Package 'calibmsm'

May 13, 2024

Title Calibration Plots for the Transition Probabilities from Multistate Models

Version 1.1.0

Description Assess the calibration of an existing (i.e. previously developed) multistate model through calibration plots.

Calibration is assessed using one of three methods. 1) Calibration methods for binary logistic regression models applied at a fixed time point in conjunction with inverse probability of censoring weights. 2) Calibration methods for multinomial logistic regression models applied at a fixed time point in conjunction with inverse probability of censoring weights. 3) Pseudo-values estimated using the Aalen-Johansen estimator of observed risk. All methods are applied in conjunction with landmarking when required. These calibration plots evaluate the calibration (in a validation cohort of interest) of the transition probabilities estimated from an existing multistate model. While package development has focused on multistate models, calibration plots can be produced for any model which utilises information post baseline to update predictions (e.g. dynamic models); competing risks models; or standard single outcome survival models, where predictions can be made at any landmark time. Please see Pate et al. (2024) <doi:10.1002/sim.10094> and Pate et al. (2024) <h >https:

//alexpate30.github.io/calibmsm/articles/Overview.html>.

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Encoding UTF-8 **RoxygenNote** 7.3.1

Imports boot, dplyr, ggplot2, ggpubr, ggExtra, gridExtra, Hmisc, mstate, rms, stats, survival, tidyr, VGAM

Depends R (>= 2.10)

LazyData true

Suggests covr, knitr, rmarkdown, R.rsp, testthat (>= 3.0.0)

VignetteBuilder knitr, R.rsp

URL https://alexpate30.github.io/calibmsm/

Config/testthat/edition 3 **NeedsCompilation** no

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```
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Repository CRAN

Date/Publication 2024-05-13 11:33:07 UTC
```

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calc_weights

Calculate inverse probability of censoring weights at time ${\sf t.}$

Description

Estimates the inverse probability of censoring weights by fitting a cox-proportial hazards model in a landmark cohort of individuals. Primarily used internally, this function has been exported to allow users to reproduce results in the vignette when estimating confidence intervals using bootstrapping manually.

Usage

```
calc_weights(
  data.mstate,
  data.raw,
  covs = NULL,
  t,
  s,
  landmark.type = "state",
  j = NULL,
  max.weight = 10,
  stabilised = FALSE,
  max.follow = NULL
)
```

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Arguments

data.mstate	Validation data in msdata format
data.raw	Validation data in data.frame (one row per individual)
covs	Character vector of variable names to adjust for when calculating inverse probability of censoring weights
t	Follow up time at which to calculate weights
S	Landmark time at which predictions were made
landmark.type	Whether weights are estimated in all individuals uncensored at time s ('all') or only in individuals uncensored and in state j at time s ('state')
j	Landmark state at which predictions were made (only required in landmark.type = 'state')
max.weight	Maximum bound for weights
stabilised	Indicates whether weights should be stabilised or not
max.follow	Maximum follow up for model calculating inverse probability of censoring weights. Reducing this to $t+1$ may aid in the proportional hazards assumption being met in this model.

Details

Estimates inverse probability of censoring weights (Hernan M, Robins J, 2020). Fits a cox proportional hazards model to individuals in a landmark cohort, predicting the probability of being censored at time t. This landmark cohort may either be all individuals uncensored at time s, or those uncensored and in state j at time s. All predictors in w.covs are assumed to have a linear effect on the hazard. Weights are estimated for all individuals in data.raw, even if they will not be used in the analysis as they do not meet the landmarking requirements. If an individual enters an absorbing state prior to t, we estimate the probability of being censored before the time of entry into the absorbing state, rather than at t. Details on all the above this are provided in vignette *overview*.

Value

A dataframe with three columns. id corresponds to the patient ids from data.raw. ipcw contains the inverse probability of censoring weights (specifically the inverse of the probability of being uncesored). pcw = 1/ipcw. If stabilised = TRUE was specified, a fourth variable ipcw.stab will be returned, which is the stabilised inverse probability of censoring weights.

References

Hernan M, Robins J (2020). "12.2 Estimating IP weights via modeling." In *Causal Inference: What If*, chapter 12.2. Chapman Hall/CRC, Boca Raton.

Examples

- # Estimate inverse probability of censoring weights for individual in cohort ebmtcal.
- # Specifically the probability of being uncensored at t = 1826 days.
- # Weights are estimated using a model fitted in all individuals uncensored at time s = 0. weights.manual <-

