# Package 'CohortMethod'

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Type Package

cli, pillar,

```
Title New-User Cohort Method with Large Scale Propensity and Outcome Models
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Maintainer Martijn Schuemie <schuemie@ohdsi.org>
Description Functions for performing new-user cohort studies
      in an observational database in the OMOP Common Data Model. Can extract the
      necessary data from a database and use a large set of covariates for both the
      propensity and outcome model, including for example all drugs, diagnoses, procedures,
      as well as age, comorbidity indexes, etc. Large scale regularized regression is used to
      fit the propensity and outcome models. Functions are included for trimming, stratifying,
      (variable and fixed ratio) matching and weighting by propensity scores, as well as
      diagnostic functions, such as propensity score distribution plots and plots showing
      covariate balance before and after matching and/or trimming. Supported outcome models
      are (conditional) logistic regression, (conditional) Poisson regression, and
      (stratified) Cox regression. Also included are Kaplan-Meier plots that can adjust for
      the stratification or matching.
License Apache License 2.0
VignetteBuilder knitr
URL https://ohdsi.github.io/CohortMethod, https://github.com/OHDSI/CohortMethod
BugReports https://github.com/OHDSI/CohortMethod/issues
Depends R (>= 4.1.0),
      DatabaseConnector (>= 6.0.0),
      Cyclops (>= 3.1.2),
      FeatureExtraction (>= 3.0.0),
      Andromeda (>= 0.6.3)
Imports methods,
      ggplot2,
      gridExtra,
      grid,
      readr,
      plyr,
      dplyr,
      rlang,
```

Contents

Se St Pe bi cl	Acpp (>= 0.11.2), qlRender (>= 1.18.0), urvival, trailelLogger (>= 3.4.2), it64, heckmate, tempiricalCalibration,
	ip
pi ki rr E w R R R	ts testthat, ROC, nitr, markdown, funomia, vithr, futils, RSQLite, ResultModelManager, DhdsiShinyAppBuilder, markdown
System	Requirements Java
Linking	gTo Rcpp
NeedsC	Compilation yes
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Roxyge	en list(markdown = TRUE)
Encodi	ng UTF-8
Cont	ents
	adjustedKm checkCmInstallation CohortMethodData-class cohortMethodDataSimulationProfile computeCovariateBalance computeEquipoise computeMdrr computePsAuc

Contents 3

createPs									
createResultsDataModel	 		 		 		 	 	25
create Stratify By Ps And Covariates Arg									
createStratifyByPsArgs									
createStudyPopulation									
create Target Comparator Outcomes  .									
createTrimByIptwArgs	 		 		 		 	 	30
createTrimByPsArgs									
createTrimByPsToEquipoiseArgs	 		 		 		 	 	31
createTruncateIptwArgs									
drawAttritionDiagram	 		 		 		 	 	32
exportToCsv	 		 		 		 	 	32
fitOutcomeModel	 		 		 		 	 	33
getAttritionTable									
getDataMigrator									
getDbCohortMethodData									
getDefaultCmTable1Specifications .	 		 		 		 	 	38
getFileReference	 		 		 		 	 	39
getFollowUpDistribution	 		 		 		 	 	39
getGeneralizabilityTable									
getInteractionResultsSummary									
getOutcomeModel									
getPsModel									
getResultsDataModelSpecifications									
getResultsSummary			 		 				43
insertExportedResultsInSqlite					 		 	 	43
isCohortMethodData									
launchResultsViewer									
launchResultsViewerUsingSqlite									
loadCmAnalysisList									
loadCohortMethodData	 	 ·	 		 	 ·	 	 	46
loadTargetComparatorOutcomesList									
matchOnPs									
matchOnPsAndCovariates									
migrateDataModel									
plotCovariateBalanceOfTopVariables									
plotCovariateBalanceScatterPlot									
plotCovariatePrevalence									
plotFollowUpDistribution									
plotKaplanMeier									
plotPs									
plotTimeToEvent									
runCmAnalyses									
saveCmAnalysisList									61
saveCohortMethodData									61
									62
saveTargetComparatorOutcomesList simulateCohortMethodData									62
									63
stratifyByPs									64
stratifyByPsAndCovariates trimByIptw									
* *									
trimByPs								 	66 66
THEORY ENTINEAUTHOUSE									

4 checkCmInstallation

truncateIptw .												 								67
upload Results												 								68

Index 69

adjustedKm

Compute a weight-adjusted Kaplan-Meier curve

# **Description**

Compute a weight-adjusted Kaplan-Meier curve

# Usage

```
adjustedKm(weight, time, y)
```

# **Arguments**

weight Vector of observation weights

time Vector of event times

y Vector outcomes (0 indicates censoring, 1 indicates event-of-interest)

checkCmInstallation

Check is CohortMethod and its dependencies are correctly installed

# **Description**

Check is CohortMethod and its dependencies are correctly installed

# Usage

```
{\tt checkCmInstallation(connectionDetails)}
```

# Arguments

connectionDetails

An R object of type

connectionDetails created using the function createConnectionDetails in the DatabaseConnector package.

# **Details**

This function checks whether CohortMethod and its dependencies are correctly installed. This will check the database connectivity, large scale regression engine (Cyclops), and large data object handling (ff).

CohortMethodData-class 5

CohortMethodData-class

Cohort Method Data

# Description

CohortMethodData is an S4 class that inherits from CoviarateData, which in turn inherits from Andromeda. It contains information on the cohorts, their outcomes, and baseline covariates. Information about multiple outcomes can be captured at once for efficiency reasons.

A CohortMethodData is typically created using getDbCohortMethodData(), can only be saved using saveCohortMethodData(), and loaded using loadCohortMethodData().

#### Usage

```
## $4 method for signature 'CohortMethodData'
show(object)
## $4 method for signature 'CohortMethodData'
summary(object)
```

### **Arguments**

object

An object of type CohortMethodData.

 $cohort {\tt MethodDataSimulationProfile}$ 

A simulation profile

# Description

A simulation profile

### Usage

data(cohortMethodDataSimulationProfile)

compute Covariate Balance

Compute covariate balance before and after PS adjustment

# **Description**

For every covariate, prevalence in treatment and comparator groups before and after matching/trimming/weighting are computed. When variable ratio matching was used the balance score will be corrected according the method described in Austin et al (2008).

#### Usage

```
computeCovariateBalance(
  population,
  cohortMethodData,
  subgroupCovariateId = NULL,
  maxCohortSize = 250000,
  covariateFilter = NULL
)
```

# **Arguments**

population A data frame containing the people that are remaining after PS adjustment. cohortMethodData

An object of type CohortMethodData as generated using getDbCohortMethodData().

subgroupCovariateId

Optional: a covariate ID of a binary covariate that indicates a subgroup of interest. Both the before and after populations will be restricted to this subgroup before computing covariate balance.

maxCohortSize

If the target or comparator cohort are larger than this number, they will be down-sampled before computing covariate balance to save time. Setting this number to 0 means no downsampling will be applied.

covariateFilter

Determines the covariates for which to compute covariate balance. Either a vector of covariate IDs, or a table 1 specifications object as generated for example using FeatureExtraction::getDefaultTable1Specifications(). If covariateFilter = NULL, balance will be computed for all variables found in the data.

#### **Details**

The population data frame should have the following three columns:

- rowId (numeric): A unique identifier for each row (e.g. the person ID).
- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group.
- propensityScore (numeric): Propensity score.

### Value

Returns a tibble describing the covariate balance before and after PS adjustment, with one row per covariate, with the same data as the covariateRef table in the CohortMethodData object, and the following additional columns:

- beforeMatchingMeanTarget: The (weighted) mean value in the target before PS adjustment.
- beforeMatchingMeanComparator: The (weighted) mean value in the comparator before PS adjustment.
- beforeMatchingSumTarget: The (weighted) sum value in the target before PS adjustment.
- beforeMatchingSumComparator: The (weighted) sum value in the comparator before PS adjustment.
- beforeMatchingSdTarget: The standard deviation of the value in the target before PS adjustment.

- beforeMatchingSdComparator: The standard deviation of the value in the comparator before PS adjustment.
- beforeMatchingMean: The mean of the value across target and comparator before PS adjustment.
- beforeMatchingSd: The standard deviation of the value across target and comparator before PS adjustment.
- afterMatchingMeanTarget: The (weighted) mean value in the target after PS adjustment.
- afterMatchingMeanComparator: The (weighted) mean value in the comparator after PS adjustment.
- afterMatchingSumTarget: The (weighted) sum value in the target after PS adjustment.
- afterMatchingSumComparator: The (weighted) sum value in the comparator after PS adjustment.
- afterMatchingSdTarget: The standard deviation of the value in the target after PS adjustment.
- afterMatchingSdComparator: The standard deviation of the value in the comparator after PS adjustment.
- afterMatchingMean: The mean of the value across target and comparator after PS adjustment.
- afterMatchingSd: The standard deviation of the value across target and comparator after PS adjustment.
- beforeMatchingStdDiff: The standardized difference of means when comparing the target to the comparator before PS adjustment.
- afterMatchingStdDiff: The standardized difference of means when comparing the target to the comparator after PS adjustment.
- targetStdDiff: The standardized difference of means when comparing the target before PS adjustment to the target after PS adjustment.
- comparatorStdDiff: The standardized difference of means when comparing the comparator before PS adjustment to the comparator after PS adjustment. -targetComparatorStdDiff: The standardized difference of means when comparing the entire population before PS adjustment to the entire population after PS adjustment.

The 'beforeMatchingStdDiff' and 'afterMatchingStdDiff' columns inform on the balance: are the target and comparator sufficiently similar in terms of baseline covariates to allow for valid causal estimation?

The 'targetStdDiff', 'comparatorStdDiff', and 'targetComparatorStdDiff' columns inform on the generalizability: are the cohorts after PS adjustment sufficiently similar to the cohorts before adjustment to allow generalizing the findings to the original cohorts?

#### References

Austin, P.C. (2008) Assessing balance in measured baseline covariates when using many-to-one matching on the propensity-score. Pharmacoepidemiology and Drug Safety, 17: 1218-1225.

8 computeMdrr

computeEquipoise

Compute fraction in equipoise

#### **Description**

Compute fraction in equipoise

#### Usage

```
computeEquipoise(data, equipoiseBounds = c(0.3, 0.7))
```

### **Arguments**

data A data frame with at least the two columns described below. equipoiseBounds

The bounds on the preference score to determine whether a subject is in equipoise.

#### **Details**

Computes the fraction of the population (the union of the target and comparator cohorts) who are in clinical equipoise (i.e. who had a reasonable chance of receiving either target or comparator, based on the baseline characteristics).

The data frame should have a least the following two columns:

- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group
- propensityScore (numeric): Propensity score

#### Value

A numeric value (fraction in equipoise) between 0 and 1.

#### References

Walker AM, Patrick AR, Lauer MS, Hornbrook MC, Marin MG, Platt R, Roger VL, Stang P, and Schneeweiss S. (2013) A tool for assessing the feasibility of comparative effectiveness research, Comparative Effective Research, 3, 11-20

computeMdrr

Compute the minimum detectable relative risk

# Description

Compute the minimum detectable relative risk

computePsAuc 9

#### Usage

```
computeMdrr(
  population,
  alpha = 0.05,
  power = 0.8,
  twoSided = TRUE,
  modelType = "cox"
)
```

# **Arguments**

population A data frame describing the study population as created using the createStudyPopulation

function. This should at least have these columns: personSeqId, treatment, out-

comeCount, timeAtRisk.

alpha Type I error.

power 1 - beta, where beta is the type II error.

twoSided Consider a two-sided test?

modelType The type of outcome model that will be used. Possible values are "logistic",

"poisson", or "cox". Currently only "cox" is supported.

#### **Details**

Compute the minimum detectable relative risk (MDRR) and expected standard error (SE) for a given study population, using the actual observed sample size and number of outcomes. Currently, only computations for Cox and logistic models are implemented. For Cox model, the computations by Schoenfeld (1983) is used. For logistic models Wald's z-test is used.

#### Value

A data frame with the MDRR and some counts.

# References

Schoenfeld DA (1983) Sample-size formula for the proportional-hazards regression model, Biometrics, 39(3), 499-503

computePsAuc

Compute the area under the ROC curve

# Description

Compute the area under the ROC curve of the propensity score.

