

Package ‘campsis’

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Type Package

Title Generic PK/PD Simulation Platform Campsis

Version 1.8.2

Description A generic, easy-to-use and intuitive pharmacokinetic/pharmacodynamic (PK/PD) simulation platform based on the R packages 'rxode2' and 'mrgsolve'. Campsis provides an abstraction layer over the underlying processes of defining a PK/PD model, assembling a custom dataset and running a simulation. The package has a strong dependency on the R package 'campsismod', which allows models to be read from and written to files, including through a JSON-based interface, and to be adapted further on the fly in the R environment. In addition, 'campsis' allows users to assemble datasets in an intuitive manner, including via a JSON-based interface to import Campsis datasets defined using formal JSON schemas distributed with the package. Once the dataset is ready, the package prepares the simulation, calls 'rxode2' or 'mrgsolve' (at the user's choice), and returns the results for the given model, dataset and desired simulation settings. The package itself is licensed under the GPL (≥ 3); the JSON schema files shipped in inst/extdata are licensed separately under the Creative Commons Attribution 4.0 International (CC BY 4.0).

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URL <https://github.com/Calvagone/campsis>,
<https://calvagone.github.io/>,
<https://calvagone.github.io/campsis.doc/>

BugReports <https://github.com/Calvagone/campsis/issues>

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Imports assertthat, digest, dplyr, furr, future, ggplot2, jsonlite, jsonvalidate, MASS, methods, progressr, purrr, rlang, stats, tibble, tidy

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Collate 'global.R' 'utilities.R' 'time_utilities.R' 'check.R'
 'generic.R' 'data.R' 'seed.R' 'distribution.R'
 'dataset_config.R' 'time_vector.R' 'time_entry.R'
 'repeated_schedule.R' 'occasion.R' 'occasions.R'
 'treatment_iov.R' 'treatment_iovs.R' 'dose_adaptation.R'
 'dose_adaptations.R' 'treatment_entry.R' 'treatment.R'
 'observations.R' 'observations_set.R' 'covariate.R'
 'covariates.R' 'bootstrap.R' 'protocol.R' 'arm.R' 'arms.R'
 'event.R' 'events.R' 'simulation_engine.R' 'dataset.R'
 'scenario_action.R' 'scenario_actions.R' 'scenario.R'
 'scenarios.R' 'event_logic.R' 'dataset_summary.R' 'outfun.R'
 'default_settings.R' 'hardware_settings.R'
 'simulation_progress.R' 'solver_settings.R' 'nocb_settings.R'
 'declare_settings.R' 'progress_settings.R'
 'internal_settings.R' 'simulation_settings.R' 'plan_setup.R'
 'json_interface.R' 'simulate_preprocess.R' 'simulate.R'
 'results_processing.R' 'default_plot.R'

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applyAction *Apply some action on the given object.*

Description

Apply some action on the given object.

Usage

```
applyAction(object, action)
```

```
## S4 method for signature 'campsis_model,replace_action'
applyAction(object, action)
```

```
## S4 method for signature 'dataset,replace_action'
applyAction(object, action)
```

Arguments

| | |
|--------|-----------------|
| object | any object |
| action | action to apply |

Value

updated object

applyCompartmentCharacteristics

Apply compartment characteristics from model. In practice, only compartment infusion duration needs to be applied.

Description

Apply compartment characteristics from model. In practice, only compartment infusion duration needs to be applied.

Usage

```
applyCompartmentCharacteristics(table, properties)
```

Arguments

| | |
|------------|-----------------------------------|
| table | current dataset |
| properties | compartment properties from model |

Value

updated dataset

| | |
|-----|--------------------------------|
| Arm | <i>Create a treatment arm.</i> |
|-----|--------------------------------|

Description

Create a treatment arm.

Usage

```
Arm(id = as.integer(NA), subjects = 1, label = as.character(NA))
```

Arguments

| | |
|----------|---|
| id | unique identifier for this arm (available through dataset), integer. If NA (default), this identifier is auto-incremented. |
| subjects | number of subjects in arm, integer |
| label | arm label, single character string. If set, this label will be output in the ARM column of CAMPSIS instead of the identifier. |

Value

an arm

| | |
|-----------|-------------------|
| arm-class | <i>Arm class.</i> |
|-----------|-------------------|

Description

Arm class.

Slots

| | |
|------------|------------------------------------|
| id | arm unique ID, integer |
| subjects | number of subjects in arm, integer |
| label | arm label, single character string |
| protocol | protocol |
| covariates | covariates |
| bootstrap | covariates to be bootstrapped |

| | |
|------------|--------------------|
| arms-class | <i>Arms class.</i> |
|------------|--------------------|

Description

Arms class.

| | |
|---------------------------------|---|
| as.numeric,time_sequence-method | <i>Time sequence to numeric vector.</i> |
|---------------------------------|---|

Description

Time sequence to numeric vector.

Usage

```
## S4 method for signature 'time_sequence'
as.numeric(x)
```

Arguments

| | |
|---|----------------------|
| x | time sequence object |
|---|----------------------|

Value

a numeric vector

| | |
|-------------------------------|---------------------------------------|
| as.numeric,time_vector-method | <i>Time vector to numeric vector.</i> |
|-------------------------------|---------------------------------------|

Description

Time vector to numeric vector.

Usage

```
## S4 method for signature 'time_vector'
as.numeric(x)
```

Arguments

| | |
|---|--------------------|
| x | time vector object |
|---|--------------------|

Value

a numeric vector

BinomialDistribution *Binomial distribution.*

Description

Binomial distribution.

Usage

BinomialDistribution(trials, prob)

Arguments

| | |
|--------|--|
| trials | number of Bernoulli trials per observation (=subject), integer |
| prob | probability of success for each trial |

Value

a binomial distribution

binomial_distribution-class
Binomial distribution class.

Description

Binomial distribution class.

Bolus *Create one or several bolus(es).*

Description

Create one or several bolus(es).

Usage

```

Bolus(
  time,
  amount,
  compartment = NULL,
  f = NULL,
  lag = NULL,
  ii = NULL,
  addl = NULL,
  wrap = TRUE,
  ref = NULL,
  rep = NULL
)

```

Arguments

| | |
|-------------|---|
| time | treatment time(s), numeric value or vector. First treatment time if used together with ii and addl. |
| amount | amount to give as bolus, single numeric value |
| compartment | compartment index or name to give the bolus(es). A vector of integers or names can be used for a complex model administration. |
| f | fraction of dose amount, list of distributions (one per compartment) |
| lag | dose lag time, list of distributions (one per compartment) |
| ii | inter-dose interval, requires argument 'time' to be a single numeric value |
| addl | number of additional doses, requires argument 'time' to be a single integer value |
| wrap | if TRUE, the bolus wrapper will be stored as is in the dataset, otherwise, it will be split into a list of boluses distinct in time. Default is TRUE. |
| ref | any reference name used to identify this bolus, single character value |
| rep | repeat the base dosing schedule several times, a 'repeated schedule' object is expected. Default is NULL (no repetition). |

Value

a single bolus or a list of boluses

bolus-class

Bolus class.

Description

Bolus class.

`bolus_wrapper-class` *Bolus wrapper class.*

Description

Bolus wrapper class.

`Bootstrap` *Create a bootstrap object.*

Description

Create a bootstrap object.

Usage

```
Bootstrap(
  data,
  id = "BS_ID",
  replacement = FALSE,
  random = FALSE,
  export_id = FALSE
)
```

Arguments

| | |
|--------------------------|--|
| <code>data</code> | data frame to be bootstrapped. It must have a unique identifier column named according to the specified argument 'id' (default value is 'BS_ID'). Other columns are covariates to bootstrap. They must all be numeric. Whatever the configuration of the bootstrap, these covariates are always read row by row and belong to a same individual. |
| <code>id</code> | unique identifier column name in data |
| <code>replacement</code> | values can be reused or not when drawn, logical |
| <code>random</code> | values are drawn randomly, logical |
| <code>export_id</code> | tell CAMPSIS if the identifier 'BS_ID' must be output or not, logical |

Value

a bootstrap object

| | |
|-----------------|-------------------------|
| bootstrap-class | <i>Bootstrap class.</i> |
|-----------------|-------------------------|

Description

Bootstrap class.

