

Package ‘mrgsolve’

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Type Package

Version 0.8.5

Title Simulate from ODE-Based Population PK/PD and Systems
Pharmacology Models

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URL <https://github.com/metrumresearchgroup/mrgsolve>

BugReports <https://github.com/metrumresearchgroup/mrgsolve/issues>

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Description Facilitates simulation from hierarchical, ordinary differential equation (ODE) based models typically employed in drug development. A model specification file is created consisting of R and C++ code that is parsed, compiled, and dynamically loaded into the R session. Input data are passed in and simulated data are returned as R objects. A dosing event engine allows interventions (bolus and infusion) to be managed separately from the model code. Differential equations are solved with the 'DLSODA' routine in 'ODEPACK' (<<https://computation.llnl.gov/casc/odepack/>>).

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Depends R (>= 3.1.2), methods

Imports Rcpp (>= 0.12.3), dplyr (>= 0.5.0), magrittr (>= 1.5),
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Collate 'RcppExports.R' 'utils.R' 'package.R' 'generics.R'
'class_modlist.R' 'class_tgrid.R' 'class_numericlist.R'
'class_matlist.R' 'class_ev.R' 'class_derived.R'
'class_mrgmod.R' 'class_mrgsims.R' 'Aaaa.R' 'altname.R'
'annot.R' 'chain.R' 'class_build.R' 'compile.R' 'complog.R'
'covset.R' 'data_set.R' 'datasets.R' 'env.R' 'events.R'

```
'example.R' 'funset.R' 'idata_set.R' 'init.R' 'knobs.R'
'library.R' 'matlist.R' 'matrix.R' 'mcache.R' 'model_include.R'
'modlib.R' 'modspec.R' 'mread.R' 'mrgindata.R' 'mrgsims.R'
'mrgsolve.R' 'nmxml.R' 'param.R' 'print.R' 'qsim.R' 'render.R'
'simtime.R' 'update.R'
```

RoxygenNote 6.0.1

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aboutsolver	<i>About the ODEPACK differential equation solver used by mrgsolve.</i>
-------------	---

Description

About the ODEPACK differential equation solver used by mrgsolve.

DLSODA

```

C-----
C This is the 12 November 2003 version of
C DLSODA: Livermore Solver for Ordinary Differential Equations, with
C     Automatic method switching for stiff and nonstiff problems.
C
C This version is in double precision.
C
C DLSODA solves the initial value problem for stiff or nonstiff
C systems of first order ODEs,
C   dy/dt = f(t,y) , or, in component form,
C   dy(i)/dt = f(i) = f(i,t,y(1),y(2),...,y(NEQ)) (i = 1,...,NEQ).

```

```

C
C This a variant version of the DLSODE package.
C It switches automatically between stiff and nonstiff methods.
C This means that the user does not have to determine whether the
C problem is stiff or not, and the solver will automatically choose the
C appropriate method. It always starts with the nonstiff method.
C
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C References:
C 1. Alan C. Hindmarsh, ODEPACK, A Systematized Collection of ODE
C    Solvers, in Scientific Computing, R. S. Stepleman et al. (Eds.),
C    North-Holland, Amsterdam, 1983, pp. 55-64.
C 2. Linda R. Petzold, Automatic Selection of Methods for Solving
C    Stiff and Nonstiff Systems of Ordinary Differential Equations,
C    Siam J. Sci. Stat. Comput. 4 (1983), pp. 136-148.
C-----

```

as_bmat

Coerce R objects to block or diagonal matrices.

Description

Coerce R objects to block or diagonal matrices.

Usage

```

as_bmat(x, ...)

## S4 method for signature 'list'
as_bmat(x, ...)

## S4 method for signature 'numeric'
as_bmat(x, pat = "*", ...)

## S4 method for signature 'data.frame'
as_bmat(x, pat = "*", cols = NULL, ...)

## S4 method for signature 'ANY'

```

```
as_bmat(x, ...)  
  
as_dmat(x, ...)  
  
## S4 method for signature 'list'  
as_dmat(x, ...)  
  
## S4 method for signature 'ANY'  
as_dmat(x, ...)  
  
## S4 method for signature 'numeric'  
as_dmat(x, pat = "*", ...)  
  
## S4 method for signature 'data.frame'  
as_dmat(x, pat = "*", cols = NULL, ...)
```

Arguments

x	an R object
...	passed along
pat	regular expression, character
cols	column names to use instead of pat

Value

A numeric matrix for list and numeric methods. For data.frames, a list of matrices are returned.

See Also

[bmat](#), [dmat](#)

Examples

```
df <- data.frame(OMEGA1.1 = c(1,2),  
                 OMEGA2.1 = c(11,22),  
                 OMEGA2.2 = c(3,4),  
                 SIGMA1.1 = 1,  
                 FOO=-1)  
  
as_bmat(df, "OMEGA")  
as_dmat(df, "SIGMA")  
as_dmat(df[1,], "OMEGA")
```

as_data_set*Create a simulatinon data set from ev objects.*

Description

Create a simulatinon data set from ev objects.

Usage

```
as_data_set(x, ...)  
  
## S4 method for signature 'ev'  
as_data_set(x, ...)
```

Arguments

x	ev objects
...	more ev objects

Details

The goal is to take a series of event objects and combine them into a single data set that can be passed to [data_set](#). Each event object is added to the data frame as an ID or set of IDs that are distinct from the IDs in the other event objects. Note that including ID argument to the [ev](#) call where `length(ID)` is greater than one will render that set of events for all of IDs that are requested.

To get a data frame with one row (event) per ID look at [expand.ev](#).

Value

a data frame suitable for passing into [data_set](#)

Examples

```
as_data_set(ev(amt=c(100,200), cmt=1, ID=1:3),  
            ev(amt=300, time=24, ID=1:2),  
            ev(amt=1000, ii=8, addl=10, ID=1:3))  
  
# Instead of this, use expand.ev  
as_data_set(ev(amt=100), ev(amt=200),ev(amt=300))
```

as_deslist

Create a list of designs from a data frame.

Description

Create a list of designs from a data frame.

Usage

```
as_deslist(data, descol = "ID")
```

Arguments

data	input data set; see details
descol	character column name to be used for design groups

Details

The input data set must have a column with the same name as the value of descol. Other column names should be start (the time of the first observation), end (the time of the last observation), delta (the time steps to take between start and end), and add (other, ad-hoc times). Note that add might be a list-column to get a vector of times for each time grid object.

Value

The function returns a list of tgrid objects, one for each unique value found in descol.

Examples

```
idata <- dplyr::data_frame(ID=1:4, end=seq(24,96,24), delta=6,
  add=list(c(122,124,135),c(111), c(99),c(88)))

idata <- dplyr::mutate(idata, GRP = ID %%2)

idata

l <- as_deslist(idata,"GRP")

l

lapply(l,stime)

lapply(as_deslist(idata, "ID"),stime)
```

as.list,mrgmod-method *Coerce a model object to list.*

Description

Coerce a model object to list.

Usage

```
## S4 method for signature 'mrgmod'
as.list(x, ...)
```

Arguments

x	mrgmod object
...	passed to other methods

assign_ev *Replicate a list of events into a data set.*

Description

Replicate a list of events into a data set.

Usage

```
assign_ev(l, idata, evgroup, join = FALSE)
```

Arguments

l	list of event objects
idata	an idata set (one ID per row)
evgroup	the character name of the column in idata that specifies event object to implement
join	if TRUE, join idata to the data set before returning.

Examples

```
ev1 <- ev(amt=100)
ev2 <- ev(amt=300, rate=100, ii=12, addl=10)

idata <- data.frame(ID=1:10)
idata$arm <- 1+(idata$ID %%2)

assign_ev(list(ev1, ev2), idata, "arm", join=TRUE)
```

BLOCK_PARSE

*Functions to parse code blocks.***Description**

Most of the basic blocks are listed in this help topic. But see also [PKMODEL](#) which has more-involved options and is documented separately.

Usage

```
PARAM(x, env, annotated = FALSE, pos = 1, ...)
```

```
FIXED(x, env, annotated = FALSE, pos = 1, ...)
```

```
THETA(x, env, annotated = FALSE, pos = 1, name = "THETA", ...)
```

```
INIT(x, env, annotated = FALSE, pos = 1, ...)
```

```
CMT(x, env, annotated = FALSE, pos = 1, ...)
```

```
CAPTURE(x, env, annotated = FALSE, pos = 1, ...)
```

Arguments

x	data
env	parse environment
annotated	logical
pos	block position
...	passed
name	block name

See Also

[PKMODEL](#)

blocks

*Return the code blocks from a model specification file.***Description**

Return the code blocks from a model specification file.

Usage

```
blocks(x, ...)

## S4 method for signature 'mrgmod'
blocks(x, ...)

## S4 method for signature 'character'
blocks(x, ...)
```

Arguments

x	model object or path to model specification file
...	passed along

Examples

```
mod <- mrgsolve:::house()
mod %>% blocks
mod %>% blocks(PARAM, TABLE)
```

bmat

*Create matrices from vector input.***Description**

Create matrices from vector input.

Usage

```
bmat(..., correlation = FALSE, digits = -1)

cmat(..., digits = -1)

dmat(...)
```

Arguments

...	matrix data
correlation	logical; if TRUE, off diagonal elements are assumed to be correlations and converted to covariances
digits	if greater than zero, matrix is passed to signif (along with digits) prior to returning

Details

bmat makes a block matrix. cmat makes a correlation matrix. dmat makes a diagonal matrix.

See Also

[as_bmat](#)
[as_dmat](#)

Examples

```
dmat(1,2,3)/10  
bmat(0.5,0.01,0.2)  
cmat(0.5, 0.87,0.2)
```

cama	<i>Run the model cama function.</i>
------	-------------------------------------

Description

Run the model cama function.

Usage

```
cama(mod, fn = "cama", ...)
```

Arguments

mod	model object
fn	function name
...	passed to update

Details

sah-mah

carry_out	<i>Select items to carry into simulated output.</i>
-----------	---

Description

When items named in this function are found in the input data set (either [data_set](#) or [idata_set](#)), they are copied into the simulated output. Special items like evid or amt or the like are not copied from the data set per se, but they are copied from datarecord objects that are created during the simulation.

Usage

```
carry_out(x, ...)
```

```
carry.out(x, ...)
```

Arguments

x	model object
...	passed along

Details

There is also a `carry.out` argument to [mrgsim](#) that can be set to accomplish the same thing as a call to `carry_out` in the pipeline.

`carry.out` and `carry_out`. Using the underscore version is now preferred.

chain	<i>Functions for chaining commands together.</i>
-------	--

Description

Use these functions with chaining commands together with the operator.

