# Package 'LW1949'

# November 5, 2015

Title An Automated Approach to Evaluating Dose-Effect Experiments

Following Litchfield and Wilcoxon (1949)

Version 0.0.0.9008
<b>Description</b> LW1949 takes the manual approach to evaluating dose-effect experiments (Litchfield and Wilcoxon 1949) and automates the steps so that the computer can do the work.
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# Description

assessfit

Assess the fit of a dose-response curve using the chi-squared statistic. The curve is described by the intercept and slope of a straight line in the log dose vs. probit effect scale.

Assess Fit of Dose-Response Curve

# Usage

```
assessfit(params, DEdata, fit = gamtable1(), simple = TRUE)
```

#### **Arguments**

params	A numeric vector of length two, with the estimated intercept and slope of the dose-effect relation on the log10 and probit scale. These parameters define the dose-response curve.
DEdata	A data frame of dose-effect data (typically, the output from dataprep) containing at least these four variables: dose, ntot, pfx, fxcateg.
fit	A model object that can be used to predict the corrected values (as proportions) from distexpprop5, the distance between the expected values (as proportions) and 0.5, default gamtable1().
simple	A logical scalar indicating if the output should be restricted to just the P value, default TRUE.

#### **Details**

This function is used to find the dose-response curve that minimizes the chi-squared statistic measuring the distance between the observed and expected values of the response (the proportion affected). Following Litchfield and Wilcoxon (1949, steps B1 and B2), records with expected effects < 0.01% or > 99.99% are deleted, and other expected effects are "corrected" using the correctval function.

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#### Value

If simple=FALSE, a list of length two. The first element, chi, is a numeric vector of length three: chistat, chi-squared statistic; df, degrees of freedom; and pval, P value. The second element, contrib, is a matrix of three numeric vectors the same length as obsn: exp, expected effects; obscorr, observed effects corrected; and contrib, contributions to the chi-squared.

If simple=TRUE, a numeric scalar, the chi-squared statistic (see details).

#### References

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

#### See Also

LWchi2 and chisq.test.

#### **Examples**

```
conc <- c(0.0625, 0.125, 0.25, 0.5, 1)

numtested <- rep(8, 5)

nalive <- c(1, 4, 4, 7, 8)

mydat <- dataprep(dose=conc, ntot=numtested, nfx=nalive)

gamfit <- gamtable1()

assessfit(log10(c(0.125, 0.5)), mydat, simple=FALSE)
```

coefprobit

Calculate the Coefficients of a Probit Regression Fit

### **Description**

Calculate the coefficients from a fitted probit regression model with confidence intervals.

# Usage

```
coefprobit(pfit, alpha = 0.05)
```

# **Arguments**

pfit An object of class glm representing a probit regression fit to dose-effect data,

typically the result of a call to fitprobit. Dose should be the only independent

variable in the model.

alpha A numeric scalar, the significance level used to generate 100\*(1 - alpha)% con-

fidence limits, default 0.05.

### Value

A numeric vector of length six, the intercept and slope of the dose-response curve, each with 100\*(1 - alpha)% confidence limits.

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#### **Examples**

```
toxdat <- data.frame(
  dose=c(0.05, 0.0625, 0.125, 0.25, 0.5, 1),
  ntot=rep(8, 6),
    nfx = c(0, 1, 4, 4, 6, 8))
myfit <- fitprobit(toxdat)
coefprobit(myfit)</pre>
```

constrain

Constrain Data to a Specified Range

# Description

Constrain data to a specified range, assigning values from the specified range to those outside the range, typically for graphing purposes.

# Usage

```
constrain(x, xrange)
```

# **Arguments**

x A numeric vector of values to constrain.

xrange A numeric vector of length two specifying the constraints, the minimum and

maximum value for x.

#### Value

A numeric vector, the same length as x, in which the minimum constraint is assigned to values of x less than the minimum, and the maximum constraint is assigned to values of x greater than the maximum.

# **Examples**

```
constrain(1:20, c(3, 19))
```

correctval

Predict the Corrected Proportional Effect

# Description

Given an expected proportional effect, calculate the "corrected" proportional effect using a model fit of Litchfield and Wilcoxon's (1949) Table 1.

