# Package 'MethodEvaluation'

December 13, 2018

```
Title Package for evaluation of estimation methods
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Description This package contains resources for the evaluation of the
      performance of methods that aim to estimate the magnitude (relative risk) of
      the effect of a drug on an outcome. These resources include reference sets for
      evaluating methods on real data, as well as functions for inserting simulated
      effects in real data based on negative control drug-outcome pairs. Further
      included are functions for the computation of the minimum detectable relative
      risks and functions for computing performance statistics such as predictive
      accuracy, error and bias.
License Apache License 2.0
VignetteBuilder knitr
URL https://github.com/OHDSI/MethodEvaluation
BugReports https://github.com/OHDSI/MethodEvaluation/issues
Depends R (>= 3.2.0),
      DatabaseConnector (>= 2.0.0),
      FeatureExtraction (>= 2.0.0),
      Cyclops (>= 1.2.2)
Imports ff,
      ffbase (>= 0.12.1),
      SqlRender (\geq 1.5.0),
      pROC,
      ggplot2,
      ParallelLogger,
      methods,
      EmpiricalCalibration
Suggests testthat,
      DT,
      shiny,
RoxygenNote 6.1.1
```

Type Package

2 computeAucs

# R topics documented:

	computeAucs	2
	computeCoverage	3
	computeMdrr	3
	computeMetrics	5
	computeMse	5
	computeOhdsiBenchmarkMetrics	6
	computeType1And2Error	6
	createReferenceSetCohorts	7
	euadrReferenceSet	8
	filterOnMdrr	9
	injectSignals	9
	launchMethodEvaluationApp	12
	MethodEvaluation	13
	ohdsiNegativeControls	13
	omopReferenceSet	14
	packageOhdsiBenchmarkResults	15
	plotCoverageInjectedSignals	16
	plotRocsInjectedSignals	16
	synthesizePositiveControls	17
Index		19

computeAucs

Compute the AUCs for various injected signal sizes

# Description

Compute the AUCs for various injected signal sizes

# Usage

```
computeAucs(logRr, trueLogRr)
```

# **Arguments**

logRr A vector containing the log of the relative risk as estimated by a method.

trueLogRr A vector containing the injected log(relative risk) for each estimate.

# Value

A data frame with per injected signal size the AUC and the 95 percent confidence interval of the AUC.

computeCoverage 3

computeCoverage	Compute the coverage

# **Description**

Compute the coverage

### Usage

```
computeCoverage(logRr, seLogRr, trueLogRr, region = 0.95)
```

# **Arguments**

logRr A numeric vector of effect estimates on the log scale.

seLogRr The standard error of the log of the effect estimates. Hint: often the standard

error = (log(<lower bound 95 percent confidence interval>) - log(<effect esti-

mate > ))/qnorm(0.025).

trueLogRr A vector of the true effect sizes.

region Size of the confidence interval. Default is .95 (95 percent).

# **Details**

Compute the fractions of estimates where the true effect size is below, above or within the confidence interval, for one or more true effect sizes.

computeMdrr	Compute minimal detectable relative risk (MDRR)	

# **Description**

computeMdrr computes the minimal detectable relative risk (MDRR) for drug-outcome pairs.

# Usage

```
computeMdrr(connectionDetails, cdmDatabaseSchema,
  oracleTempSchema = cdmDatabaseSchema, exposureOutcomePairs,
  exposureDatabaseSchema = cdmDatabaseSchema,
  exposureTable = "drug_era",
  outcomeDatabaseSchema = cdmDatabaseSchema,
  outcomeTable = "condition_era", cdmVersion = "5")
```

4 computeMdrr

#### **Arguments**

connectionDetails

An R object of type ConnectionDetails created using the function createConnectionDetails in the DatabaseConnector package.

cdmDatabaseSchema

Name of database schema that contains OMOP CDM and vocabulary.

oracleTempSchema

For Oracle only: the name of the database schema where you want all temporary tables to be managed. Requires create/insert permissions to this database.

exposureOutcomePairs

A data frame with at least two columns:

- "exposureId" or "targetId" containing the drug\_concept\_ID or cohort\_definition\_id of the exposure variable
- "outcomeId" containing the condition\_concept\_ID or cohort\_definition\_id of the outcome variable

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.