Details

Other functions that may be used in the chain of commands include: [param](#), [init](#), [update](#), [ev](#), or any other function that will take the output of the preceeding command as it's first argument.

Examples

```
mod <- mrgsolve:::house()

data(exidata)
data(exTheoph)

out <- mod %>% data_set(exTheoph) %>% mrgsim()
out <- mod %>% carry_out(evid) %>% ev(amt=100, cmt=1) %>% mrgsim()
out <- mod %>% Req(CP,RESP) %>% mrgsim()
```

c,matlist-method	<i>Operations with matlist objects.</i>
------------------	---

Description

Operations with matlist objects.

Usage

```
## S4 method for signature 'matlist'
c(x, ..., recursive = FALSE)
```

Arguments

x	a matlist object
...	other matlist objects
recursive	not used

cmt_list-class	<i>S4 cmt_list class</i>
----------------	--------------------------

Description

S4 cmt_list class

Details

cmt_list is a [numericlist-class](#)

cmtn	<i>Get the compartment number from a compartment name.</i>
------	--

Description

Get the compartment number from a compartment name.

Usage

```
cmtn(x, ...)  
  
## S4 method for signature 'mrgmod'  
cmtn(x, tag, ...)
```

Arguments

x	model object
...	passed along
tag	compartment name

Examples

```
mod <- mrgsolve:::house()  
mod %>% cmtn("CENT")
```

code	<i>Extract the code from a model.</i>
------	---------------------------------------

Description

Extract the code from a model.

Usage

```
code(x)
```

Arguments

x	an mrgsolve model object
---	--------------------------

Value

a character vector of model code

c,tgrid-method	<i>Operations with tgrid objects.</i>
----------------	---------------------------------------

Description

Operations with tgrid objects.

Usage

```
## S4 method for signature 'tgrid'
c(x, ..., recursive = FALSE)

## S4 method for signature 'tgrids'
c(x, ..., recursive = FALSE)

## S4 method for signature 'tgrid,numeric'
e1 + e2

## S4 method for signature 'tgrid,numeric'
e1 * e2

## S4 method for signature 'tgrids,numeric'
e1 + e2

## S4 method for signature 'tgrids,numeric'
e1 * e2
```

Arguments

x	mrgmod object
...	passed along to other methods
recursive	not used
e1	tgrid or tgrids object
e2	numeric value

cvec	<i>Create create character vectors.</i>
------	---

Description

Create create character vectors.

Usage

```
cvec(x, ...)
```

```
## S4 method for signature 'character'
```

```
cvec(x)
```

```
ch(...)
```

```
s(...)
```

Arguments

x	comma-separated quoted string (for cvec)
...	unquoted strings (for ch)

Examples

```
cvec("A,B,C")
ch(A,B,C)
s(A,B,C)
```

data_set	<i>Select and modify a data set for simulation.</i>
----------	---

Description

Select and modify a data set for simulation.

Usage

```
data_set(x, data, ...)
```

```
## S4 method for signature 'mrgmod,data.frame'
```

```
data_set(x, data, subset = TRUE,
```

```
  select = TRUE, object = NULL, ...)
```

```
## S4 method for signature 'mrgmod,ANY'
```

```
data_set(x, data, ...)
```

```
## S4 method for signature 'mrgmod,missing'
```

```
data_set(x, object, ...)
```

Arguments

<code>x</code>	model object
<code>data</code>	data set
<code>...</code>	passed along
<code>subset</code>	passed to <code>dplyr::filter_</code> ; retain only certain rows in the data set
<code>select</code>	passed to <code>dplyr::select_</code> ; retain only certain columns in the data set
<code>object</code>	character name of an object existing in <code>\$ENV</code> to use for the data set

Details

Input data sets are R data frames that can include columns with any valid name, however columns with selected names are treated specially by `mrugsolve` and incorporated into the simulation.

ID specifies the subject ID and is required for every input data set.

When columns have the same name as parameters (`$PARAM` in the model specification file), the values in those columns will be used to update the corresponding parameter as the simulation progresses.

Input data set may include the following columns related to PK dosing events: `time`, `cmt`, `amt`, `rate`, `ii`, `addl`, `ss`. `time` and `cmt` (and ID) are required columns in the input data set. `time` is the observation or event time, `cmt` is the compartment number (see [init](#)), `amt` is the dosing amount, `rate` is the infusion rate, `ii` is the dosing interval, `addl` specifies additional doses to administer, and `ss` is a flag for steady state dosing. These column names operate similarly to other non-linear mixed effects modeling software. Upper case PK dosing column names including `TIME`, `CMT`, `AMT`, `RATE`, `II`, `ADDL`, `SS` are also recognized. However, an error will be generated if a mix of upper case and lower case columns are found.

Only numeric data can be brought in to the problem. Any non-numeric data columns will be dropped with warning.

See [exdatasets](#) for different example data sets.

See Also

[idata_set](#), [ev](#), [valid_data_set](#), [valid_idata_set](#)

Examples

```
mod <- mrugsolve:::house()

data <- expand.ev(ID=1:3, amt=c(10,20))

mod %>% data_set(data, ID > 1) %>% mrugsim

data(extran1)
head(extran1)

mod %>% data_set(extran1) %>% mrugsim
mod %>% mrugsim(data=extran1)
```

design	<i>Set observation designs for the simulation.</i>
--------	--

Description

This function also allows you to assign different designs to different groups or individuals in a population.

Usage

```
design(x, deslist = list(), descol = character(0), ...)
```

Arguments

x	model object
deslist	a list of tgrid or tgrids objects or numeric vector to be used in place of ...
descol	the idata column name (character) for design assignment
...	not used

Details

This setup requires the use of an `idata_set`, with individual-level data passed in one ID per row. For each ID, specify a grouping variable in `idata` (`descol`). For each unique value of the grouping variable, make one `tgrid` object and pass them in order as ... or form them into a list and pass as `deslist`.

You must assign the `idata_set` before assigning the designs in the command chain (see the example below).

Examples

```
peak <- tgrid(0,6,0.1)
sparse <- tgrid(0,24,6)

des1 <- c(peak,sparse)
des2 <- tgrid(0,72,4)

data <- expand.ev(ID = 1:10, amt=c(100,300))
data$GRP <- data$amt/100

idata <- data[,c("ID", "amt")]

mod <- mrgsolve:::house()

mod %>%
  omat(dmat(1,1,1,1)) %>%
  carry_out(GRP) %>%
```

```
idata_set(idata) %>%
design(list(des1, des2),"amt") %>%
data_set(data) %>%
mrgsim %>%
plot(RES~time|GRP)
```

details	<i>Extract model details.</i>
---------	-------------------------------

Description

Extract model details.

Usage

```
details(x, complete = FALSE, values = FALSE, ...)
```

Arguments

x	a model object
complete	logical; if TRUE, un-annotated parameters and compartments will be added to the output
values	logical; if TRUE, a values column will be added to the output
...	not used

env_eval	<i>Re-evaluate the code in the ENV block.</i>
----------	---

Description

The \$ENV block is a block of R code that can realize any sort of R object that might be used in running a model.

Usage

```
env_eval(x, seed = NULL)
```

Arguments

x	model object
seed	passed to set.seed if a numeric value is supplied

See Also

[env_get](#), [env_ls](#)

env_get	<i>Return model environment.</i>
---------	----------------------------------

Description

Return model environment.

Usage

```
env_get(x, tolist = TRUE)
```

Arguments

x	model object
tolist	should the environment be coerced to list?

env_ls	<i>List objects in the model environment.</i>
--------	---

Description

Each model keeps an internal environment that allows the user to carry any R object along. Objects are coded in \$ENV.

Usage

```
env_ls(x, ...)
```

Arguments

x	model object
...	passed to ls

env_update	<i>Update objects in model environment.</i>
------------	---

Description

Update objects in model environment.

Usage

```
env_update(.x, ..., .dots = list())
```

Arguments

.x	model object
...	objects to update
.dots	list of objects to updated

ev-class	<i>S4 events class</i>
----------	------------------------

Description

S4 events class

Slots

data a data frame of events

ev_days	<i>Schedule dosing events on days of the week.</i>
---------	--

Description

This function lets you schedule doses on specific days of the week, allowing you to create dosing regimens on Monday/Wednesday/Friday, or Tuesday/Thursday, or every other day (however you want to define that) etc.

Usage

```
ev_days(ev = NULL, days = "", addl = 0, ii = 168, unit = c("hours",  
  "days"), ...)
```

Arguments

<code>ev</code>	an event object
<code>days</code>	comma- or space-separated character string of valid days of the the week (see details)
<code>addl</code>	additional doses to administer
<code>ii</code>	inter-dose interval; intended use is to keep this at the default value
<code>unit</code>	time unit; the function can only currently handle hours or days
<code>...</code>	event objects named by one the valid days of the week (see details)

Details

Valid names of the week are:

- m for Monday
- t for Tuesday
- w for Wednesday
- th for Thursday
- f for Friday
- sa for Saturday
- s for Sunday

The whole purpose of this function is to schedule doses on specific days of the week, in a repeating weekly schedule. Please do use caution when changing `ii` from it's default value.

Examples

```
# Monday, Wednesday, Friday x 4 weeks
ev_days(ev(amt=100), days="m,w,f", addl=3)

# 50 mg Tuesdays, 100 mg Thursdays x 6 months
ev_days(t=ev(amt=50), th=ev(amt=100), addl=23)
```

events

Event objects for simulating PK and other interventions.

Description

Events can either be specified when the model object is created (with `mrghmod`) or by updating an existing model object (with `update`).

Usage

```

events(x, ...)

ev(x, ...)

as.ev(x, ...)

## S4 method for signature 'mrgmod'
events(x, ...)

## S4 method for signature 'mrgmod'
ev(x, object = NULL, ...)

## S4 method for signature 'missing'
ev(time = 0, evid = 1, ID = numeric(0), cmt = 1,
    replicate = TRUE, until = NULL, realize_add1 = FALSE, ...)

## S4 method for signature 'ev'
ev(x, realize_add1 = FALSE, ...)

## S4 method for signature 'data.frame'
as.ev(x, nid = 1, ...)

## S4 method for signature 'ev'
as.matrix(x, ...)

## S4 method for signature 'ev'
as.data.frame(x, row.names = NULL, optional = FALSE, ...)

## S4 method for signature 'ev'
show(object)

## S4 method for signature 'mrgsims'
events(x, ...)