# Usage

```
correctval(val, fit)
```

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# **Arguments**

fit A model object to be used to predict the "corrected" effects (as proportions	val	A numeric vector of expected effects (as proportions).
	fit	A model object to be used to predict the "corrected" effects (as proportions) from distexpprop5, the distance between the expected effects (as proportions) and 0.5. Typically the output from gamtable 1()

# Value

A numeric vector of corrected effects (as proportions), the same length as val.

#### References

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

# **Examples**

```
gamfit <- gamtable1()
correctval(c(0.37, 0.5, 0.63), gamfit)</pre>
```

dataprep

Prepare Data

# **Description**

Prepare dose-effect data for evaluation.

# Usage

```
dataprep(dose, ntot, nfx)
```

# **Arguments**

dose	A numeric vector of unique, chemical concentrations (see Details).
ntot	A numeric vector of the number of individuals that were tested at each dose.
nfx	A numeric vector of the number of individuals that were affected at each dose.

# Details

The input data are expected to be summarized by dose. If duplicate doses are provided, an error will be thrown.

#### Value

A data frame with eight columns (ordered by dose and proportion affected), seven numeric vectors and one logical vector:

- dose = chemical concentrations.
- ntot = the number of individuals that were tested at each dose.
- nfx = the number of individuals that were affected at each dose.

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- rec = the record number corresponding to the input vectors dose, ntot, nfx.
- pfx = the proportion of individuals that were affected at each dose.
- log10dose = log transformed dose, log10(dose).
- bitpfx = probit transformed proportional affected, probit(pfx).
- fxcateg = effects category: 0 for none affected, 100 for all affected, and 50 for other proportions affected.
- LWkeep = logical vector identifying records to keep for Litchfield and Wilcoxon (1949, step A1) method.

#### References

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

#### **Examples**

```
conc <- c(0.0625, 0.125, 0.25, 0.5, 1)
numtested <- rep(8, 5)
nalive <- c(1, 4, 4, 7, 8)
dataprep(dose=conc, ntot=numtested, nfx=nalive)</pre>
```

estimable

Determine if a Dose-Effect Relation is Estimable

# **Description**

Determine if a dose-effect relation is estimable based on available data.

# Usage

```
estimable(DEdata)
```

# Arguments

DEdata

A data frame of dose-effect data (typically, the output from dataprep) containing at least two variables: dose, a numeric vector of chemical concentrations, and pfx, a numeric vector of proportional effects at each dose (see Details).

# Details

A dose-effect relation is defined to be estimable if and only if there are at least two test records and there is some (non-zero) variability in both the doses and the proportional effects. The input data are expected to be summarized by dose. If duplicate doses are provided, an error will be thrown.

#### Value

A logical scalar indicating if a dose-effect relation is estimable.

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#### **Examples**

```
conc <- c(0.0625, 0.125, 0.25, 0.5, 1)
numtested <- rep(8, 5)</pre>
nalive <- c(1, 4, 4, 7, 8)
mydat <- dataprep(dose=conc, ntot=numtested, nfx=nalive)</pre>
estimable(mydat)
nalive2 <- rep(4, 5)
mydat2 <- dataprep(dose=conc, ntot=numtested, nfx=nalive2)</pre>
estimable(mydat2)
```

fill

Fill in Missing Values

#### **Description**

Fill in missing values in a vector, using the last recorded value.

# Usage

```
fill(x, resetWhen = rep(FALSE, length(x)))
```

# **Arguments** Х

A vector, can be character, numeric, or logical.

resetWhen

A logical vector, the same length as x, indicating elements that should not be filled in.

#### **Details**

Similar to na.locf in the zoo package, but works for "" in character vectors as well.

# Value

A vector the same length as x, with all NAs or ""s replaced by the last value for the vector. Note that and missing values at the beginning of the vector will not be replaced.

# References

This is a copy of the fill function from the [jvamisc] package.

```
numvec <- c(NA, 1:5, NA, NA, NA, 10:12, NA)
newgroup \leftarrow c(1, 0, 0, 0, 1, 0, 0, 0, 1, 0, 0, 0)
fill(numvec)
fill(numvec, newgroup)
charvec <- c("", letters[1:5], "", "", "", letters[10:12], "")</pre>
fill(charvec)
```

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Determine Linear Regression Coefficients from Dose-Effect Data

#### **Description**

Determine coefficients (intercept and slope) from dose-effect data using simple linear regression on the log10 dose vs. probit effect scale.

