# Package 'puzzle'

October 14, 2022

Title Assembling Data Sets for Non-Linear Mixed Effects Modeling

Type Package

Version 0.0.1
Maintainer Mario Gonzalez Sales <mario@modelinggreatsolutions.com></mario@modelinggreatsolutions.com>
Description  To Simplify the time consuming and error prone task of assembling complex data sets for non-linear mixed effects modeling. Users are able to select from different absorption processes such as zero and first order, or a combination of both. Furthermore, data sets containing data from several entities, responses, and covariates can be simultaneously assembled.
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Encoding UTF-8
LazyData true
<b>Imports</b> utils, lubridate, stats, readxl, reshape, reshape2, sqldf, kableExtra, plyr, dplyr, tidyverse, readr
Suggests rmarkdown, knitr, devtools, testthat
RoxygenNote 6.1.1
<pre>URL https://github.com/syneoshealth/puzzle</pre>
BugReports https://github.com/syneoshealth/puzzle/issues
NeedsCompilation no
Author Olivier Barriere [aut], Mario Gonzalez Sales [aut, cre]
Repository CRAN
<b>Date/Publication</b> 2019-11-28 16:10:02 UTC
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df\_cov

A covariate data set.

## Description

A dataset containing covariate information.

## Usage

df\_cov

#### **Format**

A tibble with 12 rows and 4 variables:

ID Individual

TIME Time, in hours

VARIABLE Variable

**VALUE** Value of the variable

df\_cov\_start 3

df\_cov\_start

Starting covariate data set.

## Description

A dataset containing covariate information.

#### Usage

```
{\sf df\_cov\_start}
```

#### **Format**

A data frame with 4 rows and 3 variables:

ID Individual

VARIABLE Variable

VALUE Value of the variable

df\_cov\_time\_dependent\_start

A covariate data set to be used with time dependent covariates.

## Description

A dataset containing time dependent covariates.

## Usage

```
df_cov_time_dependent_start
```

#### **Format**

A data frame with 6 rows and 4 variables:

**ID** Individual

VARIABLE Variable

VALUE Value of the variable

TIME Time, in hours

4 df\_dose\_datetime

df\_dose

A dose data set.

## **Description**

A dataset containing dose information.

## Usage

df\_dose

#### **Format**

A data frame with 12 rows and 3 variables:

**ID** Individual

TIME Time, in weeks

AMT Dose, in mg

df\_dose\_datetime

A dose data set including datetimes.

#### **Description**

A dataset containing dose information in datetime format.

## Usage

df\_dose\_datetime

#### **Format**

A data frame with 5 rows and 12 variables:

**ID** Individual

TRT Treatment label

**DOSE** Dose, in mg

**PERIOD** Period

**DAY** Day of adminsitration

AMT Dose, in mg

**DATETIME** Dta ein datetime format

**TIMEPOINT** Timepoint

**COHORT** Cohort

FORM Drug form

TREATMENT Treatment

FOOD Food status

df\_dose\_evid4 5

df\_dose\_evid4

A dose data set to be used with EVID=4.

## Description

A dataset containing dosing information.

## Usage

df\_dose\_evid4

#### **Format**

A data frame with 418 rows and 10 variables:

**ID** Individual

PERIOD Period

TIMEPOINT Timepoint

TIME Time, in hours

AMT Dose, in mg

TRT Treatment label

**DAY** Day of adminsitration

**SEQUENCE** Sequence

TRT2 Treatment

EVID Evid value

df\_dose\_optional\_columns

A dose data set to be used with optional columns.

## Description

A dataset containing dosing information.

#### Usage

df\_dose\_optional\_columns

6 df\_extra\_times

#### **Format**

A data frame with 4 rows and 6 variables:

**ID** Individual

TIME Time, in hours

AMT Dose, in mg

OCC Occasion

TIMEPOINT Timepoint

TRT Treatment

df\_dose\_start

A dose data set example.

## Description

A dataset containing dosing information.

#### Usage

df\_dose\_start

#### **Format**

A data frame with 4 rows and 3 variables:

**ID** Individual

TIME Time, in hours

AMT Dose, in mg

 $df_extra_times$ 

An extra times data set example.

