# Package 'nlstools'

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confint2 L.minor michaelisdata michaelismodels nlsBoot nlsBootPredict						

2 confint2

Index																															19
	O2K	•	•	•	•	 •	•	•	•	•	•	 •	•	•	•	•	 	•	٠	•	•	•	•	•		•	•	•	•	•	18
	nlstools-defunct																														
	nlstools																														
	nlsResiduals																 														14
	nlsJack																 														12
	nlsContourRSS																														
	nlsConfRegions																 														9

confint2

Confidence intervals in nonlinear regression

# Description

Produces confidence intervals for the parameters in nonlinear regression model fit. The intervals can either be based large sample results or on profiling.

# Usage

```
confint2(object, parm, level = 0.95, method = c("asymptotic", "profile"), ...)
```

# Arguments

object	object of class nls.
parm	a vector character strings with names of the parameter for which to calculate confidence intervals (by default all parameters).
level	the confidence level required.
method	method to be used: "asympotic" for large sample and "profile" for profiling approach.
	additional argument(s) to pass on the method doing the profiling.

## **Details**

The profiling used is the method confint.nls.

## Value

A matrix with columns giving lower and upper confidence limits for each parameter.

## Author(s)

Christian Ritz

```
L.minor.m1 <- nls(rate ~ Vm*conc/(K+conc), data = L.minor, start = list(K=20, Vm=120))
confint2(L.minor.m1)
confint2(L.minor.m1, "K")</pre>
```

L.minor 3

L.minor

Enzyme kinetics

# Description

Enzyme kinetics

## Usage

```
data(L.minor)
```

#### **Format**

A data frame with 8 observations on the following 2 variables.

```
conc a numeric vector rate a numeric vector
```

#### **Source**

Cedergreen, N. and Madsen, T. V. (2002) Nitrogen uptake by the floating macrophyte *Lemna minor*, *New Phytologist*, **155**, 285–292.

michaelisdata

Michaelis Menten data sets

## **Description**

Michaelis Menten data sets

# Usage

```
data(vmkm)
data(vmkmki)
```

#### **Format**

```
vmkm is a data frame with 2 columns (S: concentration of substrat, v: reaction rate) vmkmki is a data frame with 3 columns (S: concentration of substrat, I: concentration of inhibitor, v: reaction rate)
```

## Source

These datasets were provided by the French research unit INRA UMR1233.

4 michaelismodels

## **Examples**

data(vmkm)
data(vmkmki)
plot(vmkm)
plot(vmkmki)

michaelismodels

Michaelis-Menten model and derived equations to model competitive and non-competitive inhibition

## Description

Formula of Michaelis-Menten model commonly used to describe enzyme kinetics, and derived formulas taking into account the effect of a competitive or a non-competitive inhibitor

## Usage

michaelis
compet\_mich
non\_compet\_mich

#### **Details**

These models describe the evolution of the reaction rate (v) as a function of the concentration of substrate (S) and the concentration of inhibitor (I) for compet\_mich and non\_compet\_mich.

michaelis is the classical Michaelis-Menten model (Dixon, 1979) with two parameters (Km, V max):

$$v = \frac{S}{S + K_m} V_{max}$$

compet\_mich is the Michaelis-Menten derived model with three parameters (Km, Vmax, Ki), describing a competitive inhibition :

$$v = \frac{S}{S + K_m(1 + \frac{I}{K_i})} V_{max}$$

non\_compet\_mich is the Michaelis-Menten derived model with three parameters (Km, Vmax, Ki), describing a non-competitive inhibition :

$$v = \frac{S}{(S + K_m)(1 + \frac{I}{Ki})} V_{max}$$

nlsBoot 5

## Value

A formula

## Author(s)

Florent Baty, Marie-Laure Delignette-Muller

#### References

Dixon M and Webb EC (1979) Enzymes, Academic Press, New York.

