Package ‘EHR’

October 7, 2021

Version 0.4-4

Date 2021-09-08

Title Electronic Health Record (EHR) Data Processing and Analysis Tool

Maintainer Leena Choi <naturechoi@gmail.com>

Description Process and analyze electronic health record (EHR) data. The ‘EHR’ package provides modules to perform diverse medication-related studies using data from EHR databases. Especially, the package includes modules to perform pharmacokinetic/pharmacodynamic (PK/PD) analyses using EHRs, as outlined in Choi, Beck, McNeer, Weeks, Williams, Niu, Abou-Khalil, Birdwell, Roden, Stein, Bejan, Denny, and Van Driest (2020) <doi:10.1002/cpt.1787>. Additional modules will be added in future. In addition, this package provides various functions useful to perform Phenome Wide Association Study (PheWAS) to explore associations between drug exposure and phenotypes obtained from EHR data, as outlined in Choi, Carroll, Beck, Mosley, Roden, Denny, and Van Driest (2018) <doi:10.1093/bioinformatics/bty306>.

Depends R (>= 2.10)

License GPL (>= 3)

URL https://choileena.github.io/

Imports stats, utils, data.table, methods, lubridate, pkdata

Suggests glmnet, logistf, medExtractR, knitr, rmarkdown, ggplot2, markdown

NeedsCompilation no

RoxygenNote 7.1.1

VignetteBuilder knitr

Author Leena Choi [aut, cre] (<https://orcid.org/0000-0002-2544-7090>), Cole Beck [aut] (<https://orcid.org/0000-0002-6849-6255>), Hannah Weeks [aut] (<https://orcid.org/0000-0002-0262-6790>), Elizabeth McNeer [aut], Nathan James [aut] (<https://orcid.org/0000-0001-7079-9151>), Michael Williams [aut]

Repository CRAN

Date/Publication 2021-10-07 04:20:02 UTC
### R topics documented:

- EHR-package .................................................. 3
- addLastDose ................................................... 4
- analysisPheWAS ............................................ 5
- buildDose ..................................................... 7
- collapseDose ................................................ 9
- dataTransformation ....................................... 10
- dd .............................................................. 10
- dd.baseline ................................................... 11
- dd.baseline.small ........................................ 11
- dd.small ...................................................... 12
- extractMed .................................................. 12
- freqNum ...................................................... 13
- idCrosswalk ................................................ 14
- lam_metadata ................................................ 15
- lam_mxr_parsed ........................................... 16
- Logistf ....................................................... 17
- makeDose .................................................... 18
- parseCLAMP ................................................ 19
- parseMedEx .................................................. 20
- parseMedExtractR ........................................ 21
- parseMedXN .................................................. 22
- processLastDose ......................................... 23
- pullFakeId .................................................. 24
- pullRealId .................................................. 25
- readTransform ............................................. 26
- run_Build_PK_IV ........................................... 27
- run_Build_PK_Oral .......................................... 29
- run_Demo .................................................... 31
- run_DrugLevel ............................................. 32
- run_Labs ..................................................... 34
- run_MedStrI ................................................ 35
- run_MedStrII ............................................... 38
- stdzDose ..................................................... 40
- stdzDoseChange .......................................... 40
- stdzDoseSchedule ....................................... 40
- stdzDuration .............................................. 41
- stdzFreq .................................................... 42
- stdzRoute .................................................. 42
- stdzStrength .............................................. 43
- tac_lab ....................................................... 43
- tac_metadata ............................................... 44
- tac_mxr_parsed ........................................... 45
- zeroOneTable .............................................. 46

Index 47
Description

The ‘EHR’ package provides modules to perform diverse medication-related studies using data from EHR databases.

Details

Package functionality:

• Process and analyze Electronic Health Record (EHR) data.
• Implement modules to perform diverse medication-related studies using data from EHR databases. Especially, the package includes modules to perform pharmacokinetic/pharmacodynamic (PK/PD) analyses using EHRs, as outlined in Choi et al. (2020).
• Implement three statistical methods for Phenome Wide Association Study (PheWAS). Contingency tables for many binary outcomes (e.g., phenotypes) and a binary covariate (e.g., drug exposure) can be efficiently generated by \texttt{zeroOneTable}, and three commonly used statistical methods to analyze data for PheWAS are implement by \texttt{analysisPheWAS}.

Author(s)

Maintainer: Leena Choi <naturechoi@gmail.com> (ORCID)

Authors:

• Cole Beck <cole.beck@vumc.org> (ORCID)
• Hannah Weeks <hannah.l.weeks@vanderbilt.edu> (ORCID)
• Elizabeth McNeer <elizabeth.mcneer@vumc.org>
• Nathan James <nathan.t.james@vanderbilt.edu> (ORCID)
• Michael Williams <michael.l.williams@vanderbilt.edu>

References

1. Development of a system for postmarketing population pharmacokinetic and pharmacodynamic studies using real-world data from electronic health records.

2. Evaluating statistical approaches to leverage large clinical datasets for uncovering therapeutic and adverse medication effects.
Choi L, Carroll RJ, Beck C, Mosley JD, Roden DM, Denny JC, Van Driest SL.
addLastDose

See Also

Useful links:

- [https://choileena.github.io/](https://choileena.github.io/)

---

**addLastDose**  
*Add Lastdose Data*

**Description**

Add lastdose data to data set from the `buildDose` process.

**Usage**

```
addLastDose(buildData, lastdoseData)
```

**Arguments**

- `buildData` data.frame, output of `buildDose` function.
- `lastdoseData` data.frame with columns filename, ld_start, lastdose, raw_time, time_type

**Details**

Lastdose is a datetime string associated with dose data. Information on time of last dose can be extracted within the `extractMed` function (i.e., `medExtractR`) using the argument `lastdose=TRUE`. Raw extracted times should first be processed using the `processLastDose` function to convert to datetime format before providing to `addLastDose`. This function then combines the processed last dose times with output from the `buildDose` process by file name to pair last dose times with dosing regimens based on position. Alternatively, the user can provide their own table of lastdose data. In this case, with position information absent, the lastdose data should be restricted to one unique last dose time per unique patient ID-date identifier.

In the case where `lastdoseData` is output from `processLastDose`, it is possible to have more than one extracted last dose time. In this case, rules are applied to determine which time should be kept. First, we give preference to an explicit time expression (e.g., "10:30pm") over a duration expression (e.g., "14 hour level"). Then, we pair last dose times with drug regimens based on minimum distance between last dose time start position and drug name start position.

See EHR Vignette for Extract-Med and Pro-Med-NLP for details.

**Value**

a data.frame with the ‘lastdose’ column added.
Examples

```r
# Get build data
data(tac_mxr_parsed)
# don’t combine lastdose at this stage
tac_build <- buildDose(tac_mxr_parsed, preserve = 'lastdose')
# Get processed last dose data
tac_mxr <- read.csv(system.file("examples", "tac_mxr.csv", package = "EHR"))
data(tac_metadata)
data(tac_lab)
ld_data <- processLastDose(tac_mxr, tac_metadata, tac_lab)

addLastDose(tac_build, ld_data)
```

### Description

Implement three commonly used statistical methods to analyze data for Phenome Wide Association Study (PheWAS)

### Usage

```r
analysisPheWAS(
  method = c("firth", "glm", "lr"),
  adjust = c("PS", "demo", "PS.demo", "none"),
  Exposure,
  PS,
  demographics,
  phenotypes,
  data
)
```

### Arguments

- **method**: define the statistical analysis method from 'firth', 'glm', and 'lr'. 'firth': Firth's penalized-likelihood logistic regression; 'glm': logistic regression with Wald test; 'lr': logistic regression with likelihood ratio test.
- **adjust**: define the adjustment method from 'PS', 'demo', 'PS.demo', and 'none'. 'PS': adjustment of PS only; 'demo': adjustment of demographics only; 'PS.demo': adjustment of PS and demographics; 'none': no adjustment.
- **Exposure**: define the variable name of exposure variable.
- **PS**: define the variable name of propensity score.
- **demographics**: define the list of demographic variables.
- **phenotypes**: define the list of phenotypes that need to be analyzed.
- **data**: define the data.
Details

Implements three commonly used statistical methods to analyze the associations between exposure (e.g., drug exposure, genotypes) and various phenotypes in PheWAS. Firth's penalized-likelihood logistic regression is the default method to avoid the problem of separation in logistic regression, which is often a problem when analyzing sparse binary outcomes and exposure. Logistic regression with likelihood ratio test and conventional logistic regression with Wald test can be also performed.

Value

- estimate: the estimate of log odds ratio.
- stdError: the standard error.
- statistic: the test statistic.
- pvalue: the p-value.