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 <> CONDITION\_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".

### **Details**

Computes the MDRR using simple power-calculations using person-level statistics stratified by age and gender.

# Value

A data frame containing the MDRRs for the given exposure-outcome pairs.

# **Examples**

computeMetrics 5

```
"cdm_truven_mdcr",
exposureOutcomePairs,
outcomeTable = "condition_era")
```

## End(Not run)

computeMetrics

Compute the AUC, coverage, MSE, and type 1 and 2 error

# **Description**

Compute the AUC, coverage, MSE, and type 1 and 2 error

# Usage

```
computeMetrics(logRr, seLogRr, trueLogRr)
```

# **Arguments**

logRr A numeric vector of effect estimates on the log scale

seLogRr The standard error of the log of the effect estimates. Hint: often the standard

error = (log(<lower bound 95 percent confidence interval>) - log(<effect esti-

mate>))/qnorm(0.025)

trueLogRr A vector of the true effect sizes

### **Details**

Compute the AUC, coverage, MSE, and type 1 and 2 error.

computeMse (	Compute the	mean so	auared	error
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# **Description**

Compute the mean squared error

### Usage

```
computeMse(logRr, trueLogRr)
```

# **Arguments**

logRr A numeric vector of effect estimates on the log scale.

trueLogRr A vector of the true effect sizes.

 ${\tt computeOhdsiBenchmarkMetrics}$ 

Generate perfomance metrics for the OHDSI Methods Benchmark

# Description

Generate perfomance metrics for the OHDSI Methods Benchmark

# Usage

```
computeOhdsiBenchmarkMetrics(exportFolder, mdrr = 1.25,
   stratum = "All", trueEffectSize = "Overall", calibrated = FALSE,
   comparative = FALSE)
```

# **Arguments**

exportFolder	The folder containing the CSV files created using the packageOhdsiBenchmarkResults function. This folder can contain results from various methods, analyses, and databases.
mdrr	The minimum detectable relative risk (MDRR). Only controls with this MDRR will be used to compute the performance metrics. Set to "All" to include all controls.
stratum	The stratum for which to compute the metrics, e.g. 'Acute Pancreatitis'. Set to 'All' to use all controls.
trueEffectSize	Should the analysis be limited to a specific true effect size? Set to "Overall" to include all.
calibrated	Should confidence intervals and p-values be empirically calibrated before computing the metrics?
comparative	Should the methods be evaluated on the task of comprative effect estimation? If FALSE, they will be evaluated on the task of effect estimation.

# Value

A data frame with the various metrics per method - analysisId - database combination.

```
computeType1And2Error Compute type 1 and 2 error
```

# Description

Compute type 1 and 2 error

# Usage

```
computeType1And2Error(logRr, seLogRr, trueLogRr, alpha = 0.05)
```

createReferenceSetCohorts 7

### **Arguments**

logRr A numeric vector of effect estimates on the log scale.

seLogRr The standard error of the log of the effect estimates. Hint: often the standard

error = (log(<lower bound 95 percent confidence interval>) - log(<effect esti-

mate>))/qnorm(0.025).

trueLogRr A vector of the true effect sizes.

alpha The alpha (expected type I error).

createReferenceSetCohorts

Create cohorts used in a reference set.

# **Description**

Create cohorts used in a reference set.