#### Usage

```
fitlinear(DEdata, constr = c(1e-04, 0.9999))
```

# **Arguments**

DEdata A data frame of dose-effect data (typically, the output from dataprep) contain-

ing at least three variables: log10dose, bitpfx, and LWkeep.

constr A numeric vector of length two, indicating the constraints (see constrain) ap-

plied to the proportional effects, default c(0.0001, 0.9999).

#### Value

A numeric vector of length two, the estimated intercept and slope.

# **Examples**

```
conc <- c(0.0625, 0.125, 0.25, 0.5, 1)

numtested <- rep(8, 5)

nalive <- c(1, 4, 4, 7, 8)

mydat <- dataprep(dose=conc, ntot=numtested, nfx=nalive)

fitlinear(mydat)
```

fitLWauto

Best Fit Using Litchfield and Wilcoxon Evaluation of Dose-Effect Experiments

#### **Description**

Use optimization to describe the best fitting line to a dose-effect experiment following the methods of Litchfield and Wilcoxon (1949).

# Usage

```
fitLWauto(DEdata)
```

# **Arguments**

DEdata

A data frame of dose-effect data (typically, the output from dataprep) containing at least eight variables: dose, ntot, nfx, pfx, log10dose, bitpfx, fxcateg, and LWkeep (see Details).

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#### **Details**

The input data are expected to be summarized by dose. If duplicate doses are provided, an error will be thrown.

#### Value

A numeric vector of length two, the estimated intercept and slope of the dose-response curve on the log10-probit scale,

#### References

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

# **Examples**

```
dose <- c(0.0625, 0.125, 0.25, 0.5, 1)

ntested <- rep(8, 5)

nalive <- c(1, 4, 4, 7, 8)

mydat <- dataprep(dose=dose, ntot=ntested, nfx=nalive)

mydat

fitLWauto(mydat)
```

fitprobit

Fit a Probit Regression to Dose-Effect Data

# **Description**

Fit a probit regression to dose-effect data, using the log10 of the dose as the response.

#### Usage

```
fitprobit(dat)
```

# **Arguments**

dat

A data frame of toxicity data, including at least three variables: dose (the concentration of the tested chemical), ntot (the number of individuals tested), and nfx (the number of affected individuals).

#### **Details**

Only those rows with dose > 0, ntot > 0, and nfx >= 0 are used in fitting the model.

### Value

A an object of class glm.

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#### **Examples**

```
toxdat <- data.frame(
  dose=c(0.05, 0.0625, 0.125, 0.25, 0.5, 1),
  ntot=rep(8, 6),
  nfx = c(0, 1, 4, 4, 6, 8))
fitprobit(toxdat)</pre>
```

fxcat

Define Effect Category

# Description

Define three effect categories, 0 for none affected, 100 for all affected, and 50 for other proportions affected.

# Usage

fxcat(dat)

#### **Arguments**

dat

A data frame of toxicity data, including at least two variables: ntot (the number of individuals tested) and nfx (the number of affected individuals).

#### Value

An integer vector the same length as prob with categories of 0, 50, or 100.

# **Examples**

```
toxdat <- data.frame(
  dose=c(0.0625, 0.125, 0.25, 0.5),
  ntot=rep(8, 4),
  nfx = c(0, 4, 6, 8))
cbind(toxdat, fxcat(toxdat))</pre>
```

gamtable1

Fit a smooth GAM to Table 1 of Litchfield and Wilcoxon (1949)

#### **Description**

Fit a smooth GAM function to replace looking up values in Table 1 of Litchfield and Wilcoxon (1949).

# Usage

```
gamtable1()
```

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#### **Details**

Note that for an expected value of 37 Table 1 gives a corrected value of 9.4, but for an expected value of 63 it gives a corrected value of 90.5. To ensure that both values add to 100, I used corrected values of 9.45 and 90.55. The expected and corrected values from Table 1 are then used to build a GAM model, which is used as input to the correctval function.

#### Value

A gamObject that can be used to predict the corrected values (as proportions) from distexpprop5, the distance bewteen the expected values (as proportions) and 0.5

#### References

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

#### **Examples**

```
fit <- gamtable1()
summary(fit)
plot(fit)</pre>
```

handLW

Best Fit Using Estimated Expected Effects

# **Description**

Use simple linear regression to describe a hand draw line fit to a dose-effect experiment.

