# Package 'iRepro'

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Title Reproducibility for Interval-Censored Data
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Author Jelena Kovacic
Maintainer Jelena Kovacic < jkovacic@imi.hr>
<b>Description</b> This package calculates intraclass correlation coefficient (ICC) for assessing repro- ducibility of interval-censored data with two repeated measurements. ICC is estimated by maxi- mum likelihood from model with one fixed and one random effect (both inter- cepts). Help in model checking (normality of subjects' means and residuals) is provided.
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```
iRepro-package
```

#### Description

This package calculates intraclass correlation coefficient (ICC) for assessing reproducibility of interval-censored data with two repeated measurements. ICC is estimated by maximum likelihood from model with one fixed and one random effect (both intercepts). Help in model checking (normality of subjects' means and residuals) is provided.

## Details

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#### Author(s)

Jelena Kovacic

Maintainer: Jelena Kovacic <jkovacic@imi.hr>

## References

Kovacic J, Varnai VM. Reproducibility of grouped data assessed by intraclass correlation coefficient. Accepted for publication in *Epidemiology*.

## Examples

```
# Data generation (grouped data)
classes <- 1:6
class.limits <- cbind(classes-0.5,classes+0.5)
r1 <- sample(classes,100,replace=TRUE) # first measurement
r2 <- sample(classes,100,replace=TRUE) # second measurement</pre>
```

summary(intervalICC(r1,r2,predefined.classes=TRUE,classes,class.limits)) # ICC estimation

intervalICC

## Description

The function calculates intraclass correlation coefficient (ICC) for interval-censored data with two repeated measurements. ICC is estimated by maximum likelihood from model with one fixed and one random effect (both intercepts).

## Usage

```
intervalICC(r1, r2, predefined.classes=FALSE, classes, c.limits, optim.method=1)
```

## Arguments

r1	data corresponding to the first measurement. If predefined.classes=TRUE (appropriate for grouped data), this is a vector of length $n$ , where each observation is one of the labels given in classes. Otherwise, if predefined.classes=FALSE, r1 is a matrix or a data frame with $n$ rows and 2 columns, with columns representing lower and upper bounds of censoring intervals (e.g., if <i>i</i> -th observation lies in the interval $[a, b]$ , then r1[i,]=c(a,b)).
r2	data corresponding to the second measurement. If predefined.classes=TRUE (appropriate for grouped data), this is a vector of length $n$ , where each observa- tion is one of the labels given in classes. Otherwise, if predefined.classes=FALSE, r2 is a matrix or a data frame with $n$ rows and 2 columns, with columns repre- senting lower and upper bounds of censoring intervals (e.g., if <i>i</i> -th observation lies in the interval $[a, b]$ , then r2[i,]=c(a,b)).
predefined.cla	asses
	logical, indicating whether observations belong to predefined classes (e.g. grouped data in questionnaires) or each observation has its own lower and upper limit (default; FALSE).
classes	a vector with unique labels for the k predefined classes. Required if predefined.classes=TRUE.
c.limits	a matrix or a data frame with $k$ rows and 2 columns, corresponding to lower and upper bounds of censoring intervals for classes. Required if predefined.classes=TRUE.
optim.method	an integer (1 or 2) specifying the optimization method to be used in maximum likelihood estimation (default is 1). Details are given below.

### Details

ICC is estimated by maximum likelihood from random effects model

$$Y_{ij} = \mu + b_i + e_{ij}$$

where  $b_i$  and  $e_{ij}$  are independent and normally distributed with means 0 and variances  $\sigma_b^2$  and  $\sigma^2$ , respectively. If data were uncensored, this would be analogous to

lme(ratings~1, random=~1|id, method="ML", data=observed)

in nlme package, where

observed=as.data.frame(rbind(cbind(r1,1:n), cbind(r2,1:n)))

and colnames(observed)=c("ratings","id"). To maximize log-likelihood, constrOptim from stats package is used (method=BFGS).

Two available optimization methods, specified by optim.method, correspond to two mathematically equivalent expressions for log-likelihood. The option optim.method=1 resulted in slightly more accurate estimates in simulations with grouped data, but optim.method=2 was more numerically stable. See the reference for more details.

#### Value

An object of class "ICCfit". The object is a list with the components:

icc	maximum likelihood estimate (MLE) of ICC
sigma2.b	MLE of between-class variance $\sigma_b^2$
sigma2.w	MLE of within-class variance $\sigma^2$
mu	MLE of mean $\mu$
loglikelihood	log-likelihood evaluated at MLE parameters

#### Note

If there are many observations with same values (i.e. with the same lower and upper bounds), it is advisable to group all observations into classes and use option predefined.classes=TRUE; this will reduce computation time.

Subjects with only one measurement are omitted from ICC calculation.

#### Author(s)

Jelena Kovacic < jkovacic@imi.hr>

## References

Kovacic J, Varnai VM. Reproducibility of grouped data assessed by intraclass correlation coefficient. Accepted for publication in *Epidemiology*.

#### See Also

print.ICCfit, summary.ICCfit

## Examples