Slots

data data frame to be bootstrapped. Column 'BS_ID' is mandatory and corresponds to the original row ID from the bootstrap. It must be numeric and unique. Other columns are covariates to be bootstrapped (row by row).

replacement values can be reused or not, logical

random values are drawn randomly, logical

export_id tell CAMPSIS if 'BS_ID' must be exported into the dataset, logical

| | |
|-----------------------|---|
| BootstrapDistribution | <i>Create a bootstrap distribution. During function sampling, CAMPSIS will generate values depending on the given data and arguments.</i> |
|-----------------------|---|

Description

Create a bootstrap distribution. During function sampling, CAMPSIS will generate values depending on the given data and arguments.

Usage

```
BootstrapDistribution(data, replacement = FALSE, random = FALSE)
```

Arguments

data values to draw, numeric vector

replacement values can be reused or not, logical

random values are drawn randomly, logical

Value

a bootstrap distribution

bootstrap_distribution-class
Bootstrap distribution class.

Description

Bootstrap distribution class.

Slots

data values to draw, numeric vector
 replacement values can be reused or not, logical
 random values are drawn randomly, logical

campsis_handler *Suggested Campsis handler for showing the progress bar.*

Description

Suggested Campsis handler for showing the progress bar.

Usage

campsis_handler()

Value

a progressr handler list

ConstantDistribution *Create a constant distribution. Its value will be constant across all generated samples.*

Description

Create a constant distribution. Its value will be constant across all generated samples.

Usage

ConstantDistribution(value)

Arguments

value covariate value, single numeric value

Value

a constant distribution (same value for all samples)

constant_distribution-class
Constant distribution class.

Description

Constant distribution class.

Slots

value covariate value, single numeric value

convertTime *Convert numeric time vector based on the provided units.*

Description

Convert numeric time vector based on the provided units.

Usage

```
convertTime(x, from, to)
```

Arguments

| | |
|------|--|
| x | numeric time vector |
| from | unit of x, single character value |
| to | destination unit, single character value |

Value

numeric vector with the converted times

| | |
|-----------|---|
| Covariate | <i>Create a non time-varying (fixed) covariate.</i> |
|-----------|---|

Description

Create a non time-varying (fixed) covariate.

Usage

```
Covariate(name, distribution)
```

Arguments

| | |
|--------------|--|
| name | covariate name, single character value |
| distribution | covariate distribution |

Value

a fixed covariate

| | |
|-----------------|-------------------------|
| covariate-class | <i>Covariate class.</i> |
|-----------------|-------------------------|

Description

Covariate class.

Slots

| | |
|--------------|--|
| name | covariate name, single character value |
| distribution | covariate distribution |

| | |
|------------------|--------------------------|
| covariates-class | <i>Covariates class.</i> |
|------------------|--------------------------|

Description

Covariates class.

| | |
|----------------|-------------------------------------|
| CyclicSchedule | <i>Cyclic schedule constructor.</i> |
|----------------|-------------------------------------|

Description

Cyclic schedule constructor.

Usage

```
CyclicSchedule(duration, repetitions)
```

Arguments

| | |
|-------------|---|
| duration | duration of the cycle, numeric value |
| repetitions | number of additional repetitions to the base pattern, integer value |

Value

a cyclic schedule

| | |
|-----------------------|-------------------------------|
| cyclic_schedule-class | <i>Cyclic schedule class.</i> |
|-----------------------|-------------------------------|

Description

Cyclic schedule class.

Slots

| | |
|-------------|---|
| duration | duration of the cycle, numeric value |
| repetitions | number of additional repetitions to the base pattern, integer value |

| | |
|---------|--------------------------|
| Dataset | <i>Create a dataset.</i> |
|---------|--------------------------|

Description

Create a dataset.

Usage

```
Dataset(subjects = NULL, label = as.character(NA), json = NULL)
```

Arguments

| | |
|----------|--|
| subjects | number of subjects in the default arm |
| label | label of the default arm, NA by default |
| json | path to JSON dataset file or JSON content in string form |

Value

a dataset

| | |
|---------------|-----------------------|
| dataset-class | <i>Dataset class.</i> |
|---------------|-----------------------|

Description

Dataset class.

Slots

| | |
|--------|--|
| arms | a list of treatment arms |
| config | dataset configuration for export |
| iiv | data frame containing the inter-individual variability (all ETAS) for the export |

| | |
|---------------|---|
| DatasetConfig | <i>Create a dataset configuration. This configuration allows CAMPSIS to know which are the default depot and observed compartments.</i> |
|---------------|---|

Description

Create a dataset configuration. This configuration allows CAMPSIS to know which are the default depot and observed compartments.

Usage

```
DatasetConfig(
  defDepotCmt = 1,
  defObsCmt = 1,
  exportTSLD = FALSE,
  exportTDOS = FALSE,
  timeUnitDataset = "hour",
  timeUnitExport = "hour"
)
```

Arguments

| | |
|-----------------|--|
| defDepotCmt | default depot compartment, integer |
| defObsCmt | default observation compartment, integer |
| exportTSLD | export column TSLD (time since last dose), logical |
| exportTDOS | export column TDOS (time of last dose), logical |
| timeUnitDataset | unit of time in dataset, character ('hour' by default) |
| timeUnitExport | unit of time in export, character ('hour' by default) |

Value

a dataset configuration

| | |
|----------------------|-------------------------------------|
| dataset_config-class | <i>Dataset configuration class.</i> |
|----------------------|-------------------------------------|

Description

Dataset configuration class.

Slots

def_depot_cmt default depot compartment, integer
 def_obs_cmt default observation compartment, integer
 export_tslD export column TSLD, logical
 export_tdos export column TDOS, logical
 time_unit_dataset unit of time in dataset, character ('hour' by default)
 time_unit_export unit of time in export, character ('hour' by default)

days *Convert days to hours.*

Description

Convert days to hours.

Usage

days(x)

Arguments

x numeric vector in days

Value

numeric vector in hours

Declare *Create declare settings.*

Description

Create declare settings.

Usage

Declare(variables = character(0))

Arguments

variables uninitialized variables to be declared, only needed with mrgsolve

Value

Declare settings

```
declare_settings-class
```

Declare settings class.

Description

Declare settings class.

Slots

variables uninitialized variables to be declared, only needed with mrgsolve

```
DefaultSettings
```

Create default settings.

Description

Create default settings.

Usage

```
DefaultSettings(
  engine = "rxode2",
  seed = NULL,
  outvars = character(),
  disabled_variabilities = character(),
  dosing = FALSE
)
```

Arguments

| | |
|------------------------|---|
| engine | simulation engine, character |
| seed | random seed number, integer (or NULL for auto-generated seed) |
| outvars | output variables, character vector |
| disabled_variabilities | variabilities to disable in the simulation, character vector |
| dosing | output dosing information, logical |

Value

default settings

default_settings-class

Default settings class.

Description

Default settings class.

Slots

engine simulation engine, character

seed random seed number, integer

outvars output variables, character vector

disabled_variabilities variabilities to disable in the simulation, character vector

dosing output dosing information, logical

DiscreteDistribution *Discrete distribution.*

Description

Discrete distribution.

Usage

```
DiscreteDistribution(x, prob, replace = TRUE)
```

Arguments

x vector of one or more integers from which to choose

prob a vector of probability weights for obtaining the elements of the vector being sampled

replace should sampling be with replacement, default is TRUE

Value

a discrete distribution

discrete_distribution-class

Discrete distribution class.

Description

Discrete distribution class.

distribution-class *Distribution class. See this class as an interface.*

Description

Distribution class. See this class as an interface.

DoseAdaptation *Create a dose adaptation.*

Description

Create a dose adaptation.

Usage

DoseAdaptation(formula, compartments = NULL)

Arguments

| | |
|--------------|--|
| formula | formula to apply, single character string, e.g. "AMT*WT" |
| compartments | compartment indexes or names where the formula needs to be applied, integer or character vector. Default is NULL (formula applied on all compartments) |

Value

a fixed covariate

dose_adaptation-class *Dose adaptation class.*

Description

Dose adaptation class.

Slots

formula formula to apply, single character string, e.g. "AMT*WT"

compartments compartment numbers where the formula needs to be applied

dose_adaptations-class
Dose adaptations class.

