Package 'PFIM'

October 23, 2024

```
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     Mentré F, Mallet A, Baccar D (1997) <doi:10.1093/biomet/84.2.429>,
     Retout S, Comets E, Samson A, Mentré F (2007) <doi:10.1002/sim.2910>,
     Bazzoli C, Retout S, Mentré F (2009) <doi:10.1002/sim.3573>,
     Le Nagard H, Chao L, Tenaillon O (2011) <doi:10.1186/1471-2148-11-326>,
     Combes FP, Retout S, Frey N, Mentré F (2013) <doi:10.1007/s11095-013-1079-3> and
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     'ModelError.R' 'Combined1.R' 'Constant.R' 'Design.R'
     'Distribution.R' 'ModelParameter.R' 'LibraryOfPDModels.R'
     'LibraryOfPKModels.R' 'LibraryOfModels.R'
     'LibraryOfPKPDModels.R' 'Model.R' 'PFIMProject.R'
     'Evaluation.R' 'OptimizationAlgorithm.R'
     'FedorovWynnAlgorithm.R' 'IndividualFim.R' 'LogNormal.R'
     'ModelODE.R' 'ModelAnalytic.R' 'ModelAnalyticBolus.R'
     'ModelAnalyticSteadyState.R' 'ModelAnalyticBolusSteadyState.R'
     'ModelInfusion.R' 'ModelAnalyticInfusion.R'
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'ModelAnalyticInfusionSteadyState.R' 'ModelBolus.R' 'ModelODEBolus.R' 'ModelODEDoseInEquations.R'
'ModelODEDoseNotInEquations.R' 'ModelODEInfusion.R'
$'Model ODE In fusion Dose In Equations. R'\ 'Multiplicative Algorithm. R'$
'Normal.R' 'Optimization.R' 'PFIM-package.R' 'PGBOAlgorithm.R'
'PSOAlgorithm.R' 'PlotEvaluation.R' 'PopulationFim.R'
'Proportional.R' 'SamplingTimeConstraints.R' 'SamplingTimes.R'
'SimplexAlgorithm.R'
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Description

Evaluate or optimize designs for nonlinear mixed effects models using the Fisher Information matrix. Methods used in the package refer to Mentré F, Mallet A, Baccar D (1997) doi: 10.1093/biomet/84.2.429, Retout S, Comets E, Samson A, Mentré F (2007) doi: 10.1002/sim.2910, Bazzoli C, Retout S, Mentré F (2009) doi: 10.1002/sim.3573, Le Nagard H, Chao L, Tenaillon O (2011) doi: 10.1186/1471214811326, Combes FP, Retout S, Frey N, Mentré F (2013) doi: 10.1007/s11095-01310793 and Seurat J, Tang Y, Mentré F, Nguyen TT (2021) doi: 10.1016/j.cmpb.2021.106126.

Description

Nonlinear mixed effects models (NLMEM) are widely used in model-based drug development and use to analyze longitudinal data. The use of the "population" Fisher Information Matrix (FIM) is a good alternative to clinical trial simulation to optimize the design of these studies. The present version, **PFIM** 6.1, is an R package that uses the S4 object system for evaluating and/or optimizing population designs based on FIM in NLMEMs.

This version of **PFIM** now includes a library of models implemented also using the object oriented system S4 of R. This library contains two libraries of pharmacokinetic (PK) and/or pharmacodynamic (PD) models. The PK library includes model with different administration routes (bolus, infusion, first-order absorption), different number of compartments (from 1 to 3), and different types of eliminations (linear or Michaelis-Menten). The PD model library, contains direct immediate models (e.g. Emax and Imax) with various baseline models, and turnover response models. The PK/PD models are obtained with combination of the models from the PK and PD model libraries. **PFIM** handles both analytical and ODE models and offers the possibility to the user to define his/her own model(s). In **PFIM 6.1**, the FIM is evaluated by first order linearization of the model assuming a block diagonal FIM as in [3]. The Bayesian FIM is also available to give shrinkage predictions [4]. **PFIM 6.1** includes several algorithms to conduct design optimization based on the D-criterion, given design constraints: the simplex algorithm (Nelder-Mead) [5], the multiplicative algorithm [6], the Fedorov-Wynn algorithm [7], PSO (*Particle Swarm Optimization*) and PGBO (*Population Genetics Based Optimizer*) [9].

Documentation

Documentation and user guide are available at http://www.pfim.biostat.fr/

Validation

PFIM 6.1 also provides quality control with tests and validation using the evaluated FIM to assess the validity of the new version and its new features. Finally, **PFIM 6.1** displays all the results with both clear graphical form and a data summary, while ensuring their easy manipulation in R. The standard data visualization package ggplot2 for R is used to display all the results with clear graphical form [10]. A quality control using the D-criterion is also provided.

Organization of the source files in the /R folder

PFIM 6.1 contains a hierarchy of S4 classes with corresponding methods and functions serving as constructors. All of the source code related to the specification of a certain class is contained in a file named [Name_of_the_class]-Class.R. These classes include:

- 1. all roxygen @include to insure the correctly generated collate for the DESCRIPTION file,
- 2. \setClass preceded by a roxygen documentation that describes the purpose and slots of the class,
- 3. specification of an initialize method,
- 4. all getter and setter, respectively returning attributes of the object and associated objects.

Content of the source code and files in the /R folder

Class Administration

- getOutcome
- setOutcome
- getTimeDose
- setTimeDose
- getDose

- setDose
- getTinf
- setTinf
- getTau
- setTau

Class AdministrationConstraints

- getOutcome
- getDose

Class Arm

- getName
- setName
- getSize
- setSize
- getAdministrations
- setAdministrations
- getSamplingTimes
- setSamplingTimes
- getInitialConditions
- setInitialConditions
- getAdministrationsConstraints
- getSamplingTimesConstraints
- getSamplingTime
- getSamplingTimeConstraint
- setSamplingTimesConstraints
- setSamplingTime
- getAdministration
- getAdministrationConstraint
- EvaluateArm

Class BayesianFim

- EvaluateFisherMatrix
- getRSE
- getConditionNumberVarianceEffects
- getShrinkage
- setShrinkage
- reportTablesFIM

• generateReportEvaluation

Class Combined1

• See class ModelError

Class Constant

• See class ModelError

Class Design

- getName
- setName
- getSize
- setSize
- setArms
- getOutcomesEvaluation
- setOutcomesEvaluation
- getOutcomesGradient
- setOutcomesGradient
- getFim
- setFim
- getNumberOfArms
- setNumberOfArms
- setArm
- EvaluateDesign
- plotOutcomesEvaluation
- plotOutcomesGradient
- reportTablesAdministration
- reportTablesDesign

Class Distribution

- getParameters
- setParameters
- getMu
- setMu
- getOmega
- setOmega
- getAdjustedGradient

Class Evaluation

- run
- reportTablesPlot
- generateTables
- Report

$Class\ {\tt FedorovWynnAlgorithm}$

- FedorovWynnAlgorithm_Rcpp
- resizeFisherMatrix
- setParameters
- optimize
- generateReportOptimization

$Class\ {\tt FedorovWynnAlgorithm}$

- FedorovWynnAlgorithm_Rcpp
- resizeFisherMatrix
- setParameters
- optimize
- generateReportOptimization

Class Fim

- EvaluateFisherMatrix
- EvaluateVarianceFIM
- getFisherMatrix
- setFisherMatrix
- getFixedEffects
- setFixedEffects
- getVarianceEffects
- setVarianceEffects
- getDeterminant
- getDcriterion
- getCorrelationMatrix
- getSE
- getRSE
- getShrinkage
- getEigenValues
- getConditionNumberFixedEffects
- getConditionNumberVarianceEffects
- getColumnAndParametersNamesFIM

- $\bullet \ {\tt getColumnAndParametersNamesFIMInLatex}$
- reportTablesFIM
- generateReportEvaluation
- setFimTypeToString

Class GenericMethods

- getName
- getNames
- getSize
- setSize
- getOutcome
- setOutcome
- getFim
- getOdeSolverParameters
- getMu
- setMu
- getOmega
- setOmega
- getParameters
- setParameters
- getModelError
- getSamplings
- getFim
- setName
- setArms
- getArms

Class IndividualFim

- EvaluateFisherMatrix
- EvaluateVarianceFIM
- getRSE
- getShrinkage
- setShrinkage
- reportTablesFIM
- generateReportEvaluation

Class LibraryOfModels

• getName

- getContent
- setContent
- addModel
- addModels
- getLibraryPKModels
- getLibraryPDModels

Class LibraryOfPKPDModels

- getPKModel
- getPDModel
- getPKPDModel

Class LogNormal

• getAdjustedGradient

Class Model

- getName
- setName
- getDescription
- setDescription
- getEquations
- setEquations
- setModelFromLibrary
- getOutcomes
- setOutcomes
- getOutcomesForEvaluation
- setOutcomesForEvaluation
- getParameters
- setParameters
- getModelError
- setModelError
- getInitialConditions
- setInitialConditions
- getOdeSolverParameters
- setOdeSolverParameters
- getModelFromLibrary
- convertPKModelAnalyticToPKModelODE
- getNumberOfParameters

- isModelODE
- isModelAnalytic
- isDoseInEquations
- isModelInfusion
- isModelSteadyState
- isModelBolus
- definePKPDModel
- definePKModel
- defineModel
- defineModelFromLibraryOfModels
- defineModelUserDefined
- defineModelType
- EvaluateModel
- parametersForComputingGradient
- EvaluateVarianceModel
- getFixedParameters
- getModelErrorParametersValues
- reportTablesModelParameters
- reportTablesModelError

Class ModelAnalytic

- EvaluateModel
- definePKModel
- definePKPDModel
- convertPKModelAnalyticToPKModelODE

Class ModelAnalyticBolus

• See class ModelAnalytic

${\bf Class\ Model Analytic Bolus Steady State}$

• See class ModelAnalyticBolus

Class ModelBolus

• See class Model

Class ModelError

- getOutcome
- getEquation

- setEquation
- getDerivatives
- setDerivatives
- getSigmaInter
- setSigmaInter
- getSigmaSlope
- setSigmaSlope
- getcError
- setcError
- getParameters
- EvaluateErrorModelDerivatives

Class ModelInfusion

- getEquationsDuringInfusion
- getEquationsAfterInfusion
- setEquationsAfterInfusion
- setEquationsDuringInfusion

Class ModelODE

• See class Model

Class ModelODEBolus

- EvaluateModel
- definePKPDModel

${\bf Class} \; {\tt ModelODEDoseInEquations}$

- EvaluateModel
- definePKModel
- definePKPDModel

${\bf Class} \; {\tt ModelODEDoseNotInE} quations$

- EvaluateModel
- definePKModel
- definePKPDModel

Class ModelODEInfusion

• See class ModelInfusion

 ${\bf Class} \ {\tt ModelODEInfusionDoseInEquations}$

- EvaluateModel
- definePKModel
- definePKPDModel

Class ModelParameter

- getName
- getDistribution
- setDistribution
- getFixedMu
- setFixedMu
- getFixedOmega
- setFixedOmega
- getMu
- setMu
- getOmega
- setOmega

Class MultiplicativeAlgorithm

- MultiplicativeAlgorithm_Rcpp
- getLambda
- getDelta
- getNumberOfIterations
- getOptimalWeights
- setOptimalWeights
- setParameters
- optimize
- getDataFrameResults
- plotWeights
- getWeightThreshold
- generateReportOptimization

Class Normal

• getAdjustedGradient

Class Optimization

- getProportionsOfSubjects
- getOptimizationResults
- setOptimizationResults

- getEvaluationFIMResults
- setEvaluationFIMResults
- setEvaluationInitialDesignResults
- getEvaluationInitialDesignResults
- getElementaryProtocols
- generateFimsFromConstraints
- run
- plotWeights
- Report

Class PFIMProject

- getName
- setModel
- getModel
- getModelEquations
- getModelParameters
- getModelError
- getDesigns
- getFim
- getOdeSolverParameters
- getOutcomes
- getOptimizer
- getOptimizerParameters
- run
- generateTables
- Report

Class PGBOAlgorithm

- setParameters
- optimize
- generateReportOptimization

Class PlotEvaluation

- plot
- plotSE
- plotRSE
- plotShrinkage

Class PopulationFim

- EvaluateFisherMatrix
- EvaluateVarianceFIM
- getRSE
- getShrinkage
- setShrinkage
- reportTablesFIM
- computeVMat
- generateReportEvaluation

Class Proportional

• See class ModelError

Class PSOAlgorithm

- setParameters
- optimize
- generateReportOptimization

Class SamplingTimeConstraints

- getOutcome
- getSamplings
- getFixedTimes
- getNumberOfTimesByWindows
- getMinSampling
- getSamplingsWindows
- getNumberOfsamplingsOptimisable
- $\bullet \ check Sampling Time Constraints For Continuous Optimization\\$
- generateSamplingsFromSamplingConstraints

Class SamplingTimes

- getOutcome
- setOutcome
- getSamplings
- setSamplings

Class SimplexAlgorithm

- setParameters
- fun.amoeba
- fisher.simplex
- optimize
- generateReportOptimization

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See Also

Useful links:

• http://www.pfim.biostat.fr/

addModel 21

addModel

Add a model to a library of models.

Description

Add a model to a library of models.

Usage

```
addModel(object, model)
## S4 method for signature 'LibraryOfModels'
addModel(object, model)
```

Arguments

object An object from the class LibraryOfModels.

model An object from the class Model.

Value

The library of models with the added model.

addModels

Add a models to a library of models.

Description

Add a models to a library of models.

Usage

```
addModels(object, models)
## S4 method for signature 'LibraryOfModels'
addModels(object, models)
```

Arguments

object An object from the class LibraryOfModels.

Models A list of object from the class Model.

Value

The library of models with the added models.

Administration-class Class "Administration"

Description

The class Administration defines information concerning the parametrization and the type of administration: single dose, multiple doses. Constraints can also be added on the allowed times, doses and infusion duration.

Objects from the class

Objects form the class Administration can be created by calls of the form Administration(...) where (...) are the parameters for the Administration objects.

Slots for Administration objects

outcome: A character string giving the name for the response of the model.

timeDose: A numeric vector giving the times when doses are given.

dose: A numeric vector giving the amount of doses.

Tinf: A numeric vector giving the infusion duration Tinf (Tinf can be null).

tau: A numeric giving the frequency.

AdministrationConstraints-class

Class "AdministrationConstraints"

Description

The class AdministrationConstraints represents the constraint of an input to the system. The class stores information concerning the constraints for the dosage regimen: response of the model, amount of dose.

Objects from the class

Objects form the class AdministrationConstraints can be created by calls of the form AdministrationConstraints(... where (...) are the parameters for the AdministrationConstraints objects.

Slots for AdministrationConstraints objects

outcome: A character string giving the name for the response of the model.

doses: A numeric vector giving the amount of doses.

Arm-class 23

Arm-class

Class "Arm"

Description

The class Arm combines the treatment and the sampling schedule.

Objects from the class

Objects form the class Arm can be created by calls of the form Arm(...) where (...) are the parameters for the Arm objects.

Slots for the Arm objects

name: A string giving the name of the arm.

size: An integer giving the number of subjects in the arm. By default set to 1.

administrations: A list of the administrations.

initialConditions: A list of the initial conditions.

samplingTimes: A list of the sampling times.

administrationsConstraints: A list of the administrations constraints.

samplingTimesConstraints: A list of the sampling times constraints.

BayesianFim-class

Class "BayesianFim"

Description

The class BayesianFim represents the population Fisher information matrix. The class BayesianFim inherits from the class Fim.

check Sampling Time Constraints For Continuous Optimization

Check for the samplingTime constraints for continuous optimization

Description

Check for the samplingTime constraints for continuous optimization

Usage

```
checkSamplingTimeConstraintsForContinuousOptimization(
  object,
  arm,
  newSamplings,
  outcome
)

## S4 method for signature 'SamplingTimeConstraints'
checkSamplingTimeConstraintsForContinuousOptimization(
  object,
  arm,
  newSamplings,
  outcome
)
```

Arguments

object An object from the class SamplingTimeConstraints.

arm An object from the class Arm.

newSamplings A vector giving the new sampling.

outcome The outcomes for the model.

Value

A list of Boolean giving true if the minimal sampling times is in the vector of sampling times & the number of sampling for each windows is respected false otherwise.