Author(s)

Leena Choi <naturechoi@gmail.com> and Cole Beck <cole.beck@vumc.org>

Examples

```r
## use small datasets to run this example
data(dataPheWASsmall)
## make dd.base with subset of covariates from baseline data (dd.baseline.small)
## or select covariates with upper code as shown below
upper.code.list <- unique(sub("[.]*([.])", "", colnames(dd.baseline.small)))
upper.code.list <- intersect(upper.code.list, colnames(dd.baseline.small))
dd.base <- dd.baseline.small[, upper.code.list]
## perform regularized logistic regression to obtain propensity score (PS)
## to adjust for potential confounders at baseline
phenos <- setdiff(colnames(dd.base), c('id', 'exposure'))
data.x <- as.matrix(dd.base[, phenos])
glmnet.fit <- glmnet::cv.glmnet(x=data.x, y=dd.base[, 'exposure'],
  family="binomial", standardize=TRUE, alpha=0.1)
dd.basePS <- c(predict(glmnet.fit, data.x, s='lambda.min'))
data.ps <- dd.base[,c('id', 'PS')]
 dd.all.ps <- merge(data.ps, dd.small, by='id')
demographics <- c('age', 'race', 'gender')
phenotypeList <- setdiff(colnames(dd.small), c('id','exposure','age','race','gender'))
## run with a subset of phenotypeList to get quicker results
phenotypeList.sub <- sample(phenotypeList, 5)
results.sub <- analysisPheWAS(method='firth', adjust='PS', Exposure='exposure',
  PS='PS', demographics=demographics, 
  phenotypes=phenotypeList.sub, data=dd.all.ps)
## run with the full list of phenotype outcomes (i.e., phenotypeList)
results <- analysisPheWAS(method='firth', adjust='PS', Exposure='exposure',
  PS='PS', demographics=demographics, 
  phenotypes=phenotypeList, data=dd.all.ps)
```
**buildDose**

---

**Combine Dose Data**

**Description**

Output from parse process is taken and converted into a wide format, grouping drug entity information together based on various steps and rules.

**Usage**

```r
buildDose(
  dat,
  dn = NULL,
  preserve = NULL,
  dist_method,
  na_penalty,
  neg_penalty,
  greedy_threshold,
  checkForRare = FALSE
)
```

**Arguments**

- **dat**: data.table object from the output of `parseMedExtractR`, `parseMedXN`, `parseMedEx`, or `parseCLAMP`.
- **dn**: Regular expression specifying drug name(s) of interest.
- **preserve**: Column names to include in output, whose values should not be combined with other rows. If present, dosechange is always preserved.
- **dist_method**: Distance method to use for calculating distance of various paths. Alternatively set the `ehr.dist_method` option, which defaults to `minEntEnd`.
- **na_penalty**: Penalty for matching extracted entities with NA. Alternatively set the `ehr.na_penalty` option, which defaults to 32.
- **neg_penalty**: Penalty for negative distances between frequency/intake time and dose amounts. Alternatively set the `ehr.neg_penalty` option, which defaults to 0.5.
- **greedy_threshold**: Threshold to use greedy matching; increasing this value too high could lead to the algorithm taking a long time to finish. Alternatively set the `ehr.greedy_threshold` option, which defaults to 1e8.
- **checkForRare**: Indicate if rare values for each entity should be found and displayed.

**Details**

The `buildDose` function takes as its main input (`dat`), a data.table object that is the output of a parse process function (`parseMedExtractR`, `parseMedXN`, `parseMedEx`, or `parseCLAMP`). Broadly, the parsed extractions are grouped together to form wide, more complete drug regimen information.
This reformatting facilitates calculation of dose given intake and daily dose in the `collapseDose` process.

The process of creating this output is broken down into multiple steps:

1. Removing rows for any drugs not of interest. Drugs of interest are specified with the `dn` argument.
2. Determining whether extractions are "simple" (only one drug mention and at most one extraction per entity) or complex. Complex cases can be more straightforward if they contain at most one extraction per entity, or require a pairing algorithm to determine the best pairing if there are multiple extractions for one or more entities.
3. Drug entities are anchored by drug name mention within the parse process. For complex cases, drug entities are further grouped together anchored at each strength (and dose with `medExtractR`) extraction.
4. For strength groups with multiple extractions for at least one entity, these groups go through a path searching algorithm, which computes the cost for each path (based on a chosen distance method) and chooses the path with the lowest cost.
5. The chosen paths for each strength group are returned as the final pairings. If route is unique within a strength group, it is standardized and added to all entries for that strength group.

The user can specify additional arguments including:

- `dist_method`: The distance method is the metric used to determine which entity path is the most likely to be correct based on minimum cost.
- `na_penalty`: NA penalties are incurred when extractions are paired with nothing (i.e., an NA), requiring that entities be sufficiently far apart from one another before being left unpaired.
- `neg_penalty`: When working with dose amount (DA) and frequency/intake time (FIT), it is much more common for the ordering to be DA followed by FIT. Thus, when we observe FIT followed by DA, we apply a negative penalty to make such pairings less likely.
- `greedy_threshold`: When there are many extractions from a clinical note, the number of possible combinations for paths can get exponentially large, particularly when the medication extraction natural language processing system is incorrect. The greedy threshold puts an upper bound on the number of entity pairings to prevent the function from stalling in such cases.

If none of the optional arguments are specified, then the `buildDose` process uses the default option values specified in the EHR package documentation. See EHR Vignette for Extract-Med and Pro-Med-NLP as well as Dose Building Using Example Vanderbilt EHR Data for details. For additional details, see McNeer, et al. 2020.

**Value**

A data.frame object that contains columns for filename (of the clinical note, inherited from the parse output object `dat`), drugname, strength, dose, route, freq, duration, and drugname_start.

**Examples**

```r
data(lam_mxr_parsed)
buildDose(lam_mxr_parsed)
```
collapseDose

**Collapse Dose Data**

**Description**

Splits drug data and calls `makeDose` to collapse at the note and date level.

**Usage**

```r
collapseDose(x, noteMetaData, naFreq = "most", ...)
```

**Arguments**

- `x`: data.frame containing the output of `buildDose`, or the output of `addLastDose` if last dose information is being incorporated.
- `noteMetaData`: data.frame containing identifying meta data for each note, including patient ID, date of the note, and note ID. Column names should be set to ‘filename’, ‘pid’, ‘date’, ‘note’. Date should have format YYYY-MM-DD.
- `naFreq`: Expression used to replace missing frequencies with, or by default use the most common.
- `...`: drug formulations to split by

**Details**

If different formulations of the drug (e.g., extended release) exist, they can be separated using a regular expression (e.g., ‘xr|er’). This function will call `makeDose` on parsed and paired medication data to calculate dose intake and daily dose and remove redundancies at the note and date level.

See EHR Vignette for Extract-Med and Pro-Med-NLP as well as Dose Building Using Example Vanderbilt EHR Data for details.

**Value**

A list containing two dataframes, one with the note level and one with the date level collapsed data.

**Examples**

```r
data(lam_mxr_parsed)
data(lam_metadata)

lam_build_out <- buildDose(lam_mxr_parsed)

lam_collapsed <- collapseDose(lam_build_out, lam_metadata, naFreq = 'most', 'xr|er')

lam_collapsed$note # Note level collapsing
lam_collapsed$date # Date level collapsing
```
**dataTransformation**  
*Data Transformation*

**Description**

Convenience function for making small modifications to a data.frame.

**Usage**

```r
dataTransformation(x, select, rename, modify)
```

**Arguments**

- `x`: a data.frame
- `select`: columns to select
- `rename`: character vector with names for all columns
- `modify`: list of expressions used to transform data set

**Value**

The modified data.frame

---

**dd**

**Description**

Simulated outcome data example from Phenome Wide Association Study (PheWAS) that examines associations between drug exposure and various phenotypes at follow-up after the drug exposure. The dataset includes 1505 variables: subject identification number (‘id’), drug exposure (‘exposure’), 3 demographic variables (‘age’, ‘race’, ‘gender’), and 1500 phenotypes.

**Usage**

```r
data(dataPheWAS, package = 'EHR')
```

**Format**

A data frame with 10000 observations on 1505 variables.

**Examples**

```r
data(dataPheWAS)
```
dd.baseline  

**Description**  
Simulated baseline data example from a Phenome Wide Association Study (PheWAS) obtained at baseline before drug exposure. The dataset includes 1505 variables: subject identification number (‘id’), drug exposure (‘exposure’), 3 demographic variables (‘age’, ‘race’, ‘gender’), and 1500 phenotypes.

**Usage**  
```r  
data(dataPheWAS, package = 'EHR')  
```

**Format**  
A data frame with 10000 observations on 1505 variables.

**Examples**  
```r  
data(dataPheWAS)  
```

---

dd.baseline.small  

**Description**  
A smaller subset of baseline data example, dd.baseline. The dataset includes 55 variables: subject identification number (‘id’), drug exposure (‘exposure’), 3 demographic variables (‘age’, ‘race’, ‘gender’), and 50 phenotypes.