# Usage

```
createReferenceSetCohorts(connectionDetails, oracleTempSchema = NULL,
  cdmDatabaseSchema, outcomeDatabaseSchema = cdmDatabaseSchema,
  outcomeTable = "outcomes", nestingDatabaseSchema = cdmDatabaseSchema,
  nestingTable = "nesting", referenceSet = "ohdsiMethodsBenchmark")
```

### **Arguments**

connectionDetails

An R object of type ConnectionDetails created using the function createConnectionDetails in the DatabaseConnector package.

oracleTempSchema

Should be used in Oracle to specify a schema where the user has write priviliges for storing temporary tables.

cdmDatabaseSchema

A database schema containing health care data in the OMOP Commond Data Model. Note that for SQL Server, botth the database and schema should be specified, e.g. 'cdm\_schema.dbo'

outcomeDatabaseSchema

The database schema where the target outcome table is located. Note that for SQL Server, both the database and schema should be specified, e.g. 'cdm\_schema.dbo'

 $\begin{tabular}{ll} \textbf{outcomeTable} & \textbf{The name of the table where the outcomes will be stored.} \\ \textbf{nestingDatabaseSchema} \end{tabular}$ 

(For the OHDSI Methods Benchmark only) The database schema where the nesting outcome table is located. Note that for SQL Server, both the database and schema should be specified, e.g. 'cdm\_schema.dbo'.

nestingTable (For the OHDSI Methods Benchmark only) The name of the table where the nesting cohorts will be stored.

referenceSet The name of the reference set for which outcomes need to be created. Currently

supported are "omopReferenceSet", "euadrReferenceSet", and "ohdsiMethods-

Benchmark".

8 euadrReferenceSet

#### **Details**

This function will create the outcomes of interest and nesting cohorts referenced in the various reference sets. The outcomes of interest are derives using information like diagnoses, procedures, and drug prescriptions. The outcomes are stored in a table on the database server.

euadrReferenceSet

The EU-ADR reference set

# **Description**

A reference set of 43 drug-outcome pairs where we believe the drug causes the outcome (positive controls) and 50 drug-outcome pairs where we believe the drug does not cause the outcome (negative controls). The controls involve 10 health outcomes of interest. Note that originally, there was an additional positive control (Nimesulide and acute liver injury), but Nimesulide is not in RxNorm, and is not available in many countries.

# Usage

data(euadrReferenceSet)

#### **Format**

A data frame with 399 rows and 10 variables:

exposureId Concept ID identifying the exposure

exposureName Name of the exposure

outcomeId Concept ID identifying the outcome

outcomeName Name of the outcome

**groundTruth** 0 = negative control, 1 = positive control

**indicationId** Concept Id identifying the (primary) indication of the drug. To be used when one wants to nest the analysis within the indication

indicationName Name of the indication

comparatorId Concept ID identifying a comparator drug that can be used as a counterfactual

comparatorName Name of the comparator drug

comparatorType How the comparator was selected

#### References

Coloma PM, Avillach P, Salvo F, Schuemie MJ, Ferrajolo C, Pariente A, Fourrier-Reglat A, Molokhia M, Patadia V, van der Lei J, Sturkenboom M, Trifiro G. A reference standard for evaluation of methods for drug safety signal detection using electronic healthcare record databases. Drug Safety 36(1):13-23, 2013

filterOnMdrr 9

filterOnMdrr	Filter data based on MDRR

# **Description**

Filters a dataset to those exposure-outcome pairs with sufficient power.

# Usage

```
filterOnMdrr(data, mdrr, threshold = 1.25)
```

### **Arguments**

data A data frame with at least two columns:

 "exposureConceptId" containing the drug\_concept\_ID or cohort\_definition\_id of the exposure variable

• "outcomeConceptId" containing the condition\_concept\_ID or cohort\_definition\_id of the outcome variable

mdrr A data frame as generated by the computeMdrr function.

threshold The required minimum detectable relative risk.