```
check Validity Sampling Constraint \\ check Validity Sampling Constraint
```

Description

Check the validity of he sampling times constraints

Usage

```
checkValiditySamplingConstraint(object)
## S4 method for signature 'Design'
checkValiditySamplingConstraint(object)
```

Arguments

object An object from the class Design.

Combined1-class 25

Value

An error message if a constraint is not valid.

Combined1-class

Class "Combined1"

Description

The class Combined1 defines the the residual error variance according to the formula g(sigmaInter, sigmaSlope, cError, f(x, theta)) = sigmaInter + sigmaSlope*f(x, theta)). The class Combined1 inherits from the class ModelError.

Objects from the class

Combined1 objects are typically created by calls to Combined1 and contain the following slots that are inherited from the class ModelError:

outcome: A string giving the name of the outcome.

equation: An symbolic expression of the model error.

derivatives: A list containing the derivatives of the model error expression.

sigmaInter: A numeric value giving the sigma inter of the error model.

sigmaSlope: A numeric value giving the sigma slope of the error model.

cError: A numeric value giving the exponant c of the error model.

computeVMat

function computeVMat

Description

function computeVMat

Usage

```
computeVMat(varParam1, varParam2, invCholV)
```

Arguments

varParam1 varParam1 varParam2 varParam2 invCholV invCholV

Value

VMat

Constant-class

Class "Constant"

Description

The class Constant defines the tresidual error variance according to the formula $g(sigma_inter, sigma_slope, c_error, f(x, theta)) = sigma_inter$. The class Constant inherits from the class ModelError.

Objects from the class

Constant objects are typically created by calls to Constant and contain the following slots that are inherited from the class ModelError:

outcome: A string giving the name of the outcome.
equation: An symbolic expression of the model error.
derivatives: A list containing the derivatives of the model error expression.
sigmaInter: A numeric value giving the sigma inter of the error model.
sigmaSlope: A numeric value giving the sigma slope of the error model.
cError: A numeric value giving the exponant c of the error model.

convertPKModelAnalyticToPKModelODE

Convert an analytic model to a ode model.

Description

Convert an analytic model to a ode model.

Usage

```
convertPKModelAnalyticToPKModelODE(object)

## S4 method for signature 'ModelAnalytic'
convertPKModelAnalyticToPKModelODE(object)

## S4 method for signature 'ModelAnalyticSteadyState'
convertPKModelAnalyticToPKModelODE(object)

## S4 method for signature 'ModelAnalyticInfusion'
convertPKModelAnalyticToPKModelODE(object)
```

Arguments

object

An object from the class Model.

dataForArmEvaluation 27

Value

A ode model.

dataForArmEvaluation dataForArmEvaluation

Description

dataForArmEvaluation

Usage

```
dataForArmEvaluation(object, arm, model)
## S4 method for signature 'Design'
dataForArmEvaluation(object, arm, model)
```

Arguments

object An object Design from the class Design.

arm .

model An object Model from the class Model.

Value

A list containing data for arm evaluation in the design.

defineModel Define a model.

Description

Define a model.

Usage

```
defineModel(object, designs)
## S4 method for signature 'Model'
defineModel(object, designs)
```

Arguments

object An object from the class Model.

designs A list of objects from the class Design.

Value

A model defined either from the library of models or user defined.

```
\label{lem:defineModelEquationsFromStringToFunction} define Model Equations From String To Function
```

Description

define Model Equations From String To Function

Usage

```
defineModelEquationsFromStringToFunction(
  object,
 parametersNames,
 outcomesWithAdministration,
  outcomesWithNoAdministration
)
## S4 method for signature 'ModelAnalytic'
define Model Equations From String To Function (
 object,
 parametersNames,
 outcomesWithAdministration,
  outcomesWithNoAdministration
)
## S4 method for signature 'ModelAnalyticSteadyState'
defineModelEquationsFromStringToFunction(
  object,
 parametersNames,
 outcomesWithAdministration,
  outcomesWithNoAdministration
)
## S4 method for signature 'ModelAnalyticInfusion'
define Model Equations From String To Function (
 object,
 parametersNames,
 outcomesWithAdministration,
  outcomesWithNoAdministration
)
## S4 method for signature 'ModelAnalyticInfusionSteadyState'
defineModelEquationsFromStringToFunction(
```

```
object,
  parametersNames,
  outcomesWithAdministration,
  outcomesWithNoAdministration
)
## S4 method for signature 'ModelODEBolus'
defineModelEquationsFromStringToFunction(
  object,
  parametersNames,
  outcomesWithAdministration,
  outcomesWithNoAdministration
## S4 method for signature 'ModelODEDoseInEquations'
defineModelEquationsFromStringToFunction(
  object,
  parametersNames,
  outcomesWithAdministration,
  outcomesWithNoAdministration
)
## S4 method for signature 'ModelODEDoseNotInEquations'
defineModelEquationsFromStringToFunction(
  object,
  parametersNames,
  outcomesWithAdministration,
  outcomesWithNoAdministration
)
## S4 method for signature 'ModelODEInfusionDoseInEquations'
defineModelEquationsFromStringToFunction(
  object,
  parametersNames,
  outcomesWithAdministration,
  \verb"outcomesWithNoAdministration"
)
```

Arguments

```
object An object from the class Model.

parametersNames

Vector of parameter names.

outcomesWithAdministration

Vector of the name of the outcome with administration.

outcomesWithNoAdministration

Vector of the name of the outcome with no administration.
```

30 defineModelType

Value

••••

 ${\tt define Model From Library Of Models}$

Define a model from the library of models.

Description

Define a model from the library of models.

Usage

```
defineModelFromLibraryOfModels(object, designs)
## S4 method for signature 'Model'
defineModelFromLibraryOfModels(object, designs)
```

Arguments

object An object from the class Model.

designs A list of objects from the class Design.

Value

A model defined from the library of models.

defineModelType

Define the type of a model.

Description

Define the type of a model.

Usage

```
defineModelType(object, designs)
## S4 method for signature 'Model'
defineModelType(object, designs)
```

Arguments

object An object from the class Model.

designs A list of objects from the class Design.

defineModelUserDefined 31

Value

Return a model defined as analytic, ode, etc.

defineModelUserDefined

Define a user defined model.

Description

Define a user defined model.

Usage

```
defineModelUserDefined(object, designs)
## S4 method for signature 'Model'
defineModelUserDefined(object, designs)
```

Arguments

object An object from the class Model.

designs A list of objects from the class Design.

Value

A model giving a user defined model.

definePKModel

Define a PK model.

Description

Define a PK model.

Usage

```
definePKModel(object, outcomes)
## S4 method for signature 'ModelAnalytic'
definePKModel(object, outcomes)
## S4 method for signature 'ModelAnalyticSteadyState'
definePKModel(object, outcomes)
## S4 method for signature 'ModelAnalyticInfusion'
```

32 definePKPDModel

```
definePKModel(object, outcomes)
## S4 method for signature 'ModelODEDoseInEquations'
definePKModel(object, outcomes)
## S4 method for signature 'ModelODE'
definePKModel(object, outcomes)
## S4 method for signature 'ModelODEInfusionDoseInEquations'
definePKModel(object, outcomes)
```

Arguments

object An object from the class Model.

outcomes A list giving the outcomes of the PK model.

Value

A model giving a PK model.

definePKPDModel

Define a PKPD model.

Description

Define a PKPD model.

Usage

```
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelAnalytic,ModelAnalytic'
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelAnalytic,ModelODE'
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelAnalyticSteadyState,ModelAnalyticSteadyState'
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelAnalyticSteadyState,ModelODE'
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelAnalyticInfusion,ModelAnalytic'
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelAnalyticInfusion,ModelODE'
```

Design-class 33

```
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelODEBolus,ModelODE'
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelODEDoseInEquations,ModelODE'
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelODEDoseNotInEquations,ModelODE'
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelODEInfusion,ModelODE'
definePKPDModel(PKModel, PDModel, outcomes)

## S4 method for signature 'ModelODEInfusionDoseInEquations,ModelODE'
definePKPDModel(PKModel, PDModel, outcomes)
```

Arguments

PKModel An object from the class Model.

PDModel An object from the class Model.

outcomes A list giving the outcomes of the PKPD model.

Value

A model giving a PKPD model.

	Design-class	Class "Design"		
--	--------------	----------------	--	--

Description

The class Design defines information concerning the parametrization of the designs.

Objects from the class

Objects form the class Design can be created by calls of the form Design(...) where (...) are the parameters for the Design objects.

Slots for the Design objects

name: A string giving the name of the design.

size: An integer giving the number of subjects in the design.

arms: A list of the arms.

outcomesEvaluation: A list of the results of the design evaluation for the outcomes.

outcomesGradient: A list of the results of the design evaluation for the sensitivity indices.

34 EvaluateArm

numberOfArms: A numeric giving the number of arms in the design.

fim: An object of the class Fim containing the Fisher Information Matrix of the design.

Distribution-class

Class "Distribution"

Description

The class defines all the required methods for a distribution object.

Objects from the class

Objects form the class Distribution can be created by calls of the form Distribution(...) where (...) are the parameters for the Distribution objects.

Slots for Distribution objects

parameters: A list containing the distribution parameters.

EvaluateArm

EvaluateArm

Description

Evaluate an arm.

Usage

```
EvaluateArm(object, model, dataForModelEvaluation, fim)
## S4 method for signature 'Arm'
EvaluateArm(object, model, dataForModelEvaluation, fim)
```

Arguments

object An object arm from the class Arm.

Model An object model from the class Model.

dataForModelEvaluation

••••

fim An object fim from the class Fim.

Value

The object fim containing the Fisher Information Matrix the two lists evaluationOutcomes, outcomesGradient containing the results of the evaluation of the outcome and the sensitivity indices.

EvaluateDesign 35

EvaluateDesign	<i>EvaluateDesign</i>

Description

Evaluate an design

Usage

```
EvaluateDesign(object, model, fim)
## S4 method for signature 'Design'
EvaluateDesign(object, model, fim)
```

Arguments

object An object Design from the class Design.

model An object model from the class Model.

fim An object fim from the class Fim.

Value

The object Design with its slot fim, evaluationOutcomes, outcomesGradient updated.

EvaluateErrorModelDerivatives

Evaluate the error model derivatives.

Description

Evaluate the error model derivatives.

Usage

```
EvaluateErrorModelDerivatives(object, evaluationOutcome)
## S4 method for signature 'ModelError'
EvaluateErrorModelDerivatives(object, evaluationOutcome)
```

Arguments

```
\begin{tabular}{ll} \beg
```

A list giving the results of the model evaluation.

36 EvaluateFisherMatrix

Value

A list giving the error variance and the Sigma derivatives.

EvaluateFisherMatrix Evaluate the Fisher matrix (population, individual and Bayesian)

Description

Evaluate the Fisher matrix (population, individual and Bayesian)

Usage

```
EvaluateFisherMatrix(object, model, arm, modelEvaluation, modelVariance)
## S4 method for signature 'BayesianFim'
EvaluateFisherMatrix(object, model, arm, modelEvaluation, modelVariance)
## S4 method for signature 'IndividualFim'
EvaluateFisherMatrix(object, model, arm, modelEvaluation, modelVariance)
## S4 method for signature 'PopulationFim'
EvaluateFisherMatrix(object, model, arm, modelEvaluation, modelVariance)
```

Arguments

object An object from the class Fim.

model An object from the class Model.

arm An object from the class Arm.

modelEvaluation

A list containing the evaluation results.

modelVariance A list containing the model variance.

Value

An object from the class Fim containing the Fisher matrix.

EvaluateModel 37

EvaluateModel

Evaluate a model.

Description

Evaluate a model.

Usage

```
EvaluateModel(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelAnalytic'
EvaluateModel(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelAnalyticSteadyState'
EvaluateModel(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelAnalyticInfusion'
EvaluateModel(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelAnalyticInfusionSteadyState'
EvaluateModel(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelODEBolus'
EvaluateModel(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelODEDoseInEquations'
EvaluateModel(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelODEDoseNotInEquations'
EvaluateModel(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelODEInfusionDoseInEquations'
EvaluateModel(object, dataForModelEvaluation, arm)
```

Arguments

```
object An object from the class Model.
dataForModelEvaluation ...
arm An object from the class Arm.
```

Value

A list giving the results of the model evaluation.

38 EvaluateModelGradient

EvaluateModelGradient Evaluate model gradient.

Description

Evaluate model gradient.

Usage

```
EvaluateModelGradient(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelAnalytic'
EvaluateModelGradient(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelAnalyticSteadyState'
EvaluateModelGradient(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelAnalyticInfusion'
EvaluateModelGradient(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelAnalyticInfusionSteadyState'
EvaluateModelGradient(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelODEBolus'
EvaluateModelGradient(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelODEDoseInEquations'
EvaluateModelGradient(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelODEDoseNotInEquations'
EvaluateModelGradient(object, dataForModelEvaluation, arm)
## S4 method for signature 'ModelODEInfusionDoseInEquations'
EvaluateModelGradient(object, dataForModelEvaluation, arm)
```

Arguments

```
object An object from the class Model.
dataForModelEvaluation ...
arm An object from the class Arm.
```

Value

A list giving the results of the model evaluation.

Evaluate Variance FIM 39

EvaluateVarianceFIM Evaluate the variance of the Fisher information matrix.

Description

Evaluate the variance of the Fisher information matrix.

Usage

```
EvaluateVarianceFIM(object, model, arm, modelEvaluation, modelVariance)
## S4 method for signature 'IndividualFim'
EvaluateVarianceFIM(object, model, arm, modelEvaluation, modelVariance)
## S4 method for signature 'PopulationFim'
EvaluateVarianceFIM(object, model, arm, modelEvaluation, modelVariance)
```

Arguments

object An object from the class Fim.

model An object from the class Model.

arm An object from the class Arm.

modelEvaluation

A list containing the evaluation results.

modelVariance A list containing the model variance.

Value

A list containing the matrices of the variance of the FIM.

EvaluateVarianceModel Evaluate the variance of a model.

Description

Evaluate the variance of a model.

```
EvaluateVarianceModel(object, arm, evaluationModel, data)
## S4 method for signature 'Model'
EvaluateVarianceModel(object, arm, evaluationModel, data)
```

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Arguments

object An object from the class Model.

An object from the class Arm.

evaluationModel

A list giving the outputs of the model evaluation.

data ...

Value

Return a list giving the results of the evaluation of the model variance.

Evaluation-class

Class "Evaluation"

Description

A class storing information concerning the evaluation of a design.

Objects from the class

Objects form the class Evaluation can be created by calls of the form Evaluation(...) where (...) are the parameters for the Evaluation objects.

Slots for the Evaluation objects

name: A string giving the name of the project.

model: A object of class Model giving the model.

modelEquations: A list giving the model equations.

modelParameters: A list giving the model parameters.

modelError: A list giving the model error for each outcome of the model.

outcomes: A list giving the model outcomes.

designs: A list giving the designs to be evaluated.

fim: An object of the class Fim containing the Fisher Information Matrix of the design.

odeSolverParameters:

FedorovWynnAlgorithm-class

Class "FedorovWynnAlgorithm"

Description

Class FedorovWynnAlgorithm represents an initial variable for ODE model.

Objects from the class FedorovWynnAlgorithm

Objects form the class FedorovWynnAlgorithm can be created by calls of the form FedorovWynnAlgorithm(...) where (...) are the parameters for the FedorovWynnAlgorithm objects.

Slots for FedorovWynnAlgorithm objects

```
elementaryProtocols: A list of vector for the initial elementary protocols.

numberOfSubjects: A vector for the number of subjects.

proportionsOfSubjects: A vector for the number of subjects.

OptimalDesign: A object Design giving the optimal Design.

showProcess: A boolean to show the process or not.