```
# Example with 6 predefined classes (grouped data)
classes <- 1:6
class.limits <- cbind(classes-0.5,classes+0.5)
r1 <- sample(classes,30,replace=TRUE)
r2 <- sample(classes,30,replace=TRUE)</pre>
```

intervalICC(r1,r2,predefined.classes=TRUE,classes,class.limits)

#### ntest.means

```
# The same result can be obtained with predefined.classes=FALSE option,
# although with slower computation time
rtg1 <- matrix(nrow=30,ncol=2)
rtg2 <- matrix(nrow=30,ncol=2)
# when predefined.classes=FALSE, ratings must be given with lower and upper bounds
# for each observation:
for(i in 1:length(classes)){
    rtg1[r1==classes[i],1] <- class.limits[i,1]
    rtg1[r1==classes[i],2] <- class.limits[i,2]
    rtg2[r2==classes[i],1] <- class.limits[i,2]
    rtg2[r2==classes[i],2] <- class.limits[i,2]
    rtg2[r2==classes[i],2] <- class.limits[i,2]
}
intervalICC(rtg1,rtg2,predefined.classes=FALSE)
```

ntest.means

Normality Check for Interval-Censored Data with Repeated Measurements - Means

#### Description

The function checks whether interval-censored data with two repeated measurements meet the normality assumption for subjects' means. This is a prerequisite for the random effects model used in ICC calculation.

## Usage

ntest.means(r1, r2, predefined.classes=FALSE, classes, c.limits, optim.method=1, bins=10)

#### Arguments

r1	argument passed to intervalICC; see documentation for that function.	
r2	argument passed to intervalICC; see documentation for that function.	
predefined.classes		
	argument passed to intervalICC; see documentation for that function.	
classes	argument passed to intervalICC; see documentation for that function.	
c.limits	argument passed to intervalICC; see documentation for that function.	
optim.method	argument passed to intervalICC; see documentation for that function.	
bins	number of categories in chi-square test; see details below (default is 10).	

#### Details

For ICC estimation the random effects data model

$$Y_{ij} = \mu + b_i + e_{ij},$$

is used, where  $b_i$  and  $e_{ij}$  are normally distributed with means 0 and variances  $\sigma_b^2$  and  $\sigma^2$ , respectively. This function assesses the assumption that the subjects' means  $0.5(Y_{i1} + Y_{i2})$  are normally distributed with mean  $\mu$  and variance  $\sigma_b^2 + 0.5\sigma^2$ , as is expected under the specified model.

To test normality, chi-square goodness-of-fit test with bins subsequent data categories is used (call to chisq.test from package stats). The categories (bins) are determined using the equidistant quantiles of expected normal distribution, with corresponding maximum likelihood parameters. Maximum likelihood estimates for parameters  $\mu$ ,  $\sigma_b^2$  and  $\sigma^2$  are obtained by calling the function intervalICC. The probability corresponding to each bin is 1/bins (expected relative frequencies; this corresponds to p = rep(1/bins,bins) in chisq.test function). Since means are interval-censored and censoring intervals overlap, the observed relative frequencies are calculated in the following way. If one of the original intervals representing subjects mean spans multiple bins, each bin receives a share of votes from the original interval. This share is calculated using the expected normal density function and it is proportional to the probability of data falling within the intersection of the original interval and bin.

### Value

An object of class "ntestMeans". The object is a list with the components:

statistic	value of chi-squared statistic; statistic in the output of chisq.test
parameter	number of degrees of freedom for chi-squared distribution; parameter in the output of chisq.test
p.value	p-value of test; p.value in the output of chisq.test
data	character string with value "means"
mu	mean of the expected normal distribution for subjects' means; equal to maximum likelihood estimate for $\mu$ from <code>intervalICC</code>
var	variance of the expected normal distribution for subjects' means; equal to maximum likelihood estimate for $\sigma_b^2 + 0.5\sigma^2$ from intervalICC
bins	number of categories in chi-square test

#### Note

This function was designed as a help in assessing goodness of model fit. However, it has not been tested in simulations nor in any other way. It is the responsibility of the user to provide appropriate number of bins; the function checks only if bins is a positive integer. Testing normality with low number of bins is unreliable. On the other hand, if the number of bins is too large, chisq.test will complain since the expected frequencies will be too low.

#### Author(s)

Jelena Kovacic < jkovacic@imi.hr>

#### ntest.res

#### References

Kovacic J, Varnai VM. Reproducibility of grouped data assessed by intraclass correlation coefficient. Accepted for publication in *Epidemiology*.

#### See Also

summary.ntestMeans, intervalICC, chisq.test

#### Examples