### Value

A subset of the data object.

injectSignals Inject signals in database

# Description

Inject signals in database

# Usage

```
injectSignals(connectionDetails, cdmDatabaseSchema,
  oracleTempSchema = cdmDatabaseSchema,
  exposureDatabaseSchema = cdmDatabaseSchema,
  exposureTable = "drug_era",
  outcomeDatabaseSchema = cdmDatabaseSchema, outcomeTable = "cohort",
  outputDatabaseSchema = outcomeDatabaseSchema,
  outputTable = outcomeTable, createOutputTable = FALSE,
  exposureOutcomePairs, modelType = "poisson",
  minOutcomeCountForModel = 100, minOutcomeCountForInjection = 25,
  covariateSettings = FeatureExtraction::createCovariateSettings(useDemographicsAgeGroup
  = TRUE, useDemographicsGender = TRUE, useDemographicsIndexYear = TRUE,
  useDemographicsIndexMonth = TRUE, useConditionGroupEraLongTerm = TRUE,
  useDrugGroupEraLongTerm = TRUE, useProcedureOccurrenceLongTerm = TRUE,
  useMeasurementLongTerm = TRUE, useObservationLongTerm = TRUE,
  useCharlsonIndex = TRUE, useDcsi = TRUE, useChads2Vasc = TRUE,
```

10 injectSignals

```
longTermStartDays = 365, endDays = 0), prior = createPrior("laplace",
exclude = 0, useCrossValidation = TRUE), control = createControl(cvType
= "auto", startingVariance = 0.1, noiseLevel = "quiet", threads = 10),
firstExposureOnly = FALSE, washoutPeriod = 183,
riskWindowStart = 0, riskWindowEnd = 0,
addExposureDaysToEnd = TRUE, addIntentToTreat = FALSE,
firstOutcomeOnly = FALSE, removePeopleWithPriorOutcomes = FALSE,
maxSubjectsForModel = 1e+05, effectSizes = c(1, 1.25, 1.5, 2, 4),
precision = 0.01, outputIdOffset = 1000,
workFolder = "./SignalInjectionTemp", cdmVersion = "5",
modelThreads = 1, generationThreads = 1)
```

#### **Arguments**

connectionDetails

An R object of type ConnectionDetails created using the function createConnectionDetails in the DatabaseConnector package.

cdmDatabaseSchema

Name of database schema that contains OMOP CDM and vocabulary.

oracleTempSchema

For Oracle only: the name of the database schema where you want all temporary tables to be managed. Requires create/insert permissions to this database.

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 table name that contains the exposure cohorts. If exposureTable <> DRUG\_ERA, then expectation is exposureTable has format of COHORT table: cohort\_concept\_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. If exposureTable = CONDITION\_ERA, exposureDatabaseSchema is not used by assumed to be cdmSchema. Requires read permissions to this database.

outcomeTable

The table name that contains the outcome cohorts. When the table name is not CONDITION\_ERA This table is expected to have the same format as the CO-HORT table: SUBJECT\_ID, COHORT\_START\_DATE, COHORT\_END\_DATE, COHORT\_CONCEPT\_ID (CDM v4) or COHORT\_DEFINITION\_ID (CDM v5 and higher).

outputDatabaseSchema

The name of the database schema that is the location of the tables containing the new outcomesRequires write permissions to this database.

outputTable The name of the table names that will contain the generated outcome cohorts. createOutputTable

Should the output table be created prior to inserting the outcomes? If TRUE and the tables already exists, it will first be deleted. If FALSE, the table is assumed to exist and the outcomes will be inserted. Any existing outcomes with the same IDs will first be deleted.

exposureOutcomePairs

A data frame with at least two columns:

injectSignals 11

 "exposureId" containing the drug\_concept\_ID or cohort\_concept\_id of the exposure variable

 "outcomeId" containing the condition\_concept\_ID or cohort\_concept\_id of the outcome variable

modelType Can be either "poisson" or "survival"

minOutcomeCountForModel

Minimum number of outcome events required to build a model.

 $\verb|minOutcomeCountForInjection| \\$ 

Minimum number of outcome events required to inject a signal.

covariateSettings

An object of type covariateSettings as created using the createCovariateSettings

function in the FeatureExtraction package.

prior The prior used to fit the outcome model. See createPrior for details.

