## Package 'gFormulaMI'

May 13, 2025

Title G-Formula for Causal Inference via Multiple Imputation

Version 1.0.1

Description Implements the G-Formula method for causal inference with time-varying treatments and confounders using Bayesian multiple imputation methods, as described by Bartlett et al (2025) <doi:10.1177/09622802251316971>. It creates multiple synthetic imputed datasets under treatment regimes of interest using the 'mice' package. These can then be analysed using rules developed for analysing multiple synthetic datasets.

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## Description

gFormulaImpute creates multiple imputed synthetic datasets of longitudinal histories under specified treatment regimes of interest, based on the G-formula.

## Usage

```
gFormulaImpute(
  data,
 M = 50,
  trtVars,
  trtRegimes,
 nSim = NULL,
 micePrintFlag = FALSE,
 silent = FALSE,
 method = NULL,
 predictorMatrix = NULL,
 missingDataCheck = TRUE
)
```

#### **Arguments**

data	The observed data frame
М	The number of imputed datasets to generate
trtVars	A vector of variable names indicating the time-varying treatment variables
trtRegimes	A vector specifying the treatment regime of interest, or a list of vectors specifying the treatment regimes of interest
nSim	The number of individuals to simulate in each imputed dataset. Defaults to number of individuals in observed data
micePrintFlag	TRUE/FALSE specifying whether the output from the call(s) to mice should be

printed

silent TRUE/FALSE indicating whether to print output to console (FALSE) or not (TRUE)

An optional method argument to pass to mice. If not specified, the default is to impute continuous variables using normal linear regression (norm), binary variables using logistic regression (logreg), polytomous regression for unordered

factors and proportional odds model for ordered factors

predictorMatrix

method

An optional predictor matrix to specify which variables to use as predictors in the imputation models. The default is to impute sequentially, i.e. impute using all variables to the left of the variable being imputed as covariates

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missingDataCheck

TRUE/FALSE indicating whether gFormulaMI checks, when passed a regular data frame, whether there any missing values.

#### **Details**

gFormulaImpute creates multiple imputed synthetic datasets of longitudinal histories under specified treatment regimes of interest, based on the G-formula, as described by Bartlett et al (2025) doi:10.1177/09622802251316971. Specifically, to the observed data frame, an additional nSim rows are added in which all variables are set to missing, except the time-varying treatment variables. The latter are set to the values as specified in the trtRegimes argument. If multiple treatment regimes are specified, nSim rows are added for each of the specified treatment regimes.

gFormulaImpute uses the mice package to impute the potential outcome values of the time-varying confounders and outcome in the synthetic datasets. Imputation is performed sequentially from left to right in the data frame. As such, the variables must be ordered in time in the input data frame, with the time-varying confounders at each time followed by the corresponding treatment variable at that time.

For the data argument, gFormulaImpute expects either a fully observed (complete) data frame, or else a set of multiple imputation stored in an object of class mids (from the mice package).

Unlike with Rubin's regular multiple imputation pooling rules, it is possible for the pooling rules developed by Raghunathan et al (2003) to give negative variance estimates. The probability of this occurring is reduced by increasing M and/or nSim.

gFormulaImpute returns an object of class mids. This can be analysed using the same methods that imputed datasets from mice can be analysed with (see examples). However, Rubin's standard pooling rules are not valid for analysis of the synthetic datasets. Instead, the synthetic variance estimator of Raghunathan et al (2003) must be used, as implemented in the syntheticPool function.

The development of the gFormulaMI package was supported by a grant from the UK Medical Research Council (MR/T023953/1).

## Value

an S3 object of class mids (multiply imputed dataset)

#### Author(s)

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#### References

Bartlett JW, Olarte Parra C, Granger E, Keogh RH., van Zwet EW and Daniel RM, 2025. G-formula with multiple imputation for causal inference with incomplete data. Statistical Methods in Medical Research.

Raghunathan TE, Reiter JP, Rubin DB. 2003. Multiple imputation for statistical disclosure limitation. Journal of Official Statistics, 19(1), p.1-16.

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#### **Examples**

simDataFullyObs

Simulated fully observed data frame

## **Description**

A simulated observational study data frame with no missing data.

## Usage

```
simDataFullyObs
```

#### **Format**

simDataFullyObs:

A data frame with 5000 rows and the following variables:

- 10 Continuous baseline confounder
- **a0** Binary baseline treatment
- 11 Continuous confounder at time 1
- **a1** Binary treatment at time 1
- 12 Continuous confounder at time 2
- **a2** Binary treatment at time 2
- y Continuous final outcome ...

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syntheticPool	Pool estimates and variances obtained by analysing multiple synthetic datasets
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## **Description**

This function pools estimates and variances which have been obtained by analysing multiple synthetic imputations (e.g. created used gFormulaImpute) using the method developed by Raghunathan et al 2003.

## Usage

```
syntheticPool(fits)
```

## Arguments

fits

Collection of model fits produced by a call of the form with(imps, lm(y~regime)) where imps is a collection of imputed datasets of class mids.

#### **Details**

The only argument to syntheticPool is a set of model fits obtained by running an analysis on an imputed dataset collection of class mids, as created for example using the mice function in the mice package.

The function returns a table containing the overall parameter estimates, the within, between and total imputation variances, 95% confidence intervals, and p-values testing the null hypothesis that the corresponding parameters equal zero.

It is possible for the variance estimator developed by Raghunathan et al 2003 to be negative. In this case syntheticPool stops and informs you to re-impute using a larger number of imputations M and/or nSim.

The development of the gFormulaMI package was supported by a grant from the UK Medical Research Council (MR/T023953/1).

#### Value

A matrix containing the pooled results.

#### Author(s)

Jonathan Bartlett < jonathan.bartlett1@lshtm.ac.uk>

#### References

Raghunathan TE, Reiter JP, Rubin DB. 2003. Multiple imputation for statistical disclosure limitation. Journal of Official Statistics, 19(1), p.1-16.

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