FisherMatrix: A vector giving the Fisher Information

optimalFrequencies: A vector of the optimal frequencies.

optimalSamplingTimes: A list of vectors for the optimal sampling times.

optimalDoses: A vector for the optimal doses.
```

FedorovWynnAlgorithm_Rcpp

Fedorov-Wynn algorithm in Rcpp.

Description

Run the FedorovWynnAlgorithm in Rcpp

```
FedorovWynnAlgorithm_Rcpp(
  protocols_input,
  ndimen_input,
  nbprot_input,
  numprot_input,
  freq_input,
  nbdata_input,
  vectps_input,
```

42 Fim-class

```
fisher_input,
nok_input,
protdep_input,
freqdep_input
```

Arguments

protocols_input

parameter protocols_input ndimen_input parameter ndimen_input nbprot_input parameter nbprot_input numprot_input parameter numprot_input freq_input parameter freq_input nbdata_input parameter nbdata_input vectps_input parameter vectps_input fisher_input parameter fisher_input nok_input parameter nok_input protdep_input parameter protdep_input freqdep_input parameter freqdep_input

Value

A list giving the results of the outputs of the FedorovWynn algorithm.

Fim-class	Class "Fim"	

Description

A class storing information regarding the Fisher matrix. Type of the Fisher information: population ("PopulationFIM"), individual ("IndividualFIM") or Bayesian ("BayesianFIM").

Objects from the class

Objects form the class Fim can be created by calls of the form Fim(...) where (...) are the parameters for the Fim objects.

Slots for Fim objects

```
fisherMatrix: A matrix giving the Fisher matrix.

fixedEffects: A matrix giving the fixed effects of the Fisher matrix.

varianceEffects: A matrix giving the variance effects of the Fisher matrix.

shrinkage: A vector giving the shrinkage value of the parameters.
```

fisher.simplex 43

fisher.simplex	Compute the fisher.simplex
----------------	----------------------------

Description

Compute the fisher.simplex

Usage

```
fisher.simplex(simplex, optimizationObject, outcomes)
```

Arguments

simplex A list giving the parameters of the simplex.

optimizationObject

An object from the class Optimization.

outcomes A vector giving the outcomes of the arms.

Value

A list giving the results of the optimization.

fun.amoeba	function fun.amoeba	
------------	---------------------	--

Description

function fun.amoeba

Usage

```
fun.amoeba(p, y, ftol, itmax, funk, outcomes, data, showProcess)
```

Arguments

р	input is a matrix p whose ndim+1 rows are ndim-dimensional vectors which are the vertices of the starting simplex.
У	vector whose components must be pre-initialized to the values of funk evaluated at the ndim+1 vertices (rows) of p.
ftol	the fractional convergence tolerance to be achieved in the function value.

itmax maximal number of iterations.

funk multidimensional function to be optimized.

outcomes A vector giving the outcomes.

data a fixed set of data.

showProcess A boolean for showing the process or not.

Value

A list containing the components of the optimized simplex. 'getColumnAndParametersNames-FIMInLatex.

generateFimsFromConstraints

Generate the fim from the constraints

Description

Generate the fim from the constraints

Usage

```
generateFimsFromConstraints(object, fims)
## S4 method for signature 'Optimization'
generateFimsFromConstraints(object)
```

Arguments

object An object from the class Optimization.

fims A list of object from the class Fim.

Value

A list giving the arms with their fims.

generateReportEvaluation

Generate the report for the evaluation

Description

Generate the report for the evaluation

Usage

```
generateReportEvaluation(
     object,
     evaluationObject,
     outputPath,
     outputFile,
     plotOptions
   )
   ## S4 method for signature 'BayesianFim'
   generateReportEvaluation(
     object,
     evaluationObject,
     outputPath,
     outputFile,
     plotOptions
   )
   ## S4 method for signature 'IndividualFim'
   generateReportEvaluation(
     object,
     evaluationObject,
     outputPath,
     outputFile,
     plotOptions
   )
   ## S4 method for signature 'PopulationFim'
   generateReportEvaluation(
     object,
     evaluationObject,
     outputPath,
     outputFile,
     plotOptions
   )
Arguments
   object
                    An object from the class Fim.
   evaluationObject
```

```
A list giving the results of the evaluation of the model.
outputPath
                   A string giving the output path.
                   A string giving the name of the output file.
outputFile
plotOptions
                   A list giving the plot options.
```

Value

Return the report for the evaluation in html.

generateReportOptimization

Generate report for the optimization.

Description

Generate report for the optimization.

```
generateReportOptimization(
  object,
  optimizationObject,
  outputPath,
  outputFile,
  plotOptions
)
## S4 method for signature 'FedorovWynnAlgorithm'
generateReportOptimization(
  object,
  optimizationObject,
  outputPath,
  outputFile,
  plotOptions
)
## S4 method for signature 'MultiplicativeAlgorithm'
generateReportOptimization(
  object,
  optimizationObject,
  outputPath,
  outputFile,
  plotOptions
)
## S4 method for signature 'PGBOAlgorithm'
generateReportOptimization(
  object,
  optimizationObject,
  outputPath,
  outputFile,
  plotOptions
)
## S4 method for signature 'PSOAlgorithm'
generateReportOptimization(
```

```
object,
optimizationObject,
outputPath,
outputFile,
plotOptions
)

## S4 method for signature 'SimplexAlgorithm'
generateReportOptimization(
  object,
  optimizationObject,
  outputPath,
  outputFile,
  plotOptions
)
```

Arguments

object An object from the class OptimizationAlgorithm.

optimizationObject

An object from the class Optimization.

outputPath A string giving the output path.

outputFile A string giving the name of the output file.

plotOptions A list giving the plot options.

Value

The report for the optimization in html.

```
generateSamplingsFromSamplingConstraints

Generate samplings from sampling constraints
```

Description

Generate samplings from sampling constraints

Usage

```
generateSamplingsFromSamplingConstraints(object)
## S4 method for signature 'SamplingTimeConstraints'
generateSamplingsFromSamplingConstraints(object)
```

Arguments

object An object from the class SamplingTimeConstraints.

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Value

A list of sampling times generated from the sampling constraints.

generateTables

Generate the tables for the report.

Description

Generate the tables for the report.

Usage

```
generateTables(object, plotOptions)
## S4 method for signature 'Evaluation'
generateTables(object, plotOptions)
## S4 method for signature 'Optimization'
generateTables(object, plotOptions)
```

Arguments

object An object from the class PFIMProject.

plotOptions A list giving the plot options.

Value

A list giving the kable able for the report (evaluation and optimization).

```
{\tt getAdjustedGradient} \qquad {\tt getAdjustedGradient}
```

Description

Get the adjusted gradient.

```
getAdjustedGradient(object, outcomesGradient)
## S4 method for signature 'LogNormal'
getAdjustedGradient(object, outcomesGradient)
## S4 method for signature 'Normal'
getAdjustedGradient(object, outcomesGradient)
```

getAdministration 49

Arguments

 $\begin{array}{ll} \text{object} & \text{An object distribution from the class } \textbf{Distribution.} \\ \text{outcomesGradient} \end{array}$

A list containing the evaluation of the outcome gradients.

Value

A list giving the adjusted gradient.

getAdministration getAdministration

Description

Get the administrations by outcome.

Usage

```
getAdministration(object, outcome)
## S4 method for signature 'Arm'
getAdministration(object, outcome)
```

Arguments

object An object Arm from the class Arm.

outcome A string giving the name of the outcome.

Value

The element of the list administrations containing the administration of the outcome outcome

```
{\tt getAdministrationConstraint} \\ {\tt getAdministrationConstraint}
```

Description

Get the administration constraints by outcome.

```
getAdministrationConstraint(object, outcome)
## S4 method for signature 'Arm'
getAdministrationConstraint(object, outcome)
```

Arguments

object An object Arm from the class Arm.

outcome A string giving the name of the outcome.

Value

The element of the list getAdministrationConstraint containing the administration constraints of the outcome outcome

getAdministrations

Description

Get all the administration for an arm.

Usage

```
getAdministrations(object)
## S4 method for signature 'Arm'
getAdministrations(object)
```

Arguments

object An object Arm from the class Arm.

Value

A list administrations of objects from the class Administration class giving the parameters of the administration for the object Arm.

 $\label{eq:getAdministrationsConstraints} getAdministrationsConstraints$

Description

Get the administrations constraints.

```
getAdministrationsConstraints(object)
## S4 method for signature 'Arm'
getAdministrationsConstraints(object)
```

getArms 51

Arguments

object

An object Arm from the class Arm.

Value

The list administrationsConstraints.

getArms

Get the arms of an object.

Description

Get the arms of an object.

Usage

```
getArms(object)
## S4 method for signature 'Design'
getArms(object)
## S4 method for signature 'OptimizationAlgorithm'
getArms(object)
```

Arguments

object

An object defined form a class of PFIM.

Value

A list containing the arms of the object.

getcError

Get the parameter c.

Description

Get the parameter c.

```
getcError(object)
## S4 method for signature 'ModelError'
getcError(object)
```

Arguments

object An object from the class ModelError.

Value

A numeric giving the parameter c.

```
getColumnAndParametersNamesFIM
```

Get the names of the names of the parameters associated to each column of the fim.

Description

Get the names of the names of the parameters associated to each column of the fim.

Usage

```
getColumnAndParametersNamesFIM(object, model)
## S4 method for signature 'BayesianFim'
getColumnAndParametersNamesFIM(object, model)
## S4 method for signature 'IndividualFim'
getColumnAndParametersNamesFIM(object, model)
## S4 method for signature 'PopulationFim'
getColumnAndParametersNamesFIM(object, model)
```

Arguments

object An object from the class Fim.

model An object from the class Model.

Value

A list giving the names of the parameters associated to each column of the fim.

```
{\tt getColumnAndParametersNamesFIMInLatex}
```

Get the names of the names of the parameters associated to each column of the fim in Latex format.

Description

Get the names of the names of the parameters associated to each column of the fim in Latex format.

Usage

```
getColumnAndParametersNamesFIMInLatex(object, model)
## S4 method for signature 'BayesianFim'
getColumnAndParametersNamesFIMInLatex(object, model)
## S4 method for signature 'IndividualFim'
getColumnAndParametersNamesFIMInLatex(object, model)
## S4 method for signature 'PopulationFim'
getColumnAndParametersNamesFIMInLatex(object, model)
```

Arguments

object An object from the class Fim.

model An object from the class Model.

Value

A list giving the names of the parameters associated to each column of the fim in Latex format.

```
getConditionNumberFixedEffects
```

Get the condition number of the matrix of the fixed effects.

Description

Get the condition number of the matrix of the fixed effects.

```
getConditionNumberFixedEffects(object)
## S4 method for signature 'Fim'
getConditionNumberFixedEffects(object)
```

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Arguments

object

An object from the class Fim.

Value

A numeric giving the condition number of the matrix of the fixed effects.

```
getConditionNumberVarianceEffects
```

Get the condition number of the matrix of the variance effects.

Description

Get the condition number of the matrix of the variance effects.

Usage

```
getConditionNumberVarianceEffects(object)
## S4 method for signature 'Fim'
getConditionNumberVarianceEffects(object)
## S4 method for signature 'BayesianFim'
getConditionNumberVarianceEffects(object)
```

Arguments

object

An object from the class Fim..

Value

A numeric giving the condition number of the matrix of the variance effects.

getContent

Get content of a library of models.

Description

Get content of a library of models.

```
getContent(object)
## S4 method for signature 'LibraryOfModels'
getContent(object)
```

getCorrelationMatrix 55

Arguments

object An object from the class LibraryOfModels.

Value

A list giving the content of the library of models.

getCorrelationMatrix Get the correlation matrix.

Description

Get the correlation matrix.

Usage

```
getCorrelationMatrix(object)

## S4 method for signature 'Fim'
getCorrelationMatrix(object)

## S4 method for signature 'Evaluation'
getCorrelationMatrix(object)

## S4 method for signature 'Optimization'
getCorrelationMatrix(object)
```

Arguments

object An object from the class Fim.

Value

The correlation matrix of the fim.

```
{\it getData} For Arm Evaluation \\ {\it getData} For Arm Evaluation
```

Description

getDataForArmEvaluation

56 getDataFrameResults

Usage

```
getDataForArmEvaluation(object)
## S4 method for signature 'Arm'
getDataForArmEvaluation(object)
```

Arguments

object

An object Arm from the class Arm.

Value

A list containing the data for arm evaluation.

getDataFrameResults

Get the dataframe of the results.

Description

Get the dataframe of the results.

Usage

```
getDataFrameResults(object)
## S4 method for signature 'FedorovWynnAlgorithm'
getDataFrameResults(object)
## S4 method for signature 'MultiplicativeAlgorithm'
getDataFrameResults(object)
## S4 method for signature 'Optimization'
getDataFrameResults(object)
```

Arguments

object

An object from the class OptimizationAlgorithm.

Value

Return the dataframe of the results.

getDcriterion 57

getDcriterion

Get the D criterion of the fim.

Description

Get the D criterion of the fim.

Usage

```
getDcriterion(object)

## S4 method for signature 'Fim'
getDcriterion(object)

## S4 method for signature 'Evaluation'
getDcriterion(object)

## S4 method for signature 'Optimization'
getDcriterion(object)
```

Arguments

object

An object from the class Fim.

Value

A numeric giving the D criterion of the fim.

getDelta

Get the parameter delta

Description

Get the parameter delta

Usage

```
getDelta(object)
## S4 method for signature 'MultiplicativeAlgorithm'
getDelta(object)
```

Arguments

object

An object from the class MultiplicativeAlgorithm.

58 getDescription

Value

A numeric giving the parameter delta.

getDerivatives

Get the derivatives of the model error equation.

Description

Get the derivatives of the model error equation.

Usage

```
getDerivatives(object)
## S4 method for signature 'ModelError'
getDerivatives(object)
```

Arguments

object

An object from the class ModelError.

Value

The derivatives of the model error equation.

getDescription

Get the description of a model.

Description

Get the description of a model.

Usage

```
getDescription(object)
## S4 method for signature 'Model'
getDescription(object)
```

Arguments

object

An object from the class Model.

Value

A list giving the description of a model.

getDesigns 59

getDesigns

Get the designs.

Description

Get the designs.

Usage

```
getDesigns(object)
## S4 method for signature 'PFIMProject'
getDesigns(object)
```

Arguments

object

An object from the class PFIMProject.

Value

A list giving the designs of the object.

getDeterminant

Get the determinant of the fim.

Description

Get the determinant of the fim.

Usage

```
getDeterminant(object)
## S4 method for signature 'Fim'
getDeterminant(object)
## S4 method for signature 'Evaluation'
getDeterminant(object)
## S4 method for signature 'Optimization'
getDeterminant(object)
```

Arguments

object

An object from the class Fim.

getDose

Value

A numeric giving the determinant of the fim.

getDistribution

Get the distribution.

Description

Get the distribution.

Usage

```
getDistribution(object)
## S4 method for signature 'ModelParameter'
getDistribution(object)
```

Arguments

object

An object from the class ModelParameter.

Value

The parameter distribution.

getDose

getDose

Description

Get the amount of doses.

Usage

```
getDose(object)
## S4 method for signature 'Administration'
getDose(object)
## S4 method for signature 'AdministrationConstraints'
getDose(object)
```

Arguments

object

An object Administration from the class Administration.

getEigenValues 61

Value

The numeric amount_dose giving the amount of doses.

getEigenValues

Get the eigenvalues of the fim.

Description

Get the eigenvalues of the fim.

Usage

```
getEigenValues(object)
## S4 method for signature 'Fim'
getEigenValues(object)
```

Arguments

object

An object from the class Fim.

Value

A vector giving the eigenvalues of the fim.

```
getElementaryProtocols
```

Get the elementary protocols.

Description

Get the elementary protocols.

Usage

```
getElementaryProtocols(object, fims)
## S4 method for signature 'Optimization'
getElementaryProtocols(object, fims)
```

Arguments

object An object from the class Optimization. fims A list of object from the class Fim.

62 getEquations

Value

A list containing the results of the evaluation of the elementary protocols giving the numberOf-Times, nbOfDimensions, totalCost, samplingTimes and the fisherMatrices

 ${\tt getEquation}$

Get the equation of a model error.

Description

Get the equation of a model error.

Usage

```
getEquation(object)
## S4 method for signature 'ModelError'
getEquation(object)
```

Arguments

object

An object from the class ModelError.

Value

An expression giving the equation of a model error.

getEquations

Get the equations of a model.

Description

Get the equations of a model.

Usage

```
getEquations(object)
## S4 method for signature 'Model'
getEquations(object)
```

Arguments

object

An object from the class Model.

Value

The list giving the equations of the model.

```
{\tt getEquationsAfterInfusion}
```

Get the equations after infusion.

Description

Get the equations after infusion.

Usage

```
getEquationsAfterInfusion(object)
## S4 method for signature 'Model'
getEquationsAfterInfusion(object)
```

Arguments

object

An object from the class Model.

Value

A list giving the equations after the infusion.

```
getEquationsDuringInfusion
```

Get the equations during infusion.

Description

Get the equations during infusion.

Usage

```
getEquationsDuringInfusion(object)
## S4 method for signature 'Model'
getEquationsDuringInfusion(object)
```

Arguments

object

An object from the class Model.

Value

A list giving the equations during the infusion.

```
getEvaluationFIMResults
```

Get the results of the evaluation.

Description

Get the results of the evaluation.

Usage

```
getEvaluationFIMResults(object)
## S4 method for signature 'Optimization'
getEvaluationFIMResults(object)
```

Arguments

object

An object from the class Optimization.

Value

An object from the class Evaluation giving the evaluation results for the optimal design.

```
{\tt getEvaluationInitialDesignResults}
```

Get the evaluation results of the initial design.

Description

Get the evaluation results of the initial design.

Usage

```
getEvaluationInitialDesignResults(object)
## S4 method for signature 'Optimization'
getEvaluationInitialDesignResults(object)
```

Arguments

object

An object from the class Optimization.

Value

The object from the class Evaluation giving the results of the evaluation of the initial design.

getFim 65

getFim getFim

Description

Get the fim of an object.

Usage

```
getFim(object)
## S4 method for signature 'Design'
getFim(object)
## S4 method for signature 'PFIMProject'
getFim(object)
## S4 method for signature 'OptimizationAlgorithm'
getFim(object)
```

Arguments

object

An object defined form a class of PFIM.

Value

The FIM of the object.

getFisherMatrix

Get the FIM.

Description

Get the FIM.