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

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

firstExposureOnly

Should signals be injected only for the first exposure? (ie. assuming an acute

effect)

washoutPeriod Number of days at the start of observation for which no signals will be injected,

but will be used to determine whether exposure or outcome is the first one, and

for extracting covariates to build the outcome model.

riskWindowStart

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

0, risk is assumed to start on the first day of exposure.

riskWindowEnd The end of the risk window relative to the start of the exposure. Note that typi-

cally the length of exposure is added to this number (when the addExposureDaysToEnd

parameter is set to TRUE).

addExposureDaysToEnd

Should length of exposure be added to the risk window?

addIntentToTreat

If true, the signal will not only be injected in the primary time at risk, but also after the time at risk (up until the obseration period end). In both time periods, the target effect size will be enforced. This allows the same positive control synthesis to be used in both on treatment and intent-to-treat analysis variants. However, this will preclude the controls to be used in self-controlled designs that consider the time after exposure. Requires firstExposureOnly = TRUE.

firstOutcomeOnly

Should only the first outcome per person be considered when modeling the outcome?

removePeopleWithPriorOutcomes

Remove people with prior outcomes?

maxSubjectsForModel

Maximum number of people used to fit an outcome model.

effectSizes A numeric vector of effect sizes that should be inserted.

precision The allowed ratio between target and injected signal size.

outputIdOffset What should be the first new outcome ID that is to be created?

workFolder Path to a folder where intermediate data will be stored.

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

modelThreads Number of parallel threads to use when fitting outcome models.

generationThreads

Number of parallel threads to use when generating outcomes.

### **Details**

This function will insert additional outcomes for a given set of drug-outcome pairs. It is assumed that these drug-outcome pairs represent negative controls, so the true relative risk before inserting any outcomes should be 1. There are two models for inserting the outcomes during the specified risk window of the drug: a Poisson model assuming multiple outcomes could occurr during a single exposure, and a survival model considering only one outcome per exposure.

It is possible to use bulk import to insert the generated outcomes in the database. This requires the environmental variable 'USE\_MPP\_BULK\_LOAD' to be set to 'TRUE'. See 'PostabaseConnector::insertTable for details on how to configure the bulk upload.

#### Value

A data frame listing all the drug-pairs in combination with requested effect sizes and the real inserted effect size (might be different from the requested effect size because of sampling error).

### References

Schuemie MJ, Hripcsak G, Ryan PB, Madigan D, Suchard MA. Empirical confidence interval calibration for population-level effect estimation studies in observational healthcare data. Proc Natl Acad Sci U S A. 2018 Mar 13;115(11):2571-2577.

launchMethodEvaluationApp

Launch the Method Evaluation Shiny app

### **Description**

Launch the Method Evaluation Shiny app

### Usage

launchMethodEvaluationApp(exportFolder, launch.browser = TRUE)

# **Arguments**

exportFolder A folder where the data files for the Method Evaluation app are stored. Use the

packageOhdsiBenchmarkResults function to populate this folder.

launch.browser Should the app be launched in your default browser, or in a Shiny window. Note:

copying to clipboard will not work in a Shiny window.

# **Details**

Launches a Shiny app that allows the user to explore the results.

MethodEvaluation 13

MethodEvaluation MethodEvaluation

# **Description**

MethodEvaluation

ohdsiNegativeControls The OHDSI Method Evaluation Benchmark - Negative Controls

# **Description**

A set of 200 negative controls, centered around four outcomes of interest (acute pancreatitis, GI bleeding, Stroke, and IBD), and 4 exposures of interest (diclofenac, ciprofloxacin, metformin, and sertraline), which 25 negative controls each. Each drug-outcome pair also includes a comparator drug (where the comparator is also a negative control), allowing for evaluation of comparative effect estimation, and a nesting cohort for evaluating methods such as the nested case-control design.

### Usage

data(ohdsiNegativeControls)

### **Format**

A data frame with 200 rows and 9 variables:

targetId Cohort ID identifying the target exposure

targetName Name of the target cohort

comparatorId Cohort ID identifying the comparator exposure

**comparatorName** Name of the comparator cohort **nestingId** Cohort ID identifying the nesting cohort

nestingName Name of the nesting cohort

outcomeId Cohort ID identifying the outcome

outcomeName Name of the outcome

type THe type of control: exposure or outcome

### **Details**

The exposure, outcome, and nesting cohorts can be created using the createReferenceSetCohorts function.