# Usage

```
computePsAuc(data, confidenceIntervals = FALSE, maxRows = 1e+05)
```

10 createCmAnalysis

### **Arguments**

data A data frame with at least the two columns described below confidenceIntervals

Compute 95 percent confidence intervals (computationally expensive for large data sets)

maxRows

Maximum number of rows to use. If the number of rows is larger, a random sample will be taken. This can increase speed, with minor cost to precision. Set to 0 to use all data.

#### **Details**

The data frame should have a least the following two columns:

- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group.
- propensityScore (numeric): Propensity score.

#### Value

A tibble holding the AUC and its 95 percent confidence interval

# **Examples**

```
treatment <- rep(0:1, each = 100)
propensityScore <- c(rnorm(100, mean = 0.4, sd = 0.25), rnorm(100, mean = 0.6, sd = 0.25))
data <- data.frame(treatment = treatment, propensityScore = propensityScore)
data <- data[data$propensityScore > 0 & data$propensityScore < 1, ]
computePsAuc(data)</pre>
```

createCmAnalysis

Create a CohortMethod analysis specification

#### **Description**

Create a CohortMethod analysis specification

### Usage

```
createCmAnalysis(
  analysisId = 1,
  description = "",
  getDbCohortMethodDataArgs,
  createStudyPopArgs,
  createPsArgs = NULL,
  trimByPsArgs = NULL,
  trimByPsToEquipoiseArgs = NULL,
  trimByIptwArgs = NULL,
  truncateIptwArgs = NULL,
  matchOnPsArgs = NULL,
  matchOnPsAndCovariatesArgs = NULL,
```

createCmAnalysis 11

```
stratifyByPsArgs = NULL,
stratifyByPsAndCovariatesArgs = NULL,
computeSharedCovariateBalanceArgs = NULL,
computeCovariateBalanceArgs = NULL,
fitOutcomeModelArgs = NULL
```

### **Arguments**

analysisId An integer that will be used later to refer to this specific set of analysis choices.

description A short description of the analysis.

 ${\tt getDbCohortMethodDataArgs}$ 

An object representing the arguments to be used when calling the getDbCohortMethodData()

function.

createStudyPopArgs

 $An object \ representing \ the \ arguments \ to \ be \ used \ when \ calling \ the \ {\tt createStudyPopulation()}$ 

function.

createPsArgs An object representing the arguments to be used when calling the createPs()

function.

trimByPsArgs An object representing the arguments to be used when calling the trimByPs()

function.

trimByPsToEquipoiseArgs

An object representing the arguments to be used when calling the trimByPsToEquipoise()

function.

 ${\tt trimByIptwArgs} \ \ An object representing the arguments to be used when calling the {\tt trimByIptw()}$ 

function.

truncateIptwArgs

An object representing the arguments to be used when calling the truncateIptw()

function.

matchOnPsArgs An object representing the arguments to be used when calling the matchOnPs()

function.

matchOnPsAndCovariatesArgs

An object representing the arguments to be used when calling the matchOnPsAndCovariates()

function.

stratifyByPsArgs

An object representing the arguments to be used when calling the stratifyByPs()

function.

stratify By Ps And Covariates Args

An object representing the arguments to be used when calling the stratifyByPsAndCovariates()

function.

computeSharedCovariateBalanceArgs

An object representing the arguments to be used when calling the computeCovariateBalance()

function per target-comparator-analysis.

 ${\tt computeCovariateBalanceArgs}$ 

 $An object \ representing \ the \ arguments \ to \ be \ used \ when \ calling \ the \ {\tt computeCovariateBalance()}$ 

function per target-comparator-outcome-analysis.

fitOutcomeModelArgs

An object representing the arguments to be used when calling the fitOutcomeModel()

function.

#### **Details**

Create a set of analysis choices, to be used with the runCmAnalyses() function.

Providing a NULL value for any of the argument applies the corresponding step will not be executed. For example, if createPsArgs = NULL, no propensity scores will be computed.

create CmDiagnostic Thresholds

Create CohortMethod diagnostics thresholds

# **Description**

Threshold used when calling exportToCsv() to determine if we pass or fail diagnostics.

# Usage

```
createCmDiagnosticThresholds(
  mdrrThreshold = 10,
  easeThreshold = 0.25,
  sdmThreshold = 0.1,
  equipoiseThreshold = 0.2,
  attritionFractionThreshold = NULL,
  generalizabilitySdmThreshold = 1
)
```

# **Arguments**

mdrrThreshold What is the maximum allowed minimum detectable relative risk (MDRR)?

easeThreshold What is the maximum allowed expected absolute systematic error (EASE).

sdmThreshold What is the maximum allowed standardized difference of mean (SDM)? If any

covariate has an SDM exceeding this threshold, the diagnostic will fail.

equipoiseThreshold

What is the minimum required equipoise?

attritionFractionThreshold

DEPRECATED. See generalizabilitySdmThreshold instead.

generalizabilitySdmThreshold

What is the maximum allowed standardized difference of mean (SDM)when comparing the population before and after PS adjustments? If the SDM is greater than this value, the diagnostic will fail.

# Value

An object of type CmDiagnosticThresholds.

createCmTable1 13

1	
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### **Description**

Creates a formatted table of cohort characteristics, to be included in publications or reports.

#### Usage

```
createCmTable1(
  balance,
  specifications = getDefaultCmTable1Specifications(),
  beforeTargetPopSize = NULL,
  beforeComparatorPopSize = NULL,
  afterTargetPopSize = NULL,
  afterComparatorPopSize = NULL,
  beforeLabel = "Before matching",
  afterLabel = "After matching",
  targetLabel = "Target",
  comparatorLabel = "Comparator",
  percentDigits = 1,
  stdDiffDigits = 2
)
```

#### **Arguments**

balance A data frame created by the compute Covariate Balance function.

specifications Specifications of which covariates to display, and how.

before Target Pop Size

The number of people in the target cohort before matching/stratification/trimming, to mention in the table header. If not provide, no number will be included in the header.

beforeComparatorPopSize

The number of people in the comparator cohort before matching/stratification/trimming, to mention in the table header. If not provide, no number will be included in the header.

afterTargetPopSize

The number of people in the target cohort after matching/stratification/trimming, to mention in the table header. If not provide, no number will be included in the header.

afterComparatorPopSize

The number of people in the comparator cohort after matching/stratification/trimming, to mention in the table header. If not provide, no number will be included in the header.

beforeLabel Label for identifying columns before matching / stratification / trimming.

Label for identifying columns after matching / stratification / trimming.

targetLabel Label for identifying columns of the target cohort.

comparatorLabel

Label for identifying columns of the comparator cohort.

#### Value

A data frame with the formatted table 1.

 $\label{lem:create} C create {\tt CohortMethodDataSimulationProfile} \\ {\tt Create\ simulation\ profile}$ 

# Description

Creates a profile based on the provided CohortMethodData object, which can be used to generate simulated data that has similar characteristics.

# Usage

```
createCohortMethodDataSimulationProfile(cohortMethodData, minCellCount = 5)
```

# **Arguments**

cohortMethodData

An object of type CohortMethodData as generated using getDbCohortMethodData().

minCellCount

If > 0, willset to zero all low-prevalence covariates in the supplied simulation table in order to prevent identification of persons.

# **Details**

The output of this function is an object that can be used by the simulateCohortMethodData() function to generate a cohortMethodData object.

#### Value

An object of type CohortDataSimulationProfile.

create Compute Covariate Balance Args

Create a parameter object for the function computeCovariateBalance

#### **Description**

Create a parameter object for the function computeCovariateBalance

### Usage

```
createComputeCovariateBalanceArgs(
  subgroupCovariateId = NULL,
  maxCohortSize = 250000,
  covariateFilter = NULL
)
```

createCreatePsArgs 15

#### **Arguments**

subgroupCovariateId

Optional: a covariate ID of a binary covariate that indicates a subgroup of interest. Both the before and after populations will be restricted to this subgroup before computing covariate balance.

maxCohortSize

If the target or comparator cohort are larger than this number, they will be down-sampled before computing covariate balance to save time. Setting this number to 0 means no downsampling will be applied.

covariateFilter

Determines the covariates for which to compute covariate balance. Either a vector of covariate IDs, or a table 1 specifications object as generated for example using FeatureExtraction::getDefaultTable1Specifications(). If covariateFilter = NULL, balance will be computed for all variables found in the data.

#### **Details**

Create an object defining the parameter values.

createCreatePsArgs

Create a parameter object for the function createPs

# **Description**

Create a parameter object for the function createPs

# Usage

#### **Arguments**

excludeCovariateIds

Exclude these covariates from the propensity model.

includeCovariateIds

Include only these covariates in the propensity model.

maxCohortSizeForFitting

If the target or comparator cohort are larger than this number, they will be down-sampled before fitting the propensity model. The model will be used to compute propensity scores for all subjects. The purpose of the sampling is to gain speed. Setting this number to 0 means no downsampling will be applied.

errorOnHighCorrelation

If true, the function will test each covariate for correlation with the treatment assignment. If any covariate has an unusually high correlation (either positive approximately), this will throw and arrow

or negative), this will throw and error.

stopOnError If an error occur, should the function stop? Else, the two cohorts will be assumed

to be perfectly separable.

prior The prior used to fit the model. See Cyclops::createPrior() for details.

control The control object used to control the cross-validation used to determine the

hyperparameters of the prior (if applicable). See Cyclops::createControl() for

details.

estimator The type of estimator for the IPTW. Options are estimator = "ate" for the average

treatment effect, estimator = "att" for the average treatment effect in the treated, and estimator = "ato" for the average treatment effect in the overlap population.

# **Details**

Create an object defining the parameter values.

createCreateStudyPopulationArgs

Create a parameter object for the function createStudyPopulation

#### **Description**

Create a parameter object for the function createStudyPopulation

# Usage

```
createCreateStudyPopulationArgs(
  firstExposureOnly = FALSE,
  restrictToCommonPeriod = FALSE,
  washoutPeriod = 0,
  removeDuplicateSubjects = "keep all",
  removeSubjectsWithPriorOutcome = TRUE,
  priorOutcomeLookback = 99999,
  minDaysAtRisk = 1,
  maxDaysAtRisk = 99999,
  riskWindowStart = 0,
  startAnchor = "cohort start",
  riskWindowEnd = 0,
  endAnchor = "cohort end",
  censorAtNewRiskWindow = FALSE
)
```

# Arguments

firstExposureOnly

Should only the first exposure per subject be included?

restrictToCommonPeriod

Restrict the analysis to the period when both exposures 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 target 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. Risk windows with fewer days

than this number are removed from the analysis.

maxDaysAtRisk The maximum allowed number of days at risk. Risk windows that are longer

will be truncated to this number of days.

riskWindowStart

The start of the risk window (in days) relative to the startAnchor.

startAnchor The anchor point for the start of the risk window. Can be "cohort start" or "cohort

end".

riskWindowEnd The end of the risk window (in days) relative to the endAnchor.

endAnchor The anchor point for the end of the risk window. Can be "cohort start" or "cohort

end".

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 an object defining the parameter values.

 ${\tt createDefaultMultiThreadingSettings}$ 

Create default CohortMethod multi-threading settings

# **Description**

Create CohortMethod multi-threading settings based on the maximum number of cores to be used.

# Usage

createDefaultMultiThreadingSettings(maxCores)

# Arguments

maxCores Maximum number of CPU cores to use.

# Value

An object of type CmMultiThreadingSettings.