```
# Example with 6 predefined classes (grouped data)
classes <- 1:6
class.limits <- cbind(classes-0.5, classes+0.5)
r1 <- sample(classes, 30, replace=TRUE)
r2 <- sample(classes, 30, replace=TRUE)
ntest.means(r1,r2, predefined.classes=TRUE, classes, class.limits, bins=10)</pre>
```

ntest.	res
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Normality Check for Interval-Censored Data with Repeated Measurements - Residuals

## Description

The function checks whether interval-censored data with two repeated measurements meet the normality assumption for subjects' residuals. This is a prerequisite for the random effects model used in ICC calculation.

### Usage

ntest.res(r1, r2, predefined.classes=FALSE, classes, c.limits, optim.method=1, bins=10)

#### Arguments

r1	argument passed to intervalICC; see documentation for that function.
r2	argument passed to intervalICC; see documentation for that function.
predefined.cla	sses
	argument passed to intervalICC; see documentation for that function.
classes	argument passed to intervalICC; see documentation for that function.
c.limits	argument passed to intervalICC; see documentation for that function.
optim.method	argument passed to intervalICC; see documentation for that function.
bins	number of categories in chi-square test; see details below (default is 10).

#### Details

For ICC estimation the random effects data model

$$Y_{ij} = \mu + b_i + e_{ij}$$

is used, where  $b_i$  and  $e_{ij}$  are normally distributed with means 0 and variances  $\sigma_b^2$  and  $\sigma^2$ , respectively. This function assesses the assumption that the subjects' "residuals"  $Y_{i1} - 0.5(Y_{i1} + Y_{i2})$  and  $Y_{i2} - 0.5(Y_{i1} + Y_{i2})$  are normally distributed with mean 0 and variance  $0.5\sigma^2$ , as is expected under the specified model.

To test normality, chi-square goodness-of-fit test with bins subsequent data categories is used (call to chisq.test from package stats). The categories (bins) are determined using the equidistant quantiles of expected normal distribution, with corresponding maximum likelihood parameters. Maximum likelihood estimates for parameters  $\mu$ ,  $\sigma_b^2$  and  $\sigma^2$  are obtained by calling the function intervalICC. The probability corresponding to each bin is 1/bins (expected relative frequencies; this corresponds to p = rep(1/bins, bins) in chisq.test function). Since residuals are interval-censored and censoring intervals overlap, the observed relative frequencies are calculated in the following way. If one of the original intervals representing subjects residual spans multiple bins, each bin receives a share of votes from the original interval. This share is calculated using the expected normal density function and it is proportional to the probability of data falling within the intersection of the original interval and bin.

Residuals for the first time point  $(Y_{i1} - 0.5(Y_{i1} + Y_{i2}))$  and residuals for the second  $(Y_{i2} - 0.5(Y_{i1} + Y_{i2}))$  are tested separately; therefore two test results in the output are given.

#### Value

An object of class "ntestRes". The object is a list with the components:

statistic.res1	
	value of chi-squared statistic corresponding to the first residual; statistic in the output of chisq.test $% \left( \frac{1}{2}\right) =0$
p.value.res1	$p\mbox{-value}$ of test corresponding to the first residual; $p\mbox{-value}$ in the output of chisq.test
statistic.res2	
	value of chi-squared statistic corresponding to the second residual; ${\tt statistic}$ in the output of chisq.test
p.value.res2	${\tt p}\mbox{-value}$ of test corresponding to the second residual; ${\tt p}\mbox{-value}$ in the output of chisq.test
parameter	number of degrees of freedom for chi-squared distribution (the same for both residuals); parameter in the output of chisq.test $% f(x) = 0$
data	character string with value "residuals"
mu	mean of the expected normal distribution for subjects' residuals; equal to 0
var	variance of the expected normal distribution for subjects' residuals; equal to maximum likelihood estimate for $0.5\sigma^2$ from <code>intervalICC</code>
bins	number of categories in chi-square test

#### Note

This function was designed as a help in assessing goodness of model fit. However, it has not been tested in simulations nor in any other way. It is the responsibility of the user to provide appropriate number of bins; the function checks only if bins is a positive integer. Testing normality with low number of bins is unreliable. On the other hand, if the number of bins is too large, chisq.test will complain since the expected frequencies will be too low.

#### Author(s)

Jelena Kovacic < jkovacic@imi.hr>

## References

Kovacic J, Varnai VM. Reproducibility of grouped data assessed by intraclass correlation coefficient. Accepted for publication in *Epidemiology*.

#### See Also

summary.ntestRes, intervalICC, chisq.test

#### Examples

