")
```

# Arguments

nma	A BUGSnetRun object produced by running nma.run().
n	Integer which limits the number of printed variables to the first n. Default is "all" which plots every variable.
thin	Thinning factor for the mcmc chains. Default is 1.
colours	An optional vector of colors, one for each chain.

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### See Also

nma.diag

parkinsons

Parkinsons contrast data

### **Description**

This dataset was used to demonstrate network meta-analysis in the NICE-DSU TSD 2 (example 7). It includes information on seven clinical trials examining the effects of five different treatments for Parkinsons on the mean off-time reduction (continuous outcome). This dataset is in the form of mean differences in order to demonstrate the contrast data.

### Usage

data(parkinsons)

#### **Format**

A dataset.

### References

Dias S, Welton NJ, Sutton AJ, Ades AE (2011). "NICE DSU technical support document 2: a generalised linear modelling framework for pairwise and network meta-analysis of randomised controlled trials." National Institute for Health and Clinical Excellence.

# Examples

data(parkinsons)

parkinsons\_arm

Parkinsons arm-based data

# Description

This dataset was used to demonstrate network meta-analysis in the NICE-DSU TSD 2 (examples 5 and 8). It includes arm-based information on three out of a total of seven clinical trials examining the effects of five different treatments for Parkinsons on the mean off-time reduction (continuous outcome). This dataset can be used in combination with rows 9 to 15 of the parkinsons dataset to demonstrate the use of shared parameter models in BUGSnet.

# Usage

data(parkinsons\_arm)

# Format

A dataset.

pma pma

### References

Dias S, Welton NJ, Sutton AJ, Ades AE (2011). "NICE DSU technical support document 2: a generalised linear modelling framework for pairwise and network meta-analysis of randomised controlled trials." National Institute for Health and Clinical Excellence.

# **Examples**

```
data(parkinsons_arm)
```

pma

Pairwise meta-analysis

# Description

implements pairwise meta-analysis via the package meta

# Usage

```
pma(
   data,
   name.trt1,
   name.trt2,
   outcome,
   N,
   sd = NULL,
   time = NULL,
   type.outcome,
   method = "MH",
   method.tau = "DL",
   sm
)
```

# **Arguments**

data	A BUGSnetData object produced by data.prep()
name.trt1	A string indicating the name of the comparator treatment (often Placebo)
name.trt2	A string indicating the name of the experimental treatment
outcome	A string indicating the name of your outcome variable
N	A string indicating the name of the variable containing the number of participants in each arm
sd	A string (only required when type.outcome="continuous") indicating variable name of the standard deviation of the outcome
time	A string required when type.outcome = "rate". Name of variable indicating person-time followup (e.g person years).
type.outcome	A string. Options are: "binomial", "continuous", "rate" (e.g $\#$ of events and $\#$ person-years reported)
method	A character string indicating what type of test was used. For more info, see $\!$ meta's documentation.

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Method.tau A character string indicating which method is used to estimate the between study variance. Either "DL", "PM", "REML", "ML", "HS", "SJ", "HE", or "EB", can be abbreviated.

Sm A character string indicating which summary measure ("RR", "OR", "RD", or ASD") is to be used for pooling of studies.

#### Value

A forest plot as produced by the package meta

raw - dataset containing summary statistics of meta-analysis (effect estimates, confidence bounds, I-squared, Q-statistic)

#### Note

Depending on the the value of type.outcome, this function will implement the functions metabin (dichotomous outcomes), metacont (continuous outcomes), or metainc (rate outcomes) from the package meta.

#### See Also

```
data.prep
```

### **Examples**

```
data(diabetes.sim)

diabetes.slr <- data.prep(
    arm.data = diabetes.sim,
    varname.t = "Treatment",
    varname.s = "Study"
)

pma(
    data = diabetes.slr,
    type.outcome = "binomial",
    sm = "OR",
    name.trt1 = "Placebo",
    name.trt2 = "Diuretic",
    outcome = "diabetes",
    N = "n"
)</pre>
```

thrombolytic

Thrombolytic data

# Description

The thrombolytic drugs dataset compiled by Boland et al. (2003) has been used to demonstrate indirect-treatment comparison methods within previous publications and other R packages such as 'gemtc'. The dataset consists of information on a binary outcome (mortality within 30 or 35 days) and eight treatments (eight different thrombolytic drugs used after a patient has had an acute myocardial infarction) extracted from 28 clinical trials (including two, three-armed trials).

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#### Usage

data(thrombolytic)

### **Format**

A dataset.

#### References

Boland A, Dundar Y, Bagust A, Haycox A, Hill R, Mota RM, Walley T, A D (2003). "Early thrombolysis for the treatment of acute myocardial infarction: a systematic review and economic evaluation." *Health Technology Assessment*, 7(15), 1–136.

Dias S, Welton NJ, Caldwell DM, AE A (2010). "Checking consistency in mixed treatment comparison meta-analysis." *Statistics in medicine*, **7-8**(29), 932–44.

Lu G, Ades AE (2006). "Assessing evidence inconsistency in mixed treatment comparisons." *Journal of the American Statistical Association*, **101**(474), 447–459.

van Valkenhoef, G, Lu, G, de Brock, B, Hillege, H, Ades, AE, Welton, NJ (2012). "Automating network meta-analysis." *Research Synthesis Methods*, **3**(4), 285-299.

# **Examples**

data(thrombolytic)

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