Description

Dose adaptations class.

dosingOnly *Filter CAMPSIS output on dosing rows.*

Description

Filter CAMPSIS output on dosing rows.

Usage

dosingOnly(x)

Arguments

x data frame, CAMPSIS output

Value

a data frame with the dosing rows

| | |
|----------------|-------------------------------------|
| DosingSchedule | <i>Dosing schedule constructor.</i> |
|----------------|-------------------------------------|

Description

Dosing schedule constructor.

Usage

DosingSchedule()

Value

a dosing schedule (schedule repeated at dose times)

| | |
|-----------------------|-------------------------------|
| dosing_schedule-class | <i>Dosing schedule class.</i> |
|-----------------------|-------------------------------|

Description

Dosing schedule class.

| | |
|-----------------|---|
| EtaDistribution | <i>Create an ETA distribution. The resulting distribution is a normal distribution, with mean=0 and sd=sqrt(OMEGA).</i> |
|-----------------|---|

Description

Create an ETA distribution. The resulting distribution is a normal distribution, with mean=0 and sd=sqrt(OMEGA).

Usage

EtaDistribution(model, omega)

Arguments

| | |
|-------|-------------------------------------|
| model | model |
| omega | corresponding THETA name, character |

Value

an ETA distribution

| | |
|-------|--------------------------------------|
| Event | <i>Create an interruption event.</i> |
|-------|--------------------------------------|

Description

Create an interruption event.

Usage

```
Event(name = NULL, times, fun, debug = FALSE)
```

Arguments

| | |
|-------|---|
| name | event name, character value |
| times | interruption times, numeric vector |
| fun | event function to apply at each interruption |
| debug | output the variables that were changed through this event |

Value

an event definition

| | |
|-------------|---------------------|
| event-class | <i>Event class.</i> |
|-------------|---------------------|

Description

Event class.

Slots

| | |
|-------|---|
| name | event name, character value |
| times | interruption times, numeric vector |
| fun | event function to apply at each interruption |
| debug | output the variables that were changed through this event |

| | |
|----------------|--|
| EventCovariate | <i>Create an event covariate. These covariates can be modified further in interruption events.</i> |
|----------------|--|

Description

Create an event covariate. These covariates can be modified further in interruption events.

Usage

```
EventCovariate(name, distribution)
```

Arguments

| | |
|--------------|----------------------------------|
| name | covariate name, character |
| distribution | covariate distribution at time 0 |

Value

a time-varying covariate

| | |
|--------|--|
| Events | <i>Create a list of interruption events.</i> |
|--------|--|

Description

Create a list of interruption events.

Usage

```
Events()
```

Value

a events object

| | |
|--------------|----------------------|
| events-class | <i>Events class.</i> |
|--------------|----------------------|

Description

Events class.

event_covariate-class *Event covariate class.*

Description

Event covariate class.

FixedDistribution *Create a fixed distribution. Each sample will be assigned a fixed value coming from vector 'values'.*

Description

Create a fixed distribution. Each sample will be assigned a fixed value coming from vector 'values'.

Usage

FixedDistribution(values)

Arguments

values covariate values, numeric vector (1 value per sample)

Value

a fixed distribution (1 value per sample)

fixed_covariate-class *Fixed covariate class.*

Description

Fixed covariate class.

fixed_distribution-class
Fixed distribution class.

Description

Fixed distribution class.

Slots

values covariate values, numeric vector (1 value per sample)

FunctionDistribution *Create a function distribution. During distribution sampling, the provided function will be responsible for generating values for each sample. If first argument of this function is not the size (n), please tell which argument corresponds to the size 'n' (e.g. list(size="n")).*

Description

Create a function distribution. During distribution sampling, the provided function will be responsible for generating values for each sample. If first argument of this function is not the size (n), please tell which argument corresponds to the size 'n' (e.g. list(size="n")).

Usage

```
FunctionDistribution(fun, args)
```

Arguments

fun function name, character (e.g. 'rnorm')

args list of arguments (e.g list(mean=70, sd=10))

Value

a function distribution

function_distribution-class
Function distribution class.

Description

Function distribution class.

Slots

fun function name, character (e.g. 'rnorm')

args list of arguments (e.g list(mean=70, sd=10))

| | |
|-------------|---|
| generateIIV | <i>Generate IIV matrix for the given Campsis model.</i> |
|-------------|---|

Description

Generate IIV matrix for the given Campsis model.

Usage

```
generateIIV(model, n, offset = 0)
```

Arguments

| | |
|--------|--|
| model | Campsis model |
| n | number of subjects |
| offset | if specified, resulting ID will be ID + offset |

Value

IIV data frame with ID column

| | |
|--------------|--|
| generateIIV_ | <i>Generate IIV matrix for the given OMEGA matrix.</i> |
|--------------|--|

Description

Generate IIV matrix for the given OMEGA matrix.

Usage

```
generateIIV_(omega, n)
```

Arguments

| | |
|-------|--------------------|
| omega | omega matrix |
| n | number of subjects |

Value

IIV data frame

`getAvailableTimeUnits` *Return the list of available time units.*

Description

Return the list of available time units.

Usage

```
getAvailableTimeUnits()
```

Value

character vector

`getCampsisOption` *Get Campsis option logic.*

Description

Get Campsis option logic.

Usage

```
getCampsisOption(name, default)
```

Arguments

| | |
|----------------------|-----------------------------------|
| <code>name</code> | option to search |
| <code>default</code> | default value if option not found |

Value

option value

| | |
|---------------|--|
| getCovariates | <i>Get all covariates (fixed / time-varying / event covariates).</i> |
|---------------|--|

Description

Get all covariates (fixed / time-varying / event covariates).

Usage

```
getCovariates(object)

## S4 method for signature 'covariates'
getCovariates(object)

## S4 method for signature 'arm'
getCovariates(object)

## S4 method for signature 'arms'
getCovariates(object)

## S4 method for signature 'dataset'
getCovariates(object)
```

Arguments

object any object

Value

all covariates from object

| | |
|--------------------|--|
| getEventCovariates | <i>Get all event-related covariates.</i> |
|--------------------|--|

Description

Get all event-related covariates.

Usage

```
getEventCovariates(object)

## S4 method for signature 'covariates'
getEventCovariates(object)

## S4 method for signature 'arm'
```

```
getEventCovariates(object)

## S4 method for signature 'arms'
getEventCovariates(object)

## S4 method for signature 'dataset'
getEventCovariates(object)
```

Arguments

object any object

Value

all event-related covariates from object

getFixedCovariates *Get all fixed covariates.*

Description

Get all fixed covariates.

Usage

```
getFixedCovariates(object)

## S4 method for signature 'covariates'
getFixedCovariates(object)

## S4 method for signature 'arm'
getFixedCovariates(object)

## S4 method for signature 'arms'
getFixedCovariates(object)

## S4 method for signature 'dataset'
getFixedCovariates(object)
```

Arguments

object any object

Value

all fixed covariates from object

| | |
|---------|-----------------------------|
| getIOVs | <i>Get all IOV objects.</i> |
|---------|-----------------------------|

Description

Get all IOV objects.

Usage

```
getIOVs(object)

## S4 method for signature 'arm'
getIOVs(object)

## S4 method for signature 'arms'
getIOVs(object)

## S4 method for signature 'dataset'
getIOVs(object)
```

Arguments

object any object

Value

all IOV's from object

| | |
|--------------|---------------------------|
| getOccasions | <i>Get all occasions.</i> |
|--------------|---------------------------|

Description

Get all occasions.

Usage

```
getOccasions(object)

## S4 method for signature 'arm'
getOccasions(object)

## S4 method for signature 'arms'
getOccasions(object)

## S4 method for signature 'dataset'
getOccasions(object)
```

Arguments

object any object

Value

all occasions from object

`getSeedForDatasetExport`

Get seed for dataset export.

Description

Get seed for dataset export.

Usage

```
getSeedForDatasetExport(seed, progress)
```

Arguments

seed original seed
progress simulation progress

Value

the seed value used to export the dataset

`getSeedForIteration`

Get seed for iteration.

Description

Get seed for iteration.

Usage

```
getSeedForIteration(seed, progress)
```

Arguments

seed original seed
progress simulation progress

Value

the seed value to be used for the given replicate number and iteration

getSeedForParametersSampling

Get seed for parameter uncertainty sampling.