**Usage**  
```r  
data(dataPheWASsmall, package = 'EHR')  
```

**Format**  
A data frame with 2000 observations on 55 variables.

**Examples**  
```r  
data(dataPheWASsmall)  
```
Description

A smaller subset of outcome data example, 'dd'. The dataset includes 55 variables: subject identification number ('id'), drug exposure ('exposure'), 3 demographic variables ('age', 'race', 'gender'), and 50 phenotypes.

Usage

data(dataPheWASsmall, package = 'EHR')

Format

A data frame with 2000 observations on 55 variables.

Examples

data(dataPheWASsmall)

extractMed

Extract medication information from clinical notes

Description

This function is an interface to the medExtractR function within the medExtractR package, and allows drug dosing information to be extracted from free-text sources, e.g., clinical notes.

Usage

extractMed(note_fn, drugnames, drgunit, windowlength, max_edit_dist = 0, ...)

Arguments

- `note_fn` File name(s) for the text file(s) containing the clinical notes. Can be a character string for an individual note, or a vector or list of file names for multiple notes.
- `drugnames` Vector of drug names for which dosing information should be extracted. Can include various forms (e.g., generic, brand name) as well as abbreviations.
- `drgunit` Unit of the drug being extracted, e.g., 'mg'
- `windowlength` Length of the search window (in characters) around the drug name in which to search for dosing entities
- `max_edit_dist` Maximum edit distance allowed when attempting to extract drugnames. Allows for capturing misspelled drug name information.
- `...` Additional arguments to medExtractR, for example lastdose=TRUE to extract time of last dose (see medExtractR package documentation for details)
freqNum

Details

Medication information, including dosing data, is often stored in free-text sources such as clinical notes. The extractMed function serves as a convenient wrapper for the medExtractR package, a natural language processing system written in R for extracting medication data. Within extractMed, the medExtractR function identifies dosing data for drug(s) of interest, specified by the drugnames argument, using rule-based and dictionary-based approaches. Relevant dosing entities include medication strength (identified using the unit argument), dose amount, dose given intake, intake time or frequency of dose, dose change keywords (e.g., 'increase' or 'decrease'), and time of last dose. After applying medExtractR to extract drug dosing information, extractMed appends the file name to results to ensure they are appropriately labeled.


Value

A data.frame with the extracted dosing information, labeled with file name as an identifier

Sample output:

<table>
<thead>
<tr>
<th>filename</th>
<th>entity</th>
<th>expr</th>
<th>pos</th>
</tr>
</thead>
<tbody>
<tr>
<td>note_file1.txt</td>
<td>DoseChange</td>
<td>decrease</td>
<td>66:74</td>
</tr>
<tr>
<td>note_file1.txt</td>
<td>DrugName</td>
<td>Prograf</td>
<td>78:85</td>
</tr>
<tr>
<td>note_file1.txt</td>
<td>Strength</td>
<td>2 mg</td>
<td>86:90</td>
</tr>
<tr>
<td>note_file1.txt</td>
<td>DoseAmt</td>
<td>1</td>
<td>91:92</td>
</tr>
<tr>
<td>note_file1.txt</td>
<td>Frequency</td>
<td>bid</td>
<td>101:104</td>
</tr>
<tr>
<td>note_file1.txt</td>
<td>LastDose</td>
<td>2100</td>
<td>121:125</td>
</tr>
</tbody>
</table>

Examples

```r
tac_fn <- list(system.file("examples", "tacpid1_2008-06-26_note1_1.txt", package = "EHR"),
               system.file("examples", "tacpid1_2008-06-26_note2_1.txt", package = "EHR"),
               system.file("examples", "tacpid1_2008-12-16_note3_1.txt", package = "EHR"))

extractMed(tac_fn,
drugnames = c("tacrolimus", "prograf", "tac", "tacro", "fk", "fk506"),
drgunit = "mg",
windowlength = 60,
max_edit_dist = 2,
lastdose=TRUE)
```

freqNum

Convert Character Frequency to Numeric

Description

This function converts the frequency entity to numeric.
Usage

freqNum(x)

Arguments

x character vector of extracted frequency values

Value

numeric vector

Examples

f <- stdzFreq(c('in the morning', 'four times a day', 'with meals'))
freqNum(f)

idCrosswalk

Create ID Crosswalk

Description

Link ID columns from multiple data sets. De-identified columns are created to make a crosswalk.

Usage

idCrosswalk(data, idcols, visit.id = "subject_id", uniq.id = "subject_uid")

Arguments

data list of data.frames
idcols list of character vectors, indicating ID columns found in each data set given in 'data'
visit.id character string indicating visit-level ID variable (default is "subject_id")
uniq.id character string indicating subject-level ID variable (default is "subject_uid")

Details

'visit.id' and 'uniq.id' may occur multiple times, but should have a one-to-one linkage defined by at least one of the input data sets. A new visit number is generated for each repeated 'uniq.id'.

Value

crosswalk of ID columns and their de-identified versions
Examples

```r
## Not run:
demo_data <- data.frame(subj_id=c(4.1,4.2,5.1,6.1),
                         pat_id=c(14872,14872,24308,37143),
                         gender=c(1,1,0,1),
                         weight=c(34,42,28,63),
                         height=c(142,148,120,167))

conc_data <- data.frame(subj_id=rep((4:6)+0.1,each=5),
                        event=rep(1:5,times=3),
                        conc.level=15*exp(-1*rep(1:5,times=3))+rnorm(15,0,0.1))

data <- list(demo_data, conc_data)
idcols <- list(c('subj_id', 'pat_id'), 'subj_id')
idCrosswalk(data, idcols, visit.id='subj_id', uniq.id='pat_id')
```  

## End(Not run)

### Description

An example of the metadata needed for the `processLastDose`, `makeDose`, and `collapseDose` functions.

### Usage

```r
data(lam_metadata, package = 'EHR')
```

### Format

A data frame with 5 observations on the following variables.

- **filename** A character vector, filename for the clinical note
- **pid** A character vector, patient ID associated with the filename
- **date** A character vector, date associated with the filename
- **note** A character vector, note ID associated with the filename

### Examples

```r
data(lam_metadata)
```
Description

The output after running `parseMedExtractR` on 4 example clinical notes.

Usage

data(lam_mxr_parsed, package = 'EHR')

Format

A data frame with 10 observations on the following variables.

- **filename**: A character vector, filename for the clinical note
- **drugname**: A character vector, drug name extracted from the clinical note along with start and stop positions
- **strength**: A character vector, strengths extracted from the clinical note along with start and stop positions
- **dose**: A character vector, dose amounts extracted from the clinical note along with start and stop positions
- **route**: A character vector, routes extracted from the clinical note along with start and stop positions
- **freq**: A character vector, frequencies extracted from the clinical note along with start and stop positions
- **dosestr**: A character vector, dose intakes extracted from the clinical note along with start and stop positions
- **dosechange**: A character vector, dose change keywords extracted from the clinical note along with start and stop positions
- **lastdose**: A character vector, last dose times extracted from the clinical note along with start and stop positions

Examples

data(lam_mxr_parsed)
Logistf

Firth’s penalized-likelihood logistic regression with more decimal places of p-value than logistf function in the R package ‘logistf’

Description

Adapted from logistf in the R package ‘logistf’, this is the same as logistf except that it provides more decimal places of p-value that would be useful for Genome-Wide Association Study (GWAS) or Phenome Wide Association Study (PheWAS).

Usage

Logistf(
  formula = attr(data, "formula"),
  data = sys.parent(),
  pl = TRUE,
  alpha = 0.05,
  control,
  plcontrol,
  firth = TRUE,
  init,
  weights,
  plconf = NULL,
  dataout = TRUE,
  ...
)

Arguments

formula a formula object, with the response on the left of the operator, and the model terms on the right. The response must be a vector with 0 and 1 or FALSE and TRUE for the outcome, where the higher value (1 or TRUE) is modeled. It is possible to include contrasts, interactions, nested effects, cubic or polynomial splines and all S features as well, e.g. \( Y \sim X1 \times X2 + ns(x3, df=4) \). From version 1.10, you may also include offset() terms.

data a data.frame where the variables named in the formula can be found, i.e. the variables containing the binary response and the covariates.

pl specifies if confidence intervals and tests should be based on the profile penalized log likelihood (pl=TRUE, the default) or on the Wald method (pl=FALSE).

alpha the significance level (1-\( \alpha \) the confidence level, 0.05 as default).

control Controls Newton-Raphson iteration. Default is control=logistf.control(maxstep, maxit, maxhs, lconv, gconv, xconv)

plcontrol Controls Newton-Raphson iteration for the estimation of the profile likelihood confidence intervals. Default is plcontrol=logistpl.control(maxstep, maxit, maxhs, lconv, xconv, ortho, pr)
**firth**

use of Firth’s penalized maximum likelihood (firth=TRUE, default) or the standard maximum likelihood method (firth=FALSE) for the logistic regression. Note that by specifying pl=True and firth=FALSE (and probably a lower number of iterations) one obtains profile likelihood confidence intervals for maximum likelihood logistic regression parameters.