# Usage

```
handLW(handDose, handPct)
```

# **Arguments**

handDose A numeric vector of doses for which expected effects are estimated from a hand

drawn line, must have at least two unique values.

handPct A numeric vector of the expected percent affected corresponding to (and the

same length as) handDose.

# Value

A numeric vector of length two, the estimated intercept and slope of the dose-response curve on the log 10-probit scale,

```
d \leftarrow c(0.0625, 0.125, 0.25, 0.5, 1)

p \leftarrow c(9.5, 34, 67, 90.5, 98.6)

handLW(handDose=d, handPct=p)
```

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invprobit

Convert Probit Scale to Proportions

# **Description**

Convert values on the probit scale to their proportions on the 0 to 1 scale.

# Usage

```
invprobit(quan)
```

# **Arguments**

quan

A numeric vector of probit quantiles.

# **Details**

Simply calls pnorm(quan).

# Value

A numeric vector of proportions the same length as quan.

# **Examples**

```
invprobit(c(-3, -1, 0, 1, 3))
```

keeponly

Eliminate Consecutive Extreme Values

# Description

Generate the index for eliminating values beyond a given maximum number of consecutive extremes allowed.

# Usage

```
keeponly(x, extremes = c(0, 100), nconsec = 2)
```

# Arguments

x A numeric vector, with no missing values.

extremes A numeric vector of length two giving the boundary limits for x, default c(0,

100).

nconsec An integer scalar, the maximum number of consecutive extreme values allowed,

default 2.

# Value

A logical vector for selecting all elements of orderedx without exceeding nconsec consecutive extreme values.

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### **Examples**

```
vec <- c(0, 0, 0, 4, 4, 4, 100, 100, 100, 100)
vec[keeponly(vec)]
# the original vector need not be ordered
vec <- c(100, 4, 100, 4, 0, 100, 0, 4, 0, 100)
keeponly(vec)</pre>
```

LW1949

Automated Litchfield and Wilcoxon (1949) Evaluation of Dose-Effect Experiments

# **Description**

**LW1949** is an automated approach to Litchfield and Wilcoxon's (1949) evaluation of dose-effect experiments. **LW1949** was first introduced by Adams et al. (*in preparation*).

#### **Details**

An example of how to use the functions in **LW1949** is given in this vignette [link]. Use dataprep to create a data frame with the results of a dose-effect experiment. Use fitLWauto and LWestimateto fit dose-effect relations. And use plotDELP and plotDEto plot the results.

*U.S. Geological Survey* (USGS) Computer Program **LW1949** version 0.0.0.9008. Written by Jean V. Adams, USGS - Great Lakes Science Center glsc.usgs.gov, Ann Arbor, Michigan, USA. Written in programming language R (R Core Team, 2015, www.R-project.org), version 3.2.2 (2015-08-14). Run on a PC with Intel(R) Core(TM) I7-4600m CPU, 2.90 GHz processor, 16.0 GB RAM, and Microsoft Windows 7 Enterprise operating system 2009 Service Pack 1. Source code is available from Jean V. Adams on GitHub, github.com/JVAdams/LW1949, *jvadams* (at) usgs (dot) gov.

*Disclaimer:* Although this program has been used by the USGS, no warranty, expressed or implied, is made by the USGS or the United States Government as to the accuracy and functioning of the program and related program material nor shall the fact of distribution constitute any such warranty, and no responsibility is assumed by the USGS in connection therewith.

#### References

Adams, JV, KS Slaght, and MA Boogaard. *In preparation*. An automated approach to Litchfield and Wilcoxon's evaluation of dose-effect experiments.

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

LWchi2

Chi-Squared Statistic

# **Description**

Calculate the chi-squared statistic from observed and expected counts using the Litchfield and Wilcoxon (1949) approach.

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#### **Usage**

```
LWchi2(obsn, expn, totn)
```

#### **Arguments**

obsn A numeric vector of observed counts.

expn A numeric vector of expected counts, the same length as obsn.
totn A numeric vector of total counts possible, the same length as obsn.