Description

Get seed for parameter uncertainty sampling.

Usage

getSeedForParametersSampling(seed)

Arguments

seed original seed

Value

the seed value used to sample parameter uncertainty

getSplittingConfiguration

Get splitting configuration for parallel export.

Description

Get splitting configuration for parallel export.

Usage

getSplittingConfiguration(dataset, hardware)

Arguments

dataset Campsis dataset to export
hardware hardware configuration

Value

splitting configuration list (if 'dataset_parallel' is enabled) or NA (if 'dataset_parallel' disabled or if the length of the dataset is less than the dataset export slice size)

getTimes *Get all distinct times for the specified object.*

Description

Get all distinct times for the specified object.

Usage

```
getTimes(object, ...)  
  
## S4 method for signature 'treatment'  
getTimes(object, unwrap = TRUE)  
  
## S4 method for signature 'observations'  
getTimes(object, doseTimes = NULL)  
  
## S4 method for signature 'observations_set'  
getTimes(object, doseTimes = NULL)  
  
## S4 method for signature 'arm'  
getTimes(object)  
  
## S4 method for signature 'arms'  
getTimes(object)  
  
## S4 method for signature 'events'  
getTimes(object)  
  
## S4 method for signature 'dataset'  
getTimes(object)
```

Arguments

| | |
|-----------|---|
| object | any object |
| ... | extra arguments like 'doseTimes' in observations or 'unwrap' in treatment |
| unwrap | unwrap treatment before accessing the times, default value is TRUE |
| doseTimes | times of the doses, only needed if a [DosingSchedule()] is referred to |

Value

numeric vector with all unique times, sorted

```
getTimeVaryingCovariates  
    Get all time-varying covariates.
```

Description

Get all time-varying covariates.

Usage

```
getTimeVaryingCovariates(object)  
  
## S4 method for signature 'covariates'  
getTimeVaryingCovariates(object)  
  
## S4 method for signature 'arm'  
getTimeVaryingCovariates(object)  
  
## S4 method for signature 'arms'  
getTimeVaryingCovariates(object)  
  
## S4 method for signature 'dataset'  
getTimeVaryingCovariates(object)
```

Arguments

object any object

Value

all time-varying covariates from object

```
Hardware            Create hardware settings.
```

Description

Create hardware settings.

Usage

```
Hardware(
  cpu = 1,
  replicate_parallel = FALSE,
  scenario_parallel = FALSE,
  slice_parallel = FALSE,
  slice_size = NULL,
  dataset_parallel = FALSE,
  dataset_slice_size = 500,
  auto_setup_plan = NULL
)
```

Arguments

`cpu` number of CPU cores to use, default is 1

`replicate_parallel` enable parallel computing for replicates, default is FALSE

`scenario_parallel` enable parallel computing for scenarios, default is FALSE

`slice_parallel` enable parallel computing for slices, default is FALSE

`slice_size` number of subjects per simulated slice, default is NULL (auto-configured by Campsis depending on the specified engine)

`dataset_parallel` enable parallelisation when exporting dataset into a table, default is FALSE

`dataset_slice_size` dataset slice size when exporting subjects to a table, default is 500. Only applicable if 'dataset_parallel' is enabled.

`auto_setup_plan` auto-setup plan with the library future, if not set (i.e. =NULL), plan will be setup automatically if the number of CPU's > 1.

Value

hardware settings

hardware_settings-class

Hardware settings class.

Description

Hardware settings class.

Slots

cpu number of CPU cores to use, default is 1
 replicate_parallel enable parallel computing for replicates, default is FALSE
 scenario_parallel enable parallel computing for scenarios, default is FALSE
 slice_parallel enable parallel computing for slices, default is FALSE
 slice_size number of subjects per simulated slice, default is NULL (auto-configured by Campsis depending on the specified engine)
 dataset_parallel enable parallelisation when exporting dataset into a table, default is FALSE
 dataset_slice_size dataset slice size when exporting subjects to a table, default is 500. Only applicable if 'dataset_parallel' is enabled.
 auto_setup_plan auto-setup plan with the library future, default is FALSE

| | |
|-------|---|
| hours | <i>Convert hours to hours (do nothing).</i> |
|-------|---|

Description

Convert hours to hours (do nothing).

Usage

hours(x)

Arguments

x numeric vector in hours

Value

numeric vector in hours

| | |
|----------|---|
| Infusion | <i>Create one or several infusion(s).</i> |
|----------|---|

Description

Create one or several infusion(s).

Usage

```

Infusion(
  time,
  amount,
  compartment = NULL,
  f = NULL,
  lag = NULL,
  duration = NULL,
  rate = NULL,
  ii = NULL,
  addl = NULL,
  wrap = TRUE,
  ref = NULL,
  rep = NULL
)

```

Arguments

| | |
|-------------|--|
| time | treatment time(s), numeric value or vector. First treatment time if used together with ii and addl. |
| amount | amount to infuse, single numeric value |
| compartment | compartment index or name to give the infusion(s). A vector of integers or names can be used for a complex model administration. |
| f | fraction of infusion amount, list of distributions (one per compartment) |
| lag | infusion lag time, , list of distributions (one per compartment) |
| duration | infusion duration, list of distributions (one per compartment) |
| rate | infusion rate, list of distributions (one per compartment) |
| ii | inter-dose interval, requires argument 'time' to be a single numeric value |
| addl | number of additional doses, requires argument 'time' to be a single integer value |
| wrap | if TRUE, the infusion wrapper will be stored as is in the dataset, otherwise, it will be split into a list of infusions distinct in time. Default is TRUE. |
| ref | any reference name used to identify this infusion, single character value |
| rep | repeat the base dosing schedule several times, a 'repeated schedule' object is expected. Default is NULL (no repetition). |

Value

a single infusion or a list of infusions.

infusion-class *Infusion class.*

Description

Infusion class.

Slots

duration infusion duration, distribution list
rate infusion rate, distribution list

infusion_wrapper-class
Infusion wrapper class.

Description

Infusion wrapper class.

internal_settings-class
Internal settings class (transient object from the simulation settings).

Description

Internal settings class (transient object from the simulation settings).

Slots

dataset_summary dataset summary
progress simulation progress
iterations list of event iterations

| | |
|-----|--|
| IOV | <i>Define inter-occasion variability (IOV) into the dataset. A new variable of name 'colname' will be output into the dataset and will vary at each dose number according to the given distribution.</i> |
|-----|--|

Description

Define inter-occasion variability (IOV) into the dataset. A new variable of name 'colname' will be output into the dataset and will vary at each dose number according to the given distribution.

Usage

```
IOV(colname, distribution, doseNumbers = NULL)
```

Arguments

| | |
|--------------|--|
| colname | name of the column that will be output in dataset |
| distribution | distribution |
| doseNumbers | dose numbers, if provided, IOV is generated at these doses only. By default, IOV is generated for all doses. |

Value

an IOV object

| | |
|-------------------|---|
| length,arm-method | <i>Return the number of subjects contained in this arm.</i> |
|-------------------|---|

Description

Return the number of subjects contained in this arm.

Usage

```
## S4 method for signature 'arm'
length(x)
```

Arguments

| | |
|---|-----|
| x | arm |
|---|-----|

Value

a number

length,cyclic_schedule-method

Return the number of repetition cycles.

Description

Return the number of repetition cycles.

Usage

```
## S4 method for signature 'cyclic_schedule'  
length(x)
```

Arguments

x schedule object

Value

a number

length,dataset-method *Return the number of subjects contained in this dataset.*

Description

Return the number of subjects contained in this dataset.

Usage

```
## S4 method for signature 'dataset'  
length(x)
```

Arguments

x dataset

Value

a number

length, repeat_at_schedule-method

Return the number of repetition cycles.

Description

Return the number of repetition cycles.

Usage

```
## S4 method for signature 'repeat_at_schedule'  
length(x)
```

Arguments

x schedule object

Value

a number

LogNormalDistribution *Create a log normal distribution.*

Description

Create a log normal distribution.

Usage

```
LogNormalDistribution(meanlog, sdlog)
```

Arguments

meanlog mean value of distribution in log domain
sdlog standard deviation of distribution in log domain

Value

a log normal distribution

lognormal_distribution-class
Log normal distribution class.