**init**

specifies the initial values of the coefficients for the fitting algorithm.

**weights**

specifies case weights. Each line of the input data set is multiplied by the corresponding element of weights.

**plconf**

specifies the variables (as vector of their indices) for which profile likelihood confidence intervals should be computed. Default is to compute for all variables.

**dataout**

If TRUE, copies the data set to the output object.

... Further arguments to be passed to logistf.

**Value**

same as logistf except for providing more decimal places of p-value.

**Author(s)**

Leena Choi <naturechoi@gmail.com> and Cole Beck <cole.beck@vumc.org>

**References**

same as those provided in the R package ‘logistf’.

**Examples**

data(dataPheWAS)
fit <- Logistf(X264.3 ~ exposure + age + race + gender, data=dd)
summary(fit)

---

**makeDose**

*Make Dose Data*

**Description**

Takes parsed and paired medication data, calculates dose intake and daily dose, and removes redundant information at the note and date level.

**Usage**

makeDose(x, noteMetaData, naFreq = "most")
Arguments

- **x**: data.frame containing the output of `buildDose`, or the output of `addLastDose` if last dose information is being incorporated.
- **noteMetaData**: data.frame containing identifying meta data for each note, including patient ID, date of the note, and note ID. Column names should be set to ‘filename’, ‘pid’, ‘date’, ‘note’. Date should have format YYYY-MM-DD.
- **naFreq**: Replacing missing frequencies with this value, or by default the most common value across the entire set in `x`.

Details

This function standardizes frequency, route, and duration entities. Dose amount, strength, and frequency entities are converted to numeric. Rows with only drug name and/or route are removed. If there are drug name changes in adjacent rows (e.g., from a generic to brand name), these rows are collapsed into one row if there are no conflicts. Missing strengths, dose amounts, frequencies, and routes are borrowed or imputed using various rules (see McNeer et al., 2020 for details). Dose given intake and daily dose are calculated. Redundancies are removed at the date and note level. If time of last dose is being used and it is unique within the level of collapsing, it is borrowed across all rows.

Value

A list containing two dataframes, one with the note level and one with the date level collapsed data.

Examples

```r
data(lam_mxr_parsed)
data(lam_metadata)

lam_build_out <- buildDose(lam_mxr_parsed)

lam_collapsed <- makeDose(lam_build_out, lam_metadata)
lam_collapsed[[1]] # Note level collapsing
lam_collapsed[[2]] # Date level collapsing
```

Description

Takes files with the raw medication extraction output generated by the CLAMP natural language processing system and converts it into a standardized format.

Usage

`parseCLAMP(filename)`
Arguments
filename File name for a single file containing CLAMP output.

Details
Output from different medication extraction systems is formatted in different ways. In order to be able to process the extracted information, we first need to convert the output from different systems into a standardized format. Extracted expressions for various drug entities (e.g., drug name, strength, frequency, etc.) each receive their own column formatted as "extracted expression::start position::stop position". If multiple expressions are extracted for the same entity, they will be separated by backticks.

CLAMP output files anchor extractions to a specific drug name extraction through semantic relations.

See EHR Vignette for Extract-Med and Pro-Med-NLP as well as Dose Building Using Example Vanderbilt EHR Data for details.

Value
A data.table object with columns for filename, drugname, strength, dose, route, and freq. The filename contains the file name corresponding to the clinical note. Each of the entity columns are of the format "extracted expression::start position::stop position".

Description
Takes files with the raw medication extraction output generated by the MedEx natural language processing system and converts it into a standardized format.

Usage
parseMedEx(filename)

Arguments
filename File name for a single file containing MedEx output.

Details
Output from different medication extraction systems is formatted in different ways. In order to be able to process the extracted information, we first need to convert the output from different systems into a standardized format. Extracted expressions for various drug entities (e.g., drug name, strength, frequency, etc.) each receive their own column formatted as "extracted expression::start position::stop position". If multiple expressions are extracted for the same entity, they will be separated by backticks.
MedEx output files anchor extractions to a specific drug name extraction.
See EHR Vignette for Extract-Med and Pro-Med-NLP as well as Dose Building Using Example Vanderbilt EHR Data for details.

Value
A data.table object with columns for filename, drugname, strength, dose, route, and freq. The filename contains the file name corresponding to the clinical note. Each of the entity columns are of the format "extracted expression::start position::stop position".

parseMedExtractR
Parse medExtractR NLP Output

Description
Takes files with the raw medication extraction output generated by the medExtractR natural language processing system and converts it into a standardized format.

Usage
parseMedExtractR(filename)

Arguments
filename
File name for a single file containing medExtractR output.

Details
Output from different medication extraction systems is formatted in different ways. In order to be able to process the extracted information, we first need to convert the output from different systems into a standardized format. Extracted expressions for various drug entities (e.g., drug name, strength, frequency, etc.) each receive their own column formatted as "extracted expression::start position::stop position". If multiple expressions are extracted for the same entity, they will be separated by backticks.

The medExtractR system returns extractions in a long table format, indicating the entity, extracted expression, and start:stop position of the extraction. To perform this initial parsing, entities are paired with the closest preceding drug name. The one exception to this is the dose change entity, which can occur before the drug name (see Weeks, et al. 2020 for details).

See EHR Vignette for Extract-Med and Pro-Med-NLP as well as Dose Building Using Example Vanderbilt EHR Data for details.

Value
A data.table object with columns for filename, drugname, strength, dose, route, freq, dosestr, dosechange and lastdose. The filename contains the file name corresponding to the clinical note. Each of the entity columns are of the format "extracted expression::start position::stop position".
parseMedXN

**Examples**
```r
mrx_output <- system.file("examples", "lam_mxr.csv", package = "EHR")
mrx_parsed <- parseMedExtractR(mrx_output)
mrx_parsed
```

<table>
<thead>
<tr>
<th>parseMedXN</th>
<th>Parse MedXN NLP Output</th>
</tr>
</thead>
</table>

**Description**
Takes files with the raw medication extraction output generated by the MedXN natural language processing system and converts it into a standardized format.

**Usage**
```r
parseMedXN(filename, begText = "^[R0-9]+_[0-9-]+_[0-9-].")
```

**Arguments**
- **filename**: File name for single file containing MedXN output.
- **begText**: A regular expression that would indicate the beginning of a new observation (i.e., extracted clinical note).

**Details**
Output from different medication extraction systems is formatted in different ways. In order to be able to process the extracted information, we first need to convert the output from different systems into a standardized format. Extracted expressions for various drug entities (e.g., drug name, strength, frequency, etc.) each receive their own column formatted as "extracted expression::start position::stop position". If multiple expressions are extracted for the same entity, they will be separated by backticks.

MedXN output files anchor extractions to a specific drug name extraction.

In MedXN output files, the results from multiple clinical notes can be combined into a single output file. The beginning of some lines of the output file can indicate when output for a new observation (or new clinical note) begins. The user should specify the argument `begText` to be a regular expression used to identify the lines where output for a new clinical note begins.

See EHR Vignette for Extract-Med and Pro-Med-NLP as well as Dose Building Using Example Vanderbilt EHR Data for details.

**Value**
A data.table object with columns for filename, drugname, strength, dose, route, freq, and duration. The filename contains the file name corresponding to the clinical note. Each of the entity columns are of the format "extracted expression::start position::stop position".
processLastDose

Examples

```r
mxn_output <- system.file("examples", "lam_medxn.csv", package = "EHR")
mxn_parsed <- parseMedXN(mxn_output, begText = "^ID\[0-9\]+_[0-9-]+")
mxn_parsed
```

Description

This function takes last dose times extracted using the `medExtractR` system and processes the times into standardized datetime objects using recorded lab data where necessary. The raw output from `extractMed` is filtered to just the LastDose extractions. Time expressions are standardized into HH:MM:SS format based on what category they fall into (e.g., a time represented with AM/PM, 24-hour military time, etc.). When the last dose time is after 12pm, it is assumed to have been taken one day previous to the note’s date. For any duration extractions (e.g., "14 hour level"), the last dose time is calculated from the labtime by extracting the appropriate number of hours. The final dataset is returned with last dose time formatted into a POSIXct variable.