```
calc_weights(data.mstate = msebmtcal,
  data.raw = ebmtcal,
  covs = c("year", "agecl", "proph", "match"),
  t = 1826,
  s = 0,
  landmark.type = "state",
  j = 1)

str(weights.manual)
```

calib_msm

Calibration plots for a multistate model'

Description

Calculates the underlying data for calibration plots of the predicted transition probabilities from a multistate model using three methods.

- 1. BLR-IPCW: Binary logistic regression with inverse probability of censoring weights.
- 2. MLR-IPCW: Multinomial logistic regression with inverse probability of censoring weights, based on the nominal calibration framework of van Hoorde et al. (2014, 2015)
- 3. Pseudo-values: Pseudo-values estimated using the Aalen-Johansen estimator (Aalen OO, Johansen S, 1978).

Usage

```
calib_msm(
  data.mstate,
  data.raw,
  j,
  s,
  t,
  tp.pred,
  tp.pred.plot = NULL,
  calib.type = "blr",
  curve.type = "rcs",
  rcs.nk = 3,
  loess.span = 0.75,
  loess.degree = 2,
  loess.surface = c("interpolate", "direct"),
  loess.statistics = c("approximate", "exact", "none"),
  loess.trace.hat = c("exact", "approximate"),
  loess.cell = 0.2,
  loess.iterations = 4,
  loess.iterTrace = FALSE,
  mlr.smoother.type = "sm.ps",
```