```

Arguments

x	mrgmodel object
...	passed on
object	passed to show
time	event time
evid	event ID
ID	subject ID
cmt	compartment
replicate	logical; if TRUE, events will be replicated for each individual in ID
until	the expected maximum observation time for this regimen

<code>realize_addl</code>	if FALSE (default), no change to addl doses. If TRUE, addl doses are made explicit with <code>realize_addl</code> .
<code>nid</code>	if greater than 1, will expand to the appropriate number of individuals
<code>row.names</code>	passed to <code>as.data.frame</code>
<code>optional</code>	passed to <code>as.data.frame</code>

Details

- Required input for creating events objects include time and cmt
- If not supplied, evid is assumed to be 1.
- If not supplied, cmt is assumed to be 1.
- If not supplied, time is assumed to be 0.
- ID may be specified as a vector.
- If replicate is TRUE (default), then the events regimen is replicated for each ID; otherwise, the number of event rows must match the number of IDs entered

Value

Returns a user-defined data frame of events that should be suitable for passing into `lsoda`. If events are stored as a data frame, `events` returns the data frame. If events are stored as a function that generates the data frame, `events` calls the function and passes return back to the user.

events object

Examples

```
mod <- mrgsolve:::house()
mod <- mod %>% ev(amt=1000, time=0, cmt=1)
events(mod)

loading <- ev(time=0, cmt=1, amt=1000)
maint <- ev(time=12, cmt=1, amt=500, ii=12, addl=10)
loading + maint

ev(ID=1:10, cmt=1, time=0, amt=100)
```

Description

Operations for ev objects.

Usage

```
## S4 method for signature 'ev,ev'
e1 + e2

e1 %then% e2

## S4 method for signature 'ev,ev'
e1 %then% e2

## S4 method for signature 'ev,numeric'
e1 + e2

## S4 method for signature 'ev'
c(x, ..., recursive = TRUE)
```

Arguments

e1	object on left hand side of operator (lhs)
e2	object on right hand side of operator (rhs)
x	an ev object
...	other ev objects to collect
recursive	not used

Details

All operations involving [mrgmod](#) objects have been deprecated.

exdatasets	<i>Example input data sets.</i>
------------	---------------------------------

Description

Example input data sets.

Usage

```
data(exidata)

data(extran1)

data(extran2)

data(extran3)

data(exTheoph)

data(exBoot)
```

Details

- exidata holds individual-level parameters and other data items, one per row
- extran1 is a "condensed" data set
- extran2 is a full dataset
- extran3 is a full dataset with parameters
- exTheoph is the theophylline data set, ready for input into mrgsolve
- exBoot a set of bootstrap parameter estimates

Examples

```
mod <- mrgsolve:::house() %>% update(end=240) %>% Req(CP)

## Full data set
data(exTheoph)
out <- mod %>% data_set(exTheoph) %>% mrgsim
out
plot(out)

## Condensed: mrgsolve fills in the observations
data(extran1)
out <- mod %>% data_set(extran1) %>% mrgsim
out
plot(out)

## Add a parameter to the data set
stopifnot(require(dplyr))
data <- extran1 %>% distinct(ID) %>% select(ID) %>%
  mutate(CL=exp(log(1.5) + rnorm(nrow(.), 0, sqrt(0.1)))) %>%
  left_join(extran1,.)

data

out <- mod %>% data_set(data) %>% carry.out(CL) %>% mrgsim
out
plot(out)

## idata
data(exidata)
out <- mod %>% idata_set(exidata) %>% ev(amt=100,ii=24,addl=10) %>% mrgsim
plot(out, CP~time|ID)
```

expand.idata

Create template data sets for simulation.

Description

Create template data sets for simulation.

Usage

```
expand.idata(...)
```

```
expand.ev(...)
```

Arguments

```
...                passed to expand.grid
```

Details

An ID column is added as 1:nrow(ans) if not supplied by the user. For expand.ev, defaults also added: cmt = 1, time = 0, evid = 1.

Examples

```
idata <- expand.idata(CL=c(1,2,3), VC=c(10,20,30))
```

```
doses <- expand.ev(amt=c(300,100), ii=c(12,24), cmt=1)
```

file_show

Show model specification and C++ files.

Description

Show model specification and C++ files.

Usage

```
file_show(x, spec = TRUE, source = TRUE, ...)
```

Arguments

x	model object
spec	logical; show the model specification file
source	logical; show the C++ file that is actually compiled
...	not used

house	<i>Return a pre-compiled, PK/PD model.</i>
-------	--

Description

Return a pre-compiled, PK/PD model.

Usage

```
house(...)
```

Arguments

... passed to update

Value

A packmod object, ready to simulate.

Examples

```
mod <- mrgsolve:::house()
see(mod)

mod %>% ev(amt=100) %>% mrgsim %>% plot
```

idata_set	<i>Select and modify a idata set for simulation.</i>
-----------	--

Description

Select and modify a idata set for simulation.

Usage

```
idata_set(x, data, ...)

## S4 method for signature 'mrgmod,data.frame'
idata_set(x, data, subset = TRUE,
  select = TRUE, object = NULL, ...)

## S4 method for signature 'mrgmod,ANY'
idata_set(x, data, ...)

## S4 method for signature 'mrgmod,missing'
idata_set(x, object, ...)
```

Arguments

<code>x</code>	model object
<code>data</code>	a data set coercable to <code>data.frame</code>
<code>...</code>	passed along
<code>subset</code>	passed to <code>dplyr::filter_</code>
<code>select</code>	passed to <code>dplyr::select_</code>
<code>object</code>	character name of an object existing in <code>\$ENV</code> to use for the data set

Details

The `idata_set` is a `data.frame` that specifies individual-level data for the problem. An ID column is required and there can be no more than one row in the data frame for each individual.

In most cases, the columns in the ‘`idata_set`’ have the same names as parameters in the [param](#) list. When this is the case, the parameter set is updated as the simulation proceeds once at the start of each individual. The ‘`idata_set`’ can also be used to set initial conditions for each individual: for a compartment called CMT, make a column in `idata_set` called `CMT_0` and make the value the desired initial value for that compartment. Note that this initial condition will be over-ridden if you also set the `CMT_0` in `$MAIN`.

The most common application of `idata_set` is to specify a population or batch of simulations to do. We commonly use `idata_set` with an event object (see [ev](#)). In that case, the event gets applied to each individual in the [idata_set](#).

It is also possible to provide both a `data_set` and a `idata_set`. In this case, the `idata_set` is used as a parameter lookup for IDs found in the `data_set`. Remember in this case, it is the `data_set` (not the `idata_set`) that determines the number of individuals in the simulation.

See Also

[data_set](#), [ev](#)

Examples

```
mod <- mrgsolve:::house()

data(exidata)

exidata

mod %>% idata_set(exidata, ID <= 2) %>% mrgsim %>% plot

mod %>% idata_set(exidata) %>% mrgsim

mod %>% mrgsim(idata=exidata)
```

`init`*Methods for working with the model compartment list.*

Description

Calling `init` with the model object as the first argument will return the model initial conditions as a `numericlist` object. See [numericlist](#) for methods to deal with `cmt_list` objects.

Usage

```
init(.x, ...)  
  
## S4 method for signature 'mrgmod'  
init(.x, .y = list(), ..., .pat = "*")  
  
## S4 method for signature 'mrgsims'  
init(.x, ...)  
  
## S4 method for signature 'missing'  
init(.x, ...)  
  
## S4 method for signature 'list'  
init(.x, ...)  
  
## S4 method for signature 'ANY'  
init(.x, ...)  
  
as.init(.x, ...)  
  
## S4 method for signature 'list'  
as.init(.x, ...)  
  
## S4 method for signature 'numeric'  
as.init(.x, ...)  
  
## S4 method for signature 'cmt_list'  
as.init(.x, ...)  
  
## S4 method for signature 'missing'  
as.init(.x, ...)  
  
## S4 method for signature '`NULL`'  
as.init(.x, ...)  
  
## S4 method for signature 'cmt_list'  
show(object)
```

Arguments

<code>.x</code>	the model object
<code>...</code>	passed along
<code>.y</code>	list to be merged into parameter list
<code>.pat</code>	a regular expression (character) to be applied as a filter when printing compartments to the screen
<code>object</code>	to show

Details

Can be used to either get a compartment list object from a mrgmod model object or to update the compartment initial conditions in a model object. For both uses, the return value is a `cmt_list` object. For the former use, `init` is usually called to print the compartment initial conditions to the screen, but the `cmt_list` object can also be coerced to a list or numeric R object.

Value

an object of class `cmt_list` (see [numericlist](#))

Examples

```
## example("init")
mod <- mrgsolve:::house()

init(mod)
init(mod, .pat="^C") ## may be useful for large models

class(init(mod))

init(mod)$CENT

as.list(init(mod))
as.data.frame(init(mod))
```

`is.mrgmod`

Check if an object is a model object.

Description

The function checks to see if the object is either `mrgmod` or `packmod`.

Usage

```
is.mrgmod(x)
```

Arguments

<code>x</code>	any object
----------------	------------

Value

TRUE if x inherits mrgsims.

is.mrgsims

Check if an object is mrgsim output.

Description

Check if an object is mrgsim output.

Usage

```
is.mrgsims(x)
```

Arguments

x any object

Value

TRUE if x inherits mrgsims.

knobs

Run sensitivity analysis on model settings.

Description

Knobs can be parameter values or PK dosing items (e.g. amt). By design, all combinations of specified knob/values are simulated.

Usage

```
knobs(x, y, ...)
```

```
## S4 method for signature 'mrgmod,missing'
knobs(x, y, ...)
```

```
## S4 method for signature 'mrgmod,batch_mrgsims'
knobs(x, y, ...)
```

```
## S4 method for signature 'batch_mrgsims'
as.data.frame(x, row.names = NULL,
  optional = FALSE, ...)
```

```
## S4 method for signature 'batch_mrgsims,ANY'
```

```
knobs(x, y, ...)

## S4 method for signature 'batch_mrgsims'
show(object)
```

Arguments

<code>x</code>	the model object
<code>y</code>	a <code>batch_mrgsims</code> object
<code>...</code>	<code>knobs</code> : named numeric vectors that identify knob names and knob values for a batch run. See details.
<code>row.names</code>	passed to <code>as.data.frame</code>
<code>optional</code>	passed to <code>as.data.frame</code>
<code>object</code>	passed to <code>show</code>

Details

Valid knob names include: any parameter name (in `param(mod)`), time variables (`start`, `end`, `delta`), PK dosing items (`amt`, `ii`, `rate`, and others ...), and solver settings (`atol`, `hmax`, etc...).

Value

An object of class `batch_mrgsims`. Most methods for `mrgsims` objects also work on `batch_mrgsims` object.