#### **Details**

The denominator of Litchfield and Wilcoxon's (1949) chi-squared estimate is the minimum of the expn and (totn - expn) following their Nomograph No. 1. This ensures that the same chi-squared value is calculated regardless of which proportion is reported (e.g., affected vs. not affected).

#### Value

A list of length two. The first element is a numeric vector of length three: chistat, chi-squared statistic; df, degrees of freedom; and pval, P value. The second element is a numeric vector the same length as obsn, containing **total** contributions to the chi-squared. To get the **individual** contributions to the chi-squared as reported in Litchfield and Wilcoxon (1949), divide by totn.

#### References

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

### See Also

```
chisq.test.
```

#### **Examples**

```
LWchi2(c(10, 8, 3), c(7, 7, 7), c(12, 12, 12))
```

LWestimate

Generate Litchfield and Wilcoxon Estimates

#### **Description**

Generate Litchfield and Wilcoxon's (1949) estimates in the evaluation of dose-effect experiments.

# Usage

LWestimate(estParams, DEdata)

#### **Arguments**

estParams A numeric vector of length 2, the estimated parameters (intercept and slope) of

a straight line on the log10-probit scale, typically output from fitLWauto.

DEdata A data frame of dose-effect data (typically, the output from dataprep) contain-

ing at least eight variables: dose, ntot, nfx, pfx, log10dose, bitpfx, fxcateg, and

LWkeep (see Details).

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#### **Details**

The input data are expected to be summarized by dose. If duplicate doses are provided, an error will be thrown.

#### Value

A list of length three:

- chi = the chi-squared statistic with associated P value and degrees of freedom,
- params = the estimated intercept and slope of the dose-response curve on the log10 probit scale,
- LWest = the Litchfield Wilcoxon estimates of ED50 with 95% confidence intervals and the number of records with partial effects (npartfx) as well as other metrics used in their step-by-step approach (ED16, ED84, S with 95% confidence intervals, N', and fED50).

#### References

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

# **Examples**

```
dose <- c(0.0625, 0.125, 0.25, 0.5, 1)
ntested <- rep(8, 5)
nalive <- c(1, 4, 4, 7, 8)
mydat <- dataprep(dose=dose, ntot=ntested, nfx=nalive)
mydat
intslope <- fitLWauto(mydat)
LWestimate(intslope, mydat)</pre>
```

LWP

User Friendly Evaluation of Dose-Effect Experiments using Litchfield-Wilcoxon and Probit Methods

# **Description**

User friendly evaluation of dose-effect experiments using automated Litchfield Wilcoxon (1949) and probit estimation methods. This function has been tailored for non-R users with input data set up in a particular way (see Details).

# Usage

```
LWP(rawfile = NULL, descroolz = 1:4, saveplots = TRUE,
   showplots = FALSE, saveresults = TRUE, showresults = TRUE,
   returnresults = FALSE)
```

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# **Arguments**

rawfile	A character scalar specifying the path of the input data as a csv file. If NULL, default, the user will be prompted to browse to a file using a menu.
descrcolz	A numeric vector, the column numbers to use as the description of the test, default 1:4.
saveplots	A logical scalar indicating if plots should be saved to a pdf file, default TRUE.
showplots	A logical scalar indicating if plots should be shown on screen, default FALSE.
saveresults	A logical scalar indicating if results should be saved to a csv file, default TRUE. The csv file is given the same name (plus the suffix "Smry") and is placed in the same directory as the input file.
showresults	A logical scalar indicating if results should be printed to the console, default TRUE. These results include the chi-squared statistic, degrees of freedom, and p-value for the Litchfield Wilcoxon method.
returnresults	A logical scalar indicating if results should be returned by the function, default FALSE.

#### **Details**

The input data must include at least these seven columns, with these names in the header row:

- Test ID = A character or numeric vector, the unique identifier for each test
- Source = A character vector, the source of the chemical
- Batch = A character or numeric vector, the batch of the chemical
- Species = A character vector, the species tested
- TFM Conc. (mg/L) = A numeric vector, the concentration of TFM in mg/L
- No. Tested = A numeric vector, the number of animals tested
- No. Dead = A numeric vector, the number of animals dead

The input data are expected to be summarized by dose. If duplicate doses are provided, an error will be thrown.