```
mlr.ps.int = 4,
 mlr.degree = 3,
 mlr.s.df = 4,
 mlr.niknots = 4,
 weights = NULL,
 w.function = NULL,
 w.covs = NULL,
 w.landmark.type = "state",
 w.max = 10,
 w.stabilised = FALSE,
 w.max.follow = NULL,
 pv.group.vars = NULL,
 pv.n.pctls = NULL,
 pv.precalc = NULL,
 pv.ids = NULL,
 CI = FALSE,
 CI.type = "bootstrap",
 CI.R.boot = NULL,
 CI.seed = 1,
  transitions.out = NULL,
 assess.moderate = TRUE,
 assess.mean = TRUE,
)
```

Arguments

data.mstate	Validation data in msdata format
data.raw	Validation data in data.frame (one row per individual)
j	Landmark state at which predictions were made
S	Landmark time at which predictions were made
t	Follow up time at which calibration is to be assessed
tp.pred	Data frame or matrix of predicted transition probabilities at time t, if in state j at time s. There must be a separate column for the predicted transition probabilities into every state, even if these predicted transition probabilities are 0.
tp.pred.plot	Data frame or matrix of predicted risks for each possible transition over which to plot the calibration curves. Argument provided to enable user to apply bootstrapping manually.
calib.type	Whether calibration plots are estimated using BLR-IPCW ('blr'), MLR-IPCW ('mlr') or pseudo-values ('pv')
curve.type	Whether calibration curves are estimated using restricted cubic splines ('rcs') or loess smoothers ('loess')
rcs.nk	Number of knots when curves are estimated using restricted cubic splines
loess.span	Span when curves are estimated using loess smoothers
loess.degree	Degree when curves are estimated. using loess smoothers

loess.surface see loess.control loess.statistics			
see loess.control loess.trace.hat			
	see loess.control		
loess.cell loess.iteratio			
loess.iterTrac	see loess.control		
10033.110111 ac	see loess.control		
mlr.smoother.t	ype		
	Type of smoothing applied. Takes values s (see s), sm.ps (see sm.ps) or sm.os (see sm.os).		
mlr.ps.int	the number of equally-spaced B spline intervals in the vector spline smoother (see $sm.ps$)		
mlr.degree	the degree of B-spline basis in the vector spline smoother (see sm.ps)		
mlr.s.df	degrees of freedom of vector spline (see s)		
mlr.niknots	number of interior knots (see sm.os)		
weights	Vector of inverse probability of censoring weights		
w.function	Custom function for estimating the inverse probability of censoring weights		
w.covs	Character vector of variable names to adjust for when calculating inverse probability of censoring weights		
w.landmark.typ			
	Whether weights are estimated in all individuals uncensored at time s ('all') or only in individuals uncensored and in state j at time s ('state')		
w.max	Maximum bound for inverse probability of censoring weights		
w.stabilised	Indicates whether inverse probability of censoring weights should be stabilised or not		
w.max.follow	Maximum follow up for model calculating inverse probability of censoring weights. Reducing this to $t+1$ may aid in the proportional hazards assumption being met in this model.		
pv.group.vars	Variables to group by before calculating pseudo-values		
pv.n.pctls	Number of percentiles of predicted risk to group by before calculating pseudo-values		
pv.precalc	Pre-calculated pseudo-values		
pv.ids	Id's of individuals to calculate pseudo-values for		
CI	Size of confidence intervals as a %		
CI.type	Method for estimating confidence interval (currently restricted to bootstrap)		
CI.R.boot	Number of bootstrap replicates when estimating the confidence interval for the calibration curve		
CI.seed	Seed for bootstrapping procedure		

transitions.out

Transitions for which to calculate calibration curves. Will do all possible transitions if left as NULL.

assess.moderate

TRUE/FALSE whether to estimate data for calibration plots

assess.mean TRUE/FALSE whether to estimate mean calibration

Extra arguments to be passed to w.function (custom function for estimating

weights)

Details

Observed event probabilities at time t are estimated for predicted transition probabilities tp.pred out of state j at time s.

calib.type = 'blr' estimates calibration curves using techniques for assessing the calibration of a binary logistic regression model (Van Calster et al., 2016). A choice between restricted cubic splines and loess smoothers for estimating the calibration curve can be made using curve.type. Landmarking (van Houwelingen HC, 2007) is applied to only assess calibration in individuals who are uncensored and in state j at time s. Calibration can only be assessed in individuals who are also uncensored at time t, which is accounted for using inverse probability of censoring weights (Hernan M, Robins J, 2020). See method BLR-IPCW from Pate et al XXXXX for a full explanation of the approach.

calib.type = 'mlr' estimates calibration scatter plots using a technique for assessing the calibration of multinomial logistic regression models, namely the nominal calibration framework of van Hoorde et al. (2014, 2015). Landmarking (van Houwelingen HC, 2007) is applied to only assess calibration in individuals who are uncensored and in state j at time s. Calibration can only be assessed in individuals who are also uncensored at time t, which is accounted for using inverse probability of censoring weights (Hernan M, Robins J, 2020). See method BLR-IPCW from Pate et al XXXX for a full explanation of the approach.

calib.type = 'pv' estimates calibration curves using using pseudo-values (Andersen PK, Pohar Perme M, 2010) calculated using the Aalen-Johansen estimator (Aalen OO, Johansen S, 1978). Calibration curves are generated by regressing the pseudo-values on the predicted transition probabilities. A choice between restricted cubic splines and loess smoothers for estimating the calibration curve can be made using curve.type. Landmarking (van Houwelingen HC, 2007) is applied to only assess calibration in individuals who are uncensored and in state j at time s. The nature of pseudo-values means calibration can be assessed in all landmarked individuals, regardless of their censoring time. See method PV from Pate et al XXXX for a full explanation of the approach.

Two datasets for the same cohort of inidividuals must be provided. Firstly, data.raw must be a data.frame with one row per individual containing the variables for the time until censoring (dtcens), and an indicator for censoring dtcens.s, where (dtcens.s = 1) if an individual is censored at time dtcens, and dtcens.s = 0 otherwise. When an individual enters an absorbing state, this prevents censoring from happening (i.e. dtcens.s = 0). data.raw must also contain the desired variables for estimating the weights. Secondly, data.mstate must be a dataset of class msdata, generated using the [mstate] package. This dataset is used to apply the landmarking and identify which state individuals are in at time t. While data.mstate can be derived from data.raw, it would be inefficient to do this within calibmsm::calib_msm due to the bootstrapping procedure, and therefore they must be inputted seperately.

Unless the user specifies the weights using weights, the weights are estimated using a cox-proportional hazard model, assuming a linear functional form of the variables defined in w. covs. We urge users to specify their own model for estimating the weights. The weights argument must be a vector with length equal to the number of rows of data.raw.

Confidence intervals cannot be produced for the calibration scatter plots (calib.type = 'mlr'). For calibration curves estimated using calib.type = 'blr', confidence intervals can only be estimated using bootstrapping (CI.type = 'bootstrap). This procedure uses the internal method for estimating weights, we therefore encourage users to specify their own bootstrapping procedure, which incorporates their own model for estimating the weights. Details on how to do this are provided in the vignette *BLR-IPCW-manual-bootstrap*. For calibration curves estimated using calib.type = 'pv', confidence intervals can be estimated using bootstrapping (CI.type = 'bootstrap) or parametric formulae (CI.type = 'parametric). For computational reasons we recommend using the parametric approach.

The calibration plots can be plotted using plot.calib_msm and plot.calib_mlr.

Value

calib_msm returns a list containing two elements: plotdata and metadata. The plotdata element contains the data for the calibration plots. This will itself be a list with each element containing calibration plot data for the transition probabilities into each of the possible states. Each list element contains patient ids (id) from data.raw, the predicted transition probabilities (pred) and the estimated observed event probabilities (obs). If a confidence interval is requested, upper (obs.upper) and lower (obs.lower) bounds for the observed event probabilities are also returned. If tp.pred.plot is specified, column (id) is not returned. The metadata element contains metadata including: a vector of the possible transitions, a vector of which transitions calibration curves have been estimated for, the size of the confidence interval, the method for estimating the calibration curve and other user specified information.

References

Aalen OO, Johansen S. An Empirical Transition Matrix for Non-Homogeneous Markov Chains Based on Censored Observations. *Scand J Stat.* 1978;5(3):141-150.

Andersen PK, Pohar Perme M. Pseudo-observations in survival analysis. *Stat Methods Med Res.* 2010;19(1):71-99. doi:10.1177/0962280209105020

Hernan M, Robins J (2020). "12.2 Estimating IP weights via modeling." In *Causal Inference: What If*, chapter 12.2. Chapman Hall/CRC, Boca Raton.

Van Calster B, Nieboer D, Vergouwe Y, De Cock B, Pencina MJ, Steyerberg EW (2016). "A calibration hierarchy for risk models was defined: From utopia to empirical data." *Journal of Clinical Epidemiology*, 74, 167–176. ISSN 18785921. doi:10.1016/j.jclinepi.2015. 12.005. URL http://dx.doi.org/10.1016/j.jclinepi.2015.12.005

Van Hoorde K, Vergouwe Y, Timmerman D, Van Huffel S, Steyerberg W, Van Calster B (2014). "Assessing calibration of multinomial risk prediction models." *Statistics in Medicine*, 33(15), 2585–2596. doi:10.1002/sim.6114.

Van Hoorde K, Van Huffel S, Timmerman D, Bourne T, Van Calster B (2015). "A spline-based tool to assess and visualize the calibration of multiclass risk predictions." *Journal of Biomedical Informatics*, 54, 283–293. ISSN 15320464. doi:10.1016/j.jbi.2014.12.016. URL http://dx.doi.org/10.1016/j.jbi.2014.12.016.

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van Houwelingen HC (2007). "Dynamic Prediction by Landmarking in Event History Analysis." *Scandinavian Journal of Statistics*, 34(1), 70–85.

Yee TW (2015). *Vector Generalized Linear and Additive Models*. 1 edition. Springer New, NY. ISBN 978-1-4939-4198-8. doi:10.1007/978-1-4939-2818-7. URL https://link.springer.com/book/10.1007/978-1-4939-2818-7.

Examples