```
getFisherMatrix(object)
## S4 method for signature 'Fim'
getFisherMatrix(object)
## S4 method for signature 'Evaluation'
getFisherMatrix(object)
## S4 method for signature 'Optimization'
getFisherMatrix(object)
```

getFixedMu

Arguments

object A

An object from the class Fim.

Value

A matrix giving the FIM.

 ${\tt getFixedEffects}$

Get the matrix of fixed effects.

Description

Get the matrix of fixed effects.

Usage

```
getFixedEffects(object)
## S4 method for signature 'Fim'
getFixedEffects(object)
```

Arguments

object

An object from the class Fim.

Value

The matrix of the fixed effects.

getFixedMu

Get the fixed effect.

Description

Get the fixed effect.

Usage

```
getFixedMu(object)
## S4 method for signature 'ModelParameter'
getFixedMu(object)
```

Arguments

object

An object from the class ModelParameter.

getFixedOmega 67

Value

A boolean giving the fixed mu.

getFixedOmega

Get the fixed variance.

Description

Get the fixed variance.

Usage

```
getFixedOmega(object)
## S4 method for signature 'ModelParameter'
getFixedOmega(object)
```

Arguments

object

An object from the class ModelParameter.

Value

A boolean giving the fixed omega.

getFixedParameters

Get the fixed parameters.

Description

Get the fixed parameters.

Usage

```
getFixedParameters(object)
## S4 method for signature 'Model'
getFixedParameters(object)
```

Arguments

object

An object from the class Model.

Value

A list giving the fixed parameters of the model.

68 getInitialConditions

getFixedTimes

Get the fixed sampling times.

Description

Get the fixed sampling times.

Usage

```
getFixedTimes(object)
## S4 method for signature 'SamplingTimeConstraints'
getFixedTimes(object)
```

Arguments

object

An object from the class SamplingTimeConstraints.

Value

A vector giving the foxed sampling times.

```
getInitialConditions getInitialConditions
```

Description

Get the initial condition for the evaluation of an ode model.

Usage

```
getInitialConditions(object)
## S4 method for signature 'Arm'
getInitialConditions(object)
## S4 method for signature 'Model'
getInitialConditions(object)
```

Arguments

object

An object Arm from the class Arm.

Value

The list initial Conditions for the object $\ensuremath{\mathsf{Arm}}.$

getIterationAndCriteria 69

```
getIterationAndCriteria
```

Get the iteration with the convergence criteria.

Description

Get the iteration with the convergence criteria.

Usage

```
getIterationAndCriteria(object)
## S4 method for signature 'OptimizationAlgorithm'
getIterationAndCriteria(object)
```

Arguments

object

An object from the class OptimizationAlgorithm.

Value

A dataframe giving the iteration with the convergence criteria.

getLambda

Get the parameter lambda.

Description

Get the parameter lambda.

Usage

```
getLambda(object)
## S4 method for signature 'MultiplicativeAlgorithm'
getLambda(object)
```

Arguments

object

An object from the class MultiplicativeAlgorithm.

Value

A numeric giving the parameter lambda.

70 getLibraryPKModels

 ${\tt getLibraryPDModels}$

Get the library of PD models.

Description

Get the library of PD models.

Usage

```
getLibraryPDModels(object)
## S4 method for signature 'LibraryOfModels'
getLibraryPDModels(object)
```

Arguments

object

An object from the class LibraryOfModels.

Value

A list giving the PD models.

getLibraryPKModels

Get the library of PK models.

Description

Get the library of PK models.

Usage

```
getLibraryPKModels(object)
## S4 method for signature 'LibraryOfModels'
getLibraryPKModels(object)
```

Arguments

object

An object from the class LibraryOfModels.

Value

A list giving the PK models.

getMinSampling 71

getMinSampling

Get the minimal sampling times.

Description

Get the minimal sampling times.

Usage

```
getMinSampling(object)
## S4 method for signature 'SamplingTimeConstraints'
getMinSampling(object)
```

Arguments

object

An object from the class SamplingTimeConstraints.

Value

A numeric giving the minimal sampling times.

getModel

Get the model.

Description

Get the model.

Usage

```
getModel(object)
## S4 method for signature 'PFIMProject'
getModel(object)
```

Arguments

object

An object from the class PFIMProject.

Value

The model of the object.

72 getModelError

getModelEquations

Get the model equations.

Description

Get the model equations.

Usage

```
getModelEquations(object)
## S4 method for signature 'PFIMProject'
getModelEquations(object)
```

Arguments

object

An object from the class PFIMProject.

Value

A list giving the model equations.

getModelError

Get the model error.

Description

Get the model error.

Usage

```
getModelError(object)

## S4 method for signature 'Model'
getModelError(object)

## S4 method for signature 'PFIMProject'
getModelError(object)
```

Arguments

object

An object defined form a class of PFIM.

Value

The model error of the object.

```
{\tt getModelErrorParametersValues}
```

Get the values of the model error parameters.

Description

Get the values of the model error parameters.

Usage

```
getModelErrorParametersValues(object)
## S4 method for signature 'Model'
getModelErrorParametersValues(object)
```

Arguments

object

An object from the class Model.

Value

A list giving the values of the model error parameters.

getModelFromLibrary

Get a model from the library of models.

Description

Get a model from the library of models.

Usage

```
getModelFromLibrary(object)
## S4 method for signature 'Model'
getModelFromLibrary(object)
```

Arguments

object

An object from the class Model.

Value

Return a model from the the library of models.

 ${\tt getModelParameters}$

Get the model parameters.

Description

Get the model parameters.

Usage

```
getModelParameters(object)
## S4 method for signature 'PFIMProject'
getModelParameters(object)
```

Arguments

object

An object from the class PFIMProject.

Value

A list giving the model parameters.

```
getModelParametersValues
```

Get the values of the model parameters.

Description

Get the values of the model parameters.

Usage

```
getModelParametersValues(object)
## S4 method for signature 'Model'
getModelParametersValues(object)
```

Arguments

object

An object from the class Model.

Value

A list giving the values of the model parameters.

getMu 75

getMu getMu

Description

Get the fixed effect of an object.

Usage

```
getMu(object)
## S4 method for signature 'Distribution'
getMu(object)
## S4 method for signature 'ModelParameter'
getMu(object)
```

Arguments

object

An object defined form a class of PFIM.

Value

The object with the updated fixed effect.

getName

getName

Description

Get the name of an object

```
getName(object)
## S4 method for signature 'Arm'
getName(object)
## S4 method for signature 'Design'
getName(object)
## S4 method for signature 'ModelParameter'
getName(object)
## S4 method for signature 'LibraryOfModels'
```

76 getNames

```
getName(object)

## S4 method for signature 'Model'
getName(object)

## S4 method for signature 'PFIMProject'
getName(object)
```

Arguments

object

An object defined form a class of PFIM.

Value

A character string name giving the name of the object.

 ${\tt getNames}$

getNames

Description

Get the names of an object.

Usage

```
getNames(object)
## S4 method for signature 'list'
getNames(object)
```

Arguments

object

An object defined form a class of PFIM.

Value

A vector giving the names of the object.

getNumberOfArms 77

getNumberOfArms

getNumberOfArms

Description

Get the number of arms in a design.

Usage

```
getNumberOfArms(object)
## S4 method for signature 'Design'
getNumberOfArms(object)
```

Arguments

object

An object Design from the class Design.

Value

A numeric numberOfArms giving the number of arms in the design.

getNumberOfIterations Get the number of iterations.

Description

Get the number of iterations.

Usage

```
getNumberOfIterations(object)
## S4 method for signature 'MultiplicativeAlgorithm'
getNumberOfIterations(object)
```

Arguments

object

An object from the class MultiplicativeAlgorithm.

Value

A numeric giving the number of iterations.

getNumberOfParameters Get the number of parameters.

Description

Get the number of parameters.

Usage

```
getNumberOfParameters(object)
## S4 method for signature 'Model'
getNumberOfParameters(object)
```

Arguments

object

An object from the class Model.

Value

A numeric giving the number of parameters of the model.

```
getNumberOfsamplingsOptimisable
```

Get the number of sampling times that are optimisable.

Description

Get the number of sampling times that are optimisable.

Usage

```
getNumberOfsamplingsOptimisable(object)
## S4 method for signature 'SamplingTimeConstraints'
getNumberOfsamplingsOptimisable(object)
```

Arguments

object

An object from the class SamplingTimeConstraints.

Value

A vector giving the number of sampling times that are optimisable.

```
{\tt getNumberOfTimesByWindows}
```

Get the number of sampling times by windows.

Description

Get the number of sampling times by windows.

Usage

```
getNumberOfTimesByWindows(object)
## S4 method for signature 'SamplingTimeConstraints'
getNumberOfTimesByWindows(object)
```

Arguments

object

An object from the class SamplingTimeConstraints.

Value

A vector giving the number of sampling times by windows.

Description

Get the parameters for the ode solvers of an object.

Usage

```
getOdeSolverParameters(object)
## S4 method for signature 'Model'
getOdeSolverParameters(object)
## S4 method for signature 'PFIMProject'
getOdeSolverParameters(object)
```

Arguments

object

An object defined form a class of PFIM.

80 getOptimalDesign

Value

The list giving the parameters for the ode solvers.

getOmega

Get the matrix omega of an object.

Description

Get the matrix omega of an object.

Usage

```
getOmega(object)
## S4 method for signature 'Distribution'
getOmega(object)
## S4 method for signature 'ModelParameter'
getOmega(object)
```

Arguments

object

An object defined form a class of PFIM.

Value

The matrix omega of an object.

getOptimalDesign

Get the optimal design.

Description

Get the optimal design.

Usage

```
getOptimalDesign(object)
## S4 method for signature 'OptimizationAlgorithm'
getOptimalDesign(object)
```

Arguments

object

An object from the class OptimizationAlgorithm.

getOptimalFrequencies

Value

The optimal design.

```
getOptimalFrequencies Get the optimal frequencies
```

Description

Get the optimal frequencies

Usage

```
getOptimalFrequencies(object)
## S4 method for signature 'FedorovWynnAlgorithm'
getOptimalFrequencies(object)
```

Arguments

object

An object from the class FedorovWynnAlgorithm.

Value

A vector giving the optimal frequencies

getOptimalWeights

Get the optimal weights.

Description

Get the optimal weights.

Usage

```
getOptimalWeights(object)
## S4 method for signature 'MultiplicativeAlgorithm'
getOptimalWeights(object)
```

Arguments

object

An object from the class Multiplicative Algorithm.

Value

A vector giving the optimal weights.

82 getOptimizer

```
getOptimizationResults
```

Get the optimization results.

Description

Get the optimization results.

Usage

```
getOptimizationResults(object)
## S4 method for signature 'Optimization'
getOptimizationResults(object)
```

Arguments

object

An object from the class Optimization.

Value

An object from the class OptimizationAlgorithm giving the optimization results.

getOptimizer

Get the optimization algorithm.

Description

Get the optimization algorithm.

Usage

```
getOptimizer(object)
## S4 method for signature 'PFIMProject'
getOptimizer(object)
```

Arguments

object

An object from the class PFIMProject.

Value

A string giving the name of the optimization algorithm.

getOptimizerParameters

```
getOptimizerParameters
```

Get the optimization parameters.

Description

Get the optimization parameters.

Usage

```
getOptimizerParameters(object)
## S4 method for signature 'PFIMProject'
getOptimizerParameters(object)
```

Arguments

object

An object from the class PFIMProject.

Value

A list giving the optimization parameters.

getOutcome

getOutcome

Description

Get the outcome of an object.

```
getOutcome(object)
## S4 method for signature 'Administration'
getOutcome(object)
## S4 method for signature 'AdministrationConstraints'
getOutcome(object)
## S4 method for signature 'ModelError'
getOutcome(object)
## S4 method for signature 'SamplingTimeConstraints'
getOutcome(object)
## S4 method for signature 'SamplingTimeConstraints'
getOutcome(object)
```

Arguments

object

An object defined from a class of PFIM.

Value

A string giving the outcome of the object.

getOutcomes

Get the outcomes of a model.

Description

Get the outcomes of a model.

Usage

```
getOutcomes(object)
## S4 method for signature 'Model'
getOutcomes(object)
## S4 method for signature 'PFIMProject'
getOutcomes(object)
```

Arguments

object

An object from the class Model.

Value

A list giving the outcomes of the model.

getOutcomesEvaluation getOutcomesEvaluation

Description

Get the results of the evaluation of the outcomes.

```
getOutcomesEvaluation(object)
## S4 method for signature 'Design'
getOutcomesEvaluation(object)
```

Arguments

object

An object Design from the class Design.

Value

The list outcomesEvaluation containing the results of the design evaluation for the outcomes.

```
getOutcomesForEvaluation
```

Get the outcomes of a model used for the evaluation (is scales outcomes).

Description

Get the outcomes of a model used for the evaluation (is scales outcomes).

Usage

```
getOutcomesForEvaluation(object)
## S4 method for signature 'Model'
getOutcomesForEvaluation(object)
```

Arguments

object

An object from the class Model.

Value

A list giving the outcomes of a model used for the evaluation (is scales outcomes).

getOutcomesGradient get

getOutcomesGradient

Description

Get the results of the evaluation of the outcome gradients.

```
getOutcomesGradient(object)
## S4 method for signature 'Design'
getOutcomesGradient(object)
```

86 getParameters

Arguments

object

An object Design from the class Design.

Value

The list outcomesGradient containing the results of the design evaluation for the outcome gradients.

getParameters

Get the parameters of an object.

Description

Get the parameters of an object.

Usage

```
getParameters(object)
## S4 method for signature 'ModelError'
getParameters(object)
## S4 method for signature 'Distribution'
getParameters(object)
## S4 method for signature 'Model'
getParameters(object)
```

Arguments

object

An object defined form a class of PFIM.

Value

Return the list of the parameters of the object.

getPDModel 87

getPDModel

Get a PD model.

Description

Get a PD model.

Usage

```
getPDModel(object, PDModelName)
## S4 method for signature 'LibraryOfPKPDModels'
getPDModel(object, PDModelName)
```

Arguments

object An object from the class LibraryOfPKPDModels.

PDModelName A string giving the name of the PD model.

Value

Return a PD model.

getPKModel

Get a PK model.

Description

Get a PK model.

Usage

```
getPKModel(object, PKModelName)
## S4 method for signature 'LibraryOfPKPDModels'
getPKModel(object, PKModelName)
```

Arguments

object An object from the class LibraryOfPKPDModels.

PKModelName A string giving the name of the PK model.

Value

Return a PK model.

88 getPlotOptions

getPKPDModel

Get a PKPD model.

Description

Get a PKPD model.

Usage

```
getPKPDModel(object, namesModel)
## S4 method for signature 'LibraryOfPKPDModels'
getPKPDModel(object, namesModel)
```

Arguments

object An object from the class LibraryOfPKPDModels.

namesModel A vector of strings giving the names of the PK and PD models.

Value

Return a PKPD model.

getPlotOptions

Get the plot options for graphs responses and SI

Description

Get the plot options for graphs responses and SI

Usage

```
getPlotOptions(plotOptions, outcomesNames)
```

Arguments

plotOptions A list giving the plots options.

outcomesNames A list giving the output names.

Value

The list containing the plot options.

```
getProportionsOfSubjects
```

Get the proportion of subjects.

Description

Get the proportion of subjects.

Usage

```
getProportionsOfSubjects(object)
## S4 method for signature 'Optimization'
getProportionsOfSubjects(object)
```

Arguments

object

An object from the class Optimization.

Value

A vector giving the proportion of subjects.

getRSE

Get the RSE

Description

Get the RSE

```
getRSE(object, model)

## S4 method for signature 'BayesianFim'
getRSE(object, model)

## S4 method for signature 'Evaluation'
getRSE(object, model)

## S4 method for signature 'IndividualFim'
getRSE(object, model)

## S4 method for signature 'Optimization'
getRSE(object, model)

## S4 method for signature 'PopulationFim'
getRSE(object, model)
```

Arguments

object An object from the class Fim.

model An object from the class Model.

Value

A vector giving the RSE.

getSamplings

Get the sampling of an object.

Description

Get the sampling of an object.

Usage

```
getSamplings(object)
## S4 method for signature 'SamplingTimeConstraints'
getSamplings(object)
## S4 method for signature 'SamplingTimes'
getSamplings(object)
```

Arguments

object

An object defined form a class of PFIM.

Value

A list of the samplings of the object.

getSamplingsWindows

Get the windows for the sampling times.

Description

Get the windows for the sampling times.

