

Package ‘RiskStratifiedEstimation’

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Type Package

Title Risk Stratified Effect Estimation

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Description RiskStratifiedEstimation is an R package for implementing risk stratified analyses of treatment effect in an observational database in the OMOP Common Data Model. It combines functionality of PatientLevelPrediction and CohortMethod R packages to extract and analyze data from a database in the OMOP Common Data Model format. Analyses are performed in two distinct steps. First, a prediction step is implemented, where personalized risks are derived from a large set of covariates applying one of various possible prediction methods. In the second step, risk stratified treatment comparisons are performed. Large scale regularized regression is used to fit the propensity model. Inverse probability of treatment weights are calculated and truncated within strata of predicted risk. Only weighted Cox regression models are supported for the outcome model.

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Encoding UTF-8

LazyData true

RoxygenNote 6.0.1

Imports PatientLevelPrediction, CohortMethod, Cyclops, OhdsiRTools, foreach, FeatureExtraction, dplyr, ff, caret, ffbase, ggplot2, gridExtra, tidyr, doSNOW, survminer

Depends survival,

Suggests knitr, rmarkdown

VignetteBuilder knitr

NeedsCompilation no

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absoluteRiskReduction	<i>Creates a data frame of absolute risk reductions within risk strata</i>
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Description

Creates a data frame of absolute risk reductions within risk strata

Usage

```
absoluteRiskReduction(dataKM, timePoint)
```

Arguments

dataKM	A list with the Kaplan-Meier estimates within risk strata
timePoint	The time point based on which the absolute risk reductions will be calculated

Value

A data frame with the absolute risk reductions along with confidence intervals

cmToPlpData	<i>Transforms a cohortMethodData object into a plpData object</i>
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Description

Transforms a cohortMethodData object, as generated from [getDbCohortMethodData](#), into a plpData object appropriate for risk stratified analysis

Usage

```
cmToPlpData(cohortMethodData)
```

Arguments

cohortMethodData	The cohortMethodData object to be transformed
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Value

A plpData object

comparisonPlot	<i>Generates a comparison plot within risk quantiles</i>
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Description

Generates a plot with relative and absolute risk reductions. It can handle one or multiple outcomes. In the case of multiple outcomes the input must be a named list.

Usage

```
comparisonPlot(dataARR, dataRRR, cases, treatmentLabel, comparatorLabel,
  ylimARR, ylimRRR, ylimCases = c(0, 1), position_dodge.w = 0.2)
```

Arguments

dataARR	A list or dataframe same as the one generated by absoluteRiskReduction
dataRRR	A list or dataframe same as the one generated by relativeRiskReduction
cases	The dataframe or list with the outcome rates within risk strata. In the case of multiple outcomes the names of the list will be used as outcome labels
treatmentLabel	The name of the treatment drug
comparatorLabel	The name of the comparator drug
ylimARR	The limits on the y-axis of the absolute risk reduction plot
ylimRRR	The limits on the y-axis of the hazard ratio plot
ylimCases	The limits on the y-axis of the outcome rates bar plot
position_dodge.w	The amount of space between bullets in the plot in the case of multiple outcomes

Value

A plot with hazard ratios and absolute risk reductions across risk strata

createIPW	<i>Creates Inverse Probability Weights</i>
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Description

Calculates inverse probability weights based on the propensity score

Usage

```
createIPW(ps, weightsType = "ATE", useStabilizedWeights = TRUE,
  truncationLevels = c(0.01, 0.99))
```

Arguments

ps	A propensity score data frame as created from createPs
weightsType	The type of the weights to be used. Allowed options are 'ATE' for average treatment effect and 'ATT' for average treatment effect on the treated weights
useStabilizedWeights	Should stabilized weights be used?
truncationLevels	The level of truncation expressed in percentiles of the propensity score.