```
# Estimate BLR-IPCW calibration curves for the predicted transition
\# probabilities at time t = 1826, when predictions were made at time
\# s = 0 in state j = 1. These predicted transition probabilities are stored in tps0.
# Extract the predicted transition probabilities out of state j = 1
tp.pred <- dplyr::select(dplyr::filter(tps0, j == 1), any_of(paste("pstate", 1:6, sep = "")))</pre>
# Now estimate the observed event probabilities for each possible transition.
dat.calib <-
calib_msm(data.mstate = msebmtcal,
data.raw = ebmtcal,
j=1,
s=0,
t = 1826,
tp.pred = tp.pred,
w.covs = c("year", "agecl", "proph", "match"))
# Summarise the output
summary(dat.calib)
```

ebmtcal

European Group for Blood and Marrow Transplantation data (one row per individual)

Description

A data frame of 2,279 individuals with blood cancer who have undergone a transplant. This data is identical to the ebmt4 data, except two extra variables have been derived, time until censoring and a censoring indicator, which are required to assess calibration using some of the methods in calibmsm. Code for the derivation of this dataset is provided in the source code for the package.

Usage

ebmtcal

Format

'ebmtcal':

A data frame with 2,279 rows and 17 columns:

10 ebmtcal.cmprsk

id Patient indentifier

rec, rec.s Time until and event indicator for recovery variable

ae, ae.s Time until and event indicator for adverse event variable

recae, recae.s Time until and event indicator for recovery + adverse event variable

rel, rel.s Time until and event indicator for relapse variable

srv, srv.s Time until and event indicator for death variable

year Year of transplant

agecl Age at transplant

proph Prophylaxis

match Donor-recipient match

dtcens Time of censoring

dtcens.s Event indicator, 1:censoring occured, 0: absorbing state entered before censoring occured

Source

This dataset was derived from data made available within the mstate package, see ebmt4. The data was originally provided by the European Group for Blood and Marrow Transplantation (https://www.ebmt.org/). We reiterate the source statement given by the developers of mstate: "We acknowledge the European Society for Blood and Marrow Transplantation (EBMT) for making available these data. Disclaimer: these data were simplified for the purpose of illustration of the analysis of competing risks and multi-state models and do not reflect any real life situation. No clinical conclusions should be drawn from these data."

References

EBMT (2023). "Data from the European Society for Blood and Marrow Transplantation." URL https://search.r-project.org/CRAN/refmans/mstate/html/EBMT-data.html.

de Wreede LC, Fiocco M, Putter H (2011). "mstate: An R Package for the Analysis of Competing Risks and Multi-State Models." *Journal of Statistical Software*, 38(7).

 ${\it ebmtcal.cmprsk} \qquad {\it European~Group~for~Blood~and~Marrow~Transplantation~data~(one~row~per~individual)}$

Description

Used in vignette/article: Comparison-with-graphical-calibration-curves-in-competing-risks-setting.

Usage

ebmtcal.cmprsk

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Format

'ebmtcal.cmprsk':

A data frame with 2,279 rows and 17 columns:

id Patient indentifier

rec, rec.s Time until and event indicator for recovery variable

ae, ae.s Time until and event indicator for adverse event variable

recae, recae.s Time until and event indicator for recovery + adverse event variable

rel, rel.s Time until and event indicator for relapse variable

srv, srv.s Time until and event indicator for death variable

year Year of transplant

agecl Age at transplant

proph Prophylaxis

match Donor-recipient match

dtcens Time of censoring

dtcens.s Event indicator, 1:censoring occured, 0: absorbing state entered before censoring occured

Details

A data frame of 2,279 individuals with blood cancer who have undergone a transplant. This data is identical to the ebmt4 data, except two extra variables have been derived, time until censoring and a censoring indicator, which are required to assess calibration using some of the methods in calibmsm. Specifically, the time until censoring ar calculated in the setting of a competing risks model out of the first state, where no further transitions can be made. This means entry into any state (as they are all absorbing states) will have the effect of preventing censoring from occurring, and dtcens and dtcens.s will be different than the values found in ebmtcal. This dataset has been designed to be used alongside dataset msebmtcal.cmprsk, when assessing the calibration of a competing risks model. Code for the derivation of this dataset is provided in the source code for the package.

Source

This dataset was derived from data made available within the mstate package, see ebmt4. The data was originally provided by the European Group for Blood and Marrow Transplantation (https://www.ebmt.org/). We reiterate the source statement given by the developers of mstate: "We acknowledge the European Society for Blood and Marrow Transplantation (EBMT) for making available these data. Disclaimer: these data were simplified for the purpose of illustration of the analysis of competing risks and multi-state models and do not reflect any real life situation. No clinical conclusions should be drawn from these data."

References

EBMT (2023). "Data from the European Society for Blood and Marrow Transplantation." URL https://search.r-project.org/CRAN/refmans/mstate/html/EBMT-data.html.

de Wreede LC, Fiocco M, Putter H (2011). "mstate: An R Package for the Analysis of Competing Risks and Multi-State Models." *Journal of Statistical Software*, 38(7).

12 msebmtcal

msdata format.	msebmtcal	European Group for Blood and Marrow Transplantation data in msdata format.
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Description

The ebmt4 data converted into msdata format (see msprep), using the processes implemented in the mstate package. Code for the derivation of this dataset is provided in the source code for the package.

Usage

msebmtcal

Format

'msebmtcal':

A data frame in msdata format (see msprep) with 15,512 rows and 8 columns:

id Patient indentifier

from transition from state

to transition to state

trans transition number

Tstart time entered state 'from'

Tstop time leaving state 'from'

time time in state 'from'

status event indicator, 1:transitioned to state 'to'

Source

This dataset was derived from data made available within the mstate package, see ebmt4. The data was originally provided by the European Group for Blood and Marrow Transplantation (https://www.ebmt.org/). We reiterate the source statement given by the developers of mstate: "We acknowledge the European Society for Blood and Marrow Transplantation (EBMT) for making available these data. Disclaimer: these data were simplified for the purpose of illustration of the analysis of competing risks and multi-state models and do not reflect any real life situation. No clinical conclusions should be drawn from these data."

References

EBMT (2023). "Data from the European Society for Blood and Marrow Transplantation." URL https://search.r-project.org/CRAN/refmans/mstate/html/EBMT-data.html.

de Wreede LC, Fiocco M, Putter H (2011). "mstate: An R Package for the Analysis of Competing Risks and Multi-State Models." *Journal of Statistical Software*, 38(7).

msebmtcal.cmprsk 13

msebmtcal.cmprsk	European Group for Blood and Marrow Transplantation data in competing risks format, for transitions out of the initial state only
	peting risks formal, for transitions out of the unital state only

Description

Used in vignette/article: Comparison-with-graphical-calibration-curves-in-competing-risks-setting.

Usage