Description

Log normal distribution class.

minutes *Convert minutes to hours.*

Description

Convert minutes to hours.

Usage

minutes(x)

Arguments

x numeric vector in minutes

Value

numeric vector in hours

months *Convert pharma months (1 month = 4 weeks) to hours.*

Description

Convert pharma months (1 month = 4 weeks) to hours.

Usage

months(x)

Arguments

x numeric vector in months

Value

numeric vector in hours

mrgsolve_engine-class *mrgsolve engine class.*

Description

mrgsolve engine class.

| | |
|--------|---|
| nhanes | <i>NHANES database (demographics and body measure data combined, from 2017-2018).</i> |
|--------|---|

Description

NHANES database (demographics and body measure data combined, from 2017-2018).

Usage

nhanes

Format

data frame

BS_ID Original identifier

SEX Sex: 1 for males, 2 for females

AGE Age in years

BW Body weight in kg

BMI Body mass index

HT Height in cm

Source

https://www.cdc.gov/Nchs/Nhanes/2017-2018/DEMO_J.XPT

https://www.cdc.gov/Nchs/Nhanes/2017-2018/BMX_J.XPT

NOCB *Create NOCB settings.*

Description

Create NOCB settings.

Usage

NOCB(enable = NULL, variables = character(0))

Arguments

| | |
|-----------|---|
| enable | enable/disable next-observation carried backward mode (NOCB), default value is TRUE for mrgsolve, FALSE for RxODE |
| variables | variable names subject to NOCB behavior (see vignette for more info) |

Value

NOCB settings

nocb_settings-class *NOCB settings class.*

Description

NOCB settings class.

Slots

| | |
|-----------|---|
| enable | enable/disable next-observation carried backward mode (NOCB), default value is TRUE for mrgsolve, FALSE for RxODE |
| variables | variable names subject to NOCB behavior (see vignette for more info) |

NormalDistribution *Create a normal distribution.*

Description

Create a normal distribution.

Usage

```
NormalDistribution(mean, sd)
```

Arguments

| | |
|------|------------------------------------|
| mean | mean value of distribution |
| sd | standard deviation of distribution |

Value

a normal distribution

normal_distribution-class
Normal distribution class.

Description

Normal distribution class.

Observations *Create an observations list. Please note that the provided 'times' will automatically be sorted. Duplicated times will be removed.*

Description

Create an observations list. Please note that the provided 'times' will automatically be sorted. Duplicated times will be removed.

Usage

```
Observations(times, compartment = NA, rep = NULL)
```

Arguments

| | |
|-------------|---|
| times | observation times, numeric vector |
| compartment | compartment index (integer) or name (character) |
| rep | repetition schedule |

Value

an observations list

observations-class *Observations class.*

Description

Observations class.

Slots

times any object that implements
compartment compartment index (integer) or name (character)
dv observed values, numeric vector (FOR EXTERNAL USE)
rep repetition schedule

observations_set-class
Observations set class.

Description

Observations set class.

| | |
|---------|---|
| obsOnly | <i>Filter CAMPSIS output on observation rows.</i> |
|---------|---|

Description

Filter CAMPSIS output on observation rows.

Usage

```
obsOnly(x)
```

Arguments

x data frame, CAMPSIS output

Value

a data frame with the observation rows

| | |
|----------|---|
| Occasion | <i>Define a new occasion. Occasions are defined by mapping occasion values to dose numbers. A new column will automatically be created in the exported dataset.</i> |
|----------|---|

Description

Define a new occasion. Occasions are defined by mapping occasion values to dose numbers. A new column will automatically be created in the exported dataset.

Usage

```
Occasion(colname, values, doseNumbers)
```

Arguments

colname name of the column that will be output in dataset
 values the occasion numbers, any integer vector
 doseNumbers the related dose numbers, any integer vector of same length as 'values'

Value

occasion object

| | |
|----------------|------------------------|
| occasion-class | <i>Occasion class.</i> |
|----------------|------------------------|

Description

Occasion class.

Slots

colname single character value representing the column name related to this occasion

values occasion values, integer vector, same length as dose_numbers

dose_numbers associated dose numbers, integer vector, same length as values

| | |
|-----------------|-------------------------|
| occasions-class | <i>Occasions class.</i> |
|-----------------|-------------------------|

Description

Occasions class.

| | |
|--------|-------------------------------------|
| Outfun | <i>Create a new output function</i> |
|--------|-------------------------------------|

Description

Create a new output function

Usage

```
Outfun(
  fun = function(x, ...) {
    x
  },
  args = list(),
  packages = NULL,
  level = "scenario"
)
```

Arguments

| | |
|----------|--|
| fun | function or purrr-style lambda formula, first argument 'x' must be the results |
| args | extra arguments, named list |
| packages | packages that must be loaded to execute the given function, character vector |
| level | either 'scenario' or 'replicate'. Default is 'scenario'. |

Value

an output function

output_function-class *Output function class.*

Description

Output function class.

Slots

fun function or purrr-style lambda formula, first argument 'x' must be the results
 args extra arguments, named list
 packages packages that must be loaded to execute the given function, character vector
 level either 'scenario' or 'replicate'. Default is 'scenario'.

ParameterDistribution *Create a parameter distribution. The resulting distribution is a log-normal distribution, with meanlog=log(THETA) and sdlog=sqrt(OMEGA).*

Description

Create a parameter distribution. The resulting distribution is a log-normal distribution, with meanlog=log(THETA) and sdlog=sqrt(OMEGA).

Usage

```
ParameterDistribution(model, theta, omega = NULL)
```

Arguments

| | |
|-------|--|
| model | model |
| theta | corresponding THETA name, character |
| omega | corresponding OMEGA name, character, NULL if not defined |

Value

a parameter distribution

PI *Compute the prediction interval summary over time.*

Description

Compute the prediction interval summary over time.

Usage

```
PI(x, output, scenarios = NULL, level = 0.9, gather = TRUE)
```

Arguments

| | |
|-----------|---|
| x | data frame |
| output | variable to show, character value |
| scenarios | scenarios, character vector, NULL is default |
| level | PI level, default is 0.9 (90% PI) |
| gather | FALSE: med, low & up columns, TRUE: metric column |

Value

a summary table

Progress *Create progress settings.*

Description

Create progress settings.

Usage

```
Progress(tick_slice = TRUE)
```

Arguments

| | |
|------------|---|
| tick_slice | tick() is called after each simulated slice, default is TRUE. In some cases, when the number of subjects per slice is low, it may be useful disable this flag, to improve performance issues. |
|------------|---|

Value

progress settings

progress_settings-class *Progress settings class.*

Description

Progress settings class.

Slots

tick_slice tick() is called after each simulated slice, default is TRUE. In some cases, when the number of subjects per slice is low, it may be useful disable this flag, to improve performance issues.

protocol-class *Protocol class.*

Description

Protocol class.

RepeatAtSchedule *'Repeat at' schedule constructor. Note that the time 0 for the base pattern will be added by default if not provided.*

Description

'Repeat at' schedule constructor. Note that the time 0 for the base pattern will be added by default if not provided.

Usage

RepeatAtSchedule(times)

Arguments

times times at which the original schedule must be repeated, numeric vector

Value

a 'repeat-at' schedule

repeated_schedule-class

Repeated schedule class. See this class as an interface.

Description

Repeated schedule class. See this class as an interface.

repeatSchedule

Repeat schedule.

Description

Repeat schedule.

Usage

```
repeatSchedule(x, schedule)
```

```
## S4 method for signature 'numeric,cyclic_schedule'  
repeatSchedule(x, schedule)
```

```
## S4 method for signature 'numeric,repeat_at_schedule'  
repeatSchedule(x, schedule)
```

```
## S4 method for signature 'numeric,undefined_schedule'  
repeatSchedule(x, schedule)
```

Arguments

| | |
|----------|-------------------------------|
| x | object to repeat the schedule |
| schedule | initial times vector |

Value

resulting times vector

repeat_at_schedule-class
'Repeat at' schedule class.

Description

'Repeat at' schedule class.

Slots

times times at which the event is repeated, numeric vector

ReplaceAction *Create a replace action.*

Description

Create a replace action.