These negative controls can form the basis to generate positive controls using the injectSignals function.

14 omopReferenceSet

omopReferenceSet

The OMOP reference set A reference set of 165 drug-outcome pairs where we believe the drug causes the outcome (positive controls) and 234 drug-outcome pairs where we believe the drug does not cause the outcome (negative controls). The controls involve 4 health outcomes of interest: acute liver injury, acute kidney injury, acute myocardial infarction, and GI bleeding.

### **Description**

The OMOP reference set A reference set of 165 drug-outcome pairs where we believe the drug causes the outcome (positive controls) and 234 drug-outcome pairs where we believe the drug does not cause the outcome (negative controls). The controls involve 4 health outcomes of interest: acute liver injury, acute kidney injury, acute myocardial infarction, and GI bleeding.

### Usage

data(omopReferenceSet)

#### **Format**

A data frame with 399 rows and 10 variables:

exposureId Concept ID identifying the exposure

exposureName Name of the exposure

outcomeId Concept ID identifying the outcome

outcomeName Name of the outcome

**groundTruth** 0 = negative control, 1 = positive control

**indicationId** Concept Id identifying the (primary) indication of the drug. To be used when one wants to nest the analysis within the indication

indicationName Name of the indication

comparatorId Concept ID identifying a comparator drug that can be used as a counterfactual

comparatorName Name of the comparator drug

comparatorType How the comparator was selected

### References

Ryan PB, Schuemie MJ, Welebob E, Duke J, Valentine S, Hartzema AG. Defining a reference set to support methodological research in drug safety. Drug Safety 36 Suppl 1:S33-47, 2013

packageOhdsiBenchmarkResults

Package results of a method on the OHDSI Methods Benchmark

### **Description**

Stores the results of a method on the OHDSI Methods Benchmark in a standardized format, for example for use in the Method Evaluation Shiny app.

### Usage

packageOhdsiBenchmarkResults(estimates, controlSummary, analysisRef, databaseName, exportFolder)

### **Arguments**

estimates A data frame containing the estimates. See details for required columns.

controlSummary A data frame with the summary of the controls as generated by the synthesizePositiveControls

function.

analysisRef A file describing the various analyses that were performed. See details for re-

quired columns.

databaseName A character string to identify the database the method was executed on.

exportFolder The folder where the output CSV files will written.

### **Details**

The estimates argument should have the following columns: "targetId", "outcomeId", "analysisId", "logRr", "seLogRr", "ci95Lb", "ci95Ub".

The analysisRef argument should have the following columns: "analysisId", "method", "comparative", "nesting", "firstExposureOnly"

The targetId and outcomeId fields identify the specific control, and should correspond to those in the controlSummary object.

The analysisId field is an integer that identifies a specific variant of the method. For example, if the method is 'CohortMethod', analysisId = 1 could identify a set of settings using propensity score matching, and analysisId = 2 could identify a set of settings using stratification.

logRr, seLogRr, ci95Lb, and ci95Ub correspond to the log of the effect estimate (e.g. the hazard ratio), the standard error, and the upper and lower bound of the effect size estimate, as produced by the method.

method is a character string identifyign the method (e.g. "CohortMethod").

comparative is a boolean indicating whether the analysis can also be considerd to perform comparative effect estimation (comparing the target to the comparator).

nesting is a boolean indicating whether the analysis is nested in the nesting cohorts identified in the gold standard.

firstExposureOnly is a boolean indicating whether only the first exposure was used in the analysis.

plotCoverageInjectedSignals

Plot the coverage

### **Description**

Plot the coverage

### Usage

```
plotCoverageInjectedSignals(logRr, seLogRr, trueLogRr, region = 0.95,
    fileName = NULL)
```

### **Arguments**

logRr A numeric vector of effect estimates on the log scale

seLogRr The standard error of the log of the effect estimates. Hint: often the standard

error = (log(<lower bound 95 percent confidence interval>) - log(<effect esti-

mate>))/qnorm(0.025)

trueLogRr A vector of the true effect sizes

region Size of the confidence interval. Default is .95 (95 percent).