Usage

```r
processLastDose(mxrData, noteMetaData, labData)
```

Arguments

- `mxrData`: data.frame containing output from the `medExtractR` system
- `noteMetaData`: data.frame with meta data (pid (patient ID) and date) for the file names contained within `mxrData`
- `labData`: data.frame that contains lab dates and times associated with the file names within `mxrData`. Must contain columns pid and date, as well as labtime. The date column must be in the same format as date in `noteMetaData`, and `labtime` must be a POSIXct

Details

See EHR Vignette for Extract-Med and Pro-Med-NLP for details.

Value

data.frame with identifying information (e.g., filename, etc) as well as processed and standardized last dose times as a POSIXct column
pullFakeId

Examples

```r
tac_mxr <- read.csv(system.file("examples", "tac_mxr.csv", package = "EHR"))
data(tac_metadata)
data(tac_lab)

processLastDose(mxrData = tac_mxr, noteMetaData = tac_metadata, labData = tac_lab)
```

---

### pullFakeId

#### Pull Fake/Mod ID

<table>
<thead>
<tr>
<th>Description</th>
</tr>
</thead>
<tbody>
<tr>
<td>Replace IDs with de-identified version pulled from a crosswalk.</td>
</tr>
</tbody>
</table>

#### Usage

```r
pullFakeId(
  dat, 
  xwalk, 
  firstCols = NULL, 
  orderBy = NULL, 
  uniq.id = "subject_uid"
)
```

#### Arguments

- **dat**
  a data.frame
- **xwalk**
  a data.frame providing linkage for each ID, e.g. output from `idCrosswalk`
- **firstCols**
  name of columns to put at front of output data set
- **orderBy**
  name of columns used to reorder output data set
- **uniq.id**
  character string indicating subject-level id variable (default is "subject_uid")

#### Value

The modified data.frame

#### Examples

```r
## Not run:
demo_data <- data.frame(subj_id=c(4.1,4.2,5.1,6.1),
                         pat_id=c(14872,14872,24308,37143),
                         gender=c(1,1,0,1),
                         weight=c(34,42,28,63),
                         height=c(142,148,120,167))

# crosswalk w/ same format as idCrosswalk() output
```
**pullRealId**

Replace de-identified IDs with identified version pulled from a crosswalk.

**Usage**

```r
pullRealId(dat, xwalk = NULL, remove.mod.id = FALSE)
```

**Arguments**

- `dat`: a data.frame
- `xwalk`: a data.frame providing linkage for each ID, e.g. output from `idCrosswalk`; if `NULL`, the crosswalk will be pulled from the `pkxwalk` option, or otherwise the unmodified data.frame.
- `remove.mod.id`: logical, should the de-identified IDs – `mod_id`, `mod_visit`, `mod_id_visit` – be removed (default=FALSE)

**Value**

The modified data.frame

**Examples**

```r
## Not run:
demo_data_deident <- pullFakeId(demo_data, xwalk, 
   firstCols = c('mod_id', 'mod_id_visit', 'mod_visit'), 
   uniq.id='pat_id')

## End(Not run)
```
readTransform <- data.frame(subj_id=c(4.1,4.2,5.1,6.1),
pat_id=c(14872,14872,24308,37143),
mod_visit=c(1,2,1,1),
mod_id=c(1,1,2,3),
mod_id_visit=c(1.1,1.2,2.1,3.1))
pullRealId(demo_data_deident, xwalk)
pullRealId(demo_data_deident, xwalk, remove.mod.id=TRUE)

## End(Not run)

readTransform  Read and Transform

Description

Convenience function for reading in a CSV file, and making small modifications to a data.frame.

Usage

readTransform(file, ...)

Arguments

file  filename of a CSV file
...
additional information passed to dataTransformation

Details

If read.csv needs additional arguments (or the file is in a different format), the user should load the data first, then directly call dataTransformation.

Value

The modified data.frame
run_Build_PK_IV

Description

This module builds PK data for intravenously (IV) administered medications.

Usage

run_Build_PK_IV(
  conc,
  conc.columns = list(),
  dose,
  dose.columns = list(),
  demo.list = NULL,
  demo.columns = list(),
  lab.list = NULL,
  lab.columns = list(),
  pk.vars = NULL,
  drugname = NULL,
  check.path = NULL,
  missdemo_fn = "-missing-demo",
  faildupbol_fn = "DuplicateBolus-",
  date.format = "%m/%d/%y %H:%M:%S",
  date.tz = "America/Chicago"
)

Arguments

conc concentration data, the output of run_DrugLevel, a filename (CSV, RData, RDS), or a correctly formatted data.frame

conc.columns a named list that should specify columns in concentration data; ‘id’, ‘datetime’, ‘druglevel’ are required. ‘idvisit’ may also be specified; ‘idvisit’ can be used when there are multiple visits (i.e., several occasions) for the same subject. ‘datetime’ is date and time for concentration measurement, which can refer to a single date-time variable (datetime = ‘date_time’) or two variables holding date and time separately (e.g., datetime = c(‘Date’, ‘Time’)).

dose dose data, the output of run_MedStrI, a filename (CSV, RData, RDS), or a correctly formatted data.frame

dose.columns a named list that should specify columns in dose data; ‘id’ is required. ‘infuseDatetime’ and ‘infuseDose’ should be set if infusion dose data is present. ‘infuseTimeExact’ may also be specified for infusion data – this variable represents an precise time, if for example the ‘infuseDatetime’ variable is rounded. ‘bolusDatetime’ and ‘bolusDose’ should be set if bolus dose data is present. A generic ‘date’ variable may be provided, agnostic to either infusion or bolus
dosing. ‘gap’ and ‘weight’ column names may also be set. Any of the date-
time variables can be specified as a single date-time variable (infuseDateTime =
‘date_time’) or two variables holding date and time separately (e.g., infuseDateTime = c(‘Date’, ‘Time’)).

demo.list demographic information, if available; the output from run_Demo or a correctly
formatted data.frame
demo.columns a named list that should specify columns in demographic data; ‘id’ is required.
‘weight’ and ‘idvisit’ may also be used to specify columns for weight or the
unique idvisit. Any other columns present in the demographic data are treated
as covariates.
lab.list lab data, if available; the output from run_Labs or a correctly formatted list
lab.columns a named list that should specify columns in lab data; ‘id’, and ‘datetime’ are
required. ‘datetime’ is the date and time when the lab data was obtained, which
can refer to a single date-time variable (datetime = ‘date_time’) or two variables
holding date and time separately (e.g., datetime = c(‘Date’, ‘Time’)). Any other
columns present in lab data are treated as lab values.
pk.vars variables to include in the returned PK data. The variable ‘date’ is a special case;
when included, it maps the ‘time’ offset to its original date-time. Other named
variables will be merged from the concentration data set. For example, rather
than being separate data sets, labs or demographics may already be present in
the concentration data. These columns should be named here.
drugname drug of interest, included in filename of check files. The default (NULL) will
produce filenames without drugname included.
check.path path to ‘check’ directory, where check files are created. The default (NULL)
will not produce any check files.
missdemo_fn filename for checking NA frequency among demographic data
faildupbol_fn filename for duplicate bolus data
date.format output format for ‘date’ variable
date.tz output time zone for ‘date’ variable

Details

See EHR Vignette for Structured Data.

Regarding the ‘gap’ variable in the dose dataset, if ‘gap’ is specified in ‘dose.columns’, it allows
a continuous infusion given when there are missing records between infusion dosing records. For
example, suppose that ‘gap’ = 60 is defined (which is typical gap size when infusion dosing is
supposed to be recorded hourly for inpatients) and time between two records (i.e., gap) are greater
than 1 hour (i.e., missing records). If the gap between the two records is less or equal to twice of the
gap (i.e., 2*60 = 120 min), a continuous infusion is assumed until the 2nd dose record; otherwise,
the first infusion is assumed to be stopped (i.e., add zero doses) after 60 min (i.e., equal to the gap
size) and a new infusion (the 2nd record) starts at its recorded time.

Value

PK data set
run_Build_PK_Oral

Examples

```r
## Not run:

# make fake data
set.seed(6543)

build_date <- function(x) as.character(seq(x, length.out=5, by="1 hour"))
dates <- unlist(lapply(rep(Sys.time(),3), build_date))

plconc <- data.frame(mod_id = rep(1:3,each=5),
                      mod_id_visit = rep(1:3,each=5)+0.1,
                      event = rep(1:5(times=3)),
                      conc.level = 15*exp(-1*rep(1:5(times=3)))+rnorm(15,0,0.1),
                      date.time = as.POSIXct(dates))

ivdose <- data.frame(mod_id = 1:3,
                      date.dose = substr(dates[seq(1,15,by=5)],1,10),
                      infuse.time.real = NA, infuse.time = NA, infuse.dose = NA,
                      bolus.time = as.POSIXct(dates[seq(1,15,by=5)])-300,
                      bolus.dose = 90,
                      maxint = 0L,
                      weight = 45)

run_Build_PK_IV(conc = plconc,
                 conc.columns = list(id = 'mod_id', datetime = 'date.time',
                                    druglevel = 'conc.level', idvisit = 'mod_id_visit'),
                 dose = ivdose,
                 dose.columns = list(id = 'mod_id', date = 'date.dose',
                                      bolusDatetime = 'bolus.time', bolusDose = 'bolus.dose',
                                      gap = 'maxint', weight = 'weight'),
                 pk.vars = 'date')

## End(Not run)
```

run_Build_PK_Oral  Build-PK-Oral Module

Description

This module builds PK data for orally administered medications.