Examples

```
## example("knobs")

mod <- mrgsolve:::house(end=72)

events <- ev(amt=1000, cmt=1, addl=3, ii=12)

out <- mod %>% ev(events) %>% knobs(CL=c(1,2,3))
plot(out)

out

out <- mod %>% ev(events) %>% knobs(CL=c(1,2,3), VC=c(5,20,50))
plot(out)
plot(out,CP~.)
plot(out, CP~time|VC, groups=CL, lty=2)

out <- knobs(mod, amt=c(100,300,500), cmt=1)
plot(out)

out <- mod %>% knobs(amt=c(100,300), CL=c(1,3), VC=c(5,20), cmt=1)
plot(out)
plot(out, CP~.)
```

```
out <- knobs(mod, CL=c(1,2,3))
out
```

```
out <- knobs(mod, CL=c(1,2,3))
out
```

lctran	<i>Convert select upper case column names to lower case to conform to mrgsolve data expectations.</i>
--------	---

Description

Convert select upper case column names to lower case to conform to mrgsolve data expectations.

Usage

```
lctran(data)
```

Arguments

data	an nmtran-like data frame
------	---------------------------

Details

Columns that will be renamed with lower case versions: AMT, II, SS, CMT, ADDL, RATE, EVID, TIME. If a lower case version of these names exist in the data set, the column will not be renamed.

Value

A data.frame with renamed columns.

loadso	<i>Load the model shared object.</i>
--------	--------------------------------------

Description

Load the model shared object.

Usage

```
loadso(x, ...)
```

```
## S4 method for signature 'mrgmod'
loadso(x, ...)
```

Arguments

x	the model object
...	passed along

lower2matrix	<i>Create a square numeric matrix from the lower-triangular elements.</i>
--------------	---

Description

Create a square numeric matrix from the lower-triangular elements.

Usage

```
lower2matrix(x, context = NULL)
```

Arguments

x	numeric data
context	the working context

Value

a square symmetric numeric matrix with column names

matlist	<i>Methods for working with matrix-list objects.</i>
---------	--

Description

Methods for working with matrix-list objects.

Usage

```
zero.re(.x, ...)
```

S4 method for signature 'mrgmod'

```
zero.re(.x, ...)
```

```
zero_re(...)
```

```
drop.re(.x, ...)
```

S4 method for signature 'mrgmod'

```
drop.re(.x, ...)
```

```
drop_re(...)
```

S4 method for signature 'matlist'

```
as.list(x, ...)
```

```
## S4 method for signature 'matlist'
as.matrix(x, ...)

## S4 method for signature 'matlist'
names(x)

## S4 method for signature 'matlist'
length(x)

## S4 method for signature 'matlist'
labels(object, ...)

## S4 method for signature 'matlist'
dim(x)

## S4 method for signature 'matlist'
nrow(x)

## S4 method for signature 'matlist'
show(object)
```

Arguments

<code>.x</code>	a matlist object
<code>...</code>	passed along
<code>x</code>	a matlist object
<code>object</code>	passed to showmatlist

matlist-class	<i>S4 class matlist.</i>
---------------	--------------------------

Description

S4 class matlist.

mcode_cache	<i>Write, compile, and load model code.</i>
-------------	---

Description

This is a convenience function that ultimately calls [mread](#).

Usage

```
mcode_cache(model, code, project = tempdir(), ...)
```

```
mcode(model, code, project = tempdir(), ...)
```

Arguments

model	model name
code	character string specifying a mrgsolve model
project	project name
...	passed to mread

Details

Note that the arguments are in slightly different order than [mread](#). The default project is `tempdir()`.

Examples

```
code <- '
$CMT DEPOT CENT
$PKMODEL ncmt=1, depot=TRUE
$MAIN
double CL = 1;
double V = 20;
double KA = 1;
'
```

```
mod <- mcode("example",code)
```

mcRNG

Set RNG to use L'Ecuyer-CMRG.

Description

Set RNG to use L'Ecuyer-CMRG.

Usage

```
mcRNG()
```

merge.list	<i>Merge two lists.</i>
------------	-------------------------

Description

Merge two lists.

Usage

```
## S3 method for class 'list'
merge(x, y, ..., open = FALSE, warn = TRUE,
      context = "object", wild = "...")
```

Arguments

x	the original list
y	the new list for merging
...	not used
open	logical indicating whether or not new items should be allowed in the list upon merging.
warn	issue warning if nothing found to update
context	description of usage context
wild	wild-card name; see details

Details

Wild-card names (wild) are always retained in x and are brought along from y only when open.

mod	<i>Return the model object.</i>
-----	---------------------------------

Description

Return the model object.

Usage

```
mod(x, ...)

## S4 method for signature 'mrgsims'
mod(x, ...)
```

Arguments

x	mrgsims object
...	passed along

modelparse	<i>Parse model specification text.</i>
------------	--

Description

Parse model specification text.

Usage

```
modelparse(txt, split = FALSE, drop_blank = TRUE, comment_re = c("//",
  "##"), ...)
```

Arguments

txt	model specification text
split	logical
drop_blank	logical; TRUE if blank lines are to be dropped
comment_re	regular expression for comments
...	arguments passed along

modlib	<i>Internal model library.</i>
--------	--------------------------------

Description

Internal model library.

Usage

```
modlib(list = FALSE)
```

Arguments

list	list available models
------	-----------------------

Details

See [modlib_details](#), [modlib_pk](#), [modlib_pkpd](#), [modlib_tmdd](#), [modlib_viral](#) for details.

Call `modlib(list=TRUE)` to list available models. Once the model is loaded (see examples below), call `mrgsolve:::code(mod)` to see model code and equations.

Examples

```
## Not run:
mod <- mread("pk1cmt", modlib())
mod <- mread("pk2cmt", modlib())
mod <- mread("pk3cmt", modlib())
mod <- mread("irm1", modlib())
mod <- mread("irm2", modlib())
mod <- mread("irm3", modlib())
mod <- mread("irm4", modlib())
mod <- mread("emax", modlib())
mod <- mread("effect", modlib())
mod <- mread("tmdd", modlib())
mod <- mread("viral1", modlib())
mod <- mread("viral2", modlib())

mrgsolve:::code(mod)

## End(Not run)
```

modlib_details	<i>modlib: PK/PD Model parameters, compartments, and output variables.</i>
----------------	--

Description

modlib: PK/PD Model parameters, compartments, and output variables.

Compartments

- EV1, EV2: extravascular dosing compartments
- CENT: central PK compartment
- PERIPH: peripheral PK compartment
- PERIPH2: peripheral PK compartment 2
- RESP: response PD compartment (irm models)

Output variables

- CP: concentration in the central compartment (CENT/VC)
- RESP: response (emax model)

PK parameters

- KA1, KA2: first order absorption rate constants from first and second extravascular compartment (1/time)
- CL: clearance (volume/time)

- VC: volume of distribution, central compartment (volume)
- VP: volume of distribution, peripheral compartment (volume)
- VP2: volume of distribution, peripheral compartment 2 (volume)
- Q: intercompartmental clearance (volume/time)
- Q2: intercompartmental clearance 2 (volume/time)
- VMAX: maximum rate, nonlinear process (mass/time)
- KM: Michaelis constant (mass/volume)
- K10: elimination rate constant (1/time); CL/VC
- K12: rate constant for transfer to peripheral compartment from central (1/time); Q/VC
- K21: rate constant for transfer to central compartment from peripheral (1/time); Q/VP

PD parameters

- E0: baseline effect (emax model)
- EMAX, IMAX: maximum effect (response)
- EC50, IC50: concentration producing 50 percent of effect (mass/volume)
- KIN: zero-order response production rate (irm models) (response/time)
- KOUT: first-order response elimination rate (irm models) (1/time)
- n: sigmoidicity factor
- KE0: rate constant for transfer to effect compartment (1/time)

modlib_pk	<i>modlib: Pharmacokinetic models.</i>
-----------	--

Description

modlib: Pharmacokinetic models.

Arguments

... passed to update

Details

See [modlib_details](#) for more detailed descriptions of parameters and compartments.

The pk1cmt model is parameterized in terms of CL, VC, KA1 and KA2 and uses compartments EV1, EV2, and CENT. The pk2cmt model adds a PERIPH compartment and parameters Q and VP to that of the one-compartment model. Likewise, the three-compartment model (pk3cmt) adds PERIPH2 and parameters Q2 and VP2 to that of the two-compartment models. All pk models also have parameters VMAX (defaulting to zero, no non-linear clearance) and KM.

Value

an object of class packmod

Model description

All pk models have two extravascular dosing compartments and potential for linear and nonlinear clearance.

- pk1cmt: one compartment pk model
- pk2cmt: two compartment pk model
- pk3cmt: three compartment pk model

modlib_pkpd

modlib: Pharmacokinetic / pharmacodynamic models.

Description

modlib: Pharmacokinetic / pharmacodynamic models.

Details

See [modlib_details](#) for more detailed descriptions of parameters and compartments.

All PK/PD models include 2-compartment PK model with absorption from 2 extravascular compartments and linear + nonlinear clearance. The PK models are parameterized with CL, VC, Q, VMAX, KM, KA1 and KA2 and implement compartments EV1, EV2, CENT, PERIPH. The indirect response models have compartment RESP and the emax model has output variable RESP. PD parameters include KIN, KOUT, IC50, EC50, IMAX, EMAX, E0, and n.

Also, once the model is loaded, use [see](#) method for mrgmod to view the model code.

Model description

- irm1 inhibition of response production
- irm2 inhibition of response loss
- irm3 stimulation of response production
- irm4 stimulation of response loss
- pd_effect effect compartment model
- emax sigmoid emax model

`modlib_tmdd`*modlib: Target mediated disposition model.*

Description

modlib: Target mediated disposition model.

Arguments

... passed to update

Parameters

- KEL: elimination rate constant
- KTP: tissue to plasma rate constant
- KPT: plasma to tissue rate constant
- VC: volume of distribution
- KA1, KA2: absorption rate constants
- KINT: internalization rate constant
- KON: association rate constant
- KOFF: dissociation rate constant
- KSYN: target synthesis rate
- KDEG: target degradation rate constant

Compartments

- CENT: unbound drug in central compartment
- TISS: unbound drug in tissue compartment
- REC: concentration of target
- RC: concentration of drug-target complex
- EV1, EV2: extravascular dosing compartments

Output variables

- CP: unbound drug in the central compartment
- TOTAL: total concentration of target (complexed and uncomplexed)

`modlib_viral`*modlib: HCV viral dynamics models.*

Description

modlib: HCV viral dynamics models.