```
msebmtcal.cmprsk
```

Format

'msebmtcal.cmprsk':

A data frame with 9,116 rows and 8 columns:

id Patient indentifier

from transition from state

to transition to state

trans transition number

Tstart time entered state 'from'

Tstop time leaving state 'from'

time time in state 'from'

status event indicator, 1:transitioned to state 'to'

Details

The ebmt4 data converted into msdata format (see msprep), where all subsequent states are considered absorbing states. i.e. only transitions out of the initial state are considered, meaning this data constitutes a competing risks model out of the initial state. Code for the derivation of this dataset is provided in the source code for the package.

Source

This dataset was derived from data made available within the mstate package, see ebmt4. The data was originally provided by the European Group for Blood and Marrow Transplantation (https://www.ebmt.org/). We reiterate the source statement given by the developers of mstate: "We acknowledge the European Society for Blood and Marrow Transplantation (EBMT) for making available these data. Disclaimer: these data were simplified for the purpose of illustration of the analysis of competing risks and multi-state models and do not reflect any real life situation. No clinical conclusions should be drawn from these data."

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References

EBMT (2023). "Data from the European Society for Blood and Marrow Transplantation." URL https://search.r-project.org/CRAN/refmans/mstate/html/EBMT-data.html.

de Wreede LC, Fiocco M, Putter H (2011). "mstate: An R Package for the Analysis of Competing Risks and Multi-State Models." *Journal of Statistical Software*, 38(7).

plot.calib_mlr

Plots calibration scatter plots for objects of class calib_mlr estimated using using calib_msm.

Description

Plots calibration scatter plots for the transition probabilities of a multistate model estimated using the MLR-IPCW approach.

Usage