#### Value

If returnresults=TRUE, a data frame with 11 rows per test and 2 more columns than the input data. Three columns from the input data are not included (TFM Conc. (mg/L), No. Tested, and No. Dead). Five columns are added: the parameter (param), the method used (method), the estimate (estimate), and the 95% confidence interval of the estimate (lower95ci and upper95ci)

#### References

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

```
## Not run:
LWP()
## End(Not run)
```

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plotDE	Plot Dose-Effect Experiments

# **Description**

Plot dose-effect experiments on the arithmetic scale.

# Usage

```
plotDE(DEdata, xlab = "Dose", ylab = "Affected (%)",
   xlim = range(DEdata$dose, na.rm = TRUE), ylim = c(0, 100), ref = c(0,
   50, 100), ...)
```

# Arguments

DEdata	A data frame of dose-effect data (typically, the output from dataprep) containing at least five variables: dose, pfx, log10dose, bitpfx, fxcateg.
xlab	A character scalar, the title for the dose (x) axis, default "Dose".
ylab	A character scalar, the title for the affected (y) axis, default "Affected (%)".
xlim	A numeric vector of length two giving the x coordinate range for dose, default range(DEdata\$dose, na.rm=TRUE).
ylim	A numeric vector of length two giving the y coordinate range for affected $(\%)$ , default $c(0.1, 99.9)$ . Observed effects beyond this range will be plotted at the limits of this range using an open symbol.
ref	A numeric vector specifying horizontal reference lines to be added to the plot, default $c(0, 50, 100)$ .
	Additional arguments to plot.

# See Also

```
predLines, plotDELP, predLinesLP
```

```
dose <- c(0.0625, 0.125, 0.25, 0.5, 1)
ntested <- rep(8, 5)
nalive <- c(1, 4, 4, 7, 8)
mydat <- dataprep(dose=dose, ntot=ntested, nfx=nalive)
plotDE(mydat)
```

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Plot Dose-Effect Experiments

# **Description**

Plot dose-effect experiments on the log10-probit scale.

# Usage

```
plotDELP(DEdata, xlab = "Dose", ylab = "Affected (%)",
    xlim = range(DEdata$dose[DEdata$dose > 0], na.rm = TRUE), ylim = c(0.1,
    99.9), ...)
```

### **Arguments**

DEdata	A data frame of dose-effect data (typically, the output from dataprep) containing at least five variables: dose, pfx, log10dose, bitpfx, fxcateg.
xlab	A character scalar, the title for the dose (x) axis, default "Dose".
ylab	A character scalar, the title for the affected (y) axis, default "Affected (%)".
xlim	A numeric vector of length two giving the x coordinate range for dose, default range(DEdata\$dose, na.rm=TRUE).
ylim	A numeric vector of length two giving the y coordinate range for affected (%), default c(0.1, 99.9). Observed effects beyond this range will be plotted at the limits of this range using an open symbol.
	Additional arguments to plot.

# See Also

```
predLinesLP, plotDE, predLines
```

# Examples

```
dose <- c(0.0625, 0.125, 0.25, 0.5, 1)

ntested <- rep(8, 5)

nalive <- c(1, 4, 4, 7, 8)

mydat <- dataprep(dose=dose, ntot=ntested, nfx=nalive)

plotDELP(mydat)
```

predlinear

Determine the Effective Dose from a Linear Regression Fit

# **Description**

Determine the effective dose for a specified percent effect from the intercept and slope of a linear regression.

# Usage

```
predlinear(pct, LWmod, simple = FALSE)
```

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#### **Arguments**

pct A numeric vector of effects (in percents) for which to estimate the effective

dose(s).

LWmod If simple=TRUE, a numeric vector of length two giving the intercept and slope of

the linear relation between the dose (x, the concentration of the applied chemical on the  $\log 10$  scale), and the proportion of affected individuals (y, on the probit scale, with 0s converted to 0.1% and 1s converted to 99.9%). If simple=FALSE, a list with the results of fitting a Litchfield and Wilcoxon model to dose-effect

data, the output from LWestimate.

simple A logical scalar indicating whether to carry out a simple estimation of effective

doses from the intercept and slope (TRUE), or an estimation of effective doses with confidence intervals from the Litchfield and Wilcoxon model (default,

FALSE).

#### **Details**

Follows methods outlined in Litchfield and Wilcoxon (1949). Specifically, for the 95% confidence intervals, see page 105, and equation 13 in the Appendix (corresponding to Nomograph 4).