Models

- `viral1`: viral dynamics model with single HCV species
- `viral2`: viral dynamics model with wild-type and mutant HCV species

Parameters

- `s`: new hepatocyte synthesis rate (cells/ml/day)
- `d`: hepatocyte death rate constant (1/day)
- `p`: viral production rate constant (copies/cell/day)
- `beta`: new infection rate constant (ml/copy/day)
- `delta`: infected cell death rate constant (1/day)
- `c`: viral clearance rate constant (1/day)
- `fit`: mutant virus fitness
- `N`: non-target hepatocytes
- `mu`: forward mutation rate
- `Tmax`: maximum number of target hepatocytes (cells/ml)
- `rho`: maximum hepatocyte regeneration rate (1/day)

Compartments

- `T`: uninfected target hepatocytes (cells/ml)
- `I`: productively infected hepatocytes (cells/ml)
- `V`: hepatitis C virus (copies/ml)
- `IM`: mutant infected hepatocytes (cells/ml)
- `VM`: mutant hepatitis C virus (copies/ml)
- `expos`: exposure metric to drive pharmacodynamic model

modlist	<i>Create a modlist object.</i>
---------	---------------------------------

Description

Create a modlist object.

Usage

```
modlist(project = ".", soloc = tempdir(), prefix = "",
        pattern = paste0(prefix, "*\\*.cpp$"), index_file = "MODLIST")
```

Arguments

project	file path to models
soloc	directory where the models will be built
prefix	leading tag for models to process
pattern	a regular expression for models to get
index_file	name of file to look for registered models

modlist-class	<i>S4 class matlist.</i>
---------------	--------------------------

Description

S4 class matlist.

Usage

```
## S4 method for signature 'modlist'
x$name
```

Arguments

x	modlist object
name	model to take; used with \$

modMATRIX

*Create a matrix.***Description**

Create a matrix.

Usage

```
modMATRIX(x, use = TRUE, block = FALSE, correlation = FALSE,
  digits = -1, context = "matlist", ...)
```

Arguments

x	data for building the matrix. Data in x are assumed to be on-diagonal elements if block is FALSE and lower-triangular elements if block is TRUE
use	logical; if FALSE, all matrix elements are set to 0
block	logical; if TRUE, try to make a block matrix; diagonal otherwise
correlation	logical; if TRUE, off diagonal elements are assumed to be correlations and converted to covariances; if correlation is TRUE, then block is set to TRUE
digits	if value of this argument is greater than zero, the matrix is passed to signif (along with digits) prior to returning
context	the working context
...	passed along

Examples

```
modMATRIX("1 2.2 333")
modMATRIX("1 1.1 2.2", block=TRUE)
modMATRIX("23 234 234 5234", use=FALSE)

ans <- modMATRIX("1.1 0.657 2.2", correlation=TRUE, block=TRUE)
ans
cov2cor(ans)
```

mread_cache

*Read a model specification file.***Description**

mread reads and parses a mrgsolve model specification file, builds the model, and returns a model object for simulation.

Usage

```
mread_cache(model, project = getwd(), code = NULL, soloc = tempdir(),
  quiet = FALSE, preclean = FALSE, ...)

mread(model = character(0), project = getwd(), code = NULL, udll = TRUE,
  ignore.stdout = TRUE, raw = FALSE, compile = TRUE, audit = TRUE,
  quiet = getOption("mrgsolve_mread_quiet", FALSE), check.bounds = FALSE,
  warn = TRUE, soloc = tempdir(), preclean = FALSE, ...)
```

Arguments

<code>model</code>	model name
<code>project</code>	location of the model specification file and any headers to be included
<code>code</code>	a character string with model specification code to be used instead of a model file
<code>soloc</code>	directory where model shared object is stored
<code>quiet</code>	don't print messages when compiling
<code>preclean</code>	logical; if TRUE, compilation artifacts are cleaned up first
<code>...</code>	passed along
<code>udll</code>	use unique name for shared object
<code>ignore.stdout</code>	passed to system call for compiling model
<code>raw</code>	if TRUE, return a list of raw output
<code>compile</code>	logical; if TRUE, the model will be built
<code>audit</code>	check the model specification file for errors
<code>check.bounds</code>	check boundaries of parameter list
<code>warn</code>	logical; if TRUE, print warning messages that may arise

Model Library

mrgsolve comes bundled with several precoded PK, PK/PD, and other systems models that are accessible via the mread interface.

Models available in the library include:

- PK models: `pk1cmt`, `pk2cmt`, `pk3cmt`, `tmdd`
- PKPD models: `irm1`, `irm2`, `irm3`, `irm4`, `emax`, `effect`
- Other models: `viral1`, `viral2`

When the library model is accessed, mrgsolve will compile and load the model as you would for any other model. It is only necessary to reference the correct model name and point the project argument to the mrgsolve model library location via [modlib](#).

For more details, see [modlib_pk](#), [modlib_pkpd](#), [modlib_tmdd](#), [modlib_viral](#), and [modlib_details](#) for more information about the state variables and parameters in each model.

Examples

```
## Not run:
code <- '
$PARAM CL = 1, VC = 5
$CMT CENT
$ODE dxdt_CENT = -(CL/VC)*CENT;
'

mod <- mcode("ex_mread",code)

mod

mod %>% init(CENT=1000) %>% mrgsim %>% plot

mod <- mread("irm3", modlib())

mod

## End(Not run)
```

mrgmod-class

S4 class for mrgsolve model object.

Description

S4 class for mrgsolve model object.

Slots

model model name <character>
project working directory; must be writeable with no spaces <character>
start simulation start time <numeric>
end simulation end time <numeric>
delta simulation time interval <numeric>
add additional simulation times <numeric-vector>
param parameter_list
fixed a parameter_list of fixed value parameters; these are not updatable from R
init cmt_list
events [events](#) object
digits significant digits in simulated output; negative integer means ignore <numeric>
hmin passed to dlsoda <numeric>
hmax passed to dlsoda <numeric>

mxhnil passed to dlsoda <numeric>
 ixpr passed to dlsoda <numeric>
 atol passed to dlsoda <numeric>
 rtol passed to dlsoda <numeric>
 maxsteps passed to dlsoda <numeric>
 preclean passed to R CMD SHLIB during compilation <logical>
 verbose print run information to screen <logical>
 tscale used to scale time in simulated output <numeric>
 omega [matlist](#) for simulating individual-level random effects
 sigma [matlist](#) for simulating residual error variates
 args <list> of arguments to be passed to [mrgsim](#)
 advan either 2, 4, or 13 <numeric>
 trans either 1, 2, 4, or 11
 request vector of compartments to request <character>
 soloc directory path for storing the model shared object <character>
 code a character vector of the model code
 mindt minimum time between simulation records <numeric>
 envir internal model environment <environment>
 annot model annotations <list>
 plugin model plugins <character>

Notes

- Spaces in paths (project and soloc) are prohibited.

\$,mrgmod-method

Create and work with parameter objects.

Description

See [numericlist](#) for methods to deal with `parameter_list` objects.

Usage

```

## S4 method for signature 'mrgmod'
x$name

param(.x, ...)

## S4 method for signature 'mrgmod'
param(.x, .y = list(), ..., .pat = "*",
```

```
.strict = FALSE)

## S4 method for signature 'mrgsims'
param(.x, ...)

## S4 method for signature 'missing'
param(..., .strict = TRUE)

## S4 method for signature 'list'
param(.x, ...)

## S4 method for signature 'ANY'
param(.x, ...)

as.param(.x, ...)

## S4 method for signature 'list'
as.param(.x, ...)

## S4 method for signature 'numeric'
as.param(.x, ...)

## S4 method for signature 'parameter_list'
as.param(.x, ...)

## S4 method for signature 'missing'
as.param(.x, ...)

## S4 method for signature 'parameter_list'
show(object)

allparam(.x)
```

Arguments

<code>x</code>	mrgmod object
<code>name</code>	parameter to take
<code>.x</code>	the model object
<code>...</code>	passed along or name/value pairs to update the parameters in a model object
<code>.y</code>	list to be merged into parameter list
<code>.pat</code>	a regular expression (character) to be applied as a filter for which parameters to show when printing
<code>.strict</code>	if TRUE, all names to be updated must be found in the parameter list
<code>object</code>	passed to show

Details

Can be used to either get a parameter list object from a `mrgmod` model object or to update the parameters in a model object. For both uses, the return value is a `parameter_list` object. For the former use, `param` is usually called to print the parameters to the screen, but the `parameter_list` object can also be coerced to a list or numeric R object.

Value

An object of class `parameter_list` (see [numericlist](#)).

Examples

```
## example("param")
mod <- mrgsolve:::house()

param(mod)
param(mod, .pat="^(C|F)") ## may be useful when large number of parameters

class(param(mod))

param(mod)$KA

as.list(param(mod))
as.data.frame(param(mod))
```

mrgsim

Simulate from a model object.

Description

This function sets up the simulation run from data stored in the model object as well as arguments passed in. Note that there are many non-formal arguments to this function that can be used to customize the simulation run and its output.

Usage

```
mrgsim(x, data = NULL, idata = NULL, nid = 1, ...)
```

Arguments

<code>x</code>	the model objects
<code>data</code>	NMTRAN-like data set
<code>idata</code>	a matrix or data frame of model parameters, one parameter per row
<code>nid</code>	integer number of individuals to simulate; only used if <code>idata</code> and <code>data</code> are missing
<code>...</code>	passed to update

Details

- Both data and idata will be coerced to numeric matrix
- carry.out can be used to insert data columns into the output data set. This is partially dependent on the nature of the data brought into the problem.
- When using data and idata together, an error is generated if an ID occurs in data but not idata. Also, when looking up data in idata, ID in idata is assumed to be uniquely keyed to ID in data. No error is generated if ID is duplicated in data; parameters will be used from the first occurrence found in idata.
- carry.out: idata is assumed to be individual-level and variables that are carried from idata are repeated throughout the individual's simulated data. Variables carried from data are carried via last-observation carry forward. NA is returned from observations that are inserted into simulated output that occur prior to the first record in data.

Value

an object of class `mrgsims`

Additional arguments

- mtime numeric vector of times where the model is evaluated (with solver reset), but results are not included in simulated output
- Request a vector of compartment or table names to take in simulated output; if this is specified, request is ignored
- obsonly omit records with evid != 0 from simulated output
- obsaug logical; when TRUE and a full data set is used, the simulated output is augmented with an observation at each time in `stime()`. When using obsaug, a flag indicating augmented observations can be requested by including `a.u.g` in `carry.out`
- recsort Default value is 1. Possible values are 1,2,3,4: 1 and 2 put doses in a data set after padded observations at the same time; 3 and 4 put those doses before padded observations at the same time. 2 and 4 will put doses scheduled through `addl` after observations at the same time; 1 and 3 put doses scheduled through `addl` before observations at the same time. `recsort` will not change the order of your input data set if both doses and observations are given.
- filbak For each ID, carry the first record data backward to start of the simulation
- tad logical; when TRUE a column is added to simulated output is added showing the time since the last dose. Only data records with `evid == 1` will be considered doses for the purposes of `tad` calculation.