Usage

ReplaceAction(object)

Arguments

object replacement object

Value

a replace action

replace_action-class *Replace action class.*

Description

Replace action class.

 retrieveParameterValue

Retrieve the parameter value (standardized) for the specified parameter name.

Description

Retrieve the parameter value (standardized) for the specified parameter name.

Usage

```
retrieveParameterValue(model, paramName, default = NULL, mandatory = FALSE)
```

Arguments

| | |
|-----------|----------------------------|
| model | model |
| paramName | parameter name |
| default | default value if not found |
| mandatory | must be in model or not |

Value

the standardized parameter value or the given default value if not found

 rxode_engine-class *RxODE/rxode2 engine class.*

Description

RxODE/rxode2 engine class.

Slots

rxode2 logical field to indicate if CAMPSIS should use rxode2 (field set to TRUE) or RxODE (field set to FALSE). Default is TRUE.

| | |
|--------|-------------------------------|
| sample | <i>Sample generic object.</i> |
|--------|-------------------------------|

Description

Sample generic object.

Usage

```
sample(object, n, ...)  
  
## S4 method for signature 'constant_distribution,integer'  
sample(object, n)  
  
## S4 method for signature 'fixed_distribution,integer'  
sample(object, n)  
  
## S4 method for signature 'function_distribution,integer'  
sample(object, n)  
  
## S4 method for signature 'bootstrap_distribution,integer'  
sample(object, n)  
  
## S4 method for signature 'bolus,integer'  
sample(object, n, ...)  
  
## S4 method for signature 'infusion,integer'  
sample(object, n, ...)  
  
## S4 method for signature 'observations,integer'  
sample(object, n, ...)  
  
## S4 method for signature 'covariate,integer'  
sample(object, n)  
  
## S4 method for signature 'bootstrap,integer'  
sample(object, n)
```

Arguments

| | |
|--------|----------------------------|
| object | generic object |
| n | number of samples required |
| ... | extra arguments |

Value

sampling result

| | |
|-------------|---------------------------------------|
| scatterPlot | <i>Scatter plot (or X vs Y plot).</i> |
|-------------|---------------------------------------|

Description

Scatter plot (or X vs Y plot).

Usage

```
scatterPlot(x, output, colour = NULL, time = NULL)
```

Arguments

| | |
|--------|--|
| x | data frame |
| output | the 2 variables to show, character vector |
| colour | variable(s) to colour |
| time | the time to look at those 2 variables, if NULL, min time is used (usually 0) |

Value

a ggplot object

| | |
|----------|----------------------------|
| Scenario | <i>Create an scenario.</i> |
|----------|----------------------------|

Description

Create an scenario.

Usage

```
Scenario(name = NULL, model = NULL, dataset = NULL)
```

Arguments

| | |
|---------|--|
| name | scenario name, single character string |
| model | either a Campsis model, a function or lambda-style formula |
| dataset | either a Campsis dataset, a function or lambda-style formula |

Value

a new scenario

| | |
|----------------|------------------------|
| scenario-class | <i>Scenario class.</i> |
|----------------|------------------------|

Description

Scenario class.

Slots

name scenario name, single character string

model either a Campsis model, a function or lambda-style formula

dataset either a Campsis dataset, a function or lambda-style formula

actions list of actions to apply

| | |
|-----------|------------------------------------|
| Scenarios | <i>Create a list of scenarios.</i> |
|-----------|------------------------------------|

Description

Create a list of scenarios.

Usage

```
Scenarios(json = NULL)
```

Arguments

json path to JSON scenarios file or JSON content in string form

Value

a scenarios object

| | |
|-----------------|-------------------------|
| scenarios-class | <i>Scenarios class.</i> |
|-----------------|-------------------------|

Description

Scenarios class.

scenario_action-class *Scenario action class.*

Description

Scenario action class.

scenario_actions-class
Scenario actions class.

Description

Scenario actions class.

seconds *Convert seconds to hours.*

Description

Convert seconds to hours.

Usage

seconds(x)

Arguments

x numeric vector in seconds

Value

numeric vector in hours

| | |
|----------|-----------------------|
| setLabel | <i>Set the label.</i> |
|----------|-----------------------|

Description

Set the label.

Usage

```
setLabel(object, x)

## S4 method for signature 'arm,character'
setLabel(object, x)

## S4 method for signature 'dataset,character'
setLabel(object, x)
```

Arguments

| | |
|--------|-----------------------------|
| object | any object that has a label |
| x | the new label |

Value

the updated object

| | |
|-------------|------------------------------------|
| setSubjects | <i>Set the number of subjects.</i> |
|-------------|------------------------------------|

Description

Set the number of subjects.

Usage

```
setSubjects(object, x)

## S4 method for signature 'arm,integer'
setSubjects(object, x)

## S4 method for signature 'dataset,integer'
setSubjects(object, x)
```

Arguments

| | |
|--------|----------------------------|
| object | any object |
| x | the new number of subjects |

Value

the updated object

Settings

Create advanced simulation settings.

Description

Create advanced simulation settings.

Usage

```
Settings(..., json = NULL)
```

Arguments

| | |
|------|--|
| ... | any user-required settings: see ?Hardware, ?Solver, ?NOCB, ?Declare, ?Progress or ?AutoReplicationSettings |
| json | path to JSON settings file or JSON content in string form |

Value

advanced simulation settings

setupPlanDefault

Setup default plan for the given simulation or hardware settings. This plan will prioritise the distribution of workers in the following order: 1) Replicates (if 'replicate_parallel' is enabled) 2) Scenarios (if 'scenario_parallel' is enabled) 3) Dataset export / slices (if 'dataset_export' or 'slice_parallel' is enabled)

Description

Setup default plan for the given simulation or hardware settings. This plan will prioritise the distribution of workers in the following order: 1) Replicates (if 'replicate_parallel' is enabled) 2) Scenarios (if 'scenario_parallel' is enabled) 3) Dataset export / slices (if 'dataset_export' or 'slice_parallel' is enabled)

Usage

```
setupPlanDefault(object)
```

Arguments

object simulation or hardware settings

Value

nothing

setupPlanSequential *Setup plan as sequential (i.e. no parallelisation).*

Description

Setup plan as sequential (i.e. no parallelisation).

Usage

```
setupPlanSequential()
```

Value

nothing

shadedPlot *Shaded plot (or prediction interval plot).*

Description

Shaded plot (or prediction interval plot).

Usage

```
shadedPlot(  
  x,  
  output,  
  colour = NULL,  
  strat_extra = NULL,  
  level = 0.9,  
  alpha = 0.25  
)
```

Arguments

| | |
|-------------|---|
| x | data frame |
| output | variable to show |
| colour | variable(s) to colour |
| strat_extra | variable(s) to stratify, but not to colour (useful for use with facet_wrap) |
| level | PI level, default is 0.9 (90% PI) |
| alpha | alpha parameter (transparency) given to geom_ribbon |

Value

a ggplot object

| | |
|----------|---------------------------|
| simulate | <i>Simulate function.</i> |
|----------|---------------------------|

Description

Simulate function.