```
getSamplingsWindows(object)
## S4 method for signature 'SamplingTimeConstraints'
getSamplingsWindows(object)
```

getSamplingTime 91

Arguments

object An object from the class SamplingTimeConstraints.

Value

A list giving the vector of the windows for the sampling times.

getSamplingTime getSam

getSamplingTime

Description

Get the sampling times by outcome.

Usage

```
getSamplingTime(object, outcome)
## S4 method for signature 'Arm'
getSamplingTime(object, outcome)
```

Arguments

object Arm from the class Arm.

outcome A string giving the name of the outcome.

Value

The element of the list samplingTimes containing the sampling times of the outcome outcome

```
\label{eq:getSamplingTimeConstraint} getSamplingTimeConstraint
```

Description

Get the sampling times constraints by outcome.

```
getSamplingTimeConstraint(object, outcome)
## S4 method for signature 'Arm'
getSamplingTimeConstraint(object, outcome)
```

Arguments

object Arm from the class Arm.

outcome A string giving the name of the outcome.

Value

The element of the list samplingTimesConstraints containing the sampling times constraints of the outcome outcome

 ${\tt getSamplingTimes}$

getSamplingTimes

Description

Get the vectors of sampling times for an arm.

Usage

```
getSamplingTimes(object)
## S4 method for signature 'Arm'
getSamplingTimes(object)
```

Arguments

object

An object Arm from the class Arm.

Value

The list samplingTimes for the object Arm.

```
getSamplingTimesConstraints
```

 ${\it get Sampling Times Constraints}$

Description

Get the sampling times constraints.

```
getSamplingTimesConstraints(object)
## S4 method for signature 'Arm'
getSamplingTimesConstraints(object)
```

getSE 93

Arguments

object

An object Arm from the class Arm.

Value

 $The \ list \ {\tt getSamplingTimesConstraints}.$

getSE

Get the SE.

Description

Get the SE.

Usage

```
getSE(object)
## S4 method for signature 'Fim'
getSE(object)
## S4 method for signature 'Evaluation'
getSE(object)
## S4 method for signature 'Optimization'
getSE(object)
```

Arguments

object

An object from the class Fim.

Value

A vector giving the SE.

getShrinkage

Get the shrinkage.

Description

Get the shrinkage.

94 getSigmaInter

Usage

```
getShrinkage(object)

## S4 method for signature 'BayesianFim'
getShrinkage(object)

## S4 method for signature 'Evaluation'
getShrinkage(object)

## S4 method for signature 'IndividualFim'
getShrinkage(object)

## S4 method for signature 'Optimization'
getShrinkage(object)

## S4 method for signature 'PopulationFim'
getShrinkage(object)
```

Arguments

object An object from the class Fim.

Value

A vector giving the shrinkage of the Bayesian fim.

getSigmaInter

Get the parameter sigma inter.

Description

Get the parameter sigma inter.

Usage

```
getSigmaInter(object)
## S4 method for signature 'ModelError'
getSigmaInter(object)
```

Arguments

object An object from the class ModelError.

Value

A numeric giving the parameter sigma inter.

getSigmaSlope 95

getSigmaSlope

Get the parameter sigma slope.

Description

Get the parameter sigma slope.

Usage

```
getSigmaSlope(object)
## S4 method for signature 'ModelError'
getSigmaSlope(object)
```

Arguments

object

An object from the class ModelError.

Value

A numeric giving the parameter sigma slope.

getSize

getSize

Description

Get the size of an object.

Usage

```
getSize(object)
## S4 method for signature 'Arm'
getSize(object)
## S4 method for signature 'Design'
getSize(object)
```

Arguments

object

An object defined form a class of PFIM.

Value

A numeric giving the size of the object.

96 getTimeDose

getTau

getTau

Description

Get the frequency tau.

Usage

```
getTau(object)
## S4 method for signature 'Administration'
getTau(object)
```

Arguments

object

An object Administration from the class Administration.

Value

The numeric tau giving the frequency tau.

getTimeDose

getTimeDose

Description

Get the times vector when doses are given.

Usage

```
getTimeDose(object)
## S4 method for signature 'Administration'
getTimeDose(object)
```

Arguments

object

An object Administration from the class Administration.

Value

The vector timeDose giving the times when the doses are given.

getTinf 97

getTinf

Get the infusion duration.

Description

Get the infusion duration.

Usage

```
getTinf(object)
## S4 method for signature 'Administration'
getTinf(object)
```

Arguments

object

An object Administration from the class Administration.

Value

The numeric Tinf giving the infusion duration Tinf.

getVariables

Return the variable of an ode model

Description

The class ModelODEBolus defines information concerning the construction of an ode model bolus. The class ModelODEBolus inherits from the class ModelBolus.

Usage

```
getVariables(object)
## S4 method for signature 'ModelODE'
getVariables(object)
## S4 method for signature 'ModelODEBolus'
getVariables(object)
## S4 method for signature 'ModelInfusion'
getVariables(object)
```

Arguments

object

An object from the class Model.

98 getWeightThreshold

Value

Return the variable of an ode model

getVarianceEffects

Get the matrix of the variance effects.

Description

Get the matrix of the variance effects.

Usage

```
getVarianceEffects(object)
## S4 method for signature 'Fim'
getVarianceEffects(object)
```

Arguments

object

An object from the class Fim.

Value

The matrix of the variance effects.

getWeightThreshold

Get the parameter weightThreshold

Description

Get the parameter weightThreshold

Usage

```
getWeightThreshold(object)
## S4 method for signature 'MultiplicativeAlgorithm'
getWeightThreshold(object)
```

Arguments

object

An object from the class Multiplicative Algorithm.

Value

A numeric giving the WeightThreshold.

IndividualFim-class 99

IndividualFim-class Class "Fim"

Description

A class storing information regarding the individual Fisher matrix. The class $IndividualFim\ inherits$ from the class Fim.

 $initialize, {\it Administration-method}\\initialize$

Description

initialize

Usage

```
## S4 method for signature 'Administration'
initialize(.Object, outcome, timeDose, dose, Tinf, tau)
```

Arguments

.Object	.Object
outcome	outcome
timeDose	timeDose
dose	dose
Tinf	Tinf
tau	tau

Value

Administration

100 initialize, Arm-method

```
initialize, {\it Administration Constraints-method} \\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'AdministrationConstraints'
initialize(.Object, outcome, doses)
```

Arguments

.Object .Object outcome outcome doses doses

initialize, Arm-method *initialize*

Description

initialize

```
## S4 method for signature 'Arm'
initialize(
   .Object,
   name,
   size,
   administrations,
   initialConditions,
   samplingTimes,
   administrationsConstraints,
   samplingTimesConstraints,
   dataForArmEvaluation
)
```

Arguments

```
.Object
                .Object
                name
name
                size
size
administrations
                administrations
initialConditions
                initialConditions
samplingTimes samplingTimes
administrationsConstraints
                administrationsConstraints
samplingTimesConstraints
                sampling Times Constraints \\
dataForArmEvaluation
                dataForArmEvaluation
```

Value

Arm

```
initialize, {\tt Combined1-method}\\ initialize
```

Description

initialize

```
## S4 method for signature 'Combined1'
initialize(
   .Object,
   outcome,
   equation,
   derivatives,
   sigmaInter,
   sigmaSlope,
   cError
)
```

Arguments

```
.Object .Object
outcome outcome
equation equation
derivatives derivatives
sigmaInter sigmaInter
sigmaSlope sigmaSlope
cError cError
```

Value

Combined1

```
initialize\,, {\tt Constant-method}\\initialize
```

Description

initialize

Usage

```
## S4 method for signature 'Constant'
initialize(
   .Object,
   outcome,
   equation,
   derivatives,
   sigmaInter,
   sigmaSlope,
   cError
)
```

Arguments

```
.Object .Object outcome equation equation derivatives sigmaInter sigmaSlope cError .Object .Ob
```

Value

Constant

```
initialize, {\tt Design-method}\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'Design'
initialize(
   .Object,
   name,
   size,
   arms,
   outcomesEvaluation,
   outcomesGradient,
   numberOfArms,
   fim
)
```

Arguments

```
.Object
                  .Object
name
                 name
size
                 size
arms
                  arms
{\tt outcomes} {\tt Evaluation}
                  outcomesEvaluation
outcomesGradient
                 outcomesGradient
numberOfArms
                 numberOfArms
fim
                 fim
```

Value

Design

```
initialize\,, {\tt Distribution-method}\\initialize
```

Description

initialize

Usage

```
## S4 method for signature 'Distribution'
initialize(.Object, parameters)
```

Arguments

```
.Object .Object parameters parameters
```

Value

Distribution

```
initialize, {\tt Evaluation-method}\\ initialize
```

Description

initialize

```
## S4 method for signature 'Evaluation'
initialize(
   .Object,
   name,
   model,
   modelEquations,
   modelParameters,
   modelError,
   outcomes,
   designs,
   fim,
   odeSolverParameters
)
```

Arguments

.Object .Object name name model model modelEquations modelEquations modelParameters modelParameters modelError modelError outcomes outcomes designs designs fim fim odeSolverParameters

odeSolverParameters

Value

Evaluation

Description

initialize

Usage

```
## S4 method for signature 'FedorovWynnAlgorithm'
initialize(
   .Object,
   elementaryProtocols,
   numberOfSubjects,
   proportionsOfSubjects,
   showProcess
)
```

Arguments

```
.Object .Object
elementaryProtocols
elementaryProtocols
numberOfSubjects
numberOfSubjects
proportionsOfSubjects
proportionsOfSubjects
showProcess
showProcess
```

Value

FedorovWynnAlgorithm

```
initialize, Fim-method initialize
```

Description

initialize

Usage

```
## S4 method for signature 'Fim'
initialize(.Object, fisherMatrix, fixedEffects, varianceEffects, shrinkage)
```

Arguments

.Object .Object fisherMatrix fixedEffects varianceEffects

varianceEffects

shrinkage shrinkage

Value

Fim

```
initialize, Library Of Models-method\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'LibraryOfModels'
initialize(.Object, name, content)
```

Arguments

.Object .Object name fisherMatrix content fixedEffects

Value

Library Of Models

```
initialize, \verb|LibraryOfPKPDModels-method|\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'LibraryOfPKPDModels'
initialize(.Object)
```

Arguments

.Object .Object

Value

LibraryOfPKPDModels

```
initialize, {\tt LogNormal-method}\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'LogNormal'
initialize(.Object, ...)
```

Arguments

```
.0bject .0bject
```

Value

LogNormal

108 initialize,Model-method

```
\verb|initialize|, \verb|Model-method||
```

initialize

Description

initialize

Usage

```
## S4 method for signature 'Model'
initialize(
   .Object,
   name,
   description,
   equations,
   outcomes,
   outcomesForEvaluation,
   parameters,
   modelError,
   initialConditions,
   odeSolverParameters,
   modelFromLibrary
)
```

Arguments

```
.Object
                .Object
name
                name
description
                description
                equations
equations
                outcomes
outcomes
outcomesForEvaluation
                outcomesForEvaluation
parameters
                parameters
modelError
                modelError
initialConditions
                initialConditions
odeSolverParameters
                odeSolverParameters
modelFromLibrary
```

model From Library

Value

Model

```
initialize, {\tt ModelAnalytic-method} \\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'ModelAnalytic'
initialize(
   .Object,
   name,
   description,
   equations,
   outcomes,
   parameters,
   modelError
)
```

Arguments

```
.Object .Object
name name
description description
equations equations
outcomes outcomes
parameters parameters
modelError modelError
```

Value

ModelAnalytic

```
initialize, {\tt ModelAnalyticBolus-method} \\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'ModelAnalyticBolus'
initialize(
   .Object,
   name,
   description,
   equations,
   outcomes,
   parameters,
   modelError
)
```

Arguments

```
.Object .Object
name name
description description
equations equations
outcomes outcomes
parameters parameters
modelError modelError
```

Value

ModelAnalyticBolus

```
initialize, {\tt ModelAnalyticBolusSteadyState-method} \\ initialize
```

Description

initialize

```
## S4 method for signature 'ModelAnalyticBolusSteadyState'
initialize(
   .Object,
   name,
   description,
   equations,
   outcomes,
   parameters,
   modelError
)
```

Arguments

.Object	.Object
name	name
description	description
equations	equations
outcomes	outcomes
parameters	parameters
modelError	modelError

Value

Model Analytic Bolus Steady State

```
initialize, {\tt ModelAnalyticInfusion-method}\\ initialize
```

Description

initialize

Usage

```
## $4 method for signature 'ModelAnalyticInfusion'
initialize(
   .Object,
   name,
   description,
   equations,
   outcomes,
   parameters,
   modelError
)
```

Arguments

```
.Object .Object
name name
description description
equations equations
outcomes outcomes
parameters parameters
modelError modelError
```

Value

ModelAnalyticInfusion

 $initialize, {\tt ModelAnalyticInfusionSteadyState-method} \\ initialize$

Description

initialize

Usage

```
## S4 method for signature 'ModelAnalyticInfusionSteadyState'
initialize(
   .Object,
   name,
   description,
   equations,
   outcomes,
   parameters,
   modelError
)
```

Arguments

```
.Object .Object
name name
description description
equations equations
outcomes outcomes
parameters parameters
modelError modelError
```

Value

Model Analytic Infusion Steady State

```
initialize, {\tt ModelAnalyticSteadyState-method}\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'ModelAnalyticSteadyState'
initialize(
   .Object,
   name,
   description,
   equations,
   outcomes,
   parameters,
   modelError
)
```

Arguments

```
.Object .Object
name name
description description
equations equations
outcomes outcomes
parameters parameters
modelError modelError
```

Value

Model Analytic Steady State

```
initialize, {\tt ModelBolus-method}\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'ModelBolus'
initialize(
   .Object,
   name,
   description,
   equations,
   outcomes,
   parameters,
   modelError,
   initialConditions,
   odeSolverParameters
)
```

Arguments

```
.Object
                 .Object
name
                name
                description
description
                equations
equations
outcomes
                outcomes
parameters
                parameters
modelError
                modelError
\\ initial Conditions
                initialConditions
odeSolverParameters
                odeSolverParameters
```

Value

ModelBolus

```
initialize, {\tt ModelError-method}\\initialize
```

Description

initialize

Usage

```
## S4 method for signature 'ModelError'
initialize(
   .Object,
   outcome,
   equation,
   derivatives,
   sigmaInter,
   sigmaSlope,
   cError
)
```

Arguments

```
.Object .Object
outcome outcome
equation equation
derivatives derivatives
sigmaInter sigmaInter
sigmaSlope sigmaSlope
cError cError
```

Value

ModelError

```
initialize, {\tt ModelInfusion-method}\\ initialize
```

Description

initialize

```
## S4 method for signature 'ModelInfusion'
initialize(
   .Object,
   name,
   description,
   equations,
   outcomes,
   parameters,
   modelError,
```

```
initialConditions,
  odeSolverParameters
)
```

Arguments

.Object .Object name name description description equations equations outcomes outcomes parameters parameters modelError modelError initial ConditionsinitialConditions odeSolverParameters

odeSolverParameters

Value

ModelInfusion

```
initialize, {\tt ModelParameter-method}\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'ModelParameter'
initialize(.Object, name, distribution, fixedMu, fixedOmega)
```

Arguments

.Object .Object name name distribution distribution fixedMu fixedOmega fixedOmega

Value

ModelParameter

```
initialize, \verb|MultiplicativeAlgorithm-method|\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'MultiplicativeAlgorithm'
initialize(
   .Object,
   arms,
   lambda,
   delta,
   numberOfIterations,
   weightThreshold,
   optimalWeights,
   optimalDesign,
   showProcess
)
```

Arguments

```
.Object
                .Object
                arms
arms
lambda
                lambda
delta
                delta
numberOfIterations
                numberOfIterations
weightThreshold
                weightThreshold
optimalWeights optimalWeights
optimalDesign
                optimalDesign
showProcess
                showProcess
```

Value

MultiplicativeAlgorithm

Description

initialize

Usage

```
## S4 method for signature 'Normal'
initialize(.Object, ...)
```

Arguments

```
.Object .Object args
```

Value

Normal

```
initialize, {\tt Optimization-method}\\ initialize
```

Description

initialize

```
## S4 method for signature 'Optimization'
initialize(
   .Object,
   name,
   model,
   modelEquations,
   modelParameters,
   modelError,
   optimizer,
   optimizerParameters,
   outcomes,
   designs,
   fim,
```

```
odeSolverParameters,
optimizationResults,
evaluationFIMResults,
evaluationInitialDesignResults)
```

Arguments

.Object .Object name name model model

modelEquations modelEquations

modelParameters

modelParameters

modelError modelError optimizer optimizeroptimizerParameters

optimizerParameters

outcomes outcomes designs designs fim fim odeSolverParameters

odeSolverParameters

optimizationResults

optimizationResults

evaluationFIMResults

evaluation FIMR esults

 $\verb|evaluationInitialDesignResults|\\$

evaluation Initial Design Results

Value

Optimization