Examples

```
## example("mrgsim")

mod <- mrgsolve:::house() %>% ev(amt=1000, cmt=1)
out <- mod %>% mrgsim()
plot(out)
```

```

out <- mod %>% mrgsim(end=22)
out

data(exTheoph)

out <- mod %>% data_set(exTheoph) %>% mrgsim()
out

out <- mod %>% mrgsim(data=exTheoph)

out <- mrgsim(mod, data=exTheoph, obsonly=TRUE)
out

out <- mod %>% mrgsim(data=exTheoph, obsaug=TRUE, carry.out="a.u.g")
out

out <- mod %>% mrgsim(req="CENT")
out

out <- mrgsim(mod, Req="CP,RESP")
out

```

mrgsims

Methods for working with mrgsims objects.

Description

These methods help the user view simulation output and extract simulated data to work with further. The methods listed here for the most part have generics defined by R or other R packages. See the `seealso` section for other methods defined by `mrgsolve` that have their own documentation pages.

Usage

```

## S4 method for signature 'mrgsims'
x$name

## S4 method for signature 'mrgsims'
tail(x, ...)

## S4 method for signature 'mrgsims'
head(x, ...)

## S4 method for signature 'mrgsims'
dim(x)

```

```
## S4 method for signature 'mrgsims'
names(x)

## S4 method for signature 'mrgsims'
as.data.frame(x, row.names = NULL, optional = FALSE,
  ...)

## S4 method for signature 'mrgsims'
as.matrix(x, ...)

## S4 method for signature 'mrgsims'
subset(x, ...)

## S4 method for signature 'mrgsims'
summary(object, ...)

## S4 method for signature 'mrgsims'
show(object)
```

Arguments

x	mrgsims object
name	name of column of simulated output to retain
...	passed to other functions
row.names	passed to as.data.frame
optional	passed to as.data.frame
object	passed to show

Details

Most methods should behave as expected according to other method commonly used in R (e.g. head, tail, as.data.frame, etc ...)

- subset coresces simulated output to data.frame and passes to subset.data.frame
- \$ selects a column in the simulated data and returns numeric
- head see [head.matrix](#); returns simulated data
- tail see [tail.matrix](#); returns simulated data
- dim, nrow, ncol returns dimensions, number of rows, and number of columns in simulated data
- as.data.frame coresces simulated data to data.frame and returns the data.frame
- as.matrix returns matrix of simulated data
- as.tbl coresces simulated to tbl_df; requires dplyr
- summary coresces simulated data to data.frame and passes to [summary.data.frame](#)
- plot plots simulated data; see [plot_mrgsims](#)

See Also[stime](#)**Examples**

```
## example("mrgsims")

mod <- mrgsolve:::house() %>% init(GUT=100)

out <- mrgsim(mod)
class(out)

out
head(out)
tail(out)

mrgsolve:::mod(out)

dim(out)
names(out)

mat <- as.matrix(out)
df <- as.data.frame(out)

df <- subset(out, time < 12) ## a data frame
out$CP

plot(out)
plot(out, CP~.)
plot(out, CP+RESP~time, scales="same", xlab="Time", main="Model sims")
```

mrgsims-class

*S4 class for mrgsolve simulation output.***Description**

S4 class for mrgsolve simulation output.

Slots

request character vector of compartments requested in simulated output
 outnames character vector of column names in simulated output coming from table step
 data matrix of simulated data
 mod the mrgmod model object

Description

Methods for handling output with dplyr verbs.

Usage

```
## S3 method for class 'mrgsims'  
as.tbl(x, ...)  
  
## S3 method for class 'mrgsims'  
filter_(.data, ..., .dots)  
  
## S3 method for class 'mrgsims'  
group_by_(.data, ..., .dots, add = FALSE)  
  
## S3 method for class 'mrgsims'  
distinct_(.data, ..., .dots, .keep_all = FALSE)  
  
## S3 method for class 'mrgsims'  
mutate_(.data, ..., .dots)  
  
summarise.each(.data, funs, ...)  
  
## S3 method for class 'mrgsims'  
summarise_(.data, ..., .dots)  
  
## S3 method for class 'mrgsims'  
do_(.data, ..., .dots)  
  
## S3 method for class 'mrgsims'  
select_(.data, ..., .dots)  
  
## S3 method for class 'mrgsims'  
slice_(.data, ...)  
  
## S3 method for class 'mrgsims'  
as_data_frame(.data_, ...)
```

Arguments

x	mrgsims object
...	passed to other methods
.data	passed to various dplyr functions

.dots	passed to various dplyr functions
add	passed to dplyr::group_by_
.keep_all	passed to dplyr::distinct_
funcs	passed to dplyr::summarise_each
.data_	mrgsims object

mrgsolve

mrgsolve

Description

mrgsolve is an R package maintained under the auspices of Metrum Research Group, LLC, that facilitates simulation from models based on systems of ordinary differential equations (ODE) that are typically employed for understanding pharmacokinetics, pharmacodynamics, and systems biology and pharmacology. mrgsolve consists of computer code written in the R and C++ languages, providing an interface to the DLSODA differential equation solver (written in FORTRAN) provided through ODEPACK - A Systematized Collection of ODE Solvers.

Example models

See [mrgsolve_example](#) to export example models into your own, writeable project directory.

Input data sets

See [data_set](#) for help creating input data sets. See [exdatasets](#) for example input data sets.

Package help

- Package [index](#), including a listing of all functions
- Reserved words in mrgsolve: [reserved](#)

About the model object

The model object has class [mrgmod](#).

Handling simulated output

See [mrgsims](#) for methods to use with simulated output.

About the solver used by mrgsolve

See: [aboutsolver](#)

Examples

```

## example("mrgsolve")

mod <- mrgsolve:::house(delta=0.1) %>% param(CL=0.5)

events <- ev(amt=1000, cmt=1, addl=5, ii=24)

events

mod

see(mod)

stime(mod)

param(mod)
init(mod)

out <- mod %>% ev(events) %>% mrgsim(end=168)

out

head(out)
tail(out)
dim(out)

plot(out, GUT+CP~.)

sims <- as.data.frame(out)

t72 <- subset(sims, time==72)
str(t72)

idata <- data.frame(ID=c(1,2,3), CL=c(0.5,1,2),VC=12)
out <- mod %>% ev(events) %>% mrgsim(end=168, idata=idata, req="")
plot(out)

out <- mod %>% ev(events) %>% mrgsim(carry.out="amt,evid,cmt,CL")
head(out)

out <-
  mod %>%
  ev() %>%
  knobs(CL=c(0.5, 1,2), amt=c(100,300,1000), cmt=1,end=48)

plot(out, CP~., scales="same")
plot(out, RESP+CP~time|amt,groups=CL)

ev1 <- ev(amt=500, cmt=2,rate=10)
ev2 <- ev(amt=100, cmt=1, time=54, ii=8, addl=10)

```

```

events <- ev1+ev2
events

out <- mod %>% ev(ev1+ev2) %>% mrgsim(end=180, req="")
plot(out)

## "Condensed" data set
data(extran1)
extran1

out <- mod %>% data_set(extran1) %>% mrgsim(end=200)

plot(out, CP~time|factor(ID))

## idata
data(exidata)
exidata

out <-
  mod %>%
    ev(amt=1000, cmt=1) %>%
    idata_set(exidata) %>%
    mrgsim(end=72)

plot(out, CP~., as="log10")

# Internal model library

mod <- mread("irm1", modlib())

mod

mod %>% ev(amt=300, ii=12, addl=3) %>% mrgsim

```

mrgsolve_example

Extract example model from system library

Description

Extract example model from system library

Usage

```
mrgsolve_example(model = c("pkExample", "pkpdExample", "firstmodeExample",
```

```
"viralExample", "popExample"), project = getwd(), overwrite = FALSE,
quiet = FALSE, ...)
```

Arguments

model	name of model
project	working directory
overwrite	passed to file.copy
quiet	don't print any status messages to the screen
...	additional arguments

mrgsolve_template	<i>Create model specification file from template</i>
-------------------	--

Description

Create model specification file from template

Usage

```
mrgsolve_template(model = "template", project = getwd(),
  writeable = FALSE, overwrite = FALSE)
```

Arguments

model	name of the model to create
project	working directory
writeable	logical; if TRUE, parameters may be overwritten in the main block
overwrite	logical; if TRUE, an existing file with same stem will be overwritten

mvgauss	<i>Simulate from a multivariate normal distribution with mean zero.</i>
---------	---

Description

Simulate from a multivariate normal distribution with mean zero.

Usage

```
mvgauss(mat, n = 10, seed = NULL)
```

Arguments

mat	a positive-definite matrix
n	number of variates to simulate
seed	if not null, passed to set.seed

nmxml

*Get THETA, OMEGA and SIGMA from a completed NONMEM run***Description**

Get THETA, OMEGA and SIGMA from a completed NONMEM run

Usage

```
nmxml(run = numeric(0), project = character(0), file = character(0),
      theta = TRUE, omega = FALSE, sigma = FALSE, olabels = NULL,
      slabels = NULL, oprefix = "", sprefix = "", tname = "THETA",
      oname = "...", sname = "...", ...)
```

Arguments

run	run number
project	project directory
file	the complete path to the run.xml file
theta	logical; if TRUE, the \$THETA vector is returned
omega	logical; if TRUE, the \$OMEGA matrix is returned
sigma	logical; if TRUE, the \$SIGMA matrix is returned
olabels	labels for \$OMEGA
slabels	labels for \$SIGMA
oprefix	prefix for \$OMEGA labels
sprefix	prefix for \$SIGMA labels
tname	name for \$THETA
oname	name for \$OMEGA
sname	name for \$SIGMA
...	passed along

Details

If run and project are supplied, the .xml file is assumed to be located in run.xml, in directory run off the project directory. If file is supplied, run and project arguments are ignored.

Value

a list with theta, omega and sigma elements, depending on what was requested

numeric2diag	<i>Create a diagonal numeric matrix from diagonal elements.</i>
--------------	---

Description

Create a diagonal numeric matrix from diagonal elements.

Usage

```
numeric2diag(x, context = NULL)
```

Arguments

x	numeric data
context	used to generate column names

Value

a numeric diagonal matrix

numericlist	<i>Methods for numericlist.</i>
-------------	---------------------------------

Description

These methods can be used to coerce param and init objects into common R data structures, extract elements from numericlists, or get attributes from numericlists.