Value

The ps data frame provided as input along with a weights column

createStudyPopulation *Create a study population*

Description

Create a study population

Usage

```
createStudyPopulation(cohortMethodData, population = NULL, outcomeId,
  firstExposureOnly = FALSE, restrictToCommonPeriod = FALSE,
  washoutPeriod = 0, removeDuplicateSubjects = FALSE,
  removeSubjectsWithPriorOutcome = TRUE, priorOutcomeLookback = 99999,
  minDaysAtRisk = 1, riskWindowStart = 0, addExposureDaysToStart = FALSE,
  riskWindowEnd = 0, addExposureDaysToEnd = TRUE,
  censorAtNewRiskWindow = FALSE)
```

Arguments

cohortMethodData	An object of type cohortMethodData as generated using getDbCohortMethodData.
population	If specified, this population will be used as the starting point instead of the cohorts in the cohortMethodData object.
outcomeId	The ID of the outcome. If not specified, no outcome-specific transformations will be performed.
firstExposureOnly	Should only the first exposure per subject be included? Note that this is typically done in the createStudyPopulation function,
restrictToCommonPeriod	Restrict the analysis to the period when both treatments are observed?
washoutPeriod	The minimum required continuous observation time prior to index date for a person to be included in the cohort.
removeDuplicateSubjects	Remove subjects that are in both the treated and comparator cohort? See details for allowed values.

removeSubjectsWithPriorOutcome	Remove subjects that have the outcome prior to the risk window start?
priorOutcomeLookback	How many days should we look back when identifying prior outcomes?
minDaysAtRisk	The minimum required number of days at risk.
riskWindowStart	The start of the risk window (in days) relative to the index date (+ days of exposure if the addExposureDaysToStart parameter is specified).
addExposureDaysToStart	Add the length of exposure the start of the risk window?
riskWindowEnd	The end of the risk window (in days) relative to the index data (+ days of exposure if the addExposureDaysToEnd parameter is specified).
addExposureDaysToEnd	Add the length of exposure the risk window?
censorAtNewRiskWindow	If a subject is in multiple cohorts, should time-at-risk be censored when the new time-at-risk starts to prevent overlap?

Details

Create a study population by enforcing certain inclusion and exclusion criteria, defining a risk window, and determining which outcomes fall inside the risk window.

The removeduplicateSubjects argument can have one of the following values:

"keep all" Do not remove subjects that appear in both target and comparator cohort

"keep first" When a subjects appear in both target and comparator cohort, only keep whichever cohort is first in time.

"remove all" Remove subjects that appear in both target and comparator cohort completely from the analysis."

Value

A data frame specifying the study population. This data frame will have the following columns:

rowId A unique identifier for an exposure

subjectId The person ID of the subject

cohortStartdate The index date

outcomeCount The number of outcomes observed during the risk window

timeAtRisk The number of days in the risk window

survivalTime The number of days until either the outcome or the end of the risk window

plotCovariateBalance	<i>Plots the covariate balance before and after balancing</i>
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Description

Plots covariate before and after weighting using the inverse of the propensity score

Usage

```
plotCovariateBalance(ps, cohortMethodData, calculateWeights = TRUE,
  weightsType = "ATE", useStabilizedWeights = TRUE, truncationLevels,
  showNotBalancedCovariateIds = TRUE)
```

Arguments

ps	A propensity score data frame as created from createPs
cohortMethodData	A cohortMethodData object
calculateWeights	Should the weights be calculated?
weightsType	The type of the weights to be used. Allowed options are 'ATE' for average treatment effect and 'ATT' for average treatment effect on the treated weights
useStabilizedWeights	Should stabilized weights be used?
truncationLevels	The level of truncation expressed in percentiles of the propensity score.
showNotBalancedCovariateIds	Show covariate ids that were not balanced after weighting?

Value

The covariate balance plot

plotWeightedKM	<i>Plots the weighted Kaplan-Meier estimate</i>
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Description

Plots the weighted Kaplan-Meier estimate

Usage

```
plotWeightedKM(dataKM, xlim, ylim, ci = TRUE, title = NULL)
```

Arguments

dataKM	A dataframe containing the weighed Kaplan-Meier estimates as computed by weightedKM
xlim	Limits on x-axis
ylim	Limits on y-axis
ci	Should confidence intervals be displayed?
title	The title on the graph

Value

A weighted Kaplan-Meier plot

`relativeRiskReduction` *Creates a data frame with hazard ratios within risk strata*

Description

Creates a data frame with hazard ratios within risk strata

Usage

```
relativeRiskReduction(ps, calculateWeights = TRUE, weightsType = "ATE",
  useStabilizedWeights = TRUE, truncationLevels)
```

Arguments

ps	A list with the propensity data frames as generated from createPs
calculateWeights	Whether to calculate the weights using createIPW
weightsType	The type of the weights to be used. Allowed options are 'ATE' for average treatment effect and 'ATT' for average treatment effect on the treated weights
useStabilizedWeights	Should stabilized weights be used?
truncationLevels	The level of truncation expressed in percentiles of the propensity score.