#### See Also

```
createMultiThreadingSettings()
```

#### **Examples**

```
settings <- createDefaultMultiThreadingSettings(10)</pre>
```

createFitOutcomeModelArgs

Create a parameter object for the function fitOutcomeModel

# **Description**

Create a parameter object for the function fitOutcomeModel

# Usage

```
createFitOutcomeModelArgs(
  modelType = "logistic",
  stratified = FALSE,
  useCovariates = FALSE,
  inversePtWeighting = FALSE,
  interactionCovariateIds = c(),
  excludeCovariateIds = c(),
  includeCovariateIds = c(),
  profileGrid = NULL,
  profileBounds = c(log(0.1), log(10)),
  prior = createPrior("laplace", useCrossValidation = TRUE),
  control = createControl(cvType = "auto", seed = 1, resetCoefficients = TRUE,
    startingVariance = 0.01, tolerance = 2e-07, cvRepetitions = 10, noiseLevel = "quiet")
)
```

#### **Arguments**

modelType The type of outcome model that will be used. Possible values are "logistic",

"poisson", or "cox".

stratified Should the regression be conditioned on the strata defined in the population

object (e.g. by matching or stratifying on propensity scores)?

useCovariates Whether to use the covariates in the cohortMethodData object in the outcome

model.

inversePtWeighting

Use inverse probability of treatment weighting (IPTW)

interactionCovariateIds

An optional vector of covariate IDs to use to estimate interactions with the main treatment effect.

excludeCovariateIds

Exclude these covariates from the outcome model.

includeCovariateIds

Include only these covariates in the outcome model.

profileGrid A one-dimensional grid of points on the log(relative risk) scale where the likeli-

hood for coefficient of variables is sampled. See details.

profileBounds The bounds (on the log relative risk scale) for the adaptive sampling of the like-

lihood function. See details.

prior The prior used to fit the model. See Cyclops::createPrior() for details.

control The control object used to control the cross-validation used to determine the

hyperparameters of the prior (if applicable). See Cyclops::createControl() for

details.

#### **Details**

Create an object defining the parameter values.

createGetDbCohortMethodDataArgs

Create a parameter object for the function getDbCohortMethodData

### **Description**

Create a parameter object for the function getDbCohortMethodData

### Usage

```
createGetDbCohortMethodDataArgs(
  studyStartDate = "",
  studyEndDate = "",
  firstExposureOnly = FALSE,
  removeDuplicateSubjects = "keep all",
  restrictToCommonPeriod = FALSE,
  washoutPeriod = 0,
  maxCohortSize = 0,
  covariateSettings
)
```

# **Arguments**

studyStartDate A calendar date specifying the minimum date that a cohort index date can ap-

pear. Date format is 'yyyymmdd'.

studyEndDate A calendar date specifying the maximum date that a cohort index date can ap-

pear. Date format is 'yyyymmdd'. Important: the study end data is also used to truncate risk windows, meaning no outcomes beyond the study end date will be

considered.

firstExposureOnly

Should only the first exposure per subject be included? Note that this is typically done in the createStudyPopulation() function, but can already be done here for efficiency reasons.

 ${\tt removeDuplicateSubjects}$ 

Remove subjects that are in both the target and comparator cohort? See details for allowed values. Note that this is typically done in the createStudyPopulation function, but can already be done here for efficiency reasons.

restrict To Common Period

Restrict the analysis to the period when both treatments are observed?

washoutPeriod T

The minimum required continuous observation time prior to index date for a person to be included in the cohort. Note that this is typically done in the createStudyPopulation function, but can already be done here for efficiency reasons.

maxCohortSize

If either the target or the comparator cohort is larger than this number it will be

sampled to this size. maxCohortSize = 0 indicates no maximum size.

covariateSettings

An object of type covariateSettings as created using the FeatureExtraction::createCovariateSettings() function.

#### **Details**

Create an object defining the parameter values.

createMatchOnPsAndCovariatesArgs

Create a parameter object for the function matchOnPsAndCovariates

# **Description**

Create a parameter object for the function matchOnPsAndCovariates

#### Usage

```
createMatchOnPsAndCovariatesArgs(
  caliper = 0.2,
  caliperScale = "standardized logit",
  maxRatio = 1,
  allowReverseMatch = FALSE,
  covariateIds
)
```

# **Arguments**

caliper

The caliper for matching. A caliper is the distance which is acceptable for any match. Observations which are outside of the caliper are dropped. A caliper of 0 means no caliper is used.

caliperScale

The scale on which the caliper is defined. Three scales are supported: caliper-Scale = 'propensity score', caliperScale = 'standardized', or caliperScale = 'standardized logit'. On the standardized scale, the caliper is interpreted in standard deviations of the propensity score distribution. 'standardized logit' is similar, except that the propensity score is transformed to the logit scale because the PS is more likely to be normally distributed on that scale (Austin, 2011).

maxRatio

The maximum number of persons in the comparator arm to be matched to each person in the treatment arm. A maxRatio of 0 means no maximum: all comparators will be assigned to a target person.

allowReverseMatch

Allows n-to-1 matching if target arm is larger

covariateIds

One or more covariate IDs in the cohortMethodData object on which subjects should be also matched.

createMatchOnPsArgs 21

#### **Details**

Create an object defining the parameter values.

createMatchOnPsArgs

Create a parameter object for the function matchOnPs

#### **Description**

Create a parameter object for the function matchOnPs

### Usage

```
createMatchOnPsArgs(
  caliper = 0.2,
  caliperScale = "standardized logit",
  maxRatio = 1,
  allowReverseMatch = FALSE,
  stratificationColumns = c()
)
```

#### **Arguments**

caliper

The caliper for matching. A caliper is the distance which is acceptable for any match. Observations which are outside of the caliper are dropped. A caliper of 0 means no caliper is used.

caliperScale

The scale on which the caliper is defined. Three scales are supported: caliper-Scale = 'propensity score', caliperScale = 'standardized', or caliperScale = 'standardized logit'. On the standardized scale, the caliper is interpreted in standard deviations of the propensity score distribution. 'standardized logit' is similar, except that the propensity score is transformed to the logit scale because the PS is more likely to be normally distributed on that scale (Austin, 2011).

maxRatio

The maximum number of persons in the comparator arm to be matched to each person in the treatment arm. A maxRatio of 0 means no maximum: all comparators will be assigned to a target person.

 $allow {\tt ReverseMatch}$ 

Allows n-to-1 matching if target arm is larger

stratificationColumns

Names or numbers of one or more columns in the data data.frame on which subjects should be stratified prior to matching. No persons will be matched with persons outside of the strata identified by the values in these columns.

### **Details**

Create an object defining the parameter values.

createMultiThreadingSettings

Create CohortMethod multi-threading settings

#### **Description**

Create CohortMethod multi-threading settings

#### Usage

```
createMultiThreadingSettings(
  getDbCohortMethodDataThreads = 1,
  createPsThreads = 1,
  psCvThreads = 1,
  createStudyPopThreads = 1,
  trimMatchStratifyThreads = 1,
  computeSharedBalanceThreads = 1,
  computeBalanceThreads = 1,
  prefilterCovariatesThreads = 1,
  fitOutcomeModelThreads = 1,
  outcomeCvThreads = 1,
  calibrationThreads = 1
)
```

#### **Arguments**

getDbCohortMethodDataThreads

The number of parallel threads to use for building the cohortMethod data objects.

createPsThreads

The number of parallel threads to use for fitting the propensity models.

psCvThreads

The number of parallel threads to use for the cross-validation when estimating the hyperparameter for the propensity model. Note that the total number of CV threads at one time could be createPsThreads \* psCvThreads.

createStudyPopThreads

The number of parallel threads to use for creating the study population.

trimMatchStratifyThreads

 $\label{threads} The \ number \ of parallel \ threads \ to \ use \ for \ trimming, \ matching \ and \ stratifying. \\ compute Shared Balance Threads$ 

 $\label{threads} The number of parallel threads to use for computing shared covariate balance. compute Balance Threads$ 

The number of parallel threads to use for computing covariate balance.

prefilterCovariatesThreads

The number of parallel threads to use for prefiltering covariates.

fitOutcomeModelThreads

The number of parallel threads to use for fitting the outcome models.

outcomeCvThreads

The number of parallel threads to use for the cross-validation when estimating the hyperparameter for the outcome model. Note that the total number of CV threads at one time could be fitOutcomeModelThreads \* outcomeCvThreads.

createOutcome 23

```
calibrationThreads
```

The number of parallel threads to use for empirical calibration.

#### Value

An object of type CmMultiThreadingSettings.

#### See Also

createDefaultMultiThreadingSettings()

createOutcome

Create outcome definition

#### **Description**

Create outcome definition

# Usage

```
createOutcome(
  outcomeId,
  outcomeOfInterest = TRUE,
  trueEffectSize = NA,
  priorOutcomeLookback = NULL,
  riskWindowStart = NULL,
  startAnchor = NULL,
  riskWindowEnd = NULL,
  endAnchor = NULL)
```

# **Arguments**

outcomeId An integer used to identify the outcome in the outcome cohort table. outcomeOfInterest

Is this an outcome of interest? If not, creation of non-essential files will be skipped, including outcome=specific covariate balance files. This could be helpful to speed up analyses with many controls, for which we're only interested in the effect size estimate.

trueEffectSize For negative and positive controls: the known true effect size. To be used for empirical calibration. Negative controls have trueEffectSize = 1. If the true

effect size is unknown, use trueEffectSize = NA
priorOutcomeLookback

How many days should we look back when identifying prior. outcomes?

riskWindowStart

The start of the risk window (in days) relative to the startAnchor.

startAnchor The anchor point for the start of the risk window. Can be "cohort start" or

"cohort end".

riskWindowEnd The end of the risk window (in days) relative to the endAnchor.

endAnchor The anchor point for the end of the risk window. Can be "cohort start" or

"cohort end".

24 createPs

#### **Details**

Any settings here that are not NULL will override any values set in createCreateStudyPopulationArgs().

#### Value

An object of type outcome, to be used in createTargetComparatorOutcomes().

createPs

Create propensity scores

# **Description**

Creates propensity scores and inverse probability of treatment weights (IPTW) using a regularized logistic regression.

# Usage

```
createPs(
  cohortMethodData,
  population = NULL,
  excludeCovariateIds = c(),
  includeCovariateIds = c(),
  maxCohortSizeForFitting = 250000,
  errorOnHighCorrelation = TRUE,
  stopOnError = TRUE,
  prior = createPrior("laplace", exclude = c(0), useCrossValidation = TRUE),
  control = createControl(noiseLevel = "silent", cvType = "auto", seed = 1,
    resetCoefficients = TRUE, tolerance = 2e-07, cvRepetitions = 10, startingVariance = 0.01),
  estimator = "att"
)
```

# Arguments

cohortMethodData

An object of type CohortMethodData as generated using getDbCohortMethodData().

population

A data frame describing the population. This should at least have a rowId column corresponding to the rowId column in the CohortMethodData covariates object and a treatment column. If population is not specified, the full population in the CohortMethodData will be used.

excludeCovariateIds

Exclude these covariates from the propensity model.

includeCovariateIds

Include only these covariates in the propensity model.

maxCohortSizeForFitting

If the target or comparator cohort are larger than this number, they will be down-sampled before fitting the propensity model. The model will be used to compute propensity scores for all subjects. The purpose of the sampling is to gain speed. Setting this number to 0 means no downsampling will be applied.

createResultsDataModel 25

errorOnHighCorrelation

If true, the function will test each covariate for correlation with the treatment assignment. If any covariate has an unusually high correlation (either positive

or negative), this will throw and error.

stopOnError If an error occur, should the function stop? Else, the two cohorts will be assumed

to be perfectly separable.

prior The prior used to fit the model. See Cyclops::createPrior() for details.

control The control object used to control the cross-validation used to determine the

hyperparameters of the prior (if applicable). See Cyclops::createControl()

for details.

estimator The type of estimator for the IPTW. Options are estimator = "ate" for the

average treatment effect, estimator = "att" for the average treatment effect in the treated, and estimator = "ato" for the average treatment effect in the

overlap population.

#### **Details**

IPTW estimates either the average treatment effect (ate) or average treatment effect in the treated (att) using stabilized inverse propensity scores (Xu et al. 2010).

#### References

Xu S, Ross C, Raebel MA, Shetterly S, Blanchette C, Smith D. Use of stabilized inverse propensity scores as weights to directly estimate relative risk and its confidence intervals. Value Health. 2010;13(2):273-277. doi:10.1111/j.1524-4733.2009.00671.x

# **Examples**

```
data(cohortMethodDataSimulationProfile)
cohortMethodData <- simulateCohortMethodData(cohortMethodDataSimulationProfile, n = 1000)
ps <- createPs(cohortMethodData)</pre>
```

createResultsDataModel

Create the results data model tables on a database server.

#### **Description**

Create the results data model tables on a database server.