## Description

A dataset containing extra times.

#### Usage

df\_extra\_times

#### **Format**

A data frame with 251 rows and 1 variable:

TIME Time, in hours

```
df_extra_times_datetime
```

An extra times data set example with datetime format.

## Description

A dataset containing extra times in datetime format.

#### Usage

```
df_extra_times_datetime
```

#### **Format**

A data frame with 20 rows and 1 variable:

**ID** Individual

**DATETIME** Datetime

**TIMEPOINT** Timepoint

```
df_extra_times_metabolite_evid4
```

An extra times metabolite data set to be used with EVID=4.

## Description

A dataset containing extra times for an hypothetical metabolite.

## Usage

```
df_extra_times_metabolite_evid4
```

#### **Format**

A data frame with 770 rows and 3 variable:

PERIOD Period

TIMEPOINT Timepoint

TIME Time, in hours

8 df\_extra\_times\_time

```
df_extra_times_parent_evid4
```

An extra times parent data set to be used with EVID=4.

## Description

A dataset containing extra times for an hypothetical parent drug.

## Usage

```
df_extra_times_parent_evid4
```

#### **Format**

A data frame with 770 rows and 3 variable:

PERIOD Period

**TIMEPOINT** Timepoint

TIME Time, in hours

## Description

A dataset containing extra times.

#### Usage

```
df_extra_times_time
```

#### **Format**

A data frame with 1040 rows and 3 variable:

**ID** Individual

TIME Time, in hours

TIMEPOINT Timepoint

df\_metabolite\_evid4 9

df\_metabolite\_evid4

A pharmacokinetic metabolite data set to be used with EVID=4.

#### **Description**

A dataset containing pharmacokinetic information for an hypothetical metabolite.

## Usage

```
df_metabolite_evid4
```

#### **Format**

A data frame with 1359 rows and 7 variables:

**ID** Individual

PERIOD Period

**TIMEPOINT** Timepoint

TIME Time, in hours

**DV** Drug concentration, in mg/L

TIMEDAY Timeday

**DAY** Day of adminsitration

df\_parent\_evid4

A pharmacokinetic parent data set to be used with EVID=4.

#### **Description**

A dataset containing pharmacokinetic information for an hypothetical parent drug.

## Usage

```
df_parent_evid4
```

#### **Format**

A data frame with 1359 rows and 7 variables:

ID Individual

PERIOD Period

TIMEPOINT Timepoint

**TIME** Time, in hours

**DV** Drug concentration, in mg/L

TIMEDAY Timeday

**DAY** Day of adminsitration

10 df\_pk

 $df_pd_start$ 

An starting pharmacoynamic data set example.

## Description

A dataset containing pharmacodynamic observations.

## Usage

```
df_pd_start
```

#### **Format**

A tibble with 6 rows and 3 variable:

**ID** Individual

TIME Time, in hours

DV Response, ng/mL

df\_pk

A pharmacokinetic data set.

## Description

A dataset containing pharmacokinetic information.

## Usage

df\_pk

#### **Format**

A tibble with 132 rows and 4 variable:

**ID** Individual

TIMEPOINT Timepoint

TIME Time, in hours

**DV** Drug concentration, ng/mL

df\_pk\_datetime 11

 $df_pk_datetime$ 

A pharmacokinetic data set example in datetime format.

## Description

A dataset containing pharmacokinetic information.

## Usage

```
df_pk_datetime
```

#### **Format**

A data frame with 65 rows and 7 variable:

**ID** Individual

DV Response, ng/mL

**DATETIME** Datetime

**TIMEPOINT** Timepoint

DAY Day

PERIOD Period

BLQ I a BLQ?

LLOQ Lower limit of quantification, ng/mL

df\_pk\_metabolite

A pharmacokinetic data set of metabolite data.

## Description

A dataset containing pharmacokinetic information for an hypothetical metabolite.

#### Usage

```
df_pk_metabolite
```

## **Format**

A data frame with 10 rows and 3 variable:

**ID** Individual

TIME Time, in hours

DV Drug concentration, ng/mL

12 df\_pk\_parent

```
df_pk_optional_columns
```

A pharmacokinetic data set to be used with optional columns.

