# Package 'NetworkMetaAnalysis'

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 $\textbf{Title} \ \ \text{Network meta analysis of target-comparator-outcomes in OHDSI network studies}$ 

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### **Description**

A package for using a collection of target-comparator-outcome estimates from OHDSI network studies to estimate relative effectiveness between all target-comparator pairs, even in the absence of head-to-head comparison.

```
Depends R (>= 3.5.0)
Imports coda (>= 0.19.3),
      DatabaseConnector (\geq 2.4.4),
      dplyr (>= 0.8.5),
      foreach (>= 1.5.0),
      gemtc (>= 0.8.4),
      ggraph (>= 2.0.2),
      ggplot2 (>= 3.3.0),
      grid (>= 3.6.3),
      igraph (>= 1.2.5),
      magrittr (>= 1.5),
      plyr (>= 1.8.6),
      rlang (>= 0.4.6),
      scales (>= 1.1.1),
      SqlRender (>= 1.6.5),
      stringr (>= 1.4),
      tibble (>= 3.0.1),
      tidyr (>= 1.0.2),
      tidyselect (>= 1.0.0)
Suggests doSNOW (>= 1.0.18),
      DT (>= 0.13),
      shiny (>= 1.4.0.2),
      shinydashboard (>= 0.7.1),
      shinythemes (>= 1.1.2),
      snow (>= 0.4.3),
      testthat
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 ${\tt computePairwiseEstimates}$ 

Compute all pairwise estimates

# Description

After fitting the model, this function pulls out all pairwise network effect (direct and indirect combined) in a tidy format, on the hazard rate scale.

# Usage

computePairwiseEstimates(fit)

# Arguments

fit

the output of fitNetwork

computeRankProbabilities

Compute the rank probabilities

# Description

To be used for, i.a., rankograms (that use the cumulative probability).

# Usage

```
computeRankProbabilities(fit, preferredDirection = 1)
```

# Arguments

```
fit object returned from GeMTC.
```

preferredDirection

scalar, preferential direction of the outcome. 1 means higher values are preferred, -1 means lower values are preferred.

### Value

Tidy data frame.

computeRhat

Compute the Rhat statistic for a given fit

# **Description**

Returns a tidy tibble with the Rhat statistic for each parameter. Also handles node-splitting models.

# Usage

```
computeRhat(fit)
```

# Arguments

fit an object of class "mtc.result" or "mtc.nodesplit".

### Value

Tidy tibble. If a node-splitting result is supplied, the first columns called modelName indicates the comparison that was split in the analysis.

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computeSucra

Compute SUCRA estimates

# Description

Computes surface under the cumulative ranking curve (SUCRA) for each treatment in the network. The ranking curve and rankogram are synonyms. Not sure this deserves its own function. Let's see down the road.

# Usage

```
computeSucra(rankProbabilities)
```

## **Arguments**

rankProbabilities

 $output\ of\ compute {\tt RankProbabilities}$ 

deriveEdges

Use to prepare data needed to draw the edges of the network

# Description

Use to prepare data needed to draw the edges of the network

## Usage

```
deriveEdges(networkData, nodes)
```

### **Arguments**

 ${\tt networkData} \qquad {\tt subset\ of\ the\ original\ network\ study\ data}.$ 

nodes output from deriveNodes

### Value

A ggplot object.

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deriveNodes

Deriving node data for drawing the network

### **Description**

Deriving node data for drawing the network

### Usage

```
deriveNodes(networkData, labelOrientation)
```

### **Arguments**

networkData

tidy data frame, subset of the full data frame with aggregate results at study arm level for specific network meta-analysis.

labelOrientation

string giving how to rotate the label. There are two allowed settings: "radial" (default) and "horisontal"/"horizontal".

### Value

A tidy data frame.

fetchParameterSummaries

Table of parameter estimates and deviance statistics for reconciled results

# Description

Table of parameter estimates and deviance statistics for reconciled results

# Usage

```
fetchParameterSummaries(
  parameterSummaries,
  devianceStatistics,
  networkMetaAnalysisId = NULL,
  nDigits = 1
)
```

### **Arguments**

```
parameterSummaries
```

output of deriveParameterQuantiles.

devianceStatistics

output of fetchDevianceStatistics.

networkMetaAnalysisId

scalar, what analysis to use. If NULL (default), the function assumes pre-filtered data to be supplied in the first two arguments.

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nDigits scalar, how many digits to show for parameter estimates. Won't affect deviance statistics.

fetchPosteriorDraws Fetches and tidies the posterior draws

### **Description**

Potentially, with some thinning to keep the amount of space required down.

### Usage