Usage

```
simulate(  
  model,  
  dataset,  
  dest = NULL,  
  events = NULL,  
  scenarios = NULL,  
  tablefun = NULL,  
  outvars = NULL,  
  outfun = NULL,  
  seed = NULL,  
  replicates = 1,  
  dosing = FALSE,  
  settings = NULL  
)  
  
## S4 method for signature  
## 'replicated_campsis_model,  
## dataset,  
## character,  
## events,  
## scenarios,  
## function,  
## character,  
## output_function,
```

```
## integer,
## integer,
## logical,
## simulation_settings'
simulate(
  model,
  dataset,
  dest = NULL,
  events = NULL,
  scenarios = NULL,
  tablefun = NULL,
  outvars = NULL,
  outfun = NULL,
  seed = NULL,
  replicates = 1,
  dosing = FALSE,
  settings = NULL
)

## S4 method for signature
## 'campsis_model,
## dataset,
## character,
## events,
## scenarios,
## function,
## character,
## output_function,
## integer,
## integer,
## logical,
## simulation_settings'
simulate(
  model,
  dataset,
  dest = NULL,
  events = NULL,
  scenarios = NULL,
  tablefun = NULL,
  outvars = NULL,
  outfun = NULL,
  seed = NULL,
  replicates = 1,
  dosing = FALSE,
  settings = NULL
)

## S4 method for signature
```

```
## 'campsis_model',
##   tbl_df,
##   character,
##   events,
##   scenarios,
##   function,
##   character,
##   output_function,
##   integer,
##   integer,
##   logical,
##   simulation_settings'
simulate(
  model,
  dataset,
  dest = NULL,
  events = NULL,
  scenarios = NULL,
  tablefun = NULL,
  outvars = NULL,
  outfun = NULL,
  seed = NULL,
  replicates = 1,
  dosing = FALSE,
  settings = NULL
)

## S4 method for signature
## 'campsis_model',
##   data.frame,
##   character,
##   events,
##   scenarios,
##   function,
##   character,
##   output_function,
##   integer,
##   integer,
##   logical,
##   simulation_settings'
simulate(
  model,
  dataset,
  dest = NULL,
  events = NULL,
  scenarios = NULL,
  tablefun = NULL,
  outvars = NULL,
```

```
    outfun = NULL,
    seed = NULL,
    replicates = 1,
    dosing = FALSE,
    settings = NULL
  )

## S4 method for signature
## 'campsis_model,
## tbl_df,
## rxode_engine,
## events,
## scenarios,
## function,
## character,
## output_function,
## integer,
## integer,
## logical,
## simulation_settings'
simulate(
  model,
  dataset,
  dest = NULL,
  events = NULL,
  scenarios = NULL,
  tablefun = NULL,
  outvars = NULL,
  outfun = NULL,
  seed = NULL,
  replicates = 1,
  dosing = FALSE,
  settings = NULL
)

## S4 method for signature
## 'campsis_model,
## tbl_df,
## mrgsolve_engine,
## events,
## scenarios,
## function,
## character,
## output_function,
## integer,
## integer,
## logical,
## simulation_settings'
```

```

simulate(
  model,
  dataset,
  dest = NULL,
  events = NULL,
  scenarios = NULL,
  tablefun = NULL,
  outvars = NULL,
  outfun = NULL,
  seed = NULL,
  replicates = 1,
  dosing = FALSE,
  settings = NULL
)

```

Arguments

| | |
|------------|--|
| model | generic CAMPSIS model |
| dataset | CAMPSIS dataset or 2-dimensional table |
| dest | destination simulation engine, default is 'RxODE' |
| events | interruption events |
| scenarios | list of scenarios to be simulated |
| tablefun | function or lambda formula to apply on exported 2-dimensional dataset |
| outvars | variables to output in resulting dataframe |
| outfun | an output function to apply on the simulation results. Type ?Outfun for more info. |
| seed | seed value |
| replicates | number of replicates, default is 1 |
| dosing | output dosing information, default is FALSE |
| settings | advanced simulation settings |

Value

dataframe with all results

SimulationProgress *Create a simulation progress object.*

Description

Create a simulation progress object.

Usage

```
SimulationProgress(
  replicates = 1,
  scenarios = 1,
  progressor = NULL,
  hardware = NULL
)
```

Arguments

| | |
|------------|--|
| replicates | total number of replicates to simulate |
| scenarios | total number of scenarios to simulate |
| progressor | progressr progressor |
| hardware | hardware settings |

Value

a progress bar

simulation_engine-class

Simulation engine class.

Description

Simulation engine class.

simulation_progress-class

Simulation progress class.

Description

Simulation progress class.

Arguments

| | |
|------------|--|
| replicates | total number of replicates to simulate |
| scenarios | total number of scenarios to simulate |
| iterations | total number of iterations to simulate |
| slices | total number of slices to simulate |
| replicate | current replicate number being simulated |

| | |
|------------|--|
| scenario | current scenario number being simulated |
| iteration | current iteration number being simulated |
| slice | current slice number being simulated |
| progressor | progressor progressor |
| hardware | hardware settings |

simulation_settings-class

Simulation settings class.

Description

Simulation settings class.

Slots

default default settings of the simulate method
hardware hardware settings object
solver solver settings object
nocb NOCB settings object
declare declare settings (mrgsolve only)
progress progress settings
replication replication settings
internal internal settings

Solver

Create solver settings.

Description

Create solver settings.

Usage

```
Solver(
  atol = 1e-08,
  rtol = 1e-08,
  hmax = NA,
  maxsteps = 70000L,
  method = "liblsoda"
)
```

Arguments

| | |
|----------|---|
| atol | absolute solver tolerance, default is 1e-08 |
| rtol | relative solver tolerance, default is 1e-08 |
| hmax | limit how big a solver step can be, default is NA |
| maxsteps | max steps between 2 integration times (e.g. when observations records are far apart), default is 70000 |
| method | solver method, for RxODE/rxode2 only: 'liblsoda' (default), 'lsoda', 'dop853', 'indLin'. Mrgsolve's method is always 'lsoda'. |

Value

solver settings

solver_settings-class *Solver settings class. See ?mrgsolve::update. See ?rxode2::rxSolve.*

Description

Solver settings class. See ?mrgsolve::update. See ?rxode2::rxSolve.

Slots

| | |
|----------|---|
| atol | absolute solver tolerance, default is 1e-08 |
| rtol | relative solver tolerance, default is 1e-08 |
| hmax | limit how big a solver step can be, default is NA |
| maxsteps | max steps between 2 integration times (e.g. when observations records are far apart), default is 70000 |
| method | solver method, for RxODE/rxode2 only: 'liblsoda' (default), 'lsoda', 'dop853', 'indLin'. Mrgsolve's method is always 'lsoda'. |

spaghettiPlot *Spaghetti plot.*

Description

Spaghetti plot.

Usage

```
spaghettiPlot(x, output, colour = NULL)
```

Arguments

| | |
|--------|-----------------------|
| x | data frame |
| output | variable to show |
| colour | variable(s) to colour |

Value

plot

| | |
|-----------------|-----------------------------------|
| standardiseTime | <i>Standardise time to hours.</i> |
|-----------------|-----------------------------------|

Description

Standardise time to hours.

Usage

```
standardiseTime(x, unit)
```

Arguments

| | |
|------|-----------------------------------|
| x | numeric time vector |
| unit | unit of x, single character value |

Value

numeric vector with the times converted to hours

| | |
|--------------|---|
| TimeSequence | <i>Instantiate a new time sequence.</i> |
|--------------|---|

Description

Instantiate a new time sequence.

Usage

```
TimeSequence(start, end, by)
```

Arguments

| | |
|-------|---------------------------|
| start | starting value |
| end | maximal value |
| by | increment of the sequence |

Value

a sequence

`TimeVaryingCovariate` *Create a time-varying covariate. This covariate will be implemented using EVID=2 rows in the exported dataset and will not use interruption events.*

Description

Create a time-varying covariate. This covariate will be implemented using EVID=2 rows in the exported dataset and will not use interruption events.

Usage

`TimeVaryingCovariate(name, table)`

Arguments

| | |
|--------------------|---|
| <code>name</code> | covariate name, character |
| <code>table</code> | data.frame, must contain the mandatory columns 'TIME' and 'VALUE'. An 'ID' column may also be specified. In that case, ID's between 1 and the max number of subjects in the dataset/arm can be used. All ID's must have a VALUE defined for TIME 0. |

Value

a time-varying covariate

`TimeVector` *Instantiate a new time vector*

Description

Instantiate a new time vector

Usage

`TimeVector(x)`

Arguments

| | |
|----------------|----------------------|
| <code>x</code> | time vector, numeric |
|----------------|----------------------|

Value

a time vector

time_sequence-class *Time sequence class.*

Description

Time sequence class.

time_varying_covariate-class
 Time-varying covariate class.

Description

Time-varying covariate class.

time_vector-class *Time vector class.*

Description

Time vector class.

treatment-class *Treatment class.*

Description

Treatment class.

treatment_iov-class *Treatment IOV class.*

Description

Treatment IOV class.

Slots

colname name of the column that will be output in dataset

distribution distribution

dose_numbers associated dose numbers, integer vector, same length as values

treatment_iovs-class *Treatment IOV's class.*

Description

Treatment IOV's class.

undefined_distribution-class

Undefined distribution class. This type of object is automatically created in method toExplicitDistribution() when the user does not provide a concrete distribution. This is because S4 objects do not accept NULL values.