```
## S3 method for class 'calib_mlr'
plot(
  х,
  . . . ,
  combine = TRUE,
 ncol = NULL,
 nrow = NULL,
  size.point = 0.5,
  size.text = 12,
  transparency.plot = 0.25,
 marg.density = FALSE,
 marg.density.size = 5,
 marg.density.type = "density",
 marg.rug = FALSE,
 marg.rug.transparency = 0.1,
  titles.include = TRUE,
  titles = NULL,
  axis.titles.x = NULL,
  axis.titles.text.x = "Predicted risk",
  axis.titles.y = NULL,
  axis.titles.text.y = "Observed risk"
)
```

Arguments

x Object of class calib_mlr generated from calib_msm
... Other

combine Whether to combine into one plot using ggarrange, or return as

Whether to combine into one plot using ggarrange, or return as a list of individual plots plot.calib_mlr 15

```
Number of columns for combined calibration plot
ncol
                  Number of rows for combined calibration plot
nrow
size.point
                  Size of points in scatter plot
size.text
                  Size of text in plot
transparency.plot
                  Degree of transparency for points in the calibration scatter plot
                  Whether to produce marginal density plots TRUE/FALSE
marg.density
marg.density.size
                  Size of the main plot relative to the density plots (see ggMarginal)
marg.density.type
                  What type of marginal plot to show (see ggMarginal)
marg.rug
                  Whether to produce marginal rug plots TRUE/FALSE
marg.rug.transparency
                  Degree of transparency for the density rug plot along each axis
titles.include Whether to include titles for each individual calibration plots
titles
                  Vector of titles for the calibration plots. Defaults to "State k" for each plot.
axis.titles.x Position of plots for which to include title on x-axis
axis.titles.text.x
                  x-axis title
axis.titles.y Position of plots for which to include title on y-axis
axis.titles.text.y
                  y-axis title
```

Value

If combine = TRUE, returns an object of classes gg, ggplot, and ggarrange, as all ggplots have been combined into one object. If combine = FALSE, returns an object of class list, each element containing an object of class gg and ggplot.

Examples

```
# Using competing risks data out of initial state (see vignette: ... -in-competing-risk-setting).
# Estimate and plot MLR-IPCW calibration scatter plots for the predicted transition
# probabilities at time t = 1826, when predictions were made at time
# s = 0 in state j = 1. These predicted transition probabilities are stored in tp.cmprsk.j0.