#### **Description**

A dataset containing pharmacokinetic information.

## Usage

```
df_pk_optional_columns
```

#### **Format**

A data frame with 12 rows and 5 variable:

ID Individual

TIME Time, in hours

**DV** Drug concentration, ng/mL

OCC Occasion

TIMEPOINT Timepoint

 $df_pk_parent$ 

A pharmacokinetic data set for an hypothetical parent drug.

## Description

A dataset containing pharmacokinetic information.

#### Usage

```
df_pk_parent
```

#### **Format**

A data frame with 12 rows and 3 variable:

ID Individual

TIME Time, in hours

DV Drug concentration, ng/mL

df\_pk\_start 13

df\_pk\_start

A pharmacokinetic data set example.

## Description

A dataset containing pharmacokinetic information.

A dataset containing pharmacokinetic information.

## Usage

```
df_pk_start
```

df\_pk\_start

#### **Format**

A tibble with 12 rows and 3 variable:

**ID** Individual

TIME Time, in hours

DV Response, ng/mL

df\_response1

A pharmacodynamic data set.

## Description

A dataset containing pharmacodynamic information for response 1.

## Usage

```
df_response1
```

#### **Format**

A data frame with 6 rows and 3 variable:

**ID** Individual

TIME Time, in hours

DV Response, ng/mL

14 df\_response3

df\_response2

A pharmacodynamic data set.

## Description

A dataset containing pharmacodynamic information for response 2.

## Usage

```
df_response2
```

#### **Format**

A data frame with 6 rows and 3 variable:

**ID** Individual

TIME Time, in hours

**DV** Response, seconds

df\_response3

A pharmacodynamic data set.

## Description

A dataset containing pharmacodynamic information for response 3.

## Usage

```
df_response3
```

## **Format**

A data frame with 6 rows and 3 variable:

**ID** Individual

TIME Time, in hours

DV Response, in hours

puzzle 15

|--|

## Description

Build pharmacometric data sets from basic tabulated files

#### Usage

```
puzzle(directory = NULL, order, coercion = list(name = NULL, sep =
  ","), optionalcolumns = NULL, pk = list(name = NULL, data = NULL),
  dose = list(name = NULL, data = NULL), cov = list(name = NULL, data =
  NULL), pd = list(name = NULL, data = NULL), extratimes = list(name =
  NULL, data = NULL), nm = list(name = NULL), fillcolumns = NULL,
  nocoercioncolumns = NULL, norepeatcolumns = NULL, initialindex = 0,
  na.strings = "N/A", arrange = "ID,TIME,CMT,desc(EVID)",
  datetimeformat = "%Y-%m-%d %H:%M:%S", timeunits = "hours",
  timezone = Sys.timezone(), ignore = "C", missingvalues = ".",
  parallel = TRUE, verbose = FALSE, username = NULL)
```

#### **Arguments**

initialindex

٤	Guinenes			
	directory	path to your directory		
	order	define the absorption order, can be $0$ , $1$ , $c(0,1)$ , or $c(1,1)$		
	coercion optionalcolumns	define name for coercion file		
		define optional columns		
	pk	define the required file containing the pk information. It can be a .csv or an .xlsx file		
	dose	define the required file containing the dose information. It can be a .csv, an .xlsx file or an R object.		
	cov	define the optional file containing the covariate information. It can be a .csv, an .xlsx file or an R object.		
	pd	define the optional file containing the pd information. It can be a .csv, or a .xlsx file.		
	extratimes	define the optional file containing the additional times. It can be a .csv, or a .xlsx file.		
	nm	name of output file generated by puzzle		
	fillcolumns nocoercioncolum	define columns to be filled ns		
		define columns to be dropped from the coercion file		
norepeatcolumns				
		define columns not to be repeated		

define the lower category of categorical covariates

16 puzzle

na.strings define value for na

arrange define how the columns should be arranged

datetimeformat define format for date times timeunits define time units if needed

timezone define timezone ignore define ignore value missingvalues define missing value

parallel define parallel zero + first order absorption

verbose define verbose

username define person performing the assembling

#### Value

a pharmacometrics ready data set

## **Examples**

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```