#### Usage

```
createResultsDataModel(
  connectionDetails = NULL,
  databaseSchema,
  tablePrefix = ""
)
```

#### **Arguments**

connectionDetails

DatabaseConnector connectionDetails instance @seealsoDatabaseConnector::createConnectionDetails

databaseSchema The schema on the server where the tables will be created.

tablePrefix (Optional) string to insert before table names for database table names

#### **Details**

Only PostgreSQL and SQLite servers are supported.

 ${\tt createStratifyByPsAndCovariatesArgs}$ 

Create a parameter object for the function stratifyByPsAndCovariates

# **Description**

Create a parameter object for the function stratifyByPsAndCovariates

# Usage

```
createStratifyByPsAndCovariatesArgs(
  numberOfStrata = 5,
  baseSelection = "all",
  covariateIds
)
```

# **Arguments**

numberOfStrata Into how many strata should the propensity score be divided? The boundaries of

the strata are automatically defined to contain equal numbers of target persons.

baseSelection What is the base selection of subjects where the strata bounds are to be deter-

mined? Strata are defined as equally-sized strata inside this selection. Possible

values are "all", "target", and "comparator".

covariateIds One or more covariate IDs in the cohortMethodData object on which subjects

should also be stratified.

# **Details**

Create an object defining the parameter values.

createStratifyByPsArgs

```
createStratifyByPsArgs
```

Create a parameter object for the function stratifyByPs

# **Description**

Create a parameter object for the function stratifyByPs

# Usage

```
createStratifyByPsArgs(
  numberOfStrata = 5,
  stratificationColumns = c(),
  baseSelection = "all"
)
```

# **Arguments**

numberOfStrata How many strata? The boundaries of the strata are automatically defined to contain equal numbers of target persons.

stratificationColumns

Names of one or more columns in the data data.frame on which subjects should also be stratified in addition to stratification on propensity score.

baseSelection

What is the base selection of subjects where the strata bounds are to be determined? Strata are defined as equally-sized strata inside this selection. Possible values are "all", "target", and "comparator".

# **Details**

Create an object defining the parameter values.

createStudyPopulation Create a study population

#### Description

Create a study population

# Usage

```
createStudyPopulation(
  cohortMethodData,
  population = NULL,
  outcomeId = NULL,
  firstExposureOnly = FALSE,
  restrictToCommonPeriod = FALSE,
  washoutPeriod = 0,
  removeDuplicateSubjects = "keep all",
  removeSubjectsWithPriorOutcome = TRUE,
```

```
priorOutcomeLookback = 99999,
minDaysAtRisk = 1,
maxDaysAtRisk = 99999,
riskWindowStart = 0,
startAnchor = "cohort start",
riskWindowEnd = 0,
endAnchor = "cohort end",
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 co-

horts in the cohortMethodData object.

outcomeId The ID of the outcome. If NULL, no outcome-specific transformations will be

performed.

firstExposureOnly

Should only the first exposure per subject be included?

restrict To Common Period

Restrict the analysis to the period when both exposures 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 target 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. Risk windows with fewer days

than this number are removed from the analysis.

maxDaysAtRisk The maximum allowed number of days at risk. Risk windows that are longer

will be truncated to this number of days.

riskWindowStart

The start of the risk window (in days) relative to the startAnchor.

startAnchor The anchor point for the start of the risk window. Can be "cohort start" or

"cohort end".

riskWindowEnd The end of the risk window (in days) relative to the endAnchor.

endAnchor The anchor point for the end of the risk window. Can be "cohort start" or

"cohort end".

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. If both cohorts start simultaneous, the person is removed from the analysis.
- "remove all": Remove subjects that appear in both target and comparator cohort completely from the analysis."

#### Value

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

- rowId: A unique identifier for an exposure.
- personSeqId: The person sequence 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.

createTargetComparatorOutcomes

Create target-comparator-outcomes combinations.

### **Description**

Create target-comparator-outcomes combinations.

#### Usage

```
createTargetComparatorOutcomes(
  targetId,
  comparatorId,
  outcomes,
  excludedCovariateConceptIds = c(),
  includedCovariateConceptIds = c()
)
```

# **Arguments**

targetId A cohort ID identifying the target exposure in the exposure table.

comparatorId A cohort ID identifying the comparator exposure in the exposure table.

outcomes A list of object of type outcome as created by createOutcome().

30 createTrimByPsArgs

 ${\tt excludedCovariateConceptIds}$ 

A list of concept IDs that cannot be used to construct covariates. This argument is to be used only for exclusion concepts that are specific to the target-comparator combination.

 $included {\tt CovariateConceptIds}$ 

A list of concept IDs that must be used to construct covariates. This argument is to be used only for inclusion concepts that are specific to the target-comparator combination.

#### **Details**

Create a set of hypotheses of interest, to be used with the runCmAnalyses() function.

#### Value

An object of type targetComparatorOutcomes.

createTrimByIptwArgs Create a parameter object for the function trimByIptw

# **Description**

Create a parameter object for the function trimByIptw

# Usage

```
createTrimByIptwArgs(maxWeight = 10)
```

# Arguments

maxWeight The maximum allowed IPTW.

#### **Details**

Create an object defining the parameter values.

createTrimByPsArgs Create a parameter object for the function trimByPs

# Description

Create a parameter object for the function trimByPs

# Usage

```
createTrimByPsArgs(trimFraction = 0.05)
```

# **Arguments**

trimFraction

This fraction will be removed from each treatment group. In the target group, persons with the highest propensity scores will be removed, in the comparator group person with the lowest scores will be removed.

# **Details**

Create an object defining the parameter values.

createTrimByPsToEquipoiseArgs

Create a parameter object for the function trimByPsToEquipoise

# **Description**

Create a parameter object for the function trimByPsToEquipoise

# Usage

```
createTrimByPsToEquipoiseArgs(bounds = c(0.3, 0.7))
```

# Arguments

bounds

The upper and lower bound on the preference score for keeping persons.

# **Details**

Create an object defining the parameter values.

createTruncateIptwArgs

Create a parameter object for the function truncateIptw

# **Description**

Create a parameter object for the function truncateIptw

# Usage

```
createTruncateIptwArgs(maxWeight = 10)
```

# **Arguments**

maxWeight

The maximum allowed IPTW.

# **Details**

Create an object defining the parameter values.

32 exportToCsv

### **Description**

drawAttritionDiagram draws the attrition diagram, showing how many people were excluded from the study population, and for what reasons.

# Usage

```
drawAttritionDiagram(
  object,
  targetLabel = "Target",
  comparatorLabel = "Comparator",
  fileName = NULL
)
```

# **Arguments**

object Either an object of type cohortMethodData, a population object generated by

functions like createStudyPopulation, or an object of type outcomeModel.

targetLabel A label to us for the target cohort.

comparatorLabel

A label to us for the comparator cohort.

fileName Name of the file where the plot should be saved, for example 'plot.png'. See the

function ggsave in the ggplot2 package for supported file formats.

# Value

A ggplot object. Use the ggsave function to save to file in a different format.

 ${\tt exportToCsv}$ 

Export cohort method results to CSV files

# **Description**

Export cohort method results to CSV files

# Usage

```
exportToCsv(
  outputFolder,
  exportFolder = file.path(outputFolder, "export"),
  databaseId,
  minCellCount = 5,
  maxCores = 1,
  cmDiagnosticThresholds = createCmDiagnosticThresholds()
)
```

fitOutcomeModel 33

#### **Arguments**

outputFolder The folder where runCmAnalyses() generated all results.

exportFolder The folder where the CSV files will written.

databaseId A unique ID for the database. This will be appended to most tables.

minCellCount To preserve privacy: the minimum number of subjects contributing to a count before it can be included in the results. If the count is below this threshold, it will be set to -minCellCount.

maxCores How many parallel cores should be used?

 ${\tt cmDiagnosticThresholds}$ 

An object of type CmDiagnosticThresholds as created using createCmDiagnosticThresholds().

#### **Details**

This requires that runCmAnalyses() has been executed first. It exports all the results in the outputFolder to CSV files for sharing with other sites.

#### Value

Does not return anything. Is called for the side-effect of populating the exportFolder with CSV files

fitOutcomeModel Create an outcome model, and compute the relative risk

# **Description**

Create an outcome model, and computes the relative risk

#### Usage

```
fitOutcomeModel(
 population,
 cohortMethodData = NULL,
 modelType = "logistic",
  stratified = FALSE,
 useCovariates = FALSE,
  inversePtWeighting = FALSE,
  interactionCovariateIds = c(),
  excludeCovariateIds = c(),
  includeCovariateIds = c(),
 profileGrid = NULL,
 profileBounds = c(log(0.1), log(10)),
 prior = createPrior("laplace", useCrossValidation = TRUE),
  control = createControl(cvType = "auto", seed = 1, resetCoefficients = TRUE,
  startingVariance = 0.01, tolerance = 2e-07, cvRepetitions = 10, noiseLevel = "quiet")
)
```

34 fitOutcomeModel

#### **Arguments**

population A population object generated by createStudyPopulation(), potentially fil-

tered by other functions.

cohortMethodData

An object of type CohortMethodData as generated using getDbCohortMethodData().

Can be omitted if not using covariates and not using interaction terms.

modelType The type of outcome model that will be used. Possible values are "logistic",

"poisson", or "cox".

stratified Should the regression be conditioned on the strata defined in the population

object (e.g. by matching or stratifying on propensity scores)?

useCovariates Whether to use the covariates in the cohortMethodData object in the outcome

model.

inversePtWeighting

Use inverse probability of treatment weighting (IPTW)

interactionCovariateIds

An optional vector of covariate IDs to use to estimate interactions with the main

treatment effect.

excludeCovariateIds

Exclude these covariates from the outcome model.

includeCovariateIds

Include only these covariates in the outcome model.

profileGrid A one-dimensional grid of points on the log(relative risk) scale where the likeli-

hood for coefficient of variables is sampled. See details.

profileBounds The bounds (on the log relative risk scale) for the adaptive sampling of the like-

lihood function. See details.

prior The prior used to fit the model. See Cyclops::createPrior() for details. The

prior is only applied to non-treatment variables, so is not used when useCovariates

= FALSE.

control The control object used to control the cross-validation used to determine the

hyperparameters of the prior (if applicable). See Cyclops::createControl()

for details.

# Details

For likelihood profiling, either specify the profileGrid for a completely user-defined grid, or profileBounds for an adaptive grid. Both should be defined on the log effect size scale. When both profileGrid and profileGrid are NULL likelihood profiling is disabled.

### Value

An object of class OutcomeModel. Generic function print, coef, and confint are available.

getAttritionTable 35

 ${\tt getAttritionTable}$ 

Get the attrition table for a population

# Description

Get the attrition table for a population

# Usage

```
getAttritionTable(object)
```

# **Arguments**

object

Either an object of type CohortMethodData, a population object generated by functions like createStudyPopulation(), or an object of type outcomeModel.

# Value

A tibble specifying the number of people and exposures in the population after specific steps of filtering.

getDataMigrator

Get database migrations instance

# **Description**

Returns ResultModelManager DataMigrationsManager instance.

# Usage

```
getDataMigrator(connectionDetails, databaseSchema, tablePrefix = "")
```

# Arguments

 ${\tt connectionDetails}$ 

DatabaseConnector connection details object

databaseSchema String schema where database schema lives

tablePrefix (Optional) Use if a table prefix is used before table names (e.g. "cd\_")

#### Value

Instance of ResultModelManager::DataMigrationManager that has interface for converting existing data models

#### **Description**

This function executes a large set of SQL statements against the database in OMOP CDM format to extract the data needed to perform the analysis.

### Usage