Usage

```r
run_Build_PK_Oral(
  x,
  idCol = "id",
  dtCol = "dt",
```

---

**Important**: The code examples above are not meant to be run directly. They are provided as an illustration of how the `run_Build_PK_Oral` function works. They should be modified and tested independently.
doseCol = "dose",
concCol = "conc",
ldCol = NULL,
first_interval_hours = 336,
imputeClosest = NULL
)

Arguments

x a data.frame or file saved as either CSV, RData, or RDS
idCol data.frame id column name
dtCol data.frame date column name
doseCol dose column name
concCol concentration column name
ldCol last-dose time column name
first_interval_hours number of hours before the first concentration to start time=0; the default is 336 hours = 14 days
imputeClosest columns to impute missing data with next observation propagated backward; this is in addition to all covariates receiving imputation using last observation carried forward

Details

See EHR Vignette for Build-PK-Oral.

Value
data.frame

Examples

```r
## Not run:
## Data Generating Function
mkdat <- function() {
  npat <- 3
  visits <- floor(runif(npat, min=2, max=6))
  id <- rep(1:npat, visits)
  dt_samp <- as.Date(sort(sample(700, sum(visits))), origin = '2019-01-01')
  tm_samp <- as.POSIXct(paste(dt_samp, '10:00:00'), tz = 'UTC')
  dt <- tm_samp + rnorm(sum(visits), 0, 1*60*60)
  dose_morn <- sample(c(2.5,5,7.5,10), sum(visits), replace = TRUE)
  conc <- round(rnorm(sum(visits), 1.5*dose_morn, 1),1)
  ld <- dt - sample(10:16, sum(visits), replace = TRUE) * 3600
  ld[rnorm(sum(visits)) < .3] <- NA
  age <- rep(sample(40:75, npat), visits)
  gender <- rep(sample(0:1, npat, replace=TRUE), visits)
  weight <- rep(round(rnorm(npat, 180, 20)),visits)
  hgb <- rep(rnorm(npat, 10, 2), visits)
```
run_Demo

data.frame(id, dt, dose_morn, conc, ld, age, gender, weight, hgb)
}

# Make raw data
set.seed(30)
dat <- mkdat()

# Process data without last-dose times
run_Build_PK_Oral(x = dat,
idCol = "id",
dtCol = "dt",
doseCol = "dose_morn",
concCol = "conc",
ldCol = NULL,
first_interval_hours = 336,
imputeClosest = NULL)

# Process data with last-dose times
run_Build_PK_Oral(x = dat, doseCol = "dose_morn", ldCol = "ld")

## End(Not run)

run_Demo

Run Demographic Data

Description

This module will load and modify demographic data.

Usage

run_Demo(demo.path, toexclude, demo.mod.list)

Arguments

demo.path filename of demographic file (CSV, RData, RDS) or data.frame
toexclude expression that should evaluate to a logical, indicating if the observation should be excluded
demo.mod.list list of expressions, giving modifications to make

Details

See EHR Vignette for Structured Data.

Value

list with two components
demo demographic data
exclude vector of excluded visit IDs
Examples

```r
set.seed(2525)
dateSeq <- seq(as.Date('2019/01/01'), as.Date('2020/01/01'), by="day")
demo <- data.frame(mod_id_visit = 1:10,
                   weight.lbs = rnorm(10, 160, 20),
                   age = rnorm(10, 50, 10),
                   enroll.date = sample(dateSeq, 10))
tmpfile <- paste0(tempfile(), '.rds')
saveRDS(demo, file = tmpfile)

# exclusion functions
exclude_wt <- function(x) x < 150
exclude_age <- function(x) x > 60
ind.risk <- function(wt, age) wt > 170 & age > 55
exclude_enroll <- function(x) x < as.Date('2019/04/01')

# make demographic data that:
# (1) excludes ids with weight.lbs < 150, age > 60, or enroll.date before 2019/04/01
# (2) creates new 'highrisk' variable for subjects with weight.lbs>170 and age>55
out <- run_Demo(demo.path = tmpfile,
                toexclude = expression(exclude_wt(weight.lbs)|exclude_age(age)|exclude_enroll(enroll.date)),
                demo.mod.list = list(highrisk = expression(ind.risk(weight.lbs, age))))

out
```

run_DrugLevel  

Run Drug Level Data

Description

This module will load and modify drug-level data.

Usage

```r
run_DrugLevel(
conc.path,
conc.select,
conc.rename,
conc.mod.list = list(mod_id_event = expression(paste(mod_id_visit, event, sep = "_"))),
samp.path = NULL,
samp.mod.list = list(mod_id_event = expression(paste(mod_id_visit, samp, sep = "_"))),
check.path = NULL,
failmiss_fn = "MissingConcDate-",
multsets_fn = "multipleSetsConc-",
```
run_DrugLevel

faildup_fn = "DuplicateConc-",
drugname = NULL,
LLOQ = NA,
demo.list = NULL
)

Arguments

conc.path filename of concentration data (CSV, RData, RDS), or data.frame
conc.select columns to select from concentration data
conc.rename new column names for concentration data
conc.mod.list list of expressions, giving modifications to make
samp.path filename of data with sampling time (CSV, RData, RDS), or data.frame
samp.mod.list list of expressions, giving modifications to make
check.path path to 'check' directory, where check files are created. The default (NULL) will not produce any check files.
failmiss_fn filename for data missing concentration date
multsets_fn filename for data with multiple concentration sets
faildup_fn filename for data with duplicate concentration observations
drugname drug of interest, included in filename of check files. The default (NULL) will produce filenames without drugname included.
LLOQ lower limit of concentration values; values below this are invalid
demo.list demographic information; if available, concentration records must have a valid demo record

Details

See EHR Vignette for Structured Data.

Value
drug-level data set

Examples

## Not run:
# concentrations
cconc.data <- data.frame(mod_id = rep(1:3,each=4),
  mod_visit = rep(c(2,1,1),each=4),
  mod_id_visit = as.numeric(paste(rep(1:3,each=4),
    rep(c(2,1,1),each=4), sep=".")),
  samp = rep(1:4,times=3),
  drug_calc_conc=15*exp(-1*rep(1:4,times=3))+rnorm(12,0,0.1))

saveRDS(conc.data,'conc_data.rds')

# sample times
```r
build_date <- function(x) as.character(seq(x, length.out=4, by="1 hour"))
dates <- unlist(lapply(rep(Sys.time(),3), build_date))
samp_data <- data.frame(mod_id = rep(1:3,each=4),
                       mod_visit = rep(c(2,1,1),each=4),
                       mod_id_visit = as.numeric(paste(rep(1:3,each=4),
                           rep(c(2,1,1),each=4), sep=".") ),
                       samp = rep(1:4,times=3),
                       Sample.Collection.Date.and.Time = dates)
saveRDS(samp_data, 'samp_data.rds')
run_DrugLevel(conc.path = 'conc_data.rds',
              conc.select = c('mod_id', 'mod_id_visit', 'samp', 'drug_calc_conc'),
              conc.rename = c(drug_calc_conc = 'conc.level', samp='event'),
              conc.mod.list = list(mod_id_event = expression(paste(mod_id_visit, event, sep = "."))),
              samp.path = 'samp_data.rds',
              samp.mod.list = list(mod_id_event = expression(paste(mod_id_visit, samp, sep = "."))),
              check.path = tempdir(),
              drugname = 'drugnm',
              LLOQ = 0.05)
# minimal example with data in required format
conc_data <- conc_data[,c('mod_id', 'mod_id_visit', 'samp', 'drug_calc_conc')]
conc_data[, 'mod_id_event'] <- paste(conc_data[, 'mod_id_visit'], conc_data[ 'samp'], sep = ".")
names(conc_data)[3:6] <- c('event', 'conc.level')
samp_data[, 'mod_id_event'] <- paste(samp_data[, 'mod_id_visit'], samp_data[ 'samp'], sep = ".")
conc_samp_link <- match(conc_data[, 'mod_id_event'], samp_data[, 'mod_id_event'])
conc_date <- samp_data[conc_samp_link, 'Sample.Collection.Date.and.Time']
conc_data[, 'date.time'] <- as.POSIXct(conc_date)
run_DrugLevel(conc_data)
```

---

**run_Labs**

**Run Lab Data**

**Description**

This module will load and modify laboratory data.