## **Examples**

```
# Example 1
data(vmkm)
nls1 <- nls(michaelis,vmkm,list(Km=1,Vmax=1))</pre>
plotfit(nls1, smooth = TRUE)
# Example 2
data(vmkmki)
def.par <- par(no.readonly = TRUE)</pre>
par(mfrow = c(2,2))
nls2_c <- nls(compet_mich, vmkmki, list(Km=1,Vmax=20,Ki=0.5))</pre>
plotfit(nls2_c, variable=1)
overview(nls2_c)
res2_c <- nlsResiduals(nls2_c)</pre>
plot(res2_c, which=1)
nls2_nc <- nls(non_compet_mich, vmkmki, list(Km=1, Vmax=20, Ki=0.5))</pre>
plotfit(nls2_nc, variable=1)
overview(nls2_nc)
res2_nc <- nlsResiduals(nls2_nc)</pre>
plot(res2_nc, which=1)
par(def.par)
```

 ${\tt nlsBoot}$ 

Bootstrap resampling

## **Description**

Bootstrap resampling

6 nlsBoot

## Usage

## **Arguments**

nls an object of class 'nls' niter number of iterations

x, object an object of class 'nlsBoot'

type type of representation (options are "pairs" or "boxplot")

mfr layout definition (number of rows and columns in the graphics device)

ask if TRUE, draw plot interactively

... further arguments passed to or from other methods

#### **Details**

Non-parametric bootstrapping is used. Mean centered residuals are bootstrapped. By default, 999 resampled data sets are created from which parameter estimates are obtained by fitting the model on each of these data sets. Whenever the fit fails to converge, a flag reports the number of nonconvergences. If the fitting procedure fails to converge in more than 50% of the cases, the procedure is interrupted with a flag and no result is given. The function summary returns the bootstrap estimates (mean and std. dev. of the bootstrapped estimates) and the median and 95 percent confidence intervals (50, 2.5, and 97.5 percentiles of the bootstrapped estimates). The bootstrapped estimate distributions can be visualized using the function plot.nlsBoot either by plotting the bootstrapped sample for each pair of parameters or by displaying the boxplot representation of the bootstrapped sample for each parameter. Notice that nlsBoot does not currently handle transformed dependent variables specified in the left side of the nls formula.

# Value

n1sBoot returns a list of 5 objects:

coefboot contains the bootstrap parameter estimates

bootCI contains the bootstrap medians and the bootstrap 95% confidence intervals estiboot contains the means and std. errors of the bootstrap parameter estimates

rse is the vector of bootstrap residual errors
nls the object of class 'nls' given in input

nlsBootPredict 7

#### Author(s)

Florent Baty, Marie-Laure Delignette-Muller

#### References

Bates DM and Watts DG (1988) Nonlinear regression analysis and its applications. Wiley, Chichester, UK.

Huet S, Bouvier A, Poursat M-A, Jolivet E (2003) Statistical tools for nonlinear regression: a practical guide with S-PLUS and R examples. Springer, Berlin, Heidelberg, New York.

## **Examples**

nlsBootPredict

Prediction from Bootstrap resampling

## **Description**

Computation of confidence intervals on predictions from Bootstrap resampling

## Usage

```
nlsBootPredict(nlsBoot, newdata, interval = c("confidence", "prediction"))
```

# **Arguments**

nlsBoot	An object of class 'nlsBoot'.
newdata	A data frame in which to look for values of independent variables for the predictions. If omitted, the data used for fitting are used.
interval	Type of interval to compute, "confidence", or "prediction".

8 nlsBootPredict

#### **Details**

nlsBootPredict produces confidence intervals on predicted values that can be obtained using predict.nls for values of the independent variable(s) defined in the data frame newdata. Non-parametric bootstrapping is used (results of nlsBoot). For confidence intervals the bootstrap sample of predictions is simply computed from the bootstrap sample of estimations of the model parameters, by evaluating the mean value of the model on each new data. For prediction intervals, to take into account the residual errors, a residual error sampled in the mean centered residuals is added to each mean predicted value. In both cases, bootstrap predictions are summarized by the median and 95 percent confidence intervals (50, 2.5, and 97.5 percentiles of the bootstrapped values).