```
getDbCohortMethodData(
  connectionDetails,
  cdmDatabaseSchema,
  tempEmulationSchema = getOption("sqlRenderTempEmulationSchema"),
  targetId,
  comparatorId,
  outcomeIds,
  studyStartDate = "",
  studyEndDate = "",
  exposureDatabaseSchema = cdmDatabaseSchema,
  exposureTable = "drug_era",
  outcomeDatabaseSchema = cdmDatabaseSchema,
  outcomeTable = "condition_occurrence",
  cdmVersion = "5",
  firstExposureOnly = FALSE,
  removeDuplicateSubjects = "keep all",
  restrictToCommonPeriod = FALSE,
  washoutPeriod = 0,
  maxCohortSize = 0,
  covariateSettings
)
```

### **Arguments**

connectionDetails

An R object of type connectionDetails created using the DatabaseConnector::createConnection function.

cdmDatabaseSchema

The name of the database schema that contains the OMOP CDM instance. Requires read permissions to this database. On SQL Server, this should specify both the database and the schema, so for example 'cdm\_instance.dbo'.

tempEmulationSchema

Some database platforms like Oracle and Impala do not truly support temp tables. To emulate temp tables, provide a schema with write privileges where temp tables can be created.

targetId

A unique identifier to define the target cohort. If exposureTable = DRUG\_ERA, targetId is a concept ID and all descendant concepts within that concept ID will be used to define the cohort. If exposureTable <> DRUG\_ERA, targetId is used to select the COHORT\_DEFINITION\_ID in the cohort-like table.

comparatorId A unique identifier to define the comparator cohort. If exposureTable = DRUG\_ERA,

comparatorId is a concept ID and all descendant concepts within that concept ID will be used to define the cohort. If exposureTable <> DRUG\_ERA, comparatorId is used to select the COHORT\_DEFINITION\_ID in the cohort-like table.

outcomeIds A list of cohort IDs used to define outcomes.

studyStartDate A calendar date specifying the minimum date that a cohort index date can ap-

pear. Date format is 'yyyymmdd'.

studyEndDate A calendar date specifying the maximum date that a cohort index date can ap-

pear. Date format is 'yyyymmdd'. Important: the study end data is also used to truncate risk windows, meaning no outcomes beyond the study end date will be

considered.

exposureDatabaseSchema

The name of the database schema that is the location where the exposure data

used to define the exposure cohorts is available.

exposureTable The tablename that contains the exposure cohorts. If exposureTable <> DRUG\_ERA,

 $then\ expectation\ is\ exposure {\tt Table}\ has\ form at\ of\ COHORT\ table:\ COHORT\_DEFINITION\_ID,$ 

SUBJECT\_ID, COHORT\_START\_DATE, COHORT\_END\_DATE.

outcomeDatabaseSchema

The name of the database schema that is the location where the data used to

define the outcome cohorts is available.

outcomeTable The tablename that contains the outcome cohorts. If outcomeTable <> CONDI-

TION\_OCCURRENCE, then expectation is outcomeTable has format of CO-HORT table: COHORT\_DEFINITION\_ID, SUBJECT\_ID, COHORT\_START\_DATE,

COHORT\_END\_DATE.

cdmVersion Define the OMOP CDM version used: currently supports "5".

firstExposureOnly

Should only the first exposure per subject be included? Note that this is typically done in the createStudyPopulation() function, but can already be done here

for efficiency reasons.

removeDuplicateSubjects

Remove subjects that are in both the target and comparator cohort? See details for allowed values. Note that this is typically done in the createStudyPopulation

function, but can already be done here for efficiency reasons.

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. Note that this is typically done in the createStudyPopulation function, but can already be done here for efficiency

reasons.

maxCohortSize If either the target or the comparator cohort is larger than this number it will be

sampled to this size. maxCohortSize = 0 indicates no maximum size.

covariateSettings

 $An \,object\,of\,type\,covariate Settings\,as\,created\,using\,the\,\textit{FeatureExtraction::createCovariate} and all the action and the covariate settings are created as a constant of the covariate settings are created as a covariate setting and the covariate settings are created using the covariate setting are covariated as a co$ 

function.

# **Details**

Based on the arguments, the treatment and comparator cohorts are retrieved, as well as outcomes occurring in exposed subjects. The treatment and comparator cohorts can be identified using the

DRUG\_ERA table, or through user-defined cohorts in a cohort table either inside the CDM schema or in a separate schema. Similarly, outcomes are identified using the CONDITION\_ERA table or through user-defined cohorts in a cohort table either inside the CDM schema or in a separate schema. Covariates are automatically extracted from the appropriate tables within the CDM.

**Important**: The target and comparator drug must not be included in the covariates, including any descendant concepts. You will need to manually add the drugs and descendants to the excludedCovariateConceptIds of the covariateSettings argument.

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."

If the covariateSettings include cohort-based covariates, and the covariateCohortTable is NULL, the covariateCohortDatabaseSchema and covariateCohortTable will be set to the exposureDatabaseSchema and exposureTable, respectively.

#### Value

A CohortMethodData object.

getDefaultCmTable1Specifications

Get the default table 1 specifications

# **Description**

Loads the default specifications for a table 1, to be used with the createCmTable1 function.

Important: currently only works for binary covariates.

# Usage

getDefaultCmTable1Specifications()

#### Value

A specifications objects.

getFileReference 39

getFileReference Get file reference

#### **Description**

Get file reference

# Usage

```
getFileReference(outputFolder)
```

# **Arguments**

outputFolder Name of the folder where all the outputs have been written to.

#### Value

A tibble containing file names of artifacts generated for each target-comparator-outcome-analysis combination.

getFollowUpDistribution

Get the distribution of follow-up time

# **Description**

Get the distribution of follow-up time

# Usage

```
getFollowUpDistribution(population, quantiles = c(0, 0.25, 0.5, 0.75, 1))
```

# Arguments

population A data frame describing the study population as created using the createStudyPopulation

function. This should at least have these columns: treatment, timeAtRisk.

quantiles The quantiles of the population to compute minimum follow-up time for.

# **Details**

Get the distribution of follow-up time as quantiles. Follow-up time is defined as time-at-risk, so not censored at the outcome.

# Value

A data frame with per treatment group at each quantile the amount of follow-up time available.

getGeneralizabilityTable

Get information on generalizability

# **Description**

to assess generalizability we compare the distribution of covariates before and after any (propensity score) adjustments. We compute the standardized difference of mean as our metric of generalizability. (Lipton et al., 2017)

Depending on our target estimand, we need to consider a different base population for generalizability. For example, if we aim to estimate the average treatment effect in thetreated (ATT), our base population should be the target population, meaning we should consider the covariate distribution before and after PS adjustment in the target population only. By default this function will attempt to select the right base population based on what operations have been performed on the population. For example, if PS matching has been performed we assume the target estimand is the ATT, and the target population is selected as base.

Requires running computeCovariateBalance()' first.

## Usage

getGeneralizabilityTable(balance, baseSelection = "auto")

#### Arguments

balance A data frame created by the computeCovariateBalance function.

baseSelection The selection of the population to consider for generalizability. Options are

"auto", "target", "comparator", and "both". The "auto" option will attempt to use the balance meta-data to pick the most appropriate population based on the

target estimator.

# Value

A tibble with the following columns:

- covariateId: The ID of the covariate. Can be linked to the covariates and covariateRef tables in the CohortMethodData object.
- covariateName: The name of the covariate.
- beforeMatchingMean: The mean covariate value before any (propensity score) adjustment.
- afterMatchingMean: The mean covariate value after any (propensity score) adjustment.
- stdDiff: The standardized difference of means between before and after adjustment.

The tibble also has a 'baseSelection' attribute, documenting the base population used to assess generalizability.

## References

Tipton E, Hallberg K, Hedges LV, Chan W (2017) Implications of Small Samples for Generalization: Adjustments and Rules of Thumb, Eval Rev. Oct;41(5):472-505.

getInteractionResultsSummary

Get a summary report of the analyses results

# Description

Get a summary report of the analyses results

# Usage

getInteractionResultsSummary(outputFolder)

# **Arguments**

outputFolder

Name of the folder where all the outputs have been written to.

#### Value

A tibble containing summary statistics for each target-comparator-outcome-analysis combination.

getOutcomeModel

Get the outcome model

# Description

Get the full outcome model, so showing the betas of all variables included in the outcome model, not just the treatment variable.

# Usage

```
getOutcomeModel(outcomeModel, cohortMethodData)
```

## **Arguments**

outcomeModel

An object of type OutcomeModel as generated using he fitOutcomeModel() function.

cohortMethodData

An object of type CohortMethodData as generated using getDbCohortMethodData().

# Value

A tibble.

getPsModel

Get the propensity model

# Description

Returns the coefficients and names of the covariates with non-zero coefficients.

# Usage

```
getPsModel(propensityScore, cohortMethodData)
```

# Arguments

propensityScore

The propensity scores as generated using the createPs() function.

 ${\tt cohortMethodData}$ 

 $An object of type Cohort Method Data as generated using \verb|getDbCohortMethodData()|.$ 

# Value

A tibble.

 ${\tt getResultsDataModelSpecifications}$ 

Get specifications for CohortMethod results data model

# Description

Get specifications for CohortMethod results data model

# Usage

getResultsDataModelSpecifications()

# Value

A tibble data frame object with specifications

getResultsSummary 43

getResultsSummary	Get a summary report of the analyses result.
gethesurtssummar y	Get a summary report of the analyses result

#### **Description**

Get a summary report of the analyses results

## Usage

```
getResultsSummary(outputFolder)
```

# **Arguments**

outputFolder Name of the folder where all the outputs have been written to.

# Value

A tibble containing summary statistics for each target-comparator-outcome-analysis combination.

insertExportedResultsInSqlite

Insert exported results into a SQLite database

# **Description**

Insert exported results into a SQLite database

# Usage

insertExportedResultsInSqlite(sqliteFileName, exportFolder, cohorts)

# Arguments

sqliteFileName The name of the SQLite file to store the results in. If the file does not exist it

will be created.

exportFolder The folder containing the CSV files to upload, as generated using the exportToCsv()

function.

cohorts A data frame describing the cohorts used in the study. Should include the target,

comparator, and outcome of interest cohorts. The data frame should at least have

a cohortDefinitionId and cohortName columns.

# Value

Does not return anything. Called for the side effect of inserting data into the SQLite database.

44 launchResultsViewer

isCohortMethodData

Check whether an object is a CohortMethodData object

# Description

Check whether an object is a CohortMethodData object

# Usage

```
isCohortMethodData(x)
```

# Arguments

Х

The object to check.

#### Value

A logical value.

launchResultsViewer

Launch Shiny app using

# Description

Launch Shiny app using

# Usage

launchResultsViewer(connectionDetails, databaseSchema)

# **Arguments**

 ${\tt connectionDetails}$ 

An R object of type connectionDetails created using the DatabaseConnector::createConnection function.

databaseSchema The name of the database schema where the results were written using uploadResults().

# Value

Does not return anything. Is called for the side-effect of launching the Shiny app.

 $launch {\tt Results Viewer Using Sqlite}$ 

Launch Shiny app using a SQLite database

# **Description**

Launch Shiny app using a SQLite database

# Usage

launchResultsViewerUsingSqlite(sqliteFileName)

# **Arguments**

sqliteFileName The name of the SQLite file where the results were stored using the insertExportedResultsInSqlifunction.

# Value

Does not return anything. Is called for the side-effect of launching the Shiny app.

 $load {\tt CmAnalysisList}$ 

Load a list of cmAnalysis from file

# Description

Load a list of objects of type cmAnalysis from file. The file is in JSON format.

# Usage

loadCmAnalysisList(file)

# **Arguments**

file

The name of the file

# Value

A list of objects of type cmAnalysis.

loadCohortMethodData Load the cohort method data from a file

# **Description**

Loads an object of type CohortMethodData from a file in the file system.

# Usage

loadCohortMethodData(file)

# **Arguments**

file

The name of the file containing the data.

## Value

An object of class CohortMethodData.

loadTargetComparatorOutcomesList

Load a list of targetComparatorOutcomes from file

# Description

Load a list of objects of type targetComparatorOutcomes from file. The file is in JSON format.

# Usage

loadTargetComparatorOutcomesList(file)

# Arguments

file

The name of the file

#### Value

A list of objects of type targetComparatorOutcomes.

matchOnPs 47

ma	tchC	)nPs

Match persons by propensity score

#### **Description**

Use the provided propensity scores to match target to comparator persons.

#### Usage