```
initialize, Optimization Algorithm-method\\ initialize
```

Description

initialize

```
## S4 method for signature 'OptimizationAlgorithm'
initialize(.Object, name, parameters)
```

Arguments

.Object .Object name

parameters parameters

Value

OptimizationAlgorithm

```
initialize, {\tt PFIMProject-method}\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'PFIMProject'
initialize(.Object, name, description)
```

Arguments

.Object .Object name

description description

Value

PFIMProject

```
initialize, {\tt PGBOAlgorithm-method}\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'PGBOAlgorithm'
initialize(
   .Object,
   N,
   muteEffect,
   maxIteration,
   purgeIteration,
   seed,
   showProcess,
   optimalDesign,
   iterationAndCriteria
)
```

Arguments

.Object .Object N muteEffect muteEffect maxIteration maxIteration purgeIteration purgeIteration seed seed showProcess showProcess optimalDesign optimalDesign iteration And CriteriaiterationAndCriteria

Value

PGBOAlgorithm

```
initialize, {\tt Proportional-method} \\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'Proportional'
initialize(
   .Object,
   outcome,
   equation,
   derivatives,
   sigmaInter,
   sigmaSlope,
   cError
)
```

Arguments

```
.Object .Object
outcome outcome
equation equation
derivatives derivatives
sigmaInter sigmaSlope
sigmaSlope cError cError
```

Value

Proportional

```
initialize, {\tt PSOAlgorithm-method}\\ initialize
```

Description

initialize

```
## S4 method for signature 'PSOAlgorithm'
initialize(
   .Object,
   maxIteration,
   populationSize,
   personalLearningCoefficient,
   globalLearningCoefficient,
   seed,
   showProcess,
```

```
optimalDesign,
  iterationAndCriteria
)
```

Arguments

```
.Object
                 .Object
                 maxIteration
maxIteration
populationSize populationSize
personalLearningCoefficient
                 personal Learning Coefficient \\
{\tt globalLearningCoefficient}
                 global Learning Coefficient \\
seed
                 seed
showProcess
                 showProcess
optimalDesign
                 optimalDesign
iteration And Criteria
                 iterationAndCriteria
```

Value

PSOAlgorithm

```
initialize, {\tt SamplingTimeConstraints-method}\\ initialize
```

Description

initialize

```
## S4 method for signature 'SamplingTimeConstraints'
initialize(
   .Object,
   outcome,
   initialSamplings,
   fixedTimes,
   numberOfsamplingsOptimisable,
   samplingsWindows,
   numberOfTimesByWindows,
   minSampling
)
```

Arguments

.Object .Object outcome

initialSamplings

initialSamplings

fixedTimes
numberOfsamplingsOptimisable

numberOfsamplingsOptimisable

samplingsWindows

samplingsWindows

numberOfTimesByWindows

number Of Times By Windows

minSampling minSampling

Value

SamplingTimeConstraints

Description

initialize

Usage

```
## S4 method for signature 'SamplingTimes'
initialize(.Object, outcome, samplings)
```

Arguments

.Object .Object outcome samplings samplings

Value

SamplingTimes

```
initialize, {\tt SimplexAlgorithm-method}\\ initialize
```

Description

initialize

Usage

```
## S4 method for signature 'SimplexAlgorithm'
initialize(
   .Object,
   pctInitialSimplexBuilding,
   maxIteration,
   tolerance,
   optimalDesigns,
   iterationAndCriteria,
   showProcess
)
```

Arguments

Value

SimplexAlgorithm

isModelAnalytic

isDoseInEquations

Test if the dose is in the equations of the model.

Description

Test if the dose is in the equations of the model.

Usage

```
isDoseInEquations(object)
## S4 method for signature 'Model'
isDoseInEquations(object)
```

Arguments

object

An object from the class Model.

Value

Return a Boolean giving if the dose is in the equations of the model.

is Model Analytic

Test if a mode is analytic.

Description

Test if a mode is analytic.

Usage

```
isModelAnalytic(object)
## S4 method for signature 'Model'
isModelAnalytic(object)
```

Arguments

object

An object from the class Model.

Value

Return a Boolean giving if the mode is analytic or not.

isModelBolus 127

isModelBolus

Test if a mode is bolus.

Description

Test if a mode is bolus.

Usage

```
isModelBolus(object, designs)
## S4 method for signature 'Model'
isModelBolus(object, designs)
```

Arguments

object An object from the class Model.

designs A list of objects from the class Design.

Value

Return a Boolean giving if the mode is bolus or not.

is Model Infusion

Test if a mode is infusion

Description

Test if a mode is infusion

Usage

```
isModelInfusion(object)
## S4 method for signature 'Model'
isModelInfusion(object)
```

Arguments

object

An object from the class Model.

Value

Return a Boolean giving if the mode is infusion or not.

128 isModelSteadyState

isModelODE

Test if a mode is ode.

Description

Test if a mode is ode.

Usage

```
isModelODE(object)
## S4 method for signature 'Model'
isModelODE(object)
```

Arguments

object

An object from the class Model.

Value

Return a Boolean giving if the mode is ode or not.

isModelSteadyState

Test if a mode is steady state.

Description

Test if a mode is steady state.

Usage

```
isModelSteadyState(object)
## S4 method for signature 'Model'
isModelSteadyState(object)
```

Arguments

object

An object from the class Model.

Value

Return a Boolean giving if the mode is steady state or not.

LibraryOfModels-class 129

LibraryOfModels-class Class "LibraryOfModels"

Description

The class LibraryOfModels represents the library of models.

Objects from the class

Objects form the class LibraryOfModels can be created by calls of the form LibraryOfModels(...) where (...) are the parameters for the LibraryOfModels objects.

Slots for LibraryOfModels **objects**

name: A string giving the name of the library of models.

content: A list giving the content of the library of model.

LibraryOfPDModels

Library of the PK models

Description

Library of the PK models

Usage

LibraryOfPDModels()

LibraryOfPKModels

Library of the PK models

Description

Library of the PK models

Usage

LibraryOfPKModels()

Model-class

LibraryOfPKPDModels-class

Class "LibraryOfPKPDModels"

Description

The class LibraryOfPKPDModels represents the library of PKPD models. The class LibraryOfPKPDModels inherits from the class LibraryOfModels.

LogNormal-class

Class "LogNormal"

Description

The class defines all the required methods for a LogNormal distribution object. The class LogNormal inherits from the class Distribution.

Model-class

Class "Model"

Description

The class Model defines information concerning the construction of a model.

Objects from the class

Objects form the class Model can be created by calls of the form Model(...) where (...) are the parameters for the Model objects.

Slots for Administration objects

name: A string giving the name of the model.

description: A list of string giving the description of the model.

equations: A list giving the equations of the model.

outcomes: A list giving the outcomes of the model.

outcomesForEvaluation: A list giving the outcomes used for the evaluation of the model.

parameters: A list giving the parameters of the model.

modelError: A list giving the model error of the model.

initialConditions: A list giving the initial conditions of the model.

odeSolverParameters: A list giving the parameters for the solver of the model.

modelFromLibrary: A list giving the model equations when the model is constructed from the library of model.

ModelAnalytic-class 131

ModelAnalytic-class Class "ModelAnalytic"

Description

The class Model defines information concerning the construction of an analytical model. The class ModelAnalytic inherits from the class Model.

ModelAnalyticBolus-class

Class "ModelAnalyticBolus"

Description

The class Model defines information concerning the construction of an analytical bolus model. The class ModelAnalyticBolus inherits from the class ModelAnalytic.

 ${\tt ModelAnalyticBolusSteadyState-class}$

Class "ModelAnalyticBolusSteadyState"

Description

The class Model defines information concerning the construction of an analytical model in steady state. The class ModelAnalyticBolusSteadyState inherits from the class ModelAnalyticSteadyState.

ModelAnalyticInfusion-class

Class "ModelAnalyticInfusion"

Description

The class Model defines information concerning the construction of an analytical model in infusion. The class ModelAnalyticInfusion inherits from the class ModelInfusion.

132 ModelInfusion-class

 ${\tt ModelAnalyticInfusionSteadyState-class}$

Class "ModelAnalyticInfusionSteadyState"

Description

The class Model defines information concerning the construction of an analytical model in infusion in steady state. The class ModelAnalyticInfusionSteadyState inherits from the class ModelAnalyticInfusion.

ModelAnalyticSteadyState-class

Class "ModelAnalyticSteadyState"

Description

The class ModelAnalyticSteadyState defines information concerning the construction of an analytical model steady state. The class ModelAnalyticSteadyState inherits from the class ModelAnalytic.

ModelBolus-class	Class "ModelBolus"	

Description

•••

ModelError-class	Class "ModelError" representing a Model error.
------------------	--

Description

...

ModelInfusion-class Class "ModelInfusion"

Description

...

ModelODE-class 133

ModelODE-class

Class "ModelODE"

Description

The class ModelODE defines information concerning the construction of an ode model. The class ModelODE inherits from the class Model.

ModelODEDoseInEquations-class

Class "ModelODEDoseInEquations"

Description

The class ModelODEDoseInEquations defines information concerning the construction of an ode model where the dose is in the model equations. The class ModelODEDoseInEquations inherits from the class ModelODE.

ModelODEDoseNotInEquations-class

Class "ModelODEDoseNotInEquations"

Description

...

ModelODEInfusion-class

Class "ModelODEInfusion"

Description

The class ModelODEInfusion defines information concerning the construction of an ode model in infusion. The class ModelODEInfusion inherits from the class ModelInfusion.

ModelODEInfusionDoseInEquations-class

Class "ModelODEInfusionDoseInEquations"

Description

The class ModelODEInfusionDoseInEquations defines information concerning the construction of an ode model in infusion where the dose is in the model equations. The class ModelODEInfusionDoseInEquations inherits from the class ModelODEInfusion.

ModelParameter-class Class "ModelParameter"

Description

The class ModelParameter defines information concerning the model parameters.

Objects from the class

Objects form the class ModelParameter can be created by calls of the form ModelParameter(...) where (...) are the parameters for the ModelParameter objects.

Slots for ModelParameter objects

name: A string giving the name of the parameter.

distribution: An object from the class Distribution giving the distribution of the parameter.

fixedMu: A boolean giving if mu is fixed or not.

fixedOmega: A boolean giving if omega is fixed or not.

MultiplicativeAlgorithm-class

Class "MultiplicativeAlgorithm"

Description

The class MultiplicativeAlgorithm implements the multiplicative algorithm.

Objects from the class

Objects form the class MultiplicativeAlgorithm can be created by calls of the form MultiplicativeAlgorithm(...) where (...) are the parameters for the MultiplicativeAlgorithm objects.

Slots for MultiplicativeAlgorithm objects

```
arms: A list giving the arms.

lambda: A numeric giving the lambda parameter of the multiplicative algorithm.

delta: A numeric giving the delta parameter of the multiplicative algorithm.

numberOfIterations: A numeric giving the maximal number iteration of the optimization process.

weightThreshold: A numeric giving the threshold of the weights.

optimalWeights: A vector giving the optimal weights.

optimalDesign: An object of the class Design giving the optimal design.

showProcess: A boolean for showing or not the process of optimization.
```

MultiplicativeAlgorithm_Rcpp

Function MultiplicativeAlgorithm_Rcpp

Description

Run the MultiplicativeAlgorithm_Rcpp in Rcpp

Usage

```
MultiplicativeAlgorithm_Rcpp(
   fisherMatrices_input,
   numberOfFisherMatrices_input,
   weights_input,
   numberOfParameters_input,
   dim_input,
   lambda_input,
   delta_input,
   iterationInit_input
)
```

Arguments

```
fisherMatrices_input
fisherMatrices_input
numberOfFisherMatrices_input
numberOfFisherMatrices_input
weights_input weights_input
numberOfParameters_input
numberOfParameters_input
dim_input dim_input
lambda_input lambda_input
```

136 Optimization-class

delta_input delta_input
iterationInit_input
 iterationInit_input

Normal-class

Class "Normal"

Description

The class defines all the required methods for a Normal distribution object. The class Normal inherits from the class Distribution.

Optimization-class

Class "Optimization"

Description

A class storing information concerning the design optimization.

Objects from the class

Objects form the class Optimization can be created by calls of the form Optimization(...) where (...) are the parameters for the Optimization objects.

Slots for Administration objects

name: A character string giving the name of the optimization process.

model: A object of class Model giving the model.

modelEquations: A list giving the model equations.

modelParameters: A list giving the model parameters.

modelError: A list giving the model error.

optimizer: A object of class OptimizationAlgorithm giving the optimization algorithm.

optimizerParameters: A list giving the parameters of the optimization algorithm.

outcomes: A list giving the outcomes of the model.

designs: A list giving the designs to be optimized.

fim: A object of class FIM giving the Fisher information matrix.

odeSolverParameters: A list giving the parameters for the ode solver.

optimizationResults: A object of class OptimizationAlgorithm giving the results of the optimization.

evaluationFIMResults: A object of class Evaluation giving the results of the evaluation of the optimal design.

evaluationInitialDesignResults: A object of class Evaluation giving the results of the evaluation of the initial design.

```
OptimizationAlgorithm-class

Class "OptimizationAlgorithm"
```

Description

A class storing information concerning the optimization algorithm.

Objects from the class

Objects form the class OptimizationAlgorithm can be created by calls of the form OptimizationAlgorithm(...) where (...) are the parameters for the OptimizationAlgorithm objects.

Slots for Administration objects

```
name: A character string giving the name of the optimization algorithm. parameters: A list giving the parameters of the optimization algorithm.
```

optimize

Optimize a design.

Description

Optimize a design.

```
optimize(object, optimizerParameters, optimizationObject)
## S4 method for signature 'FedorovWynnAlgorithm'
optimize(object, optimizerParameters, optimizationObject)
## S4 method for signature 'MultiplicativeAlgorithm'
optimize(object, optimizerParameters, optimizationObject)
## S4 method for signature 'PGBOAlgorithm'
optimize(object, optimizationObject)
## S4 method for signature 'PSOAlgorithm'
optimize(object, optimizationObject)
## S4 method for signature 'SimplexAlgorithm'
optimize(object, optimizerParameters, optimizationObject)
```

Arguments

Value

A list giving the results if the optimization.

```
parametersForComputingGradient
```

Define the parameters for computing the gradients of a model.

Description

Define the parameters for computing the gradients of a model.

Usage

```
parametersForComputingGradient(object, valuePars)
## S4 method for signature 'Model'
parametersForComputingGradient(object, valuePars)
```

Arguments

object An object from the class Model.

valuePars Vector of parameter values

Value

A list giving the parameters for computing the gradients of a model.

PFIMProject-class 139

PFIMProject-class

Class "PFIMProject"

Description

A class storing information concerning a PFIM project.

Objects from the class

Objects form the class PFIMProject can be created by calls of the form PFIMProject(...) where (...) are the parameters for the PFIMProject objects.

Slots for PFIMProject objects

name: A character string giving the name of the PFIM project. description: A list giving the description of the PFIM project.

PGBOAlgorithm-class

Class "PGBOAlgorithm"

Description

The class "PGBOAlgorithm" implements the PGBO algorithm: Population Genetics Based Optimizer, developed by Hervé Le Nagard [1].

Objects from the Class PGBOAlgorithm

Objects form the Class PGBOAlgorithm can be created by calls of the form PGBOAlgorithm(...) where (...) are the parameters for the PGBOAlgorithm objects.

Slots for PGBOAlgorithm objects

N: A numeric giving the population size.

muteEffect: A numeric giving the mutation effect.

maxIteration: A numeric giving the maximum number of iterations.

seed: A numeric giving the seed.

showProcess: A boolean to show or not the process.

optimalDesign: A Design object giving the optimal design.

iterationAndCriteria: A list giving the optimal criteria at each iteration.

References

[1] Rebecca Bauer, France Mentré, Halima Kaddouri, Jacques Le Bras, Hervé Le Nagard, Benefits of a new Metropolis-Hasting based algorithm, in non-linear regression for estimation of ex vivo antimalarial sensitivity in patients infected with two strains, Computers in Biology and Medicine, Volume 55, 2014, Pages 16-25, ISSN 0010-4825

plotFrequencies

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Graphs of the results of the evaluation.

Description

Graphs of the results of the evaluation.

Usage

```
plotEvaluation(object, plotOptions)
## S4 method for signature 'Evaluation'
plotEvaluation(object, plotOptions)
```

Arguments

object An object from the class Evaluation.

plotOptions A list giving the plot options.

Value

A list giving the graphs for the evaluation of the responses and sensitivity indices.

PlotEvaluation-class Class "PlotEvaluation"

Description

A class storing information concerning the design evaluation. The class PlotEvaluation inherits from the class Evaluation.

plotFrequencies

Graph of the frequencies for the FW algorithm.

Description

Graph of the frequencies for the FW algorithm.

plotOutcomesEvaluation 141

Usage

```
plotFrequencies(object)
## S4 method for signature 'FedorovWynnAlgorithm'
plotFrequencies(object)
## S4 method for signature 'Optimization'
plotFrequencies(object)
```

Arguments

object

An object from the class OptimizationAlgorithm.

Value

The graphs of the frequencies for the FW algorithm.