#### Value

A numeric vector the same length as pct giving the estimated dose at the specified percent effect.

If simple=TRUE, a numeric vector the same length as pct with the estimated effective doses. If simple=FALSE, an n\*4 numeric matrix with the given effects (pct), the effective doses (ED), and Litchfield and Wilcoxon's (1949) 95% confidence intervals for the effective doses (lower and upper). The number of rows of the matrix, n, is the length of pct.

#### References

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

#### **Examples**

```
predlinear(c(16, 50, 84, 99.9), c(1.700875, 2.199559), simple=TRUE)

dose <- c(0.0625, 0.125, 0.25, 0.5, 1)

ntested <- rep(8, 5)

nalive <- c(1, 4, 4, 7, 8)

mydat <- dataprep(dose=dose, ntot=ntested, nfx=nalive)
fLW <- LWestimate(fitLWauto(mydat), mydat)
predlinear(c(25, 50, 99.9), fLW)</pre>
```

predLines

Add Litchfield and Wilcoxon Predictions to a Plot

# Description

Add predictions from a Litchfield and Wilcoxon model fit to a plot of the results of a dose-effect experiment on the arithmetic scale.

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#### Usage

```
predLines(fit)
```

# Arguments

fit

A list of length three containing the result of a Litchfield and Wilcoxon model fit, typically the output from LWestimate.

#### Value

A solid fitted line is added to the plot. Dashed lines are added to the plot representing the **horizontal** 95 for the predicted dose to elicit a given percent affected.

#### See Also

```
plotDE, plotDELP, predLinesLP
```

# **Examples**

```
dose <- c(0.0625, 0.125, 0.25, 0.5, 1)
ntested <- rep(8, 5)
nalive <- c(1, 4, 4, 7, 8)
mydat <- dataprep(dose=dose, ntot=ntested, nfx=nalive)
plotDE(mydat)
myfit <- LWestimate(fitLWauto(mydat), mydat)
predLines(myfit)</pre>
```

predLinesLP

Add Litchfield and Wilcoxon Predictions to a Plot

# Description

Add predictions from a Litchfield and Wilcoxon model fit to a plot of the results of a dose-effect experiment on the log10-probit scale.

# Usage

```
predLinesLP(fit)
```

# **Arguments**

fit

A list of length three containing the result of a Litchfield and Wilcoxon model fit, typically the output from LWestimate.

# Value

A solid fitted line is added to the plot. Dashed lines are added to the plot representing the **horizontal** 95 for the predicted dose to elicit a given percent affected.

#### See Also

```
plotDELP, plotDE, predLines
```

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#### **Examples**

```
dose <- c(0.0625, 0.125, 0.25, 0.5, 1)

ntested <- rep(8, 5)

nalive <- c(1, 4, 4, 7, 8)

mydat <- dataprep(dose=dose, ntot=ntested, nfx=nalive)

plotDELP(mydat)

myfit <- LWestimate(fitLWauto(mydat), mydat)

predLinesLP(myfit)
```

predprobit

Determine the Effective Dose from a Probit Regression Fit

# **Description**

Determine the effective dose for a specified percent effect from a fitted probit regression model.

# Usage

```
predprobit(pct, pfit, alpha = 0.05, logbase = 10)
```

# **Arguments**

pct	A numeric scalar of the effect (as a percent) for which to estimate the effective dose.
pfit	An object of class glm representing a probit regression fit to dose-effect data, typically the result of a call to fitprobit.
alpha	A numeric scalar, the significance level used to generate $100*(1 - alpha)\%$ confidence limits, default $0.05$ .
logbase	A numeric or logical scalar, the base of the log transformation used for dose in pfit, default 10. Use logbase=FALSE, if the dose was not log transformed.

# Value

A numeric vector of length three, the effective dose and the lower and upper 100\*(1 - alpha)% confidence limits.

```
toxdat <- data.frame(
  dose=c(0.05, 0.0625, 0.125, 0.25, 0.5, 1),
  ntot=rep(8, 6),
  nfx = c(0, 1, 4, 4, 6, 8))
myfit <- fitprobit(toxdat)
predprobit(50, myfit)</pre>
```

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nre	tt.	log

Pretty Breakpoints on Log Scale

#### **Description**

Compute a sequence of "round" values which cover the range of x on the log scale.