#### Value

nlsBoot returns a matrix of predictions with three columns respectively corresponding to the 50, 2.5 and 97.5 percentiles of bootstrap predictions.

## Author(s)

Florent Baty, Marie-Laure Delignette-Muller

#### References

Huet S, Bouvier A, Poursat M-A, Jolivet E (2003) Statistical tools for nonlinear regression: a practical guide with S-PLUS and R examples. Springer, Berlin, Heidelberg, New York.

#### See Also

See nlsBoot and predict.nls.

```
formulaExp <- as.formula(V02 \sim (t \le 5.883) * V02res + (t > 5.883) *
                         (VO2res + (VO2peak - VO2res) *
                         (1 - \exp(-(t - 5.883) / mu))))
O2K.nls1 <- nls(formulaExp, start = list(VO2res = 400, VO2peak = 1600, mu = 1), data = O2K)
niter <- 200
### To reach stable prediction intervals use far greater niter (>> 1000)
02K.boot1 <- nlsBoot(02K.nls1, niter = niter)</pre>
newdata \leftarrow data.frame(t = seq(0, 12, length.out = 50))
(pred.clim <- nlsBootPredict(02K.boot1, newdata = newdata, interval = "confidence"))</pre>
(pred.plim <- nlsBootPredict(02K.boot1, newdata = newdata, interval = "prediction"))</pre>
plotfit(02K.nls1, smooth = TRUE, ylim = c(200, 1800))
lines(newdata$t, pred.clim[, 2], col = "red")
lines(newdata$t, pred.clim[, 3], col = "red")
lines(newdata$t, pred.plim[, 2], col = "blue")
lines(newdata$t, pred.plim[, 3], col = "blue")
### An example without giving newdata
# plot of data
```

nlsConfRegions 9

```
plot(02K$t, 02K$V02)

# add of predictions computed using predict.nls()
pred <- predict(02K.nls1)
points(02K$t, pred, pch = 16)

# add of prediction intervals using nlsBootPredict()
(pred.plim <- nlsBootPredict(02K.boot1, interval = "prediction"))
segments(02K$t, pred.plim[, 2], 02K$t, pred.plim[, 3], col = "blue")</pre>
```

nlsConfRegions

Confidence regions

#### Description

Draws parameter values in the Beale's 95 percent unlinearized confidence region

## Usage

```
nlsConfRegions (nls, length = 1000, exp = 1.5)
## S3 method for class 'nlsConfRegions'
plot(x, bounds = FALSE, ask = FALSE, ...)
## S3 method for class 'nlsConfRegions'
print(x, ...)
```

## **Arguments**

nls	an object of class 'nls'
length	number of points to draw in the confidence region
exp	expansion factor of the hypercube in which random values of parameters are drawn
X	an object of class 'nlsConfRegions'
bounds	logical defining whether bounds of the drawing hypercube are plotted
ask	if TRUE, draw plot interactively
	further arguments passed to or from other methods

## Details

A sample of points in the 95 percent confidence region is computed according to Beale's criterion (Beale, 1960). This region is also named the joint parameter likelihood region (Bates and Watts, 1988). The method used consists in a random sampling of parameters values in a hypercube centered on the least squares estimate and rejecting the parameters values whose residual sum of squares do not verify the Beale criterion. The confidence region is plotted by projection of the sampled points in each plane defined by a couple of parameters. Bounds of the hypercube in which random values of parameters are drawn may be plotted in order to check if the confidence region was totally included in the hypercube defined by default. If not the hypercube should be expanded in order to obtain the full confidence region

10 nlsContourRSS

#### Value

nlsConfRegions returns a list of four objects:

a data frame containing the sample drawn in the Beale's confidence region

a vector containing the residual sums of squares corresponding to cr

the 95 percent residual sum of squares threshold according to Beale (1960)

lower and upper bounds of the hypercube in which random values of parameters

have been drawn

#### Author(s)

Florent Baty, Marie-Laure Delignette-Muller

#### References

Beale EML (1960) Confidence regions in non-linear estimations. *Journal of the Royal Statistical Society*, **22B**, 41-88.