```
plotOutcomesEvaluation
```

plotOutcomesEvaluation

Description

Plot the evaluation of the outcomes.

Usage

```
plotOutcomesEvaluation(
   object,
   outcomesEvaluationInitialDesign,
   model,
   plotOptions
)

## S4 method for signature 'Design'
plotOutcomesEvaluation(
   object,
   outcomesEvaluationInitialDesign,
   model,
   plotOptions
)
```

Arguments

```
object An object Design from the class Design. outcomesEvaluationInitialDesign
```

A list containing the evaluation of the initial design.

142 plotRSE

```
model An object model from the class Model.
plotOptions A list containing the plot options.
```

Value

A list containing the plots the evaluation of the outcomes.

```
plotOutcomesGradient plotOutcomesGradient
```

Description

Plot the evaluation of the outcome gradients.

Usage

```
plotOutcomesGradient(object, outcomesGradientInitialDesign, model, plotOptions)
## S4 method for signature 'Design'
plotOutcomesGradient(object, outcomesGradientInitialDesign, model, plotOptions)
```

Arguments

object An object design from the class Design.

outcomesGradientInitialDesign

A list with the evaluation of the gradient for the initial design.

model An object model from the class Model.
plotOptions A list containing the plot options.

Value

A list containing the plots the evaluation of the outcome gradients..

plotRSE Graph of the RSE.

Description

Graph of the RSE.

```
plotRSE(object, plotOptions)
## S4 method for signature 'PFIMProject'
plotRSE(object, plotOptions)
```

plotSE 143

Arguments

object An object from the class Evaluation.

Value

A graph of the RSE.

plotOptions

plotSE

Graph the SE.

A list giving the plot options.

Description

Graph the SE.

Usage

```
plotSE(object, plotOptions)
## S4 method for signature 'PFIMProject'
plotSE(object, plotOptions)
```

Arguments

object An object from the class Evaluation.

plotOptions A list giving the plot options.

Value

A graph of the SE.

plotSensitivityIndice Graphs of the results of the evaluation.

Description

Graphs of the results of the evaluation.

```
plotSensitivityIndice(object, plotOptions)
## S4 method for signature 'Evaluation'
plotSensitivityIndice(object, plotOptions)
```

144 plotWeights

Arguments

object An object from the class Evaluation.
plotOptions A list giving the plot options.

Value

A list giving the graphs for the evaluation of the responses and sensitivity indices.

plotShrinkage

Graph of the shrinkage.

Description

Graph of the shrinkage.

Usage

```
plotShrinkage(object, plotOptions)
## S4 method for signature 'PFIMProject'
plotShrinkage(object, plotOptions)
```

Arguments

object An object from the class Evaluation.
plotOptions A list giving the plot options.

Value

A graph of the shrinkage.

plotWeights

Graph of the weights for the multiplicative algorithm.

Description

Graph of the weights for the multiplicative algorithm.

```
plotWeights(object)
## S4 method for signature 'MultiplicativeAlgorithm'
plotWeights(object)
## S4 method for signature 'Optimization'
plotWeights(object)
```

PopulationFim-class 145

Arguments

object

An object from the class OptimizationAlgorithm.

Value

The graphs of the weights for the multiplicative algorithm.

PopulationFim-class

Class "PopulationFim"

Description

A class storing information regarding the population Fisher matrix. The class PopulationFim inherits from the class Fim.

Proportional-class

Class "Proportional"

Description

The Class "Proportional" defines the tresidual error variance according to the formula $g(sigma_inter, sigma_slope, c_error, f(x, theta)) = sigma_slope*f(x, theta).$

Objects from the Class Proportional

Objects are typically created by calls to Proportional and contain the following slots that are inherited from the class Combined1:

Slots for the Proportional objects

.Object: An object of the Class Proportional

sigma_inter: A numeric value giving the sigma inter of the error model

sigma_slope: A numeric value giving the sigma slope of the error model

Report Report

PSOAlgorithm-class

Class "PSOAlgorithm"

Description

The class "PSOAlgorithm" implements the PSO algorithm.

Objects from the class PSOAlgorithm

Objects form the class PSOAlgorithm can be created by calls of the form PSOAlgorithm(...) where (...) are the parameters for the PSOAlgorithm objects.

Slots for PSOAlgorithm objects

maxIteration: A numeric giving the maximum of iterations.

populationSize: A numeric giving the population size.

seed: A numeric giving the seed.

personalLearningCoefficient: A numeric giving the personal learning coefficient.

globalLearningCoefficient: A numeric giving the global learning coefficient.

showProcess: A boolean to show or not the process.

optimalDesign: A Design object giving the optimal design.

iterationAndCriteria: A list giving the optimal criteria at each iteration.

Report

Report

Description

Report

Usage

```
Report(object, outputPath, outputFile, plotOptions)
## S4 method for signature 'Evaluation'
Report(object, outputPath, outputFile, plotOptions)
## S4 method for signature 'Optimization'
Report(object, outputPath, outputFile, plotOptions)
```

Arguments

object An object from the class PFIMProject.

outputPath A string giving the output path.

outputFile A string giving the name of the output file.

plotOptions A list giving the plot options.

Value

The report in html.

reportTablesAdministration

report Tables Administration

Description

Generate table for the report.

Usage

```
reportTablesAdministration(object)
## S4 method for signature 'Design'
reportTablesAdministration(object)
```

Arguments

object An object design from the class Design.

Value

A table of the administration parameters for the report.

reportTablesDesign reportTablesDesign

Description

Generate table for the report.

Usage

```
reportTablesDesign(object)
## S4 method for signature 'Design'
reportTablesDesign(object)
```

148 reportTablesFIM

Arguments

object An object design from the class Design.

Value

A table of the design parameters for the report.

reportTablesFIM

Generate the tables for the report.

Description

Generate the tables for the report.

Usage

```
reportTablesFIM(object, evaluationObject)
## S4 method for signature 'BayesianFim'
reportTablesFIM(object, evaluationObject)
## S4 method for signature 'IndividualFim'
reportTablesFIM(object, evaluationObject)
## S4 method for signature 'PopulationFim'
reportTablesFIM(object, evaluationObject)
```

Arguments

```
object An object from the class Fim. evaluationObject
```

A list giving the results of the evaluation of the model.

Value

A list giving the table in kable format for the report.

reportTablesModelError

Generate the tables for model errors for the evaluation report.

Description

Generate the tables for model errors for the evaluation report.

Usage

```
reportTablesModelError(object)
## S4 method for signature 'Model'
reportTablesModelError(object)
```

Arguments

object

An object from the class Model.

Value

A kable table for the evaluation report.

```
{\tt reportTablesModelParameters}
```

Generate the tables for model parameters for the evaluation report.

Description

Generate the tables for model parameters for the evaluation report.

Usage

```
reportTablesModelParameters(object)
## S4 method for signature 'Model'
reportTablesModelParameters(object)
```

Arguments

object

An object from the class Model.

Value

A kable table for the evaluation report.

reportTablesPlot

reportTablesPlot

Description

Generate all the table for the evaluation report

Usage

```
reportTablesPlot(object, plotOptions)
## S4 method for signature 'Evaluation'
reportTablesPlot(object, plotOptions)
```

Arguments

object An object evaluation from the class Evaluation.

plotOptions A list containing the options for the plots.

Value

The list tables containing the tables for the evaluation report.

```
report Tables Sampling Constraints \\ report Tables Sampling Constraints
```

Description

Generate table for the report.

Usage

```
reportTablesSamplingConstraints(object)
## S4 method for signature 'Design'
reportTablesSamplingConstraints(object)
```

Arguments

object An object design from the class Design.

Value

A table of the sampling constraints parameters for the report.

resizeFisherMatrix 151

resizeFisherMatrix

Resize the fisher Matrix from a vector to a matrix.

Description

Resize the fisher Matrix from a vector to a matrix.

Usage

```
resizeFisherMatrix(nbOfDimensions, fisherMatrix)
## S4 method for signature 'ANY'
resizeFisherMatrix(nbOfDimensions, fisherMatrix)
```

Arguments

nbOfDimensions : a numeric for the dimensions of the fisher matrix.

fisherMatrix : a vector that contain the low triangular Fisher matrix + its main diagonal.

Value

The Fisher matrix of size nbOfDimensions*nbOfDimensions

run run

Description

run

Usage

```
run(object)
## S4 method for signature 'Evaluation'
run(object)
## S4 method for signature 'Optimization'
run(object)
```

Arguments

object An object from the class PFIMProject.

Value

A list giving the results of evaluation or optimization.

152 SamplingTimes-class

SamplingTimeConstraints-class

Class "SamplingTimeConstraints"

Description

The class "SamplingTimeConstraints" implements the constraints for the sampling times.

Objects from the class SamplingTimeConstraints

Objects form the class SamplingTimeConstraints can be created by calls of the form SamplingTimeConstraints(...) where (...) are the parameters for the SamplingTimeConstraints objects.

Slots for SamplingTimeConstraints objects

outcome: A string giving the outcome.

initialSamplings: A vector giving the sampling times.

fixedTimes: A vector giving the fixed sampling times.

numberOfsamplingsOptimisable: A vector giving the sampling times to be optimized.

samplingsWindows: A list giving the windows for the sampling times.

numberOfTimesByWindows: A vector giving the number of sampling times by windows.

minSampling: A numeric giving the minimal sampling times.

SamplingTimes-class Class "SamplingTimes"

Description

The class "SamplingTimes" implements the sampling times.

Objects from the class SamplingTimes

Objects form the class SamplingTimes can be created by calls of the form SamplingTimes(...) where (...) are the parameters for the SamplingTimes objects.

Slots for SamplingTimes objects

outcome: A string giving the outcome.

samplings: A vector giving the sampling times.

setAdministrations 153

setAdministrations

setAdministrations

Description

Set all the administration for an arm.

Usage

```
setAdministrations(object, administrations)
## S4 method for signature 'Arm'
setAdministrations(object, administrations)
```

Arguments

object Arm from the class Arm. administrations

A list administrations of objects from the class Administration class giving the parameters of the administration for the object Arm.

Value

The object Arm with the list administrations of objects from the class Administration class giving the parameters of the administration for the object Arm.

setArm

setArm

Description

Set the arms in a design.

Usage

```
setArm(object, arm)
## S4 method for signature 'Design'
setArm(object, arm)
```

Arguments

object An object Design from the class Design.

arm A list of object Arm giving the arms of the design.

Value

An object Design with the list Arm updated.

154 setcError

setArms

Set the arms of an object.

Description

Set the arms of an object.

Usage

```
setArms(object, arms)
## S4 method for signature 'Design'
setArms(object, arms)
## S4 method for signature 'OptimizationAlgorithm'
setArms(object, arms)
```

Arguments

object An object defined form a class of PFIM.

arms A list of arms.

Value

The object with the updated arms.

setcError

Set the parameter c.

Description

Set the parameter c.

Usage

```
setcError(object, cError)
## S4 method for signature 'ModelError'
setcError(object, cError)
```

Arguments

object An object from the class ModelError. cError A numeric giving the parameter c.

setContent 155

Value

The model error with the parameter c.

setContent

Set content of a library of models.

Description

Set content of a library of models.

Usage

```
setContent(object, content)
## S4 method for signature 'LibraryOfModels'
setContent(object, content)
```

Arguments

object An object from the class LibraryOfModels.

content A list giving the content of the library of models.

Value

The library of models with the updated content.

```
setDataForArmEvaluation
```

setDataForArmEvaluation

Description

setDataForArmEvaluation

Usage

```
setDataForArmEvaluation(object, data)
## S4 method for signature 'Arm'
setDataForArmEvaluation(object, data)
```

Arguments

object An object Arm from the class Arm.

data A list containing the data for arm evaluation

Value

Set the list containing the data for arm evaluation.

setDataForModelEvaluation

Generate the table of dose, time dose etc. for model evaluation

Description

Generate the table of dose, time dose etc. for model evaluation

Usage

```
setDataForModelEvaluation(object, arm)
## S4 method for signature 'ModelAnalytic'
setDataForModelEvaluation(object, arm)
## S4 method for signature 'ModelAnalyticSteadyState'
setDataForModelEvaluation(object, arm)
## S4 method for signature 'ModelAnalyticInfusion'
setDataForModelEvaluation(object, arm)
## S4 method for signature 'ModelAnalyticInfusionSteadyState'
setDataForModelEvaluation(object, arm)
## S4 method for signature 'ModelODEBolus'
setDataForModelEvaluation(object, arm)
## S4 method for signature 'ModelODEDoseInEquations'
setDataForModelEvaluation(object, arm)
## S4 method for signature 'ModelODEDoseNotInEquations'
setDataForModelEvaluation(object, arm)
## S4 method for signature 'ModelODEInfusionDoseInEquations'
setDataForModelEvaluation(object, arm)
```

Arguments

object An object from the class Model.

An object from the class Arm.

Value

Return a dataframe with all the data for model evaluation

setDerivatives 157

setDerivatives

Set the derivatives of the model error equation.

Description

Set the derivatives of the model error equation.

Usage

```
setDerivatives(object, derivatives)
## S4 method for signature 'ModelError'
setDerivatives(object, derivatives)
```

Arguments

object An object from the class ModelError.

derivatives The derivatives of the model error equation.

Value

The model error with the updated model error equation.

setDescription

Set the description of a model.

Description

Set the description of a model.

Usage

```
setDescription(object, description)
## S4 method for signature 'Model'
setDescription(object, description)
```

Arguments

object An object from the class Model.

description A list giving the description of a model.

Value

The model with the updated description.

158 setDistribution

 $\operatorname{setDesigns}$

Set the designs.

Description

Set the designs.

Usage

```
setDesigns(object, designs)
## S4 method for signature 'Optimization'
setDesigns(object, designs)
```

Arguments

object An object from the class Optimization.
designs A list of objects from the class Design.

Value

The object with the new designs.

setDistribution

Set the distribution.

Description

Set the distribution.

Usage

```
setDistribution(object, distribution)
## S4 method for signature 'ModelParameter'
setDistribution(object, distribution)
```

Arguments

object An object from the class ModelParameter.
distribution An object from the class Distribution.

Value

The model parameter with the updated distribution.

setDose 159

setDose

Set the amount of dose

Description

Set the amount of dose

Usage

```
setDose(object, dose)
## S4 method for signature 'Administration'
setDose(object, dose)
```

Arguments

object An object Administration from the class Administration.

dose A numeric value of the amount of dose.

Value

The numeric amount_dose giving the new value of the amount of dose.

setEquation

Set the equation of a model error.

Description

Set the equation of a model error.

Usage

```
setEquation(object, equation)
## S4 method for signature 'ModelError'
setEquation(object, equation)
```

Arguments

object An object from the class ModelError.

equation An expression giving the equation of a model error.

Value

The model error with the updated equation.

setEquations

Set the equations of a model.

Description

Set the equations of a model.

Usage

```
setEquations(object, equations)
## S4 method for signature 'Model'
setEquations(object, equations)
```

Arguments

object An object from the class Model.

equations A list giving the equations of the model.

Value

The model with the updated equations.

```
setEquationsAfterInfusion
```

Set the equations after infusion.

Description

Set the equations after infusion.

Usage

```
setEquationsAfterInfusion(object, equations)
## S4 method for signature 'Model'
setEquationsAfterInfusion(object, equations)
```

Arguments

object An object from the class Model.

equations A list giving the equations after the infusion.

Value

The model with the updated equations after the infusion.

setEquationsDuringInfusion

Set the equations during infusion.

Description

Set the equations during infusion.

Usage

```
setEquationsDuringInfusion(object, equations)
## S4 method for signature 'Model'
setEquationsDuringInfusion(object, equations)
```

Arguments

object An object from the class Model.

equations A list giving the equations during the infusion.

Value

The model with the updated equations during the infusion.