```
matchOnPs(
  population,
  caliper = 0.2,
  caliperScale = "standardized logit",
  maxRatio = 1,
  allowReverseMatch = FALSE,
  stratificationColumns = c()
)
```

## **Arguments**

population A data frame with the three columns described below.

caliper The caliper for matching. A caliper is the distance which is acceptable for any

match. Observations which are outside of the caliper are dropped. A caliper of

0 means no caliper is used.

caliperScale The scale on which the caliper is defined. Three scales are supported: caliperScale

= 'propensity score', caliperScale = 'standardized', or caliperScale = 'standardized logit'. On the standardized scale, the caliper is interpreted in standard deviations of the propensity score distribution. 'standardized logit' is similar, except that the propensity score is transformed to the logit scale because the PS is more likely to be normally distributed on that scale (Austin, 2011).

maxRatio

The maximum number of persons in the comparator arm to be matched to each person in the treatment arm. A maxRatio of 0 means no maximum: all com-

parators will be assigned to a target person.

allowReverseMatch

Allows n-to-1 matching if target arm is larger

stratification Columns

Names or numbers of one or more columns in the data data.frame on which subjects should be stratified prior to matching. No persons will be matched with persons outside of the strata identified by the values in these columns.

# **Details**

The data frame should have the following three columns:

- rowId (numeric): A unique identifier for each row (e.g. the person ID).
- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group.
- propensityScore (numeric): Propensity score.

The default caliper (0.2 on the standardized logit scale) is the one recommended by Austin (2011).

#### Value

Returns a date frame with the same columns as the input data plus one extra column: stratumId. Any rows that could not be matched are removed

# References

Rassen JA, Shelat AA, Myers J, Glynn RJ, Rothman KJ, Schneeweiss S. (2012) One-to-many propensity score matching in cohort studies, Pharmacoepidemiology and Drug Safety, May, 21 Suppl 2:69-80.

Austin, PC. (2011) Optimal caliper widths for propensity-score matching when estimating differences in means and differences in proportions in observational studies, Pharmaceutical statistics, March, 10(2):150-161.

# **Examples**

```
rowId <- 1:5
treatment <- c(1, 0, 1, 0, 1)
propensityScore <- c(0, 0.1, 0.3, 0.4, 1)
age_group <- c(1, 1, 1, 1, 1)
data <- data.frame(
   rowId = rowId,
    treatment = treatment,
   propensityScore = propensityScore,
   age_group = age_group
)
result <- matchOnPs(data, caliper = 0, maxRatio = 1, stratificationColumns = "age_group")</pre>
```

matchOnPsAndCovariates

Match by propensity score as well as other covariates

# **Description**

Use the provided propensity scores and a set of covariates to match target to comparator persons.

# Usage

```
matchOnPsAndCovariates(
  population,
  caliper = 0.2,
  caliperScale = "standardized logit",
  maxRatio = 1,
  allowReverseMatch = FALSE,
  cohortMethodData,
  covariateIds
)
```

#### **Arguments**

population A data frame with the three columns described below.

caliper The caliper for matching. A caliper is the distance which is acceptable for any

match. Observations which are outside of the caliper are dropped. A caliper of

0 means no caliper is used.

caliperScale The scale on which the caliper is defined. Three scales are supported: caliperScale

= 'propensity score', caliperScale = 'standardized', or caliperScale = 'standardized logit'. On the standardized scale, the caliper is interpreted in standard deviations of the propensity score distribution. 'standardized logit' is similar, except that the propensity score is transformed to the logit scale because the PS is more likely to be normally distributed on that scale (Austin, 2011).

maxRatio The maximum number of persons in the comparator arm to be matched to each

person in the treatment arm. A maxRatio of 0 means no maximum: all com-

parators will be assigned to a target person.

allowReverseMatch

Allows n-to-1 matching if target arm is larger

cohortMethodData

An object of type CohortMethodData as generated using getDbCohortMethodData().

covariateIds One or more covariate IDs in the cohortMethodData object on which subjects

should be also matched.

#### **Details**

The data frame should have the following three columns:

- rowId (numeric): A unique identifier for each row (e.g. the person ID).
- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group.
- propensityScore (numeric): Propensity score.

The default caliper (0.2 on the standardized logit scale) is the one recommended by Austin (2011).

#### Value

Returns a tibble with the same columns as the input data plus one extra column: stratumId. Any rows that could not be matched are removed

#### References

Rassen JA, Shelat AA, Myers J, Glynn RJ, Rothman KJ, Schneeweiss S. (2012) One-to-many propensity score matching in cohort studies, Pharmacoepidemiology and Drug Safety, May, 21 Suppl 2:69-80.

Austin, PC. (2011) Optimal caliper widths for propensity-score matching when estimating differences in means and differences in proportions in observational studies, Pharmaceutical statistics, March, 10(2):150-161.

migrateDataModel

Migrate Data model

# **Description**

Migrate data from current state to next state

It is strongly advised that you have a backup of all data (either sqlite files, a backup database (in the case you are using a postgres backend) or have kept the csv/zip files from your data generation.

# Usage

```
migrateDataModel(connectionDetails, databaseSchema, tablePrefix = "")
```

#### **Arguments**

```
connectionDetails
DatabaseConnector connection details object

databaseSchema String schema where database schema lives

tablePrefix (Optional) Use if a table prefix is used before table names (e.g. "cd_")
```

```
plotCovariateBalanceOfTopVariables
```

Plot variables with largest imbalance

# **Description**

Create a plot showing those variables having the largest imbalance before matching, and those variables having the largest imbalance after matching. Requires running computeCovariateBalance first.

#### Usage

```
plotCovariateBalanceOfTopVariables(
  balance,
  n = 20,
  maxNameWidth = 100,
  title = NULL,
  fileName = NULL,
  beforeLabel = "before matching",
  afterLabel = "after matching"
)
```

#### **Arguments**

balance A data frame created by the computeCovariateBalance function.

n (Maximum) count of covariates to plot.

maxNameWidth Covariate names longer than this number of characters are truncated to create a

nicer plot.

title Optional: the main title for the plot.

fileName Name of the file where the plot should be saved, for example 'plot.png'. See the

function ggsave in the ggplot2 package for supported file formats.

beforeLabel Label for identifying data before matching / stratification / trimming.

afterLabel Label for identifying data after matching / stratification / trimming.

#### Value

A ggplot object. Use the ggplot2::ggsave function to save to file in a different format.

plotCovariateBalanceScatterPlot

Create a scatterplot of the covariate balance

# Description

Create a scatterplot of the covariate balance, showing all variables with balance before and after matching on the x and y axis respectively. Requires running computeCovariateBalance first.

#### Usage

```
plotCovariateBalanceScatterPlot(
  balance,
  absolute = TRUE,
  threshold = 0,
  title = "Standardized difference of mean",
  fileName = NULL,
  beforeLabel = "Before matching",
  afterLabel = "After matching",
  showCovariateCountLabel = FALSE,
  showMaxLabel = FALSE
)
```

#### **Arguments**

balance A data frame created by the computeCovariateBalance function.

absolute Should the absolute value of the difference be used?

threshold Show a threshold value for after matching standardized difference.

title The main title for the plot.

fileName Name of the file where the plot should be saved, for example 'plot.png'. See the

function ggsave in the ggplot2 package for supported file formats.

beforeLabel Label for the x-axis.

```
afterLabel Label for the y-axis.

showCovariateCountLabel
Show a label with the number of covariates included in the plot?

showMaxLabel Show a label with the maximum absolute standardized difference after match-
```

ing/stratification?

#### Value

A ggplot object. Use the ggplot2::ggsave function to save to file in a different format.

```
plotCovariatePrevalence
```

Plot covariate prevalence

#### **Description**

Plot prevalence of binary covariates in the target and comparator cohorts, before and after matching. Requires running computeCovariateBalance first.

## Usage

```
plotCovariatePrevalence(
  balance,
  threshold = 0,
  title = "Covariate prevalence",
  fileName = NULL,
  beforeLabel = "Before matching",
  afterLabel = "After matching",
  targetLabel = "Target",
  comparatorLabel = "Comparator"
)
```

#### **Arguments**

balance A data frame created by the computeCovariateBalance function.

threshold A threshold value for standardized difference. When exceeding the threshold,

covariates will be marked in a different color. If threshold = 0, no color coding

will be used.

title The main title for the plot.

fileName Name of the file where the plot should be saved, for example 'plot.png'. See the

function ggsave in the ggplot2 package for supported file formats.

beforeLabel Label for the before matching / stratification panel.

Label for the after matching / stratification panel.

targetLabel Label for the x-axis.

comparatorLabel

Label for the y-axis.

#### Value

A ggplot object. Use the ggplot2::ggsave function to save to file in a different format.

```
plotFollowUpDistribution
```

Plot the distribution of follow-up time

# **Description**

Plot the distribution of follow-up time

#### Usage

```
plotFollowUpDistribution(
  population,
  targetLabel = "Target",
  comparatorLabel = "Comparator",
  yScale = "percent",
  logYScale = FALSE,
  dataCutoff = 0.95,
  title = NULL,
  fileName = NULL
)
```

# **Arguments**

population A data frame describing the study population as created using the createStudyPopulation

function. This should at least have these columns: treatment, timeAtRisk.

targetLabel A label to us for the target cohort.

comparatorLabel

A label to us for the comparator cohort.

yScale Should be either 'percent' or 'count'.
logYScale Should the Y axis be on the log scale?

dataCutoff Fraction of the data (number censored) after which the graph will not be shown.

title The main title of the plot.

fileName Name of the file where the plot should be saved, for example 'plot.png'. See the

function ggsave in the ggplot2 package for supported file formats.

## **Details**

Plot the distribution of follow-up time, stratified by treatment group. Follow-up time is defined as time-at-risk, so not censored at the outcome.

#### Value

A ggplot object. Use the ggsave function to save to file in a different format.

54 plotKaplanMeier

plotKaplanMeier

Plot the Kaplan-Meier curve

# **Description**

plotKaplanMeier creates the Kaplan-Meier (KM) survival plot. Based (partially) on recommendations in Pocock et al (2002).

When variable-sized strata are detected, an adjusted KM plot is computed to account for stratified data, as described in Galimberti eta al (2002), using the closed form variance estimator described in Xie et al (2005).

# Usage

```
plotKaplanMeier(
   population,
   censorMarks = FALSE,
   confidenceIntervals = TRUE,
   includeZero = FALSE,
   dataTable = TRUE,
   dataCutoff = 0.9,
   targetLabel = "Treated",
   comparatorLabel = "Comparator",
   title = NULL,
   fileName = NULL
)
```

# Arguments

population A population object generated by createStudyPopulation, potentially filtered

by other functions.

censorMarks Whether or not to include censor marks in the plot.

confidenceIntervals

Plot 95 percent confidence intervals? Default is TRUE, as recommended by

Pocock et al.

includeZero Should the y axis include zero, or only go down to the lowest observed survival?

The default is FALSE, as recommended by Pocock et al.

dataTable Should the numbers at risk be shown in a table? Default is TRUE, as recom-

mended by Pocock et al.

dataCutoff Fraction of the data (number censored) after which the graph will not be shown.

The default is 90 percent as recommended by Pocock et al.

targetLabel A label to us for the target cohort.

comparatorLabel

A label to us for the comparator cohort.

title The main title of the plot.

fileName Name of the file where the plot should be saved, for example 'plot.png'. See the

function ggsave in the ggplot2 package for supported file formats.

plotPs 55

#### Value

A ggplot object. Use the ggsave function to save to file in a different format.

#### References

Pocock SJ, Clayton TC, Altman DG. (2002) Survival plots of time-to-event outcomes in clinical trials: good practice and pitfalls, Lancet, 359:1686-89.

Galimberti S, Sasieni P, Valsecchi MG (2002) A weighted Kaplan-Meier estimator for matched data with application to the comparison of chemotherapy and bone-marrow transplant in leukaemia. Statistics in Medicine, 21(24):3847-64.

Xie J, Liu C. (2005) Adjusted Kaplan-Meier estimator and log-rank test with inverse probability of treatment weighting for survival data. Statistics in Medicine, 26(10):2276.

plotPs

Plot the propensity score distribution

# **Description**

Plots the propensity (or preference) score distribution.

# Usage