Bates DM and Watts DG (1988) Nonlinear regression analysis and its applications. Wiley, Chichester, UK.

## See Also

ellipse.nls in the ellipse library

#### **Examples**

nlsContourRSS

Surface contour of RSS

## Description

Provides residual sum of squares (RSS) contours

nlsContourRSS 11

#### Usage

### **Arguments**

nls	an object of class 'nls'
lseq	length of the sequences of parameters
exp	expansion factor of the parameter intervals defining the grids
nlev	number of contour levels to add to the likelihood contour at level 95 percent
col	logical. Contours are plotted with colors if TRUE
col.pal	Palette of colors. Colors to be used as background (default is terrain.colors(100); unused if col is FALSE)
x	an object of class 'nlsContourRSS'
ask	if TRUE, draw plot interactively (default is FALSE)
useRaster	a bitmap raster is used to plot the image instead of polygons (default is TRUE)
	further arguments passed to or from other methods

## **Details**

The aim of these functions is to plot the residual sum of squares (RSS) contours which correspond to likelihood contours for a Gaussian model. For each pair of parameters the RSS is calculated on a grid centered on the least squares estimates of both parameters, the other parameters being fixed to their least square estimates. The contours of RSS values are then plotted for each pair of parameters. For each pair of parameters, one of this contour corresponds to a section of the 95 percent Beale's confidence region in the plane of these parameters. This contour is plotted in a different color.

#### Value

nlsContourRSS returns a list of three objects:

seqPara	a matrix with the sequence of grid values for each parameter
lrss	a list of matrices with logarithm values of RSS in the grid for each pair of parameters
lrss95	the logarithm of the 95 percent residual sum of squares threshold according to Beale (1960)

#### Author(s)

Florent Baty, Marie-Laure Delignette-Muller

12 nlsJack

#### References

Beale EML (1960) Confidence regions in non-linear estimations. *Journal of the Royal Statistical Society*, **22B**, 41-88.

Bates DM and Watts DG (1988) Nonlinear regression analysis and its applications. Wiley, Chichester, UK.

## **Examples**

nlsJack

Jackknife resampling

## **Description**

Jackknife resampling

## Usage

```
nlsJack (nls)
## S3 method for class 'nlsJack'
plot(x, mfr = c(nrow(x$reldif),1), ask = FALSE, ...)
## S3 method for class 'nlsJack'
print(x, ...)
## S3 method for class 'nlsJack'
summary(object, ...)
```

## Arguments

```
nls an object of class 'nls'

x, object an object of class 'nlsJack'

mfr layout definition, default is k rows (k: number of parameters) and 1 column ask if TRUE, draw plot interactively

... further arguments passed to or from other methods
```

nlsJack 13

#### **Details**

A jackknife resampling procedure is performed. Each observation is sequentially removed from the initial data set using a leave-one-out strategy. A data set with n observations provides thus n resampled data sets of n-l observations. The jackknife estimates with confidence intervals are calculated as described by Seber and Wild (1989) from the results of n new fits of the model on the n jackknife resampled data sets. The leave-one-out procedure is also employed to assess the influence of each observation on each parameter estimate. An observation is empirically defined as influential for one parameter if the difference between the estimate of this parameter with and without the observation exceeds twice the standard error of the estimate divided by sqrt(n). This empirical method assumes a small curvature of the nonlinear model. For each parameter, the absolute relative difference (in percent of the estimate) of the estimates with and without each observation is plotted. An asterisk is plotted for each influential observation.

#### Value

nlsJack returns a list with 7 objects:

estijack a data frame with jackknife estimates and bias
coefjack a data frame with the parameter estimates for each jackknife sample
reldif a data frame with the absolute relative difference (in percent of the estimate) of
the estimates with and without each observation

dfb a data frame with dfbetas for each parameter and each observation
jackCI a data frame with jackknife confidence intervals
rse a vector with residual standard error for each jackknife sample

residual a vector with residual sum of squares for each jackknife sample

## Author(s)

rss

Florent Baty, Marie-Laure Delignette-Muller

#### References

Seber GAF, Wild CJ (1989) Nonlinear regression. Wiley, New York.