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 fractions of estimates where the true effect size is below, above or within the confidence interval, for one or more true effect sizes.

```
plotRocsInjectedSignals
```

Plot the ROC curves for various injected signal sizes

### **Description**

Plot the ROC curves for various injected signal sizes

# Usage

```
plotRocsInjectedSignals(logRr, trueLogRr, showAucs, fileName = NULL)
```

# **Arguments**

logRr A vector containing the log of the relative risk as estimated by a method.
trueLogRr A vector containing the injected log(relative risk) for each estimate.

showAucs Should the AUCs be shown in 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.

#### Value

A Ggplot object. Use the ggsave function to save to file.

synthesizePositiveControls

Synthesize positive controls based on negative controls

### **Description**

Synthesize positive controls based on negative controls

### Usage

```
synthesizePositiveControls(connectionDetails, cdmDatabaseSchema,
  oracleTempSchema = NULL, outcomeDatabaseSchema = cdmDatabaseSchema,
  outcomeTable = "cohort", exposureDatabaseSchema = cdmDatabaseSchema,
  exposureTable = "drug_era", referenceSet = "ohdsiMethodsBenchmark",
  maxCores = 1, workFolder, summaryFileName = file.path(workFolder,
  "allControls.csv"))
```

### **Arguments**

connectionDetails

An R object of type ConnectionDetails created using the function createConnectionDetails in the DatabaseConnector package.

cdmDatabaseSchema

A database schema containing health care data in the OMOP Commond Data Model. Note that for SQL Server, botth the database and schema should be specified, e.g. 'cdm\_schema.dbo'

oracleTempSchema

Should be used in Oracle to specify a schema where the user has write priviliges for storing temporary tables.

outcomeDatabaseSchema

The database schema where the target outcome table is located. Note that for SQL Server, both the database and schema should be specified, e.g. 'cdm\_schema.dbo'

outcomeTable The name of the table where the outcomes will be stored. 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 and assumed to be cdmDatabaseSchema.

Requires read permissions to this database.

then expectation is exposure Table has format of COHORT table: COHORT\_DEFINITION\_ID,

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

referenceSet The name of the reference set for which positive controls need to be synthesized.

Currently supported are "ohdsiMethodsBenchmark".

maxCores How many parallel cores should be used? If more cores are made available this

can speed up the analyses.

workFolder

Name of local folder to place intermediary results; make sure to use forward slashes (/). Do not use a folder on a network drive since this greatly impacts performance.

summaryFileName

The name of the CSV file where to store the summary of the final set of positive and negative controls.

### **Details**

This function will synthesize positive controls for a given reference set based on the real negative controls. Data from the database will be used to fit outcome models for each negative control outcome, and these models will be used to sample additional synthetic outcomes during eposure to increase the true hazard ratio.

The positive control outcome cohorts will be stored in the same database table as the negative control outcome cohorts.

A summary file will be created listing all positive and negative controls. This list should then be used as input for the method under evaluation.

# **Index**

```
*Topic datasets
    euadrReferenceSet, 8
    ohdsiNe gative {\tt Controls}, 13
    omopReferenceSet, 14
computeAucs, 2
computeCoverage, 3
computeMdrr, 3, 9
computeMetrics, 5
computeMse, 5
computeOhdsiBenchmarkMetrics, 6
computeType1And2Error, 6
createControl, 11
createPrior, 11
createReferenceSetCohorts, 7, 13
euadrReferenceSet, 8
filterOnMdrr, 9
injectSignals, 9, 13
launchMethodEvaluationApp, 12
MethodEvaluation, 13
MethodEvaluation-package
        (MethodEvaluation), 13
ohdsiNe gative Controls, \\ 13
omopReferenceSet, 14
packageOhdsiBenchmarkResults, 6, 12, 15
plotCoverageInjectedSignals, 16
plotRocsInjectedSignals, 16
synthesizePositiveControls, 15, 17
```