```
plotPs(
  data,
  unfilteredData = NULL,
  scale = "preference",
  type = "density",
  binWidth = 0.05,
  targetLabel = "Target",
  comparatorLabel = "Comparator",
  showCountsLabel = FALSE,
  showAucLabel = FALSE,
  showEquiposeLabel = FALSE,
  equipoiseBounds = c(0.3, 0.7),
  unitOfAnalysis = "subjects",
  title = NULL,
  fileName = NULL
)
```

## **Arguments**

data

A data frame with at least the two columns described below

unfilteredData To be used when computing preference scores on data from which subjects have already been removed, e.g. through trimming and/or matching. This data frame should have the same structure as data.

scale

The scale of the graph. Two scales are supported: scale = 'propensity' or scale = 'preference'. The preference score scale is defined by Walker et al (2013).

56 plotPs

type = 'histogramCount', or type = 'histogramProportion'. 'histogram'

defaults to 'histogramCount'.

binWidth For histograms, the width of the bins

targetLabel A label to us for the target cohort.

comparatorLabel

A label to us for the comparator cohort.

showCountsLabel

Show subject counts?

showAucLabel Show the AUC?

showEquiposeLabel

Show the percentage of the population in equipoise?

equipoiseBounds

The bounds on the preference score to determine whether a subject is in equipoise.

unitOfAnalysis The unit of analysis in the input data. Defaults to 'subjects'.

title Optional: the main title for the plot.

fileName Name of the file where the plot should be saved, for example 'plot.png'. See the

function ggplot2::ggsave() for supported file formats.

#### **Details**

The data frame should have a least the following two columns:

- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group
- propensityScore (numeric): Propensity score

#### Value

A ggplot object. Use the ggplot2::ggsave() function to save to file in a different format.

# References

Walker AM, Patrick AR, Lauer MS, Hornbrook MC, Marin MG, Platt R, Roger VL, Stang P, and Schneeweiss S. (2013) A tool for assessing the feasibility of comparative effectiveness research, Comparative Effective Research, 3, 11-20

#### **Examples**

```
treatment <- rep(0:1, each = 100)
propensityScore <- c(rnorm(100, mean = 0.4, sd = 0.25), rnorm(100, mean = 0.6, sd = 0.25))
data <- data.frame(treatment = treatment, propensityScore = propensityScore)
data <- data[data$propensityScore > 0 & data$propensityScore < 1, ]
plotPs(data)</pre>
```

plotTimeToEvent 57

plotTimeToEvent

Plot time-to-event

#### **Description**

Plot time-to-event

# Usage

```
plotTimeToEvent(
  cohortMethodData,
  population = NULL,
  outcomeId = NULL,
  firstExposureOnly = FALSE,
  restrictToCommonPeriod = FALSE,
  washoutPeriod = 0,
  removeDuplicateSubjects = "keep all",
  minDaysAtRisk = 1,
  riskWindowStart = 0,
  startAnchor = "cohort start",
  riskWindowEnd = 0,
  endAnchor = "cohort end",
  censorAtNewRiskWindow = FALSE,
  periodLength = 7,
  numberOfPeriods = 52,
  highlightExposedEvents = TRUE,
  includePostIndexTime = TRUE,
  showFittedLines = TRUE,
  targetLabel = "Target",
  comparatorLabel = "Comparator",
  title = NULL,
  fileName = NULL
)
```

#### **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 co-

horts in the cohortMethodData object.

outcomeId The ID of the outcome. If NULL, no outcome-specific transformations will be

performed.

firstExposureOnly

(logical) Should only the first exposure per subject be included?

restrict To Common Period

(logical) Restrict the analysis to the period when both exposures are observed?

washoutPeriod The minimum required continuous observation time prior to index date for a

person to be included in the cohort.

58 plotTimeToEvent

removeDuplicateSubjects

Remove subjects that are in both the target and comparator cohort? See details

for allowed values.

minDaysAtRisk The minimum required number of days at risk.

riskWindowStart

The start of the risk window (in days) relative to the startAnchor.

startAnchor The anchor point for the start of the risk window. Can be "cohort start" or

"cohort end".

riskWindowEnd The end of the risk window (in days) relative to the endAnchor.

endAnchor The anchor point for the end of the risk window. Can be "cohort start" or

"cohort end".

censorAtNewRiskWindow

If a subject is in multiple cohorts, should time-at-risk be censored when the new

time-at-risk starts to prevent overlap?

periodLength The length in days of each period shown in the plot.

numberOfPeriods

Number of periods to show in the plot. The periods are equally divided before

and after the index date.

highlightExposedEvents

(logical) Highlight event counts during exposure in a different color?

includePostIndexTime

(logical) Show time after the index date?

showFittedLines

(logical) Fit lines to the proportions and show them in the plot?

targetLabel A label to us for the target cohort.

comparatorLabel

A label to us for the comparator cohort.

title Optional: the main title for the plot.

fileName Name of the file where the plot should be saved, for example 'plot.png'. See

ggplot2::ggsave() for supported file formats.

#### **Details**

Creates a plot showing the number of events over time in the target and comparator cohorts, both before and after index date. The plot also distinguishes between events inside and outside the time-at-risk period. This requires the user to (re)specify the time-at-risk using the same arguments as the createStudyPopulation() function. Note that it is not possible to specify that people with the outcome prior should be removed, since the plot will show these prior events.

#### Value

A ggplot object. Use the ggplot2::ggsave() function to save to file in a different format.

runCmAnalyses 59

runCmAnalyses

Run a list of analyses

#### **Description**

Run a list of analyses

## Usage

```
runCmAnalyses(
  connectionDetails,
  cdmDatabaseSchema,
  tempEmulationSchema = getOption("sqlRenderTempEmulationSchema"),
  exposureDatabaseSchema = cdmDatabaseSchema,
  exposureTable = "drug_era",
  outcomeDatabaseSchema = cdmDatabaseSchema,
  outcomeTable = "condition_occurrence",
  cdmVersion = "5"
  outputFolder = "./CohortMethodOutput",
  cmAnalysisList,
  targetComparatorOutcomesList,
  analysesToExclude = NULL,
  refitPsForEveryOutcome = FALSE,
  refitPsForEveryStudyPopulation = TRUE,
 multiThreadingSettings = createMultiThreadingSettings()
)
```

# **Arguments**

connectionDetails

An R object of type connectionDetails created using the DatabaseConnector::createConnection function.

cdmDatabaseSchema

The name of the database schema that contains the OMOP CDM instance. Requires read permissions to this database. On SQL Server, this should specify both the database and the schema, so for example 'cdm\_instance.dbo'.

tempEmulationSchema

Some database platforms like Oracle and Impala do not truly support temp tables. To emulate temp tables, provide a schema with write privileges where temp tables can be created.

exposureDatabaseSchema

The name of the database schema that is the location where the exposure data used to define the exposure cohorts is available. If exposureTable = DRUG\_ERA, exposureDatabaseSchema is not used by assumed to be cdmSchema. Requires read permissions to this database.

exposureTable

The tablename that contains the exposure cohorts. If exposureTable <> DRUG\_ERA, then expectation is exposureTable has format of COHORT table: COHORT\_DEFINITION\_ID, SUBJECT\_ID, COHORT\_START\_DATE, COHORT\_END\_DATE.

60 runCmAnalyses

outcomeDatabaseSchema

The name of the database schema that is the location where the data used to define the outcome cohorts is available. If exposureTable = CONDITION\_ERA, exposureDatabaseSchema is not used by assumed to be cdmSchema. Requires

read permissions to this database.

outcomeTable The tablename that contains the outcome cohorts. If outcomeTable <> CONDI-

TION\_OCCURRENCE, then expectation is outcomeTable has format of COHORT table: COHORT\_DEFINITION\_ID, SUBJECT\_ID, COHORT\_START\_DATE,

COHORT END DATE.

cdmVersion Define the OMOP CDM version used: currently support "4" and "5".

outputFolder Name of the folder where all the outputs will written to.

 $\verb|cmAnalysisList| A list of objects of type \verb|cmAnalysis| as created using the `createCmAnalysis| \\$ 

function.

targetComparatorOutcomesList

A list of objects of type targetComparatorOutcomes as created using the createTargetComparatorOutcomes function.

analysesToExclude

Analyses to exclude. See the Analyses to Exclude section for details.

refitPsForEveryOutcome

Should the propensity model be fitted for every outcome (i.e. after people who already had the outcome are removed)? If false, a single propensity model will be fitted, and people who had the outcome previously will be removed afterwards.

refitPsForEveryStudyPopulation

Should the propensity model be fitted for every study population definition? If false, a single propensity model will be fitted, and the study population criteria will be applied afterwards.

multiThreadingSettings

An object of type CmMultiThreadingSettings as created using the createMultiThreadingSettin or createDefaultMultiThreadingSettings() functions.

#### **Details**

Run a list of analyses for the target-comparator-outcomes of interest. This function will run all specified analyses against all hypotheses of interest, meaning that the total number of outcome models is length(cmAnalysisList) \* length(targetComparatorOutcomesList) (if all analyses specify an outcome model should be fitted). When you provide several analyses it will determine whether any of the analyses have anything in common, and will take advantage of this fact. For example, if we specify several analyses that only differ in the way the outcome model is fitted, then this function will extract the data and fit the propensity model only once, and re-use this in all the analysis.

After completion, a tibble containing references to all generated files can be obtained using the getFileReference() function. A summary of the analysis results can be obtained using the getResultsSummary() function.

## Analyses to Exclude:

Normally, runCmAnalyses will run all combinations of target-comparator-outcome-analyses settings. However, sometimes we may not need all those combinations. Using the analysesToExclude argument, we can remove certain items from the full matrix. This argument should be a data frame with at least one of the following columns:

• targetId

saveCmAnalysisList 61

- · comparatorId
- · outcomeId
- · analysisId

This data frame will be joined to the outcome model reference table before executing, and matching rows will be removed. For example, if one specifies only one target ID and analysis ID, then any analyses with that target and that analysis ID will be skipped.

#### Value

A tibble describing for each target-comparator-outcome-analysisId combination where the intermediary and outcome model files can be found, relative to the outputFolder.

save Cm Analysis List

Save a list of cmAnalysis to file

#### **Description**

Write a list of objects of type cmAnalysis to file. The file is in JSON format.

# Usage

```
saveCmAnalysisList(cmAnalysisList, file)
```

# **Arguments**

cmAnalysisList The cmAnalysis list to be written to file

file The name of the file where the results will be written

saveCohortMethodData Save the cohort method data to file

# **Description**

Saves an object of type CohortMethodData to a file.

# Usage

 $save {\tt CohortMethodData}, \ {\tt file})$ 

# Arguments

cohortMethodData

An object of type CohortMethodData as generated using getDbCohortMethodData().

file

The name of the file where the data will be written. If the file already exists it will be overwritten.

## Value

Returns no output.

saveTargetComparatorOutcomesList

Save a list of targetComparatorOutcomes to file

# **Description**

Write a list of objects of type targetComparatorOutcomes to file. The file is in JSON format.

#### Usage

saveTargetComparatorOutcomesList(targetComparatorOutcomesList, file)

#### Arguments

targetComparatorOutcomesList

The targetComparatorOutcomes list to be written to file

The name of the file where the results will be written

simulateCohortMethodData

Generate simulated data

#### **Description**

Creates a CohortMethodData object with simulated data.

# Usage

simulateCohortMethodData(profile, n = 10000)

# Arguments

profile An object of type CohortMethodDataSimulationProfile as generated using

 $the \ create Cohort Method Data Simulation Profile () \ function.$ 

n The size of the population to be generated.

# **Details**

This function generates simulated data that is in many ways similar to the original data on which the simulation profile is based. The contains same outcome, comparator, and outcome concept IDs, and the covariates and their 1st order statistics should be comparable.

#### Value

An object of type CohortMethodData.

stratifyByPs 63

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Stratify persons by propensity score

# **Description**

Use the provided propensity scores to stratify persons. Additional stratification variables for stratifications can also be used.

# Usage

```
stratifyByPs(
 population,
 numberOfStrata = 5,
  stratificationColumns = c(),
 baseSelection = "all"
)
```

# **Arguments**

population

A data frame with the three columns described below

numberOfStrata How many strata? The boundaries of the strata are automatically defined to contain equal numbers of target persons.

stratificationColumns

Names of one or more columns in the data data.frame on which subjects should also be stratified in addition to stratification on propensity score.

baseSelection

What is the base selection of subjects where the strata bounds are to be determined? Strata are defined as equally-sized strata inside this selection. Possible values are "all", "target", and "comparator".

# **Details**

The data frame should have the following three columns:

- rowId (numeric): A unique identifier for each row (e.g. the person ID).
- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group.
- propensityScore (numeric): Propensity score.

# Value

Returns a tibble with the same columns as the input data plus one extra column: stratumId.

# **Examples**