```
fetchPosteriorDraws(fit, drawsThin = 10)
```

## **Arguments**

fit the result of fitNetwork.

drawsThin scalar, thinning factor. The default value (10) keeps every tenth observation.

This is mainly used when producing output to be saved to the server, to keep the

storage requirements at a reasonable level.

### Value

Tidy tibble with columns for chain and MCMC iteration indicators and one column per parameter.

fetchTcoEstimates

Load target-comparator-outcome estimates from server

### **Description**

Load TCO estimates from the server and wrangle them into a format appropriate for network visualisation and analysis.

## Usage

```
fetchTcoEstimates(
  conn,
  resultsCdm,
  resultsTable,
  excludedDatabases = "Meta-analysis"
)
```

### **Arguments**

conn a connection to a database, e.g., the result of calling DatabaseConnector::connect.

resultsCdm string, the name of the schema in which the table lives. resultsTable string, the name of the table holding the estimates.

excludedDatabases

character vector, names of the databases whose results should be ignored. The default ("Meta-analysis") is obvious and should be kept if other databases are added.

fetchTidyNodesplitResults

Get the node-splitting analyses results in tidy format

# Description

Returns a tidy data frame with direct, indirect and consistency (network) effect estimates for each target-comparator pair along with the probability of inconsistency between the direct and indirect estimates.

# Usage

```
fetchTidyNodesplitResults(fit)
```

### **Arguments**

fit

the result of calling fitNetwork with includeNodesplittingAnalysis = TRUE.

#### Value

Tidy data frame.

fitNetwork

Run model on network to estimate parameters

### **Description**

Runs a GeMTC-powered network meta-analysis on a subset of the results of calling fetchTcoEstimates (i.e., setting analysisId and outcomeId) and

# Usage

```
fitNetwork(
  networkData,
  includeNodesplittingAnalysis = FALSE,
  modelType = "random",
  nWarmup = 1000,
  nIter = 3000,
  fitThin = 1,
  nChains = 4
)
```

## **Arguments**

networkData

a dataframe holding the subset of a target-comparator-outcome estimates from the result of fetchTcoEstimates, with specified outcomeId and analysisId.

include Node splitting Analysis

boolean, should also node-splitting analyses be run? NB: This can be quite time-consuming, so is by default set to FALSE.

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modelType	string defining whether to run a random-effects ("random", default) or a fixed-effects ("fixed") mode.
nWarmup	scalar, the number of iterations the sampler should do before it starts sampling from the posterior. This should be sufficient to the chains to have converged.
nIter	scalar, the number of posterior samples to draw per chain.
fitThin	scalar, thinning factor. 1 (default) keeps all samples whereas e.g. 10 would keep only every tenth sample.
nChains	scalar, the number of JAGS chains to run

launchShinyApp

Launch a Shiny app to explore the results

# Description

Launch a Shiny app to explore the results

### Usage

```
launchShinyApp(results, rstudio = TRUE, ...)
```

# **Arguments**

results	the output of reconcileResults directly or loadFromDb if the results live on a server.
rstudio	boolean, should the app run in the Viewer window (TRUE, default) or in an internet browser (FALSE)?
	passed on to shiny::runApp

loadFromDb

Loads results stored in a database following its data model

# Description

Loads results stored in a database following its data model

# Usage

```
loadFromDb(conn, schema)
```

# Arguments

conn connection to database, e.g. result of calling DatabaseConnector::connect

schema string, the name of the schema where results tables live.

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plotForest

Draw forest plot of total network estimates

### **Description**

Using a fitNetwork object, this functions draws a classic forest plot of the combined (direct + indirect) effects estimated.

### Usage

```
plotForest(
  relativeEffectsEstimates,
  reference = NULL,
  lineSize = 0.5,
  pointSize = 1
)
```

### **Arguments**

```
relativeEffectsEstimates
the output of computeRelativeEffects.

reference string, which exposure to use as the reference. Choice is arbitrary. If nothing is supplied, the first value in the data frame is used.

lineSize, pointSize scalars
```

plotNetwork

Visualise network

# Description

Draw the head-to-head comparisons as a network. The function relies on **ggraph** to do the heavy lifting.

### Usage