14 nlsResiduals

nlsResiduals NLS residuals

#### **Description**

Provides several plots and tests for the analysis of residuals

## Usage

```
nlsResiduals (nls)
## S3 method for class 'nlsResiduals'
plot(x, which = 0, ...)
test.nlsResiduals (x)
## S3 method for class 'nlsResiduals'
print(x, ...)
```

## **Arguments**

nls an object of class 'nls'

x an object of class 'nlsResiduals'

which an integer:

0 = 4 graphs of residuals (types 1, 2, 4 and 6)

1 = non-transformed residuals against fitted values

2 = standardized residuals against fitted values

3 = sqrt of absolute value of standardized residuals against fitted values

4 = auto-correlation residuals (i+1th residual against ith residual)

5 = histogram of the residuals

6 = qq-plot of the residuals

further arguments passed to or from other methods

#### **Details**

Several plots and tests are proposed to check the validity of the assumptions of the error model based on the analysis of residuals.

The function plot.nlsResiduals proposes several plots of residuals from the nonlinear fit: plot of non-transformed residuals against fitted values, plot of standardized residuals against fitted values, plot of square root of absolute value of standardized residuals against fitted values, auto-correlation plot of residuals (i+1th residual against ith residual), histogram of the non-transformed residuals and normal Q-Q plot of standardized residuals.

test.nlsResiduals tests the normality of the residuals with the Shapiro-Wilk test (shapiro.test in package stats) and the randomness of residuals with the runs test (Siegel and Castellan, 1988). The runs.test function used in nlstools is the one implemented in the package tseries.

nlstools 15

## Value

nlsResiduals returns a list of five objects:

std95	the Student value for alpha=0.05 (bilateral) and the degree of freedom of the model
resi1	a matrix with fitted values vs. non-transformed residuals
resi2	a matrix with fitted values vs. standardized residuals
resi3	a matrix with fitted values vs. sqrt(abs(standardized residuals))
resi4	a matrix with ith residuals vs. i+1th residuals

## Author(s)

Florent Baty, Marie-Laure Delignette-Muller

## References

Bates DM and Watts DG (1988) Nonlinear regression analysis and its applications. Wiley, Chichester, UK.

Siegel S and Castellan NJ (1988) Non parametric statistics for behavioral sciences. McGraw-Hill international, New York.

# **Examples**

nlstools

Nonlinear least squares fit

## **Description**

Tools to help the fit of nonlinear models with nls

16 nlstools

#### Usage

## **Arguments**

formula	formula of a non-linear model
data	a data frame with header matching the variables given in the formula
start	a list of parameter starting values which names match the parameters given in the formula
variable	index of the variable to be plotted against the predicted values; default is the first independent variable as it appears in the original dataset
X	an object of class 'nls'
smooth	a logical value, default is FALSE. If smooth is TRUE, a plot of observed values is plotted as a function of 1000 values continuously taken in the range interval [min(variable),max(variable)]. This option can only be used if the number of controlled variables is 1.
xlab	X-label
ylab	Y-label
pch.obs	type of point of the observed values
pch.fit	type of point of the fitted values (not applicable if smooth=TRUE)
lty	type of line of the smoothed fitted values (if smooth=TRUE)
lwd	thickness of line of the smoothed fitted values (if smooth=TRUE)
col.obs	color of the observed points
col.fit	color of the fitted values
• • •	further arguments passed to or from other methods

## **Details**

The function preview helps defining the parameter starting values prior fitting the model. It provides a superimposed plot of observed (circles) and predicted (crosses) values of the dependent variable versus one of the independent variables with the model evaluated at the starting values of the parameters. The function overview returns the parameters estimates, their standard errors as well as their asymptotic confidence intervals and the correlation matrix (alternately, the function confint provides better confidence interval estimates whenever it converges). plotfit displays a superimposed plot of the dependent variable versus one the independent variables together with the fitted model.