```
setEvaluationFIMResults
```

Set the evaluation results.

Description

Set the evaluation results.

Usage

```
setEvaluationFIMResults(object, value)
## S4 method for signature 'Optimization'
setEvaluationFIMResults(object, value)
```

Arguments

object An object from the class Optimization.

value An object from the class Evaluation giving the evaluation results.

Value

The object with the updated object from the class Evaluation.

162 setFim

```
setEvaluationInitialDesignResults
```

Set the evaluation results of the initial design.

Description

Set the evaluation results of the initial design.

Usage

```
setEvaluationInitialDesignResults(object, value)
## S4 method for signature 'Optimization'
setEvaluationInitialDesignResults(object, value)
```

Arguments

object An object from the class Optimization.

value An object from the class Evaluation giving the evaluation results of the initial

design.

Value

The object with the updated object from the class Evaluation.

setFim setFim

Description

Set the fim of the design.

Usage

```
setFim(object, fim)
## S4 method for signature 'Design'
setFim(object, fim)
```

Arguments

object An object Design from the class Design.
fim An object fim from the class Fim.

Value

An object Design with the fim updated.

setFimTypeToString 163

 ${\tt setFimTypeToString}$

Convert the type of the object fim to a string.

Description

Convert the type of the object fim to a string.

Usage

```
setFimTypeToString(object)
## S4 method for signature 'Fim'
setFimTypeToString(object)
```

Arguments

object

An object from the class Fim.

Value

The type of the object fim convert as a string.

setFisherMatrix

Set the FIM.

Description

Set the FIM.

Usage

```
setFisherMatrix(object, value)
## S4 method for signature 'Fim'
setFisherMatrix(object, value)
```

Arguments

object An object from the class Fim. value A matrix giving the FIM.

Value

The object from the class Fim with the FIM updated.

164 setFixedMu

setFixedEffects

Set the fixed effects.

Description

Set the fixed effects.

Usage

```
setFixedEffects(object)
## S4 method for signature 'Fim'
setFixedEffects(object)
```

Arguments

object

An object from the class Fim.

Value

Update the matrix of the fixed effects.

setFixedMu

Set the mu as fixed or not.

Description

Set the mu as fixed or not.

Usage

```
setFixedMu(object, value)
## S4 method for signature 'ModelParameter'
setFixedMu(object, value)
```

Arguments

object An object from the class ModelParameter.

value A Boolean if fixed or not.

Value

The mode parameter with the mu updated as fixed or not.

setFixedOmega 165

setFixedOmega

Set the omega as fixed of not.

Description

Set the omega as fixed of not.

Usage

```
setFixedOmega(object, value)
## S4 method for signature 'ModelParameter'
setFixedOmega(object, value)
```

Arguments

object An object from the class ModelParameter.

value A Boolean fixed or not.

Value

The model parameter with the omega updated as fixed or not.

```
setInitialConditions setInitialConditions
```

Description

Set the initial conditions of a ode model.

Usage

```
setInitialConditions(object, initialConditions)
## S4 method for signature 'Arm'
setInitialConditions(object, initialConditions)
## S4 method for signature 'Model'
setInitialConditions(object, initialConditions)
```

Arguments

```
\begin{tabular}{ll} \beg
```

A list giving the initial conditions.

166 setModel

Value

The model with the updated initial conditions.

```
setIterationAndCriteria
```

Set the iteration with the convergence criteria.

Description

Set the iteration with the convergence criteria.

Usage

```
setIterationAndCriteria(object, value)
## S4 method for signature 'OptimizationAlgorithm'
setIterationAndCriteria(object, value)
```

Arguments

object An object from the class OptimizationAlgorithm.

value A dataframe giving the iteration with the convergence criteria.

Value

A dataframe giving the iteration with the convergence criteria.

setModel

Set the model.

Description

Set the model.

Usage

```
setModel(object, model)
## S4 method for signature 'PFIMProject'
setModel(object, model)
```

Arguments

object An object from the class PFIMProject.

model An object from the class Model.

setModelError 167

Value

The object with the updated model.

setModelError

Set the model error.

Description

Set the model error.

Usage

```
setModelError(object, modelError)
## S4 method for signature 'Model'
setModelError(object, modelError)
```

Arguments

object An object from the class Model.

modelError An object from the class ModelError.

Value

The model with the updated model error.

setModelFromLibrary

Set a model from the library of model

Description

Set a model from the library of model

Usage

```
setModelFromLibrary(object, modelFromLibrary)
## S4 method for signature 'Model'
setModelFromLibrary(object, modelFromLibrary)
```

Arguments

```
object An object from the class Model. modelFromLibrary
```

An object from the class Model.

168 setName

Value

The model with the updated model from library of models.

setMu

Set the value of the fixed effect mu of an object.

Description

Set the value of the fixed effect mu of an object.

Usage

```
setMu(object, value)
## S4 method for signature 'Distribution'
setMu(object, value)
## S4 method for signature 'ModelParameter'
setMu(object, value)
```

Arguments

object An object defined form a class of PFIM. value The value of the fixed effect mu.

Value

The object with the updated fixed effect mu.

setName

Set the name of an object.

Description

Set the name of an object.

Usage

```
setName(object, name)
## S4 method for signature 'Arm'
setName(object, name)
## S4 method for signature 'Design'
setName(object, name)
## S4 method for signature 'Model'
setName(object, name)
```

setNumberOfArms 169

Arguments

object An object defined form a class of PFIM.

name A string giving the name of the object.

Value

The object with the updated name.

setNumberOfArms

setNumberOfArms

Description

Set the number of arms in a design.

Usage

```
setNumberOfArms(object, numberOfArms)
## S4 method for signature 'Design'
setNumberOfArms(object, numberOfArms)
```

Arguments

object An object Design from the class Design.

numberOfArms A numeric numberOfArms giving the new number of arms in the design.

Value

An object Design with the numberOfArms updated.

setOdeSolverParameters

Set the parameters of the ode solver.

Description

Set the parameters of the ode solver.

Usage

```
setOdeSolverParameters(object, odeSolverParameters)
## S4 method for signature 'Model'
setOdeSolverParameters(object, odeSolverParameters)
```

170 setOmega

Arguments

```
object An object from the class Model.
odeSolverParameters
A list giving the parameters of the ode solver.
```

Value

The model with the updated parameters of the ode solver.

setOmega

Set the matrix omega of an object.

Description

Set the matrix omega of an object.

Usage

```
setOmega(object, value)
## S4 method for signature 'Distribution'
setOmega(object, value)
## S4 method for signature 'ModelParameter'
setOmega(object, value)
```

Arguments

object An object defined form a class of PFIM.

value The matrix omega.

Value

The object with the updated matrix omega.

setOptimalDesign 171

setOptimalDesign

Set the optimal design.

Description

Set the optimal design.

Usage

```
setOptimalDesign(object, optimalDesign)
## S4 method for signature 'OptimizationAlgorithm'
setOptimalDesign(object, optimalDesign)
```

Arguments

object An object from the class OptimizationAlgorithm.
optimalDesign An object from the class Design.

Value

The object with the updated optimal design.

setOptimalWeights

Set the optimal weights.

Description

Set the optimal weights.

Usage

```
setOptimalWeights(object, optimalWeights)
## S4 method for signature 'MultiplicativeAlgorithm'
setOptimalWeights(object, optimalWeights)
```

Arguments

```
object An object from the class MultiplicativeAlgorithm. optimalWeights A vector giving the optimal weights.
```

Value

The object with the updated optimal weights.

172 setOutcome

```
{\tt setOptimizationResults}
```

Set the optimization results.

Description

Set the optimization results.

Usage

```
setOptimizationResults(object, value)
## S4 method for signature 'Optimization'
setOptimizationResults(object, value)
```

Arguments

object An object from the class Optimization.

value An object from the class OptimizationAlgorithm giving the optimization results.

Value

The object with the updated object from the class OptimizationAlgorithm.

setOutcome setOutcome

Description

Set the outcome of an object.

Usage

```
setOutcome(object, outcome)
## S4 method for signature 'Administration'
setOutcome(object, outcome)
## S4 method for signature 'SamplingTimes'
setOutcome(object, outcome)
```

Arguments

object An object defined form a class of PFIM.

outcome A string defined the outcome.

setOutcomes 173

Value

A string giving the updated outcome of the object.

setOutcomes

Set the outcomes of a model.

Description

Set the outcomes of a model.

Usage

```
setOutcomes(object, outcomes)
## S4 method for signature 'Model'
setOutcomes(object, outcomes)
```

Arguments

object An object from the class Model.

outcomes A list giving the outcomes of the model.

Value

The model with the updated outcomes.

```
setOutcomesEvaluation setOutcomesEvaluation
```

Description

Set the results of the evaluation of the outcomes.

Usage

```
setOutcomesEvaluation(object, outcomesEvaluation)
## S4 method for signature 'Design'
setOutcomesEvaluation(object, outcomesEvaluation)
```

Arguments

```
\begin{array}{ll} \mbox{object} & \mbox{An object Design from the class } \mbox{Design.} \\ \mbox{outcomesEvaluation} \end{array}
```

A list containing the evaluation of the outcomes.

174 setOutcomesGradient

Value

An object Design with the list outcomesEvaluation updated.

```
setOutcomesForEvaluation
```

Set the outcomes of a model used for the evaluation (is scales outcomes).

Description

Set the outcomes of a model used for the evaluation (is scales outcomes).

Usage

```
setOutcomesForEvaluation(object, outcomes)
## S4 method for signature 'Model'
setOutcomesForEvaluation(object, outcomes)
```

Arguments

object An object from the class Model.

outcomes A list giving the outcomes of a model used for the evaluation (is scales out-

comes).

Value

The model with the updated outcomes for the evaluation.

setOutcomesGradient setOutcomesGradient

Description

Set the results of the evaluation of the outcomes.

Usage

```
setOutcomesGradient(object, outcomesGradient)
## S4 method for signature 'Design'
setOutcomesGradient(object, outcomesGradient)
```

setParameters 175

Arguments

```
object An object Design from the class Design.
outcomesGradient
A list containing the evaluation of the outcome gradients.
```

Value

An object Design with the list outcomesGradient updated.

setParameters

Set the parameters of an object.

Description

Set the parameters of an object.

Usage

```
setParameters(object, parameters)

## S4 method for signature 'Distribution'
setParameters(object, parameters)

## S4 method for signature 'Model'
setParameters(object, parameters)

## S4 method for signature 'FedorovWynnAlgorithm'
setParameters(object, parameters)

## S4 method for signature 'MultiplicativeAlgorithm'
setParameters(object, parameters)

## S4 method for signature 'PGBOAlgorithm'
setParameters(object, parameters)

## S4 method for signature 'PSOAlgorithm'
setParameters(object, parameters)

## S4 method for signature 'SimplexAlgorithm'
setParameters(object, parameters)
```

Arguments

object An object defined form a class of PFIM.

parameters A list of parameters.

176 setSamplings

Value

The object with the updated list of parameters.

```
set Sampling Constraint For Optimization \\ set Sampling Constraint For Optimization
```

Description

Set the sampling times constraint for optimization with PSO, PGBO and Simplex

Usage

```
setSamplingConstraintForOptimization(object)
## S4 method for signature 'Design'
setSamplingConstraintForOptimization(object)
```

Arguments

object An object from the class Design.

Value

The arms with the sampling times constraints.

setSamplings

Set the sampling times.

Description

Set the sampling times.

Usage

```
setSamplings(object, samplings)
## S4 method for signature 'SamplingTimes'
setSamplings(object, samplings)
```

Arguments

object An object from the class SamplingTimes. samplings A vector giving the sampling times.

Value

The updated sampling times.

setSamplingTime 177

setSamplingTime

setSamplingTime

Description

Set the sampling time of an arm.

Usage

```
setSamplingTime(object, samplingTime)
## S4 method for signature 'Arm'
setSamplingTime(object, samplingTime)
```

Arguments

object Arm from the class Arm.

samplingTime An object samplingTime from the class SamplingTimes.

Value

An object Arm from the class Arm with the new sampling time samplingTime.

setSamplingTimes

setSamplingTimes

Description

Set the vectors of sampling times for an arm.

Usage

```
setSamplingTimes(object, samplingTimes)
## S4 method for signature 'Arm'
setSamplingTimes(object, samplingTimes)
```

Arguments

object Arm from the class Arm.

samplingTimes The list containing the new sampling times.

Value

An object $\mbox{\it Arm}$ from the class $\mbox{\it Arm}$ with the new sampling times sampling $\mbox{\it Times}.$

178 setShrinkage

```
set Sampling Times Constraints \\ set Sampling Times Constraints
```

Description

Set the sampling times constraints.

Usage

```
setSamplingTimesConstraints(object, samplingTimesConstraints)
## S4 method for signature 'Arm'
setSamplingTimesConstraints(object, samplingTimesConstraints)
```

Arguments

```
object An object Arm from the class Arm.

samplingTimesConstraints

An object SamplingTimeConstraints from the class SamplingTimeConstraints.
```

Value

The arm with the new sampling time constraints.

setShrinkage

Set the shrinkage.

Description

Set the shrinkage.

Usage

```
setShrinkage(object, value)
## S4 method for signature 'BayesianFim'
setShrinkage(object, value)
## S4 method for signature 'IndividualFim'
setShrinkage(object, value)
## S4 method for signature 'PopulationFim'
setShrinkage(object, value)
```

setSigmaInter 179

Arguments

object An object from the class Fim.

value A vector giving the shrinkage of the Bayesian fim.

Value

The object with the updated shrinkage.

setSigmaInter

Set the parameter sigma inter.

Description

Set the parameter sigma inter.

Usage

```
setSigmaInter(object, sigmaInter)
## S4 method for signature 'ModelError'
setSigmaInter(object, sigmaInter)
```

Arguments

object An object from the class ModelError.

sigmaInter A numeric giving the parameter sigma inter.

Value

The model error with the updated sigma inter.

setSigmaSlope

Set the parameter sigma slope.

Description

Set the parameter sigma slope.

Usage

```
setSigmaSlope(object, sigmaSlope)
## S4 method for signature 'ModelError'
setSigmaSlope(object, sigmaSlope)
```

180 setSize

Arguments

object An object from the class ModelError.

sigmaSlope A numeric giving the parameter sigma slope.

Value

The model error with the updated sigma slope.

setSize

setSize

Description

Set the size of an object.

Set the size of an arm.

Usage

```
setSize(object, size)
setSize(object, size)
## S4 method for signature 'Arm'
setSize(object, size)
## S4 method for signature 'Design'
setSize(object, size)
```

Arguments

object Arm from the class Arm.

size A numeric giving the new size of the object Arm.

Value

The object with its size updated.

The object Arm object with its new size.

setTau 181

setTau setTau

Description

Set the frequency tau.

Usage

```
setTau(object, tau)
## S4 method for signature 'Administration'
setTau(object, tau)
```

Arguments

object An object Administration from the class Administration.

tau A numeric value for the infusion lag tau.

Value

The object Administration object with its new value of the infusion lag tau.

setTimeDose setTimeDose

Description

Set the times vector when doses are given.

Usage

```
setTimeDose(object, timeDose)
## S4 method for signature 'Administration'
setTimeDose(object, timeDose)
```

Arguments

object An object Administration from the class Administration.

timeDose A numeric value of the time dose.

Value

The object Administration with its new times vector for doses.

182 setVarianceEffects

setTinf

Set the infusion duration.

Description

Set the infusion duration.

Usage

```
setTinf(object, Tinf)
## S4 method for signature 'Administration'
setTinf(object, Tinf)
```

Arguments

object An object Administration from the class Administration.

Tinf A numeric value for the infusion duration Tinf.

Value

The object Administration with its new value of the infusion duration Tinf.

setVarianceEffects

Set the matrix of the variance effects.

Description

Set the matrix of the variance effects.

Usage

```
setVarianceEffects(object)
## S4 method for signature 'Fim'
setVarianceEffects(object)
```

Arguments

object

An object from the class Fim.

Value

Update the matrix of the variance effects.

show, Design-method 183

show, Design-method show

Description

show

show

show

show

show

show

show

show

Usage

```
## S4 method for signature 'Design'
show(object)
## S4 method for signature 'Evaluation'
show(object)
## S4 method for signature 'FedorovWynnAlgorithm'
show(object)
## S4 method for signature 'MultiplicativeAlgorithm'
show(object)
## S4 method for signature 'Optimization'
show(object)
## S4 method for signature 'PGBOAlgorithm'
show(object)
## S4 method for signature 'PSOAlgorithm'
show(object)
## S4 method for signature 'SimplexAlgorithm'
show(object)
```

Arguments

object object

SimplexAlgorithm-class

Class "SimplexAlgorithm"

Description

Class "Simplex Algorithm" implements the Multiplicative algorithm.

Objects from the class SimplexAlgorithm

Objects form the class SimplexAlgorithm can be created by calls of the form SimplexAlgorithm(...) where (...) are the parameters for the SimplexAlgorithm objects.

Slots for SamplingTimes objects

pctInitialSimplexBuilding: A numeric giving the percentage of the initial simplex.

maxIteration: A numeric giving the number of maximum iteration.

tolerance: A numeric giving the tolerance threshold.

showProcess: A boolean to show or not the process.

optimalDesign: A Design object giving the optimal design.

iterationAndCriteria: A list giving the optimal criteria at each iteration.

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