Usage

```
## S4 method for signature 'numericlist'
as.list(x, ...)

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

## S4 method for signature 'numericlist'
as.data.frame(x, row.names = NULL, optional = FALSE,
  ...)

## S4 method for signature 'numericlist'
length(x)

## S4 method for signature 'numericlist'
names(x)
```

```
## S4 method for signature 'numericlist'
x$name

## S4 method for signature 'numericlist'
x[i, j, ..., drop = TRUE]
```

Arguments

x	object
...	passed along to other methods
row.names	passed to as.data.frame
optional	passed to as.data.frame
name	column to take
i	elements to keep
j	not used
drop	not used

numericlist-class	<i>S4 class numeric list.</i>
-------------------	-------------------------------

Description

S4 class numeric list.

Arguments

data	list of data
pattern	character of length 1 containing regular expression to be used as a filter when printing data to the console

obsaug	<i>Augment observations in the simulated output.</i>
--------	--

Description

Augment observations in the simulated output.

Usage

```
obsaug(x, value = TRUE, ...)
```


Arguments

x	model object
value	the value for obsaug
...	passed along There is also a obsaug argument to mrgsim that can be set to accomplish the same thing as a call to obsaug in the pipeline.

obsonly	<i>Collect only observations in the simulated output.</i>
---------	---

Description

Collect only observations in the simulated output.

Usage

```
obsonly(x, value = TRUE, ...)
```

Arguments

x	model object
value	the value for obsonly
...	passed along

Details

There is also a obsonly argument to [mrgsim](#) that can be set to accomplish the same thing as a call to obsonly in the pipeline.

omega	<i>Manipulate OMEGA matrices.</i>
-------	-----------------------------------

Description

The primary function is `omat` that can be used to both get the \$OMEGA matrices out of a model object and to update \$OMEGA matrices in a model object.

Usage

```

omat(.x, ...)

## S4 method for signature 'missing'
omat(.x, ...)

## S4 method for signature 'matrix'
omat(.x, ..., labels = list())

## S4 method for signature ``NULL``
omat(.x, ...)

## S4 method for signature 'list'
omat(.x, ...)

## S4 method for signature 'omegalist'
omat(.x, ...)

## S4 method for signature 'mrgmod'
omat(.x, ..., make = FALSE, open = FALSE)

## S4 method for signature 'mrgsims'
omat(.x, make = FALSE, ...)

```

Arguments

<code>.x</code>	a matrix, list of matrices or matlist object
<code>...</code>	passed to other functions, including modMATRIX
<code>labels</code>	character vector of names for \$OMEGA elements; must be equal to number of rows/columns in the matrix
<code>make</code>	logical; if TRUE, matrix list is rendered into a single matrix
<code>open</code>	passed to merge.list
<code>x</code>	matlist object

Examples

```

## example("omega")
mat1 <- matrix(1)
mat2 <- diag(c(1,2,3))
mat3 <- matrix(c(0.1, 0.002, 0.002, 0.5), 2,2)
mat4 <- dmat(0.1, 0.2, 0.3, 0.4)

omat(mat1)
omat(mat1, mat2, mat3)
omat(A=mat1, B=mat2, C=mat3)

mod <- mrgsolve:::house() %>% omat(mat4)

```

```
omat(mod)
omat(mod, make=TRUE)

## Not run:

$OMEGA
1 2 3

$OMEGA @block
1 0.1 2

$OMEGA \@cor
\@ prefix ETA_
\@ labels CL VC KA
0.1
0.67 0.2
0 0 0.3

## End(Not run)
```

parameter_list-class	<i>S4 parameter_list class</i>
----------------------	--------------------------------

Description

S4 parameter_list class

Details

parameter_list is a [numericlist-class](#)

pkmodel	<i>Simulate from 1- or 2-compartment PK model.</i>
---------	--

Description

This is an R function that returns model objects based on \$PKMODEL.

Usage

```
pkmodel(ncmt = 1, depot = FALSE, ...)
```

Arguments

ncmt	passed to PKMODEL
depot	passed to PKMODEL
...	passed to update

Details

Once the model object is generated, use `param` to check names of the parameters in the model and `init` to check the names of the compartments in the model. Calculations for the amounts in each compartment are done via analytical solutions, not differential equations. A subject-level random effect is also provided for each PK parameter; use `omat` to see the names of those random effects. All random effect variances have initial value of zero and may be updated via `omat`.

Value

An object of class `mrmod-class`

Examples

```
## Not run:
mod <- pkmodel(1)

mod %>% ev(amt=1000, ii=24, addl=3) %>% mrgsim(end=120)
mod <- pkmodel(1,TRUE)
mod <- pkmodel(2)
mod <- pkmodel(2,TRUE)

## End(Not run)
```

PKMODEL

Parse PKMODEL BLOCK data.

Description

Parse PKMODEL BLOCK data.

Usage

```
PKMODEL(ncmt = 1, depot = FALSE, cmt = NULL, trans = pick_trans(ncmt,
  depot), env = list(), pos = 1, ...)
```

Arguments

<code>ncmt</code>	number of compartments; must be 1 (one-compartment, not including a depot dosing compartment) or 2 (two-compartment model, not including a depot dosing compartment)
<code>depot</code>	logical indicating whether to add depot compartment
<code>cmt</code>	compartment names as comma-delimited character

trans	the parameterization for the PK model; must be 1, 2, 4, or 11
env	parse environment
pos	block position number
...	not used

Details

When using \$PKMODEL, certain symbols must be defined in the model specification depending on the value of ncmt, depot and trans.

- ncmt 1, depot FALSE, trans 2: CL, V
- ncmt 1, depot TRUE, trans 2: CL, V, KA
- ncmt 2, depot FALSE, trans 4: CL, V1, Q, V2
- ncmt 2, depot TRUE, trans 4: CL, V2, Q, V3, KA

If trans=11 is specified, use the symbols listed above for the ncmt / depot combination, but append i at the end (e.g. CLi or Qi or KAi).

If trans=1, the user must utilize the following symbols:

- pred_CL for clearance
- pred_V or pred_V2 for central compartment volume of distribution
- pred_Q for intercompartmental clearance
- pred_V3 for peripheral compartment volume of distribution
- pred_KA for absorption rate constant

See Also

[BLOCK_PARSE](#)

plot, batch_mrgsims, missing-method

Plot method for mrgsims objects.

Description

Plot method for mrgsims objects.

Usage

```
## S4 method for signature 'batch_mrgsims,missing'
plot(x, yval = variables(x),
     auto.key = list(), mincol = 3, ...)

## S4 method for signature 'batch_mrgsims,formula'
plot(x, y, show.grid = TRUE, lwd = 2,
     type = "l", yval = variables(x), auto.key = list(columns = 1),
     scales = list(y = list(relation = "free")), ...)
```

Arguments

x	mrsims object
yval	y variables to plot
auto.key	passed to xyplot
mincol	minimum number of columns in key
...	arguments passed to xyplot
y	a formula passed to xyplot
show.grid	print grid in the plot
lwd	passed to xyplot
type	passed to xyplot
scales	passed to xyplot

plot_mrgsims	<i>Generate a quick plot of simulated data.</i>
--------------	---

Description

Generate a quick plot of simulated data.

Usage

```
## S4 method for signature 'mrgsims,missing'
plot(x, limit = 16, ...)

## S4 method for signature 'mrgsims,formula'
plot(x, y, limit = 16, show.grid = TRUE,
     outer = TRUE, type = "l", lwd = 2, ylab = "value", groups = ID,
     scales = list(y = list(relation = "free")), ...)
```

Arguments

x	mrgsims object
limit	limit the the number of panels to create
...	other arguments passed to xyplot
y	formula used for plotting
show.grid	logical indicating whether or not to draw panel.grid
outer	passed to xyplot
type	passed to xyplot
lwd	passed to xyplot
ylab	passed to xyplot
groups	passed to xyplot
scales	passed to xyplot

Details

Values for as argument: ; raw: raw simulated output;

Examples

```
mod <- mrgsolve:::house(end=48, delta=0.2) %>% init(GUT=1000)

out <- mrgsim(mod)

plot(out)

plot(out, subset=time <=24)

plot(out, GUT+CP~.)

plot(out, CP+RESP~time, col="black", scales="same", lty=2)
```

 qsim

A quick simulation function.

Description

A quick simulation function.

Usage

```
qsim(x, e, idata, req = NULL, tgrid = NULL)
```

Arguments

x	model object
e	event object
idata	individual data set
req	compartments to request
tgrid	tgrid object; used if e is an ev object

Examples

```
mod <- mrgsolve:::house()

des <- tgrid(0,2400,1)

data <- recmatrix(ev(amt=1000, ii=24, addl=100),des)

out <- mod %>% qsim(data)
```

realize_addl	<i>Make addl doses explicit in an event object or data set.</i>
--------------	---

Description

Make addl doses explicit in an event object or data set.

Usage

```
realize_addl(x, ...)

## S3 method for class 'data.frame'
realize_addl(x, ...)

## S3 method for class 'ev'
realize_addl(x, ...)
```

Arguments

x	a data_set data frame or an ev object (see details)
...	not used

Details

Required data elements: addl and ii.

recmatrix	<i>Create a matrix of events for simulation.</i>
-----------	--

Description

This function is for use with [qsim](#) only.

Usage

```
recmatrix(x, times, c_indexing = TRUE)
```

Arguments

x	an events object
times	object that can be coerced to numeric with stime
c_indexing	if TRUE, compartment numbers will be decremented by 1

relocate	<i>Update model or project in an model object.</i>
----------	--

Description

Update model or project in an model object.

Usage

```
relocate(x, ...)

## S4 method for signature 'mrgmod'
relocate(x, model = NULL, project = NULL)
```

Arguments

x	mrgmod object
...	passed along
model	model name
project	project directory

Value

updated model object

rename_cols	<i>rename columns from vector for new names</i>
-------------	---

Description

rename columns from vector for new names

Usage

```
rename_cols(.df, new_names)
```

Arguments

.df	dataframe to rename
new_names	vector of names using syntax "<newname>" = "<oldname>"

Examples

```
rename_cols(Theoph, c("dv" = "conc", "ID" = "Subject"))
```

render	<i>Render a model to a document.</i>
--------	--------------------------------------

Description

Render a model to a document.

Usage

```
render(x, ...)

## S4 method for signature 'character'
render(x, project, ...)

## S4 method for signature 'mrgmod'
render(x, ...)

dorender(model, project, template = NULL, compile = TRUE, ...)
```

Arguments

x	model object or the model name
...	passed to <code>rmarkdown::render</code>
project	the directory containing the <code>.cpp</code> model file
model	model name
template	template document
compile	logical; if true, the model will be compiled to run

Examples

```
## Not run:
mod <- mrgsolve:::house()
mrgsolve:::render(mod)
mrgsolve:::render("irm2", modlib())

## End(Not run)
```

Req	<i>Request simulated output.</i>
-----	----------------------------------

Description

Use this function to select, by name, either compartments or derived variables that have been captured (see [CAPTURE](#)).