### Author(s)

Florent Baty, Marie-Laure Delignette-Muller

nlstools-defunct 17

#### References

Baty F, Ritz C, Charles S, Brutsche M, Flandrois J-P, Delignette-Muller M-L (2015). A Toolbox for Nonlinear Regression in R: The Package nlstools. *Journal of Statistical Software*, **66**(5), 1-21.

Bates DM and Watts DG (1988) Nonlinear regression analysis and its applications. Wiley, Chichester, UK.

#### See Also

nls in the stats library and confint.nls in the package MASS

## **Examples**

nlstools-defunct

Defunct Functions in Package nlstools

## **Description**

The models or data sets listed here are no longer part of package **nlstools**. In order to access these models and data set in the future, please load the additional package **nlsMicrobio**.

## **Details**

```
Defunct functions are:
geeraerd
geeraerd_without_Nres
geeraerd_without_Sl
mafart
albert
trilinear
bilinear_without_Nres
bilinear_without_Nres
bilinear_without_Sl
baranyi
baranyi_without_Nmax
baranyi_without_lag
buchanan
buchanan_without_lag
```

18 O2K

```
gompertzm
jameson_buchanan
jameson_baranyi
jameson_without_lag
cpm_T
cpm_pH_4p
cpm_pH_3p
cpm_aw_3p
cpm_aw_2p
cpm_T_pH_aw
competition1
competition2
growthcurve1
growthcurve2
growthcurve3
growthcurve4
ross
survivalcurve1
survivalcurve2
survivalcurve3
```

02K

Oxygen kinetics during 6-minute walk test data set

## **Description**

Oxygen uptake kinetics during a 6-minute walking test in a patient with pulmonary disease. The first 5.83 minutes correspond to the resting phase prior to exercise.

# Usage

```
data(02K)
```

### **Format**

02K is a data frame with 2 columns (t: time, VO2: oxygen uptake)

## Source

This data set was provided by the Cantonal Hospital St. Gallen, Switzerland.

```
data(02K)
plot(02K)
```

# **Index**

* datasets	<pre>plot.nlsBoot (nlsBoot), 5</pre>
L.minor, 3	<pre>plot.nlsConfRegions(nlsConfRegions), 9</pre>
michaelisdata, 3	plot.nlsContourRSS(nlsContourRSS), 10
02K, 18	plot.nlsJack(nlsJack), 12
* models	plot.nlsResiduals(nlsResiduals), 14
confint2, 2	plotfit (nlstools), 15
michaelismodels, 4	predict.nls,8
* nonlinear	preview (nlstools), 15
confint2, 2	<pre>print.nlsBoot(nlsBoot), 5</pre>
nlsBoot, 5	<pre>print.nlsConfRegions (nlsConfRegions), 9</pre>
nlsBootPredict, 7	<pre>print.nlsContourRSS (nlsContourRSS), 10</pre>
nlsConfRegions, 9	print.nlsJack(nlsJack), 12
nlsContourRSS, 10	<pre>print.nlsResiduals(nlsResiduals), 14</pre>
nlsJack, 12	
nlsResiduals, 14	<pre>summary.nlsBoot(nlsBoot), 5</pre>
nlstools, 15	summary.nlsJack(nlsJack), 12
<pre>compet_mich (michaelismodels), 4</pre>	test.nlsResiduals(nlsResiduals), 14
confint.nls, 2	
confint2, 2	vmkm (michaelisdata), 3
551111162, 2	vmkmki (michaelisdata), 3
L.minor, 3	
michaelis (michaelismodels), 4	
michaelisdata, 3	
michaelismodels, 4	
nls, 2	
nlsBoot, 5, 8	
nlsBootPredict, 7	
nlsConfRegions, 9	
nlsContourRSS, 10	
nlsJack, 12	
nlsResiduals, 14	
nlstools, 15	
nlstools-defunct, 17	
non_compet_mich (michaelismodels), 4	
02K, 18	
overview (nlstools), 15	