Description

Undefined distribution class. This type of object is automatically created in method toExplicitDistribution() when the user does not provide a concrete distribution. This is because S4 objects do not accept NULL values.

undefined_schedule-class

Undefined schedule class.

Description

Undefined schedule class.

UniformDistribution *Create an uniform distribution.*

Description

Create an uniform distribution.

Usage

UniformDistribution(min, max)

Arguments

| | |
|-----|-----------|
| min | min value |
| max | max value |

Value

an uniform distribution

uniform_distribution-class
Uniform distribution class.

Description

Uniform distribution class.

unwrapTreatment *Unwrap treatment.*

Description

Unwrap treatment.

Usage

```
unwrapTreatment(object)

## S4 method for signature 'bolus'
unwrapTreatment(object)

## S4 method for signature 'infusion'
unwrapTreatment(object)

## S4 method for signature 'bolus_wrapper'
unwrapTreatment(object)

## S4 method for signature 'infusion_wrapper'
unwrapTreatment(object)

## S4 method for signature 'treatment'
unwrapTreatment(object)

## S4 method for signature 'arm'
unwrapTreatment(object)
```

```
## S4 method for signature 'arms'
unwrapTreatment(object)

## S4 method for signature 'dataset'
unwrapTreatment(object)
```

Arguments

object any object

Value

updated object

| | |
|------------|--|
| updateADDL | <i>Update the number of additional doses (ADDL).</i> |
|------------|--|

Description

Update the number of additional doses (ADDL).

Usage

```
updateADDL(object, addl, ref = NULL)

## S4 method for signature 'bolus_wrapper,integer,character'
updateADDL(object, addl, ref = NULL)

## S4 method for signature 'infusion_wrapper,integer,character'
updateADDL(object, addl, ref = NULL)

## S4 method for signature 'bolus,integer,character'
updateADDL(object, addl, ref = NULL)

## S4 method for signature 'infusion,integer,character'
updateADDL(object, addl, ref = NULL)

## S4 method for signature 'treatment,integer,character'
updateADDL(object, addl, ref = NULL)

## S4 method for signature 'arm,integer,character'
updateADDL(object, addl, ref = NULL)

## S4 method for signature 'arms,integer,character'
updateADDL(object, addl, ref = NULL)

## S4 method for signature 'dataset,integer,character'
updateADDL(object, addl, ref = NULL)
```

Arguments

| | |
|--------|--------------------------------|
| object | generic object |
| add1 | new number of additional doses |
| ref | reference treatment name |

Value

updated object

| | |
|--------------|-----------------------|
| updateAmount | <i>Update amount.</i> |
|--------------|-----------------------|

Description

Update amount.

Usage

```
updateAmount(object, amount, ref)
```

```
## S4 method for signature 'bolus,numeric,character'
updateAmount(object, amount, ref)
```

```
## S4 method for signature 'infusion,numeric,character'
updateAmount(object, amount, ref)
```

```
## S4 method for signature 'bolus_wrapper,numeric,character'
updateAmount(object, amount, ref)
```

```
## S4 method for signature 'infusion_wrapper,numeric,character'
updateAmount(object, amount, ref)
```

```
## S4 method for signature 'treatment,numeric,character'
updateAmount(object, amount, ref)
```

```
## S4 method for signature 'arm,numeric,character'
updateAmount(object, amount, ref)
```

```
## S4 method for signature 'arms,numeric,character'
updateAmount(object, amount, ref)
```

```
## S4 method for signature 'dataset,numeric,character'
updateAmount(object, amount, ref)
```

Arguments

| | |
|--------|--------------------------|
| object | generic object |
| amount | new amount |
| ref | reference treatment name |

Value

updated object

| | |
|----------|---|
| updateII | <i>Update the inter-dose interval (II).</i> |
|----------|---|

Description

Update the inter-dose interval (II).

Usage

```
updateII(object, ii, ref = NULL)

## S4 method for signature 'bolus_wrapper,numeric,character'
updateII(object, ii, ref = NULL)

## S4 method for signature 'infusion_wrapper,numeric,character'
updateII(object, ii, ref = NULL)

## S4 method for signature 'bolus,numeric,character'
updateII(object, ii, ref = NULL)

## S4 method for signature 'infusion,numeric,character'
updateII(object, ii, ref = NULL)

## S4 method for signature 'treatment,numeric,character'
updateII(object, ii, ref = NULL)

## S4 method for signature 'arm,numeric,character'
updateII(object, ii, ref = NULL)

## S4 method for signature 'arms,numeric,character'
updateII(object, ii, ref = NULL)

## S4 method for signature 'dataset,numeric,character'
updateII(object, ii, ref = NULL)
```

Arguments

| | |
|--------|--------------------------|
| object | generic object |
| ii | new inter-dose interval |
| ref | reference treatment name |

Value

updated object

| | |
|--------------|---|
| updateRepeat | <i>Update the repeat field (argument 'rep' in Bolus and Infusion constructors).</i> |
|--------------|---|

Description

Update the repeat field (argument 'rep' in Bolus and Infusion constructors).

Usage

```
updateRepeat(object, rep, ref = NULL)

## S4 method for signature 'bolus_wrapper,repeated_schedule,character'
updateRepeat(object, rep, ref = NULL)

## S4 method for signature 'infusion_wrapper,repeated_schedule,character'
updateRepeat(object, rep, ref = NULL)

## S4 method for signature 'bolus,repeated_schedule,character'
updateRepeat(object, rep, ref = NULL)

## S4 method for signature 'infusion,repeated_schedule,character'
updateRepeat(object, rep, ref = NULL)

## S4 method for signature 'treatment,repeated_schedule,character'
updateRepeat(object, rep, ref = NULL)

## S4 method for signature 'arm,repeated_schedule,character'
updateRepeat(object, rep, ref = NULL)

## S4 method for signature 'arms,repeated_schedule,character'
updateRepeat(object, rep, ref = NULL)

## S4 method for signature 'dataset,repeated_schedule,character'
updateRepeat(object, rep, ref = NULL)
```

Arguments

| | |
|--------|--|
| object | generic object |
| rep | repeated dosing schedule (definition) object |
| ref | reference treatment name |

Value

updated object

| | |
|-----|--|
| VPC | <i>Compute the VPC summary. Input data frame must contain the following columns: - replicate: replicate number - low: low percentile value in replicate (and in scenario if present) - med: median value in replicate (and in scenario if present) - up: up percentile value in replicate (and in scenario if present) - any scenario column</i> |
|-----|--|

Description

Compute the VPC summary. Input data frame must contain the following columns: - replicate: replicate number - low: low percentile value in replicate (and in scenario if present) - med: median value in replicate (and in scenario if present) - up: up percentile value in replicate (and in scenario if present) - any scenario column

Usage

```
VPC(x, scenarios = NULL, level = 0.9)
```

Arguments

| | |
|-----------|--|
| x | data frame |
| scenarios | scenarios, character vector, NULL is default |
| level | PI level, default is 0.9 (90% PI) |

Value

VPC summary with columns TIME, <scenarios> and all combinations of low, med, up (i.e. low_low, low_med, low_up, etc.)

| | |
|---------|------------------|
| vpcPlot | <i>VPC plot.</i> |
|---------|------------------|

Description

VPC plot.

Usage

```
vpcPlot(x, scenarios = NULL, level = 0.9, alpha = 0.15)
```

Arguments

| | |
|-----------|---|
| x | data frame, output of CAMPSIS with replicates |
| scenarios | scenarios, character vector, NULL is default |
| level | PI level, default is 0.9 (90% PI) |
| alpha | alpha parameter (transparency) given to geom_ribbon |

Value

a ggplot object

| | |
|-------|--------------------------------|
| weeks | <i>Convert weeks to hours.</i> |
|-------|--------------------------------|

Description

Convert weeks to hours.

Usage

```
weeks(x)
```

Arguments

| | |
|---|-------------------------|
| x | numeric vector in weeks |
|---|-------------------------|

Value

numeric vector in hours

years

*Convert pharma years (1 year = 12*4 weeks) to hours.*

Description

Convert pharma years (1 year = 12*4 weeks) to hours.

Usage

years(x)

Arguments

x numeric vector in years

Value

numeric vector in hours

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