Value

A data frame with hazard ratios along with confidence intervals

runRiskStratifiedEstimation

Runs a risk stratified analysis

Description

Runs a risk stratified analysis

Usage

```
runRiskStratifiedEstimation(cohortMethodData, population, modelSettings, save,
  testSplit = "person", testFraction = 0.3, nfold = 10, riskStrata = 4,
  weightsType = "ATE", useStabilizedWeights = TRUE, truncationLevels,
  timePoint, excludeCovariateIds = NULL, binary = TRUE,
  includeAllOutcomes = TRUE, requireTimeAtRisk = TRUE,
  savePlpPlots = FALSE, psThreads = 1, priorType = "laplace",
  verbosity = "INFO", analysisId = NULL, savePlpResult = TRUE,
  saveMapMatrix = TRUE, savePs = TRUE, saveDataKM = TRUE,
  saveAbsoluteRiskReduction = TRUE, saveRelativeRiskReduction = TRUE)
```

Arguments

cohortMethodData	A cohortMethodData object
population	The study population to perform the analysis
modelSettings	The model settings for the prediction step
save	The save directory
testSplit	The type of split for the cross validation. Should be either 'person' or 'time'
testFraction	The size of the test set
nfold	The number of folds for cross validation
riskStrata	The number of risk strata on which to perform the analysis
weightsType	The type of weights for the balancing of covariates. Should be either 'ATE' or 'ATT'
useStabilizedWeights	Should stabilized weights be used?
truncationLevels	The level of truncation expressed in percentiles of the propensity score.
timePoint	The time point of interest for the calculation of the absolute risk reduction
excludeCovariateIds	Covariate Ids to be excluded from calculation of propensity scores
binary	Forces the outcomeCount to be 0 or 1 in the prediction step
includeAllOutcomes	(binary) indicating whether to include people with outcomes who are not observed for the whole at risk period
requireTimeAtRisk	Should subjects without time at risk be removed at the prediction step?
savePlpPlots	(binary) Should plots for the prediction step be generated?

psThreads	The number of cores to use for the estimation of the propensity score. If 1 then serial approach is implemented
priorType	The prior for the propensity score model
verbosity	Sets the level of the verbosity. If the log level is at or higher in priority than the logger threshold, a message will print. The levels are: <ul style="list-style-type: none"> • DEBUGHighest verbosity showing all debug statements • TRACEShowing information about start and end of steps • INFOShow informative information (Default) • WARNShow warning messages • ERRORShow error messages • FATALBe silent except for fatal errors
analysisId	Identifier of the analysis
savePlpResult	Should the prediction result be saved?
saveMapMatrix	Should the map matrix with the risk stratum allocations be saved?
savePs	Should the propensity scores be saved?
saveDataKM	Should the weighted Kaplan-Meier estimates be saved?
saveAbsoluteRiskReduction	Should the absolute risk reduction estimates be saved?
saveRelativeRiskReduction	Should the hazard ratios be saved?

Value

ps	The propensity scores within risk strata along with patient weights
mapMatrix	The matrix that maps the patients to risk strata
dataKM	The weighted Kaplan-Meier estimates within risk strata
absoluteRiskReduction	The absolute risk reduction within risk strata
relativeRiskReduction	The relative risk reduction within risk strata
predictionResult	The result of the prediction step

weightedKM

Calculates and plots weighted Kaplan-Meier estimates

Description

Calculates the weighted Kaplan-Meier estimates based on: Xie J, Liu C. Adjusted Kaplan-Meier estimator and log-rank test with inverse probability of treatment weighting for survival data. *Statistics in Medicine* 2005; 24:3089–3110.

Usage

```
weightedKM(ps, calculateWeights = TRUE, weightsType = "ATE",
  useStabilizedWeights = TRUE, truncationLevels)
```

Arguments

<code>ps</code>	A data frame including the propensity scores as generated from <code>createPs</code>
<code>calculateWeights</code>	Whether to calculate the weights using <code>createIPW</code>
<code>weightsType</code>	The type of the weights to be used. Allowed options are 'ATE' for average treatment effect and 'ATT' for average treatment effect on the treated weights
<code>useStabilizedWeights</code>	Should stabilized weights be used?
<code>truncationLevels</code>	The level of truncation expressed in percentiles of the propensity score.

Value

A data frame with the Kaplan-Meier estimates

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