Usage

```
Req(x, ...)

## S4 method for signature 'mrgmod'
Req(x, ...)

req(x, ...)

## S4 method for signature 'mrgmod'
req(x, ...)
```

Arguments

x	model object
...	unquoted names of compartments or tabled items

There is also a Req argument to [mrgsim](#) that can be set to accomplish the same thing as a call to Req in the pipeline.

Note the difference between req and Req: the former only selects compartments to appear in output while the latter selects both compartments and captured items. Also, when there are items explicitly listed in Req, all other compartments or captured items not listed there are ignored. But when compartments are selected with req all of the captured items are returned. Remember that req is strictly for compartments.

Examples

```
mod <- mrgsolve:::house()

mod %>% Req(CP,RESP) %>% ev(amt=1000) %>% mrgsim
```

reserved	<i>Reserved words.</i>
----------	------------------------

Description

Reserved words.

Usage

```
reserved()
```

Details

Note: this function is not exported; you must go into the `mrgsolve` namespace by using the `mrgsolve::` prefix.

Examples

```
mrgsolve::reserved()
```

revar	<i>Get model random effect variances and covariances.</i>
-------	---

Description

Get model random effect variances and covariances.

Usage

```
revar(x, ...)
```

```
## S4 method for signature 'mrgmod'
```

```
revar(x, ...)
```

Arguments

<code>x</code>	model object
<code>...</code>	passed along

%>%	<i>Forward pipe.</i>
-----	----------------------

Description

Forward pipe.
Tee.

scrape_and_call	<i>Scrape options and pass to function.</i>
-----------------	---

Description

Scrape options and pass to function.

Usage

```
scrape_and_call(x, env, pass, ...)
```

Arguments

x	data
env	parse environment
pass	function to call
...	dots

Details

Attributes of x are also scraped and merged with options.

scrape_opts	<i>Scrape options from a code block.</i>
-------------	--

Description

Scrape options from a code block.

Usage

```
scrape_opts(x, envir = list(), def = list(), all = TRUE, marker = "=",  
            narrow = TRUE)
```

Arguments

<code>x</code>	data
<code>envir</code>	environment from <code>\$ENV</code>
<code>def</code>	default values
<code>all</code>	return all options, even those that are not in <code>def</code>
<code>marker</code>	assignment operator; used to locate lines with options
<code>narrow</code>	logical; if <code>TRUE</code> , only get options on lines starting with <code>>></code>

Value

list with elements `x` (the data without options) and named options as specified in the block.

see	<i>Print model code to the console.</i>
-----	---

Description

Print model code to the console.

Usage

```
see(x, ...)
```

S4 method for signature 'mrgmod'

```
see(x, raw = FALSE, ...)
```

Arguments

<code>x</code>	model object
<code>...</code>	passed along
<code>raw</code>	return the raw code

Value

invisible `NULL`

show,modlist-method	<i>Show a modlist object.</i>
---------------------	-------------------------------

Description

Show a modlist object.

Usage

```
## S4 method for signature 'modlist'  
show(object)
```

Arguments

object	modlist object
--------	----------------

show,mrgmod-method	<i>Print model details.</i>
--------------------	-----------------------------

Description

Print model details.

Usage

```
## S4 method for signature 'mrgmod'  
show(object)
```

Arguments

object	the model object
--------	------------------

sigma

*Manipulate SIGMA matrices.***Description**

The primary function is `smat` that can be used to both get the `$SIGMA` matrices out of a model object and to update `$SIGMA` matrices in a model object.

Usage

```
smat(.x, ...)
```

S4 method for signature 'missing'

```
smat(.x, ...)
```

S4 method for signature 'matrix'

```
smat(.x, ..., labels = list())
```

S4 method for signature 'list'

```
smat(.x, ...)
```

S4 method for signature 'sigmalist'

```
smat(.x, ...)
```

S4 method for signature 'mrgmod'

```
smat(.x, ..., make = FALSE, open = FALSE)
```

S4 method for signature '`NULL`'

```
smat(.x, ...)
```

S4 method for signature 'mrgsims'

```
smat(.x, make = FALSE, ...)
```

Arguments

<code>.x</code>	a matrix, list of matrices or <code>matlist</code> object
<code>...</code>	passed to other functions, including modMATRIX
<code>labels</code>	character vector of names for <code>\$SIGMA</code> elements; must be equal to number of rows/columns in the matrix
<code>make</code>	logical; if <code>TRUE</code> , matrix list is rendered into a single matrix
<code>open</code>	passed to merge.list
<code>x</code>	<code>matlist</code> object

Examples

```
## example("sigma")
mat1 <- matrix(1)
mat2 <- diag(c(1,2))
mat3 <- matrix(c(0.1, 0.002, 0.002, 0.5), 2,2)
mat4 <- dmat(0.1, 0.2, 0.3, 0.4)

smat(mat1)
smat(mat1, mat2, mat3)
smat(A=mat1, B=mat2, C=mat3)

mod <- mrgsolve:::house() %>% smat(mat1)

smat(mod)
smat(mod, make=TRUE)
```

simargs

Access or clear arguments for calls to mrgsim.

Description

Access or clear arguments for calls to mrgsim.

Usage

```
simargs(x, ...)
```

```
## S3 method for class 'mrgmod'
simargs(x, clear = FALSE, ...)
```

Arguments

x	model object
...	passed along
clear	logical indicating whether or not clear args from the model object

Value

If clear is TRUE, the argument list is cleared and the model object is returned. Otherwise, the argument list is returned.

Examples

```
mod <- mrgsolve:::house()
mod %>% Req(CP,RESP) %>% carry_out(evid,WT,FLAG) %>% simargs
```

soloc	<i>Return the location of the model shared object.</i>
-------	--

Description

Return the location of the model shared object.

Usage

```
soloc(x, short = FALSE)
```

Arguments

x	model object
short	logical; if TRUE, soloc will be rendered with a short path name

Examples

```
mod <- mrgsolve:::house()
soloc(mod)
```

stime	<i>Get the times at which the model will be evaluated.</i>
-------	--

Description

Get the times at which the model will be evaluated.

Usage

```
stime(x, ...)
```

Arguments

x	object of class mrgmod
...	passed on

Details

Simulation times include the sequence of times created from start, end, and delta and the vector of times found in add. Making end negative will omit any start / end / delta sequence. Negative values are discarded from the result.

Value

a sorted vector of unique times

Examples

```
## example("stime", package="mrgsolve")

mod <- mrgsolve:::house(end=12, delta=2, add=c(11,13,15))

stime(mod)
```

stime, mrgmod-method *Create a simtime object.*

Description

simtime objects allow the user to specify simulation start and end times, along with the simulation time step.

Usage

```
## S4 method for signature 'mrgmod'
stime(x, ...)

tgrid(start = 0, end = 24, delta = 1, add = numeric(0), .offset = 0,
       .scale = 1, ...)

## S4 method for signature 'tgrid'
stime(x, ...)

## S4 method for signature 'tgrids'
stime(x, ...)

## S4 method for signature 'numeric'
stime(x, ...)

## S4 method for signature 'tgrid'
show(object)

## S4 method for signature 'tgrids'
show(object)
```

Arguments

x	tgrid object
...	passed on to other methods
start	simulation start time

end	simulation end time
delta	simulation time step
add	addition simulation times
.offset	the resulting set of times will be adjusted by this amount
.scale	the resulting set of times will be scaled by this factor
object	passed to show

Examples

```
peak <- tgrid(0,6,0.2)
sparse <- tgrid(0,24,4)

day1 <- c(peak,sparse)

design <- c(day1, day1+72, day1+240)

## Not run:
mod <- mrgsolve:::house()

out <- mod %>% ev(amt=1000, ii=24, addl=10) %>% mrgsim(tgrid=design)

plot(out,CP~., type='b')

## End(Not run)
```

touch_funs	<i>Get inits from compiled function.</i>
------------	--

Description

Get inits from compiled function.

Usage

```
touch_funs(x, keep_pointers = TRUE)
```

Arguments

x	mrgmod model object
keep_pointers	should function pointers be returned?

tscale	<i>Rescale time in the simulated output.</i>
--------	--

Description

Rescale time in the simulated output.

Usage

```
tscale(x, value = 1, ...)
```

Arguments

x	model object
value	value by which time will be scaled
...	passed along

Details

There is also a `tscale` argument to `mrgsim` that can be set to accomplish the same thing as a call to `tscale` in the pipeline.

Examples

```
# The model is in hours:
mod <- mrgsolve:::house()

# The output is in days:
mod %>% tscale(1/24) %>% mrgsim
```

update	<i>Get all names from a model object.</i>
--------	---

Description

Get all names from a model object.

After the model object is created, update various attributes.

Usage

```
## S4 method for signature 'mrgmod'
names(x)

## S4 method for signature 'mrgmod'
update(object, ..., merge = TRUE, open = FALSE,
       data = list())

## S4 method for signature 'omegalist'
update(object, y, ...)

## S4 method for signature 'sigmalist'
update(object, y, ...)

## S4 method for signature 'parameter_list'
update(object, y, ...)

## S4 method for signature 'ev'
update(object, y, ...)
```

Arguments

x	the model object
object	a model object
...	passed to other functions
merge	logical indicating to merge (rather than replace) new and existing attributes.
open	logical; used only when merge is TRUE and parameter list or initial conditions list is being updated; if FALSE, no new items will be added; if TRUE, the parameter list may expand.
data	a list of items to update; not used for now
y	another object involved in update

Value

The updated model object is returned.

Examples

```
mod <- mrgsolve:::house()
names(mod)

## Not run:
mod <- mrgsolve:::house()

mod <- update(mod, end=120, delta=4, param=list(CL=19.1))

## End(Not run)
```

valid_data	<i>Validate and prepare data sets for simulation.</i>
------------	---

Description

Validate and prepare data sets for simulation.

Validate and prepare idata data sets for simulation.

Usage

```
valid_data_set(x, ...)  
  
## Default S3 method:  
valid_data_set(x, ...)  
  
## S3 method for class 'data.frame'  
valid_data_set(x, m = NULL, verbose = FALSE,  
  quiet = FALSE, ...)  
  
valid_idata_set(x, verbose = FALSE, quiet = FALSE, ...)  
  
## S3 method for class 'matrix'  
valid_data_set(x, verbose = FALSE, ...)
```

Arguments

x	data.frame or matrix
...	additional arguments
m	a model object
verbose	logical
quiet	if TRUE, messages will be suppressed

Value

a matrix with non-numeric columns dropped; if x is a data.frame with character cmt column comprised of valid compartment names and m is a model object, the cmt column will be converted to the corresponding compartment number.

A numeric matrix with class `valid_idata_set`.

See Also

[idata_set](#), [data_set](#), [valid_data_set](#)

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