# Usage

```
prettylog(x, lead = c(1, 5), extra = 5)
```

# **Arguments**

x A numeric vector.

lead An integer vector giving the desired lead digits of pretty values on the log scale,

default c(1, 5).

extra An integer scalar giving the desired number of additional non-log scale values

to include, default 5.

#### Value

A numeric vector of pretty values covering the range of x on the log scale.

# References

This is a copy of the prettylog function from the [jvamisc] package.

# **Examples**

```
vals <- rlnorm(100, 6)
summary(vals)
prettylog(vals, 1, 0)
prettylog(vals, 1)
prettylog(vals, c(1, 2, 5))</pre>
```

probit

Convert Proportions to the Probit Scale

# **Description**

Convert proportions to the probit scale.

### Usage

```
probit(prob)
```

# **Arguments**

prob

A numeric vector of proportions.

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#### **Details**

Simply calls qnorm(prob).

#### Value

A numeric vector the same length as prob with quantiles on the probit scale.

# **Examples**

```
probit(c(0.001, 0.01, 0.1, 0.5, 0.9, 0.99, 0.999))
```

relPotency

Relative Potency of Two Toxins

# **Description**

Estimate of relative potency of two toxins using Litchfield and Wilcoxon's (1949) approach to evaluating dose-effect experiments.

### Usage

```
relPotency(ED50nS1, ED50nS2, vec = FALSE)
```

# Arguments

ED50nS1	Either the list output from LWestimate (vec = FALSE) or a numeric vector of length four (vec = TRUE) with the estimated ED50, fED50, S, and fS from a Litchfield and Wilcoxon fit to dose-effect data for the first toxin.
ED50nS2	Either the list output from LWestimate (vec = FALSE) or a numeric vector of length four (vec = TRUE) with the estimated ED50, fED50, S, and fS from a Litchfield and Wilcoxon fit to dose-effect data for the second toxin.
vec	A logical scalar indicating whether the inputs ED50nS1 and ED50nS2 are both numeric vectors (TRUE) or both lists (FALSE, the default).

# **Details**

The ratios reported (both for slope and potency) have the first toxin in the numerator and the second toxin in the denominator, but the test results (both for parallelism and relative potency) are based on the ratios of the larger values over the smaller values.

No relative potency is estimated if the two dose-effect curves differ significantly from parallelism (with 95% confidence).

#### Value

A list with two elements, SR with three elements:

- r = a numeric vector of length three with the estimated slope ratio with 95% confidence limits,
- f = a numeric scalar with the f of the slope ratio, and
- parallel = a logical scalar indicating whether the two curves differ significantly from parallelism (FALSE).

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and PR with one (just difPotency if parallel=FALSE) or three (if parallel=TRUE) elements:

- r = a numeric vector of length three with the estimated potency ratio with 95% confidence limits,
- f = a numeric scalar with the f of the potency ratio, and
- difPotency = a logical scalar indicating whether the two toxins differ significantly in potency (FALSE).

#### References

Litchfield, JT Jr. and F Wilcoxon. 1949. A simplified method of evaluating dose-effect experiments. Journal of Pharmacology and Experimental Therapeutics 96(2):99-113. [link].

```
# Example starting from raw tox data
dose <- c(0.0625, 0.125, 0.25, 0.5, 1)
ntested <- rep(8, 5)
nalive1 <- c(1, 4, 4, 7, 8)
mydat1 <- dataprep(dose=dose, ntot=ntested, nfx=nalive1)
nalive2 <- c(0, 1, 2, 6, 6)
mydat2 <- dataprep(dose=dose, ntot=ntested, nfx=nalive2)
fit1 <- LWestimate(fitLWauto(mydat1), mydat1)
fit2 <- LWestimate(fitLWauto(mydat2), mydat2)
relPotency(fit1, fit2)

# Example from Litchfield and Wilcoxon (1949)
# comparing Tagathen and Pyribenzamine
relPotency(c(0.18, 1.72, 2.20, 1.60), c(0.60, 1.60, 2.34, 1.57), vec=TRUE)
# Example in which curves differ significantly from parallelism.
relPotency(c(0.18, 1.72, 2.20, 1.60), c(0.60, 1.60, 4.34, 1.57), vec=TRUE)</pre>
```

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