# To minimise example time we reduce the datasets to 150 individuals.
# Extract the predicted transition probabilities out of state j = 1 for first 150 individuals
tp.pred <- tp.cmprsk.j0 |>
dplyr::filter(id %in% 1:150) |>
dplyr::select(any_of(paste("pstate", 1:6, sep = "")))
# Reduce ebmtcal to first 150 individuals
ebmtcal <- ebmtcal |> dplyr::filter(id %in% 1:150)
# Reduce msebmtcal.cmprsk to first 150 individuals
msebmtcal.cmprsk <- msebmtcal.cmprsk |> dplyr::filter(id %in% 1:150)
```

plot.calib_msm

```
# Now estimate the observed event probabilities for each possible transition.
dat.calib <-
calib_msm(data.mstate = msebmtcal.cmprsk,
    data.raw = ebmtcal,
    j=1,
    s=0,
    t = 1826,
    tp.pred = tp.pred,
    calib.type = "mlr",
    w.covs = c("year", "agecl", "proph", "match"),
    mlr.ps.int = 2,
    mlr.degree = 2)

# These are then plotted
    plot(dat.calib, combine = TRUE, nrow = 2, ncol = 3)</pre>
```

plot.calib_msm

Plots calibration curves estimated using calib_msm.

Description

Plots calibration curves for the transition probabilities of a multistate model estimated using BLR-IPCW and pseudo-value approaches.

Usage

```
## S3 method for class 'calib_msm'
plot(
  х,
  combine = TRUE,
  ncol = NULL,
  nrow = NULL,
  size.line = 0.5,
  size.text = 12,
  marg.density = FALSE,
 marg.density.size = 5,
  marg.density.type = "density",
 marg.rug = FALSE,
  marg.rug.transparency = 0.1,
  titles.include = TRUE,
  titles = NULL,
  axis.titles.x = NULL,
  axis.titles.text.x = "Predicted risk",
  axis.titles.y = NULL,
  axis.titles.text.y = "Observed risk",
  legend.include = TRUE,
```

plot.calib_msm 17

```
legend.seperate = FALSE,
legend.title = NULL,
legend.position = "bottom"
)
```

Arguments

x	Object of class 'calib_msm' generated from calib_msm.	
	Other	
combine	Whether to combine into one plot using ggarrange, or return as a list of individual plots	
ncol	Number of columns for combined calibration plot	
nrow	Number of rows for combined calibration plot	
size.line	Size of line plots	
size.text	Size of text in plot	
marg.density	Whether to produce marginal density plots TRUE/FALSE	
marg.density.si	ize	
	Size of the main plot relative to the density plots (see ggMarginal)	
marg.density.ty	/pe	
	What type of marginal plot to show (see ggMarginal)	
marg.rug	Whether to produce marginal rug plots TRUE/FALSE	
marg.rug.transp		
	Degree of transparency for the density rug plot along each axis	
titles.include	Whether to include titles for each individual calibration plots	
titles	Vector of titles for the calibration plots. Defaults to "State k" for each plot.	
axis.titles.x	Position of plots for which to include title on x-axis	
axis.titles.tex	xt.x	
	x-axis title	
axis.titles.y	Position of plots for which to include title on y-axis	
axis.titles.tex	ct.y	
	y-axis title	
legend.include	Whether to produce a legend	
legend.seperate		
	= Whether to include legend in plot (FALSE) or as a seperate object (TRUE)	
legend.title	Title of legend	
legend.position		
	Position of legend	

Value

If combine = TRUE, returns an object of classes gg, ggplot, and ggarrange, as all ggplots have been combined into one object. If combine = FALSE, returns an object of class list, each element containing an object of class gg and ggplot.

18 tp.cmprsk.j0

Examples

```
# Estimate and plot BLR-IPCW calibration curves for the predicted transition
\# probabilities at time t = 1826, when predictions were made at time
\# s = 0 in state j = 1. These predicted transition probabilities are stored in tps0.
# Extract the predicted transition probabilities out of state j = 1
tp.pred <- dplyr::select(dplyr::filter(tps0, j == 1), any_of(paste("pstate", 1:6, sep = "")))</pre>
# Now estimate the observed event probabilities for each possible transition.
dat.calib <-
calib_msm(data.mstate = msebmtcal,
data.raw = ebmtcal,
j=1,
s=0,
t = 1826,
 tp.pred = tp.pred,
 w.covs = c("year", "agecl", "proph", "match"))
 # These are then plotted
 plot(dat.calib, combine = TRUE, nrow = 2, ncol = 3)
```

tp.cmprsk.j0

Predicted risks for a competing risks model out of state j = 0

Description

Used in vignette/article: Comparison-with-graphical-calibration-curves-in-competing-risks-setting.

Usage