```
rowId <- 1:200
treatment \leftarrow rep(0:1, each = 100)
propensityScore <- c(runif(100, min = 0, max = 1), runif(100, min = 0, max = 1))
data <- data.frame(rowId = rowId, treatment = treatment, propensityScore = propensityScore)</pre>
result <- stratifyByPs(data, 5)</pre>
```

```
stratifyByPsAndCovariates
```

Stratify persons by propensity score and other covariates

# **Description**

Use the provided propensity scores and covariates to stratify persons.

# Usage

```
stratifyByPsAndCovariates(
  population,
  numberOfStrata = 5,
  baseSelection = "all",
  cohortMethodData,
  covariateIds
)
```

# Arguments

population A data frame with the three columns described below

numberOfStrata Into how many strata should the propensity score be divided? The boundaries of

the strata are automatically defined to contain equal numbers of target persons.

baseSelection What is the base selection of subjects where the strata bounds are to be deter-

mined? Strata are defined as equally-sized strata inside this selection. Possible

values are "all", "target", and "comparator".

cohortMethodData

An object of type CohortMethodData as generated using getDbCohortMethodData().

covariateIds One or more of

One or more covariate IDs in the cohortMethodData object on which subjects

should also be stratified.

# **Details**

The data frame should have the following three columns:

- rowId (numeric): A unique identifier for each row (e.g. the person ID).
- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group.
- propensityScore (numeric): Propensity score.

#### Value

Returns a date frame with the same columns as the input population plus one extra column: stratumId. trimByIptw 65

trimByIptw

Remove subjects with a high IPTW

# **Description**

Remove subjects having a weight higher than the user-specified threshold.

# Usage

```
trimByIptw(population, maxWeight = 10)
```

# **Arguments**

population A data frame with at least the two columns described in the details

 ${\tt maxWeight} \qquad \qquad {\tt The \ maximum \ allowed \ IPTW}.$ 

## **Details**

The data frame should have the following two columns:

- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group.
- iptw (numeric): Propensity score.

## Value

Returns a tibble with the same columns as the input.

# Examples

```
rowId <- 1:2000
treatment <- rep(0:1, each = 1000)
iptw <- 1 / c(runif(1000, min = 0, max = 1), runif(1000, min = 0, max = 1))
data <- data.frame(rowId = rowId, treatment = treatment, iptw = iptw)
result <- trimByIptw(data)</pre>
```

trimByPs

Trim persons by propensity score

# Description

Use the provided propensity scores to trim subjects with extreme scores.

# Usage

```
trimByPs(population, trimFraction = 0.05)
```

## **Arguments**

population A data frame with the three columns described below

trimFraction This fraction will be removed from each treatment group. In the target group,

persons with the highest propensity scores will be removed, in the comparator

group person with the lowest scores will be removed.

#### **Details**

The data frame should have the following three columns:

- rowId (numeric): A unique identifier for each row (e.g. the person ID).
- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group.
- propensityScore (numeric): Propensity score.

## Value

Returns a tibble with the same three columns as the input.

# **Examples**

```
rowId <- 1:2000
treatment <- rep(0:1, each = 1000)
propensityScore \leftarrow c(runif(1000, min = 0, max = 1), runif(1000, min = 0, max = 1))
data <- data.frame(rowId = rowId, treatment = treatment, propensityScore = propensityScore)</pre>
result <- trimByPs(data, 0.05)</pre>
```

trimByPsToEquipoise

Keep only persons in clinical equipoise

## Description

Use the preference score to trim subjects that are not in clinical equipoise

# Usage

```
trimByPsToEquipoise(population, bounds = c(0.3, 0.7))
```

#### **Arguments**

population

A data frame with at least the three columns described below.

bounds The upper and lower bound on the preference score for keeping persons.

#### **Details**

The data frame should have the following three columns:

- rowId (numeric): A unique identifier for each row (e.g. the person ID).
- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group.
- propensityScore (numeric): Propensity score.

truncateIptw 67

#### Value

Returns a tibble with the same three columns as the input.

#### References

Walker AM, Patrick AR, Lauer MS, Hornbrook MC, Marin MG, Platt R, Roger VL, Stang P, and Schneeweiss S. (2013) A tool for assessing the feasibility of comparative effectiveness research, Comparative Effective Research, 3, 11-20

## **Examples**

```
rowId <- 1:2000
treatment <- rep(0:1, each = 1000)
propensityScore <- c(runif(1000, min = 0, max = 1), runif(1000, min = 0, max = 1))
data <- data.frame(rowId = rowId, treatment = treatment, propensityScore = propensityScore)
result <- trimByPsToEquipoise(data)</pre>
```

truncateIptw

Truncate IPTW values

# **Description**

Set the inverse probability of treatment weights (IPTW) to the user-specified threshold if it exceeds said threshold.

# Usage

```
truncateIptw(population, maxWeight = 10)
```

# **Arguments**

population A data frame with at least the two columns described in the details

maxWeight The maximum allowed IPTW.

# **Details**

The data frame should have the following two columns:

- treatment (integer): Column indicating whether the person is in the target (1) or comparator (0) group.
- iptw (numeric): Propensity score.

### Value

Returns a tibble with the same columns as the input.

68 uploadResults

#### **Examples**

```
rowId <- 1:2000
treatment \leftarrow rep(0:1, each = 1000)
iptw \leftarrow 1 / c(runif(1000, min = 0, max = 1), runif(1000, min = 0, max = 1))
data <- data.frame(rowId = rowId, treatment = treatment, iptw = iptw)</pre>
result <- truncateIptw(data)</pre>
```

uploadResults

Upload results to the database server.

#### **Description**

Requires the results data model tables have been created using the createResultsDataModel function.

### Usage

```
uploadResults(
  connectionDetails,
  schema,
  zipFileName,
  forceOverWriteOfSpecifications = FALSE,
  purgeSiteDataBeforeUploading = TRUE,
  tempFolder = tempdir(),
  tablePrefix = "",
)
```

#### Arguments

connectionDetails

An object of type connectionDetails as created using the createConnectionDetails

function in the DatabaseConnector package.

schema The schema on the server where the tables have been created.

zipFileName The name of the zip file. forceOverWriteOfSpecifications

> If TRUE, specifications of the phenotypes, cohort definitions, and analysis will be overwritten if they already exist on the database. Only use this if these specifications have changed since the last upload.

purgeSiteDataBeforeUploading

If TRUE, before inserting data for a specific databaseId all the data for that site will be dropped. This assumes the input zip file contains the full data for that

data site.

tempFolder A folder on the local file system where the zip files are extracted to. Will be

cleaned up when the function is finished. Can be used to specify a temp folder on a drive that has sufficient space if the default system temp space is too limited.

tablePrefix (Optional) string to insert before table names for database table names

See ResultModelManager::uploadResults

# Index

* datasets	<pre>createOutcome(), 29</pre>	
cohortMethodDataSimulationProfile,	createPs, 24	
5	createPs(), <i>11</i> , <i>42</i>	
	createResultsDataModel, 25, 68	
adjustedKm, 4	<pre>createStratifyByPsAndCovariatesArgs,</pre>	
Andromeda, 5	26	
	createStratifyByPsArgs,27	
checkCmInstallation, 4	createStudyPopulation, $9, 27, 39, 53$	
CohortMethodData, 6, 14, 24, 28, 34, 35, 38,	createStudyPopulation(), 11, 34, 35, 37,	
41, 42, 46, 49, 57, 61, 62, 64	58	
CohortMethodData	createTargetComparatorOutcomes, $29,60$	
(CohortMethodData-class), 5	<pre>createTargetComparatorOutcomes(), 24</pre>	
CohortMethodData-class, 5	createTrimByIptwArgs, 30	
cohortMethodDataSimulationProfile, 5	createTrimByPsArgs, 30	
computeCovariateBalance, 5	createTrimByPsToEquipoiseArgs, 31	
computeCovariateBalance(), 11, 40	<pre>createTruncateIptwArgs, 31</pre>	
computeEquipoise, 8	Cyclops::createControl(), 25, 34	
computeMdrr, 8	Cyclops::createPrior(), 25, 34	
computePsAuc, 9		
CoviarateData, 5	<pre>DatabaseConnector::createConnectionDetails,</pre>	
createCmAnalysis, 10, 60	26	
createCmDiagnosticThresholds, 12	<pre>DatabaseConnector::createConnectionDetails(),</pre>	
createCmDiagnosticThresholds(), 33	36, 44, 59	
createCmTable1, 13, 38	drawAttritionDiagram, 32	
createCohortMethodDataSimulationProfile,		
14	exportToCsv, 32	
${\it createCohortMethodDataSimulationProfile()}, \\ {\it 62}$	exportToCsv(), <i>12</i> , <i>43</i>	
createComputeCovariateBalanceArgs, 14	FeatureExtraction::createCovariateSettings(),	
createConnectionDetails, 68	37	
createCreatePsArgs, 15	FeatureExtraction::getDefaultTable1Specifications()	
createCreateStudyPopulationArgs, 16	6	
<pre>createCreateStudyPopulationArgs(), 24</pre>	fitOutcomeModel, 33	
createDefaultMultiThreadingSettings, 17	<pre>fitOutcomeModel(), 11, 41</pre>	
<pre>createDefaultMultiThreadingSettings(),</pre>	<pre>getAttritionTable, 35</pre>	
23, 60	getDataMigrator, 35	
<pre>createFitOutcomeModelArgs, 18</pre>	getDbCohortMethodData, 36	
createGetDbCohortMethodDataArgs, 19	getDbCohortMethodData(), $5$ , $6$ , $11$ , $14$ , $24$ ,	
$\verb createMatchOnPsAndCovariatesArgs , 20 $	28, 34, 41, 42, 49, 57, 61, 64	
createMatchOnPsArgs, 21	<pre>getDefaultCmTable1Specifications, 38</pre>	
<pre>createMultiThreadingSettings, 22</pre>	getFileReference, 39	
<pre>createMultiThreadingSettings(), 18,60</pre>	getFileReference(), $60$	
createOutcome, 23	getFollowUpDistribution,39	

70 INDEX

getGeneralizabilityTable,40 getInteractionResultsSummary,41 getOutcomeModel,41
getPsModel, 42
getResultsDataModelSpecifications, 42
getResultsSummary, 43
getResultsSummary(), 60
ggplot2::ggsave, 51, 52
ggplot2::ggsave(), 56, 58 ggsave, 32, 53, 55
ggsave, 32, 33, 33
<pre>insertExportedResultsInSqlite, 43</pre>
<pre>insertExportedResultsInSqlite(), 45</pre>
isCohortMethodData,44
launchResultsViewer, 44
launchResultsViewerUsingSqlite,45
loadCmAnalysisList, 45
loadCohortMethodData, 46
loadCohortMethodData(), 5
<pre>loadTargetComparatorOutcomesList, 46</pre>
matchOnPs, 47
matchOnPs(), <i>11</i>
matchOnPsAndCovariates,48
matchOnPsAndCovariates(), 11
migrateDataModel, 50
plotCovariateBalanceOfTopVariables, 50
plotCovariateBalanceScatterPlot, 51
plotCovariatePrevalence, 52
plotFollowUpDistribution, 53
plotKaplanMeier,54
plotPs, 55
plotTimeToEvent, 57
runCmAnalyses, 59
runCmAnalyses(), <i>12</i> , <i>30</i> , <i>33</i>
saveCmAnalysisList, 61
saveCohortMethodData, 61
${\sf saveCohortMethodData()}, {\sf 5}$
<pre>saveTargetComparatorOutcomesList, 62</pre>
show,CohortMethodData-method
(CohortMethodData-class), $5$
simulateCohortMethodData,62
simulateCohortMethodData(), 14
stratifyByPs, 63
stratifyByPs(), 11
stratifyByPsAndCovariates, 64
stratifyByPsAndCovariates(), 11
summary, CohortMethodData-method
(CohortMethodData-class), 5

trimByIptw, 65
trimByIptw(), 11
trimByPs, 65
trimByPs(), 11
trimByPsToEquipoise, 66
trimByPsToEquipoise(), 11
truncateIptw, 67
truncateIptw(), 11
uploadResults, 68
uploadResults(), 44