**Usage**

```r
run_Labs(lab.path, lab.select, lab.mod.list)
```
run_MedStrI

Arguments

lab.path filename of a lab file (CSV, RData, RDS), or data.frame
lab.select columns to select
lab.mod.list list of expressions giving modifications to make; passed to dataTransformation

Details

See EHR Vignette for Structured Data.

Value

lab data set

Examples

```r
## Not run:
lab_data <- data.frame(mod_id=rep(1:3,each=3),
                       date=rep(c("01/12/17","05/05/18","11/28/16"),each=3),
                       time=rep(c("1:30","2:30","3:30"),3),
                       creat=rnorm(9,0.5,0.05))

saveRDS(lab_data, 'lab_data.rds')

run_Labs('lab_data.rds', lab.mod.list=list(log_creat=expression(log(creat))))

## End(Not run)
```
wgt.columns = list(),
check.path = NULL,
failflow_fn = "FailFlow",
failunit_fn = "Unit",
failnowgt_fn = "NoWgt",
infusion.unit = "mcg/kg/hr",
bolus.unit = "mcg",
bol.rate.thresh = Inf,
rateunit = "mcg/hr",
ratewgtunit = "mcg/kg/hr",
weightunit = "kg",
drugname = NULL
)

Arguments

mar.path filename of MAR data (CSV, RData, RDS), or data.frame
mar.columns a named list that should specify columns in MAR data: ‘id’, ‘datetime’ and ‘dose’ are required. ‘drug’, ‘weight’, ‘given’ may also be specified. ‘datetime’ is date and time for data measurement, which can refer to a single date-time variable (datetime = ‘date_time’) or two variables holding date and time separately (e.g., datetime = c(‘Date’, ‘Time’)). ‘dose’ can also be given as a single variable or two variables. If given as a single column, the column’s values should contain dose and units such as ‘25 mcg’. If given as two column names, the dose column should come before the unit column (e.g., dose = c(‘doseamt’, ‘unit’)). ‘drug’ can provide list of acceptable drug names. If ‘drug’ is present, the ‘medchk.path’ argument should also be provided. The ‘given’ is a variable that flags whether the medication (inpatient) was given. When it is given, values should be “Given”; should be used in conjunction with the ‘medGivenReq’ argument.
medGivenReq if TRUE, values in ‘given’ column in MAR data should equal “Given”; if this is FALSE (the default), NA values are also acceptable.
flow.path filename of flow data (CSV, RData, RDS), or data.frame
flow.columns a named list that should specify columns in flow data; ‘id’, ‘datetime’, ‘finalunits’, ‘unit’, ‘rate’, ‘weight’ are required. ‘idvisit’ may also be specified. ‘datetime’ is date and time for data measurement, which can refer to a single date-time variable (datetime = ‘date_time’) or two variables holding date and time separately (e.g., datetime = c(‘Date’, ‘Time’)).
medchk.path filename containing data set (CSV, RData, RDS), or data.frame; should have the column ‘medname’ with list of acceptable drug names (e.g., brand and generic name, abbreviations) to subset drugs of interest using ‘drug’ column in MAR data. This argument can be used when MAR data contains different drugs that should be excluded.
demo.list demographic information; if available, the output from ‘run_Demo’ or a correctly formatted data.frame, which can be used to impute weight when missing
demo.columns a named list that should specify columns in demographic data; ‘id’, ‘datetime’, and ‘weight’ are required. ‘datetime’ is the date and time when the demographic data were obtained, which can refer to a single date-time variable (datetime =
`date_time`) or two variables holding date and time separately (e.g., `datetime = c('Date', 'Time')).

**missing.wgt.path**
filename containing additional weight data (CSV, RData, RDS), or data.frame. The variables in this file should be defined in the `wgt.columns` argument.

**wgt.columns**
a named list that should specify columns in additional weight data; 'id', 'date-time', and 'weight' are required. 'datetime' is date and time for weight measurement, which can refer to a single date-time variable (datetime = `date_time`) or two variables holding date and time separately (e.g., `datetime = c('Date', 'Time')).

**check.path**
path to 'check' directory, where check files are created. The default (NULL) will not produce any check files.

**failflow_fn**
filename for duplicate flow data with rate zero

**failunit_fn**
filename for MAR data with invalid unit

**failnowgt_fn**
filename for infusion data with missing weight where unit indicates weight is required

**infusion.unit**
acceptable unit for infusion data

**bolus.unit**
acceptable unit for bolus data

**bol.rate.thresh**
upper limit for bolus rate; values above this are invalid

**rateunit**
acceptable unit for hourly rate; defaults to 'mcg/hr'

**ratewgtunit**
acceptable unit for hourly rate by weight; defaults to 'mcg/kg/hr'

**weightunit**
acceptable unit for weight; defaults to 'kg'

**drugname**
drug of interest, included in filename of check files. The default (NULL) will produce filenames without drugname included.

### Details
See EHR Vignette for Structured Data.

### Value
structured data set

### Examples
```r
## Not run:
# flow data for 'Fakedrug1'
flow <- data.frame(mod_id=c(1,1,2,2,2),
                    mod_id_visit=c(46723,46723,84935,84935,84935),
                    record.date=c("7/5/2019 5:25", "7/5/2019 6:01",
                        "9/4/2020 5:32"),
                    Final.Weight=c(6.75,6.75,4.5,4.5,4.5),
                    Final.Rate=c(rep("1 mcg/kg/hr",2),
                                rep("0.5 mcg/kg/hr",3)),
```
run_MedStrII

```r
Final.Units = c("3.375", "6.5", "2.25", "2.25", "2.25")
flow[, 'Perform.Date'] <- pkdata::parse_dates(EHR:::fixDates(flow[, 'record.date']))
flow[, 'unit'] <- sub('.*[,]', '', flow[, 'Final.Rate'])
flow[, 'rate'] <- as.numeric(sub('[0-9.]+.*', '\1', flow[, 'Final.Rate']))

saveRDS(flow, 'flow.rds')

# mar data for 4 fake drugs
mar <- data.frame(mod_id=rep(1,5),
                  Date=rep("2019-07-05",5),
                  Time=c("07:12", "07:31", "08:47", "09:16", "10:22"),
                  'med:nDrug'=c("Fakedrug2", "Fakedrug1", "Fakedrug2",
                                "Fakedrug3", "Fakedrug4"),
                  'med:dosage'=c("30 mg", "0.5 mcg", "1 mg",
                                 "20 mg", "3 mcg/kg/min"),
                  'med:route'=rep("IV", 5),
                  'med:given'=rep("Given", 5),
                  check.names=FALSE)

saveRDS(mar, 'mar.rds')

# medcheck file for drug of interest ('Fakedrug1')
medcheck <- data.frame(medname="Fakedrug1", freq=4672)
write.csv(medcheck, 'medcheck.csv')

run_MedStrI(mar.path='mar.rds',
            mar.columns = list(id = 'mod_id',
                               datetime = c('Date','Time'),
                               dose = 'med:dosage',
                               drug = 'med:nDrug',
                               given = 'med:given'),
            flow.path='flow.rds',
            flow.columns = list(id = 'mod_id',
                                datetime = 'Perform.Date',
                                finalunits = 'Final.Units',
                                unit = 'unit',
                                rate = 'rate',
                                weight = 'Final.Weight'),
            medchk.path='medcheck.csv',
            check.path=tempdir(),
            drugname='fakedrug1')

## End(Not run)
```

**Description**

This module will load and modify structured e-prescription data.
run_MedStrII

Usage

run_MedStrII(file, dat.columns = list())

Arguments

file filename of prescription data (CSV, RData, RDS), or data.frame
dat.columns a named list that should specify columns in data; 'id', 'dose', 'freq', 'date', and 'str' are required. 'desc' may also be specified.

Details

See EHR Vignette for Structured Data.

Value

str data set

Examples

```r
## Not run:
erx_data <- data.frame(GRID=paste0("ID",c(1,1,2,2,2,2)),
MED_NAME=c("fakedrug","fakedrug","fakedrug",
"Brandname","fakedrug","fakedrug"),
RX_DOSE=c(1,2,1,'2 tabs',1,'1+1.5+1'),
FREQUENCY=c(rep("bid",3),"qam","bid",
"brkfst,lunch,dinner"),
ENTRY_DATE=c("2018-02-15","2018-03-14","2017-07-01",
"2017-07-01","2017-09-15","2017-11-01"),
STRENGTH_AMOUNT=c("100","100","200",
"100mg","100","100"),
DESCRIPTION=c("fakedrug 100 mg tablet","fakedrug 100 mg tablet",
"fakedrug 200 mg tablet (also known as brandname)
"Brandname 100mg tablet", "fakedrug 100 mg tablet",
"fakedrug 100 mg tablet")

write.csv(erx_data, 'erx_data.csv')

run_MedStrII('erx_data.csv', list(id = 'GRID', dose = 'RX_DOSE', freq = 'FREQUENCY',
date = 'ENTRY_DATE', str = 'STRENGTH_AMOUNT', desc = 'DESCRIPTION'))

## End(Not run)
```
**stdzDose**  
*Standardize Dose Entity*

**Description**
This function standardizes the dose entity.

**Usage**

```r
stdzDose(x)
```

**Arguments**

- `x`  
  character vector of extracted dose values

**Details**
Some dose strings may include multiple values and additional interpretation may be needed. For example ‘2-1’ likely indicates a dose of 2 followed by a dose of 1. Currently it would be converted to the average of 1.5.