```
plotNetwork(
  edges,
  nodes,
  maxNodeRadius = 0.1,
  nodeColour = "dodgerblue",
  labelOrientation = "radial",
  labelColour = "black",
  edgeColour = "dodgerblue",
  edgeAlpha = 0.2,
  treatmentMetaData = NULL,
  edgeMetaData = NULL
)
```

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### **Arguments**

edges, nodes data frames with data on edges and nodes in the study graph.

maxNodeRadius the radius of the largest node in the network on the original coordinate scales (=

[-1, [1). Default = 0.1.

labelOrientation

string giving how to rotate the label. There are two allowed settings: "radial" (default) and "horisontal"/"horizontal".

edgeColour, nodeColour, labelColour

strings given the colours of edges, nodes and label text.

edgeAlpha scalar in [0, 1] giving the opacity of the edges (default: 0.2). 1 = complete

opaque, 0 =completely transparent.

treatmentMetaData

 $NOT\ IN\ USE\ a$  data frame with information about each node in the network,

e.g., name and databases with data for this treatment.

edgeMetaData NOT IN USE a dataframe with information about the head-to-head compar-

isons. Note that the number of head-to-head comparisons is computed so doesn't

need to be supplied.

plotNodesplitResults Visualise results of node-splitting analysis

# Description

Pending, but could be useful for getting an overview.

### Usage

plotNodesplitResults(nodesplitResults)

### **Arguments**

 ${\tt nodesplitResults}$ 

tidy data frame with consistency, direct and indirect HR estimates with CrIs.

### **Description**

Some customisation possible, but this is still pretty basic.

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#### Usage

```
plotPairwiseEstimates(
  tidyEstimates,
  nDigits = 1,
  colour_low = "blue",
  colour_high = "red",
  contrastThreshold = 0.5,
  mid = "hr",
  lo = "cri95Lb"
  hi = "cri95Ub",
  textSize = 12,
  sucraEstimates = NULL
)
```

### **Arguments**

tidyEstimates data frame/tibble with all pairwise combinations to feature in the heatmap. scalar, how many digits should the shown estimates have? Default is 1 nDigits

colour\_low, colour\_high

valid colour specifications for the lower and higher bounds of the diverging colour scale

contrastThreshold

scalar, absolute values on the log-scale above this threshold will be white, otherwise they will be black

mid, lo, hi

strings, the names of the columns in the tidyEstimates data frame holding the point estimate (usually median) and lower and upper bounds of the credibility intervals (usually 2.5 and 97.5 percentiles)

textSize

scalar.

sucraEstimates data frame with two columns: exposureId and SUCRA estimate as numeric value. Output from computeSucra. Default: NULL. If valid data supplied, the SUCRA estimates will be shown in the diagonal, and the exposures will be ordered by descending SUCRA estimate.

plotPosteriorDensities

Produces density plots of parameter estimates from the fitted models

### **Description**

Produces a ggplot2 object so can be further customised as desired. If the results of a node-splitting analysis is supplied, the output is a list of plots, one for each analysis and one for the consistency fit.

### Usage

```
plotPosteriorDensities(
  posteriorDraws,
  separateChains = TRUE,
  lineAlpha = 0.7,
```

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```
lineSize = 0.2,
densityResolution = 256,
wrapByModelName = FALSE
)
```

### **Arguments**

posteriorDraws tidy data frame, all posterior draws with (at least) the following columns: modelName,

chainId, mcmcIterationId, parameterName, parameterValue.

separateChains boolean, should chains be plotted as separate distributions (TRUE, default) or as

one (FALSE)? Not plotting chains separately somewhat defeats the purpose of

the visualisation but is anyway left to the user.

lineAlpha the opacity of drawn lines. If alpha = 1, lines will be entirely opaque masking

overlain lines, so the default is 0.7.

lineSize scalar, thickness of the plot lines.

densityResolution

scalar, the number of grid points over which to estimate the densities. Higher

values yield smoother curves but produces a larger object.

wrapByModelName

boolean, if the modelName column contains more than one distinct value, this should be set to TRUE to show separate plots for each model. Otherwise, the plot will combine samples for the same parameter from different models, which will

give misleading results.

### Value

A ggplot object, or a list of ggplot objects.

plotRankogram

Rank treatments by effectiveness

# Description

Produces a **ggplot2** object so can be further customised as desired. If the results of a node-splitting analysis is supplied, the output is a list of plots, one for each analysis and one for the consistency fit.

#### Usage

```
plotRankogram(rankProbabilities, lineSize = 1, lineAlpha = 0.9)
```

### **Arguments**

rankProbabilities

output of computeRankProbabilities

lineSize scalar, thickness of the plot lines.

lineAlpha the opacity of drawn lines. If alpha = 1, lines will be entirely opaque masking

overlain lines, so the default is 0.7.

### Value

A ggplot object, or a list of ggplot objects.