```
tp.cmprsk.j0
```

Format

'tp.cmprsk.j0':

A data frame with 2,279 rows and 13 columns:

id Patient indentifier

pstate1, pstate2, pstate3, pstate4, pstate5, pstate6 Predicted transition probabilities of transitions into states 1 to 6

se1, se2, se3, se4, se5, se6 Standard error of the predicted transition probabilities of transitions into states 1 to 6

tps0

Details

Data frame containing the predicted transition probabilities out of state j=1 made at time s=0, for a competing risks model out of the initial state (see msebmtcal.cmprsk). The predicted transition probabilities were estimated by fitting a competing risks model to the msebmtcal.cmprsk data using a leave-one-out approach. Code for deriving this dataset is provided in the source code for calibmsm. Code for the derivation of this dataset is provided in the source code for the package.

Source

This dataset was derived from data made available within the mstate package, see ebmt4. The data was originally provided by the European Group for Blood and Marrow Transplantation (https://www.ebmt.org/). We reiterate the source statement given by the developers of mstate: "We acknowledge the European Society for Blood and Marrow Transplantation (EBMT) for making available these data. Disclaimer: these data were simplified for the purpose of illustration of the analysis of competing risks and multi-state models and do not reflect any real life situation. No clinical conclusions should be drawn from these data."

References

EBMT (2023). "Data from the European Society for Blood and Marrow Transplantation." URL https://search.r-project.org/CRAN/refmans/mstate/html/EBMT-data.html.

de Wreede LC, Fiocco M, Putter H (2011). "mstate: An R Package for the Analysis of Competing Risks and Multi-State Models." *Journal of Statistical Software*, 38(7).

tps0 Predicted transition probabilities out of transplant state made at time s=0

Description

Data frame containing the predicted transition probabilities out of state j = 1 made at time s = 0. The predicted transition probabilities were estimated by fitting a multistate model to the ebmt4 data using a leave-one-out approach. Code for deriving this dataset is provided in the source code for calibmsm. Code for the derivation of this dataset is provided in the source code for the package.

Usage

tps0

Format

'tps0':

A data frame with 13,674 (CHANGE) rows and 14 columns:

id Patient indentifier

pstate1, pstate2, pstate3, pstate4, pstate5, pstate6 Predicted transition probabilities of transitions into states 1 to 6

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se1, se2, se3, se4, se5, se6 Standard error of the predicted transition probabilities of transitions into states 1 to 6

j State from which the predicted transition probabilities are estimated from

Source

This dataset was derived from data made available within the mstate package, see ebmt4. The data was originally provided by the European Group for Blood and Marrow Transplantation (https://www.ebmt.org/). We reiterate the source statement given by the developers of mstate: "We acknowledge the European Society for Blood and Marrow Transplantation (EBMT) for making available these data. Disclaimer: these data were simplified for the purpose of illustration of the analysis of competing risks and multi-state models and do not reflect any real life situation. No clinical conclusions should be drawn from these data."

References

EBMT (2023). "Data from the European Society for Blood and Marrow Transplantation." URL https://search.r-project.org/CRAN/refmans/mstate/html/EBMT-data.html.

de Wreede LC, Fiocco M, Putter H (2011). "mstate: An R Package for the Analysis of Competing Risks and Multi-State Models." *Journal of Statistical Software*, 38(7).

tps100 Predicted tran

Predicted transition probabilities out of every state made at time s = 100

Description

Data frame containing the predicted transition probabilities out of states 1 (transplant), 2 (adverse event), 3 (recovery) and 4 (adverse event + recovery), made at time s = 100. The predicted transition probabilities were estimated by fitting a multistate model to the ebmt4 data using a leave-one-out approach. Code for deriving this dataset is provided in the source code for calibmsm. Code for derivation of this dataset is provided in the source code for the package.

Usage

tps100

Format

'tps100':

A data frame with 13,674 (CHANGE) rows and 14 columns:

id Patient indentifier

pstate1, pstate2, pstate3, pstate4, pstate5, pstate6 Predicted transition probabilities of transitions into states 1 to 6

se1, se2, se3, se4, se5, se6 Standard error of the predicted transition probabilities of transitions into states 1 to 6

j State from which the predicted transition probabilities are estimated from

Source

This dataset was derived from data made available within the mstate package, see ebmt4. The data was originally provided by the European Group for Blood and Marrow Transplantation (https://www.ebmt.org/). We reiterate the source statement given by the developers of mstate: "We acknowledge the European Society for Blood and Marrow Transplantation (EBMT) for making available these data. Disclaimer: these data were simplified for the purpose of illustration of the analysis of competing risks and multi-state models and do not reflect any real life situation. No clinical conclusions should be drawn from these data."

References

EBMT (2023). "Data from the European Society for Blood and Marrow Transplantation." URL https://search.r-project.org/CRAN/refmans/mstate/html/EBMT-data.html.

de Wreede LC, Fiocco M, Putter H (2011). "mstate: An R Package for the Analysis of Competing Risks and Multi-State Models." *Journal of Statistical Software*, 38(7).

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