**Value**
numeric vector

**Examples**

```r
stdzDose(c("one tablet", "1/2 pill", "1-3 tabs"))
```

---

**stdzDoseChange**  
*Standardize Dose Change Entity*

**Description**
This function standardizes the dose change entity.

**Usage**

```r
stdzDoseChange(x)
```

**Arguments**

- `x`  
  character vector of extracted dose change values

**Value**
character vector
**stdzDoseSchedule**

**Examples**

```r
stdzDoseChange(c('decreasing', 'dropped', 'increased'))
```

---

**Description**

This function standardizes the dose schedule entity.

**Usage**

```r
stdzDoseSchedule(x)
```

**Arguments**

- `x`: character vector of extracted dose schedule values

**Value**

character vector

**Examples**

```r
stdzDoseSchedule(c('tapered', 'weaned', 'TAPER'))
```

---

**stdzDuration**

**Standardize Duration Entity**

**Description**

This function standardizes the duration entity.

**Usage**

```r
stdzDuration(x)
```

**Arguments**

- `x`: character vector of extracted duration values

**Value**

character vector

**Examples**

```r
stdzDuration(c('1 month', 'three days', 'two-weeks'))
```
### stdzFreq

**Standardize Frequency Entity**

**Description**
This function standardizes the frequency entity.

**Usage**
```
stdzFreq(x)
```

**Arguments**
- `x` character vector of extracted frequency values

**Value**
character vector

**Examples**
```
stdzFreq(c("in the morning", "four times a day", "with meals"))
```

### stdzRoute

**Standardize Route Entity**

**Description**
This function standardizes the route entity.

**Usage**
```
stdzRoute(x)
```

**Arguments**
- `x` character vector of extracted route values

**Value**
character vector

**Examples**
```
stdzRoute(c("oral", "po", "subcut"))
```
stdzStrength

**Standardize Strength Entity**

**Description**

This function standardizes the strength entity.

**Usage**

`stdzStrength(str, freq)`

**Arguments**

- `str` character vector of extracted strength values
- `freq` character vector of extracted frequency values

**Details**

Some strength strings may include multiple values and additional interpretation may be needed. For example '2-1' likely indicates a strength of 2 followed by a strength of 1. Thus a single element may need to be standarized into two elements. This can only happen if the frequency entity is missing or in agreement ('bid' for example). See the 'addl_data' attribute of the returned vector.

**Value**

numeric vector

**Examples**

```r
stdzStrength(c('1.5', '1/2', '1/1/1'))
stdzStrength(c('1.5', '1/2', '1/1/1'), c('am', 'daily', NA))
stdzStrength(c('1.5', '1/2', '1/1/1'), FALSE)
```

tac_lab

**Example of Lab Time Data for Tacrolimus**

**Description**

An example dataset used in `processLastDose` that contains lab time data. This dataset should have one row per patient ID-date pair, and contain the time a lab was performed as a datetime variable.

**Usage**

```r
data(tac_lab, package = 'EHR')
```
Format

A data frame with 2 observations on the following variables.

- **pid** A character vector, patient ID associated with the lab value
- **date** A character vector, date associated with the lab value
- **labtime** A POSIXct vector, datetime at which the lab was performed formatted as YYYY-MM-DD HH:MM:SS

Examples

```r
data(tac_lab)
```

---

**tac_metadata**

*Example of Metadata for Tacrolimus Data*

Description

An example of the metadata needed for the `processLastDose`, `makeDose`, and `collapseDose` functions.

Usage

```r
data(tac_metadata, package = 'EHR')
```

Format

A data frame with 5 observations on the following variables.

- **filename** A character vector, filename for the clinical note
- **pid** A character vector, patient ID associated with the filename
- **date** A character vector, date associated with the filename
- **note** A character vector, note ID associated with the filename

Examples

```r
data(tac_metadata)
```
Example of Tacrolimus Output from 'parseMedExtractR'

Description

The output after running parseMedExtractR on 3 example clinical notes.

Usage

data(tac_mxr_parsed, package = 'EHR')

Format

A data frame with 7 observations on the following variables.

filename  A character vector, filename for the clinical note
drugname  A character vector, drug name extracted from the clinical note along with start and stop positions
strength   A character vector, strengths extracted from the clinical note along with start and stop positions
dose      A character vector, dose amounts extracted from the clinical note along with start and stop positions
route     A character vector, routes extracted from the clinical note along with start and stop positions
freq      A character vector, frequencies extracted from the clinical note along with start and stop positions
dosestr   A character vector, dose intakes extracted from the clinical note along with start and stop positions
dosechange A character vector, dose change keywords extracted from the clinical note along with start and stop positions
lastdose  A character vector, last dose times extracted from the clinical note along with start and stop positions

Examples

data(tac_mxr_parsed)
zeroOneTable

Make Zero One Contingency Tables

Description

Make contingency tables for many binary outcomes and a binary covariate

Usage

zeroOneTable(EXPOSURE, phenotype)

Arguments

EXPOSURE binary covariate (e.g., exposure).
phenotype binary outcome (e.g., phenotype).

Details

Generates frequency and contingency tables for many binary outcomes (e.g., large number of phenotypes) and a binary covariate (e.g., drug exposure, genotypes) more efficiently.

Value

\( t_{00} \) frequency for non-exposed group and non-case outcome.
\( t_{01} \) frequency for non-exposed group and case outcome.
\( t_{10} \) frequency for exposed group and non-case outcome.
\( t_{11} \) frequency for exposed group and case outcome.

Author(s)

Leena Choi <naturechoi@gmail.com> and Cole Beck <cole.beck@vumc.org>

Examples

```r
## full example data
data(dataPheWAS)
demo.covariates <- c('id', 'exposure', 'age', 'race', 'gender')
phenotypeList <- setdiff(colnames(dd), demo.covariates)

tablePhenotype <- matrix(NA, ncol=4, nrow=length(phenotypeList),
dimnames=list(phenotypeList, c("n.nocase.nonexp", "n.case.nonexp",
"n.nocase.exp", "n.case.exp")))
for(i in seq_along(phenotypeList)) {
    tablePhenotype[i, ] <- zeroOneTable(dd[, 'exposure'], dd[, phenotypeList[i]])
}
```
Index

* EHR
  EHR-package, 3
* PheWAS
  EHR-package, 3
* datasets
  dd, 10
  dd.baseline, 11
  dd.baseline.small, 11
  dd.small, 12
  lam_metadata, 15
  lam_mxr_parsed, 16
  tac_lab, 43
  tac_metadata, 44
  tac_mxr_parsed, 45
* process
  EHR-package, 3
  addLastDose, 4, 9, 19
  analysisPheWAS, 3, 5
  buildDose, 4, 7, 9, 19
  collapseDose, 8, 9, 15, 44
  dataTransformation, 10, 26, 35
  dd, 10
  dd.baseline, 11
  dd.baseline.small, 11
  dd.small, 12
  EHR (EHR-package), 3
  EHR-package, 3
  extractMed, 4, 12, 23
  freqNum, 13
  idCrosswalk, 14, 24, 25
  lam_metadata, 15
  lam_mxr_parsed, 16
  Logistf, 17
  makeDose, 9, 15, 18, 44
  medExtractR, 4, 12, 13, 23
  parseCLAMP, 7, 19
  parseMedEx, 7, 20
  parseMedExtractR, 7, 16, 21, 45
  parseMedXN, 7, 22
  processLastDose, 4, 15, 23, 43, 44
  pullFakeId, 24
  pullRealId, 25
  read.csv, 26
  readTransform, 26
  run_Build_PK_IV, 27
  run_Build_PK_Oral, 29
  run_Demo, 28, 31
  run_DrugLevel, 27, 32
  run_Labs, 28, 34
  run_MedStrI, 27, 35
  run_MedStrII, 38
  stdzDose, 40
  stdzDoseChange, 40
  stdzDoseSchedule, 41
  stdzDuration, 41
  stdzFreq, 42
  stdzRoute, 42
  stdzStrength, 43
  tac_lab, 43
  tac_metadata, 44
  tac_mxr_parsed, 45
  zeroOneTable, 3, 46