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plotRhat

Plot the Rhat values of the fitted model

### **Description**

Instead of looking at the Rhat values in table format, it's usually easier to quickly scan a plot, and this function allows you to do just that.

# Usage

```
plotRhat(
   rhat,
   threshold = 1.05,
   showParameters = FALSE,
   pointSize = 1,
   lineSize = 0.5
)
```

### Arguments

rhat output from computeRhat

threshold scalar, what is the threshold below which you want all Rhat values to lie? De-

fault is 1.05. Set to NULL to remove the indicator in the plot.

showParameters boolean, should parameters be distinguishable with colours? Potentially useful

if any are above the threshold, so by default set to FALSE.

pointSize scalar, size of points representing Rhat values.

lineSize scalar, thickness of vertical indicator of threshold, if shown.

### Value

A ggplot object.

plotTraces

Produces density plots of parameter estimates from the fitted models

### **Description**

Produces a **ggplot2** object so can be further customised as desired. If the results of a node-splitting analysis is supplied, the output is a list of plots, one for each analysis and one for the consistency fit.

### Usage

```
plotTraces(
  posteriorDraws,
  separateChains = TRUE,
  lineAlpha = 0.7,
  lineSize = 0.2,
  wrapByModelName = FALSE
)
```

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### **Arguments**

posteriorDraws tidy data frame, all posterior draws with (at least) the following columns: modelName,

 $chain Id, \verb|mcmcIterationId|, \verb|parameterName|, \verb|parameterValue|.$ 

separateChains boolean, should chains be plotted as separate distributions (TRUE, default) or as

one (FALSE)? Not plotting chains separately somewhat defeats the purpose of

the visualisation but is anyway left to the user.

lineAlpha the opacity of drawn lines. If alpha = 1, lines will be entirely opaque masking

overlain lines, so the default is 0.7.

lineSize scalar, thickness of the plot lines.

wrapByModelName

boolean, if the modelName column contains more than one distinct value, this should be set to TRUE to show separate plots for each model. Otherwise, the plot will combine samples for the same parameter from different models, which will give misleading results.

#### Value

A ggplot object, or a list of ggplot objects.

runAnalyses

Runs the analysis and returns results ready to be stored in database or used directly

### **Description**

Runs the analysis and returns results ready to be stored in database or used directly

# Usage

```
runAnalyses(
   aggregatedResults,
   includeNodesplittingAnalysis = FALSE,
   nCores = 4,
   nWarmup = 1000,
   nIter = 3000,
   nChains = 4,
   fitThin = 1,
   referenceTreatmentId = NULL,
   preferredDirection = 1,
   modelType = "random",
   drawsThin = 10,
   nmaIds = NULL
)
```

## **Arguments**

```
aggregatedResults
```

tidy data frame, result of calling fetchTcoEstimates.

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include Node splitting Analysis

boolean, should also node-splitting analyses be run? NB: This can be quite

time-consuming, so is by default set to FALSE.

nCores scalar, how many cores to use when running the analyses. If > 1, **foreach** is used

to parallel processing.

nWarmup scalar, the number of iterations the sampler should do before it starts sampling

from the posterior. This should be sufficient to the chains to have converged.

nIter scalar, the number of posterior samples to draw per chain.

nChains scalar, the number of JAGS chains to run

fitThin scalar, thinning factor. 1 (default) keeps all samples whereas e.g. 10 would keep

only every tenth sample.

referenceTreatmentId

string, the name of the exposure to use as the reference when computing pairwise

estimates. If none given, the first in alphabetical order will be used.

preferredDirection

scalar, preferential direction of the outcome. 1 means higher values are pre-

ferred, -1 means lower values are preferred.

modelType string defining whether to run a random-effects ("random", default) or a fixed-

effects ("fixed") mode.

drawsThin, scalar, thinning factor used when fetching posterior draws to keep in the database.

Used to produce further derived quantities, density plots of posterior distribu-

tions and trace plots.

nmaIds scalar vector, if only specific network meta analyses should be run, their id's are

supplied here. If none supplied, all will be run.

saveToDatabase

Saves the reconciled results to database

### **Description**

If desired, this function takes the reconciled results and saves them to the database for downstream use.

### Usage

saveToDatabase(results, conn, schema, overwriteExistingTables = FALSE)

### **Arguments**

results output of runAnalyses.

conn connection object.

schema string, the name of the schema where the tables will live. Should probably be a

dedicated schema to prevent conflicting names.

overwriteExistingTables

boolean, should tables be overridden? This is a bit dangerous and so set to FALSE

by default. See note on dedicated schema above.

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