```
# Example with 6 predefined classes (grouped data)
classes <- 1:6
class.limits <- cbind(classes-0.5,classes+0.5)
r1 <- sample(classes,30,replace=TRUE)
r2 <- sample(classes,30,replace=TRUE)
ntest.res(r1,r2,predefined.classes=TRUE,classes,class.limits,bins=10)</pre>
```

summary.ICCfit Summary for ICCfit Objects

#### Description

The function summarizes the results of ICC estimation.

#### Usage

```
## S3 method for class 'ICCfit'
summary(object, ...)
```

#### Arguments

object	object of the class ICCfit (output of the intervalICC function)
	additional arguments passed to the function (they do not affect the summary produced)

#### Details

For more details about ICC estimation and output values shortly described below, please refer to the documentation for intervalICC.

## Value

An object of class "summary.ICCfit". The object is a list with the components:

estimates	a data frame containing maximum likelihood estimates for ICC, mean and vari-
	ance components
loglikelihood	log-likelihood evaluated at maximum likelihood estimates

#### Author(s)

Jelena Kovacic <jkovacic@imi.hr>

#### References

Kovacic J, Varnai VM. Reproducibility of grouped data assessed by intraclass correlation coefficient. Accepted for publication in *Epidemiology*.

## See Also

intervalICC

#### Examples

```
# Example with 6 predefined classes (grouped data)
classes <- 1:6
class.limits <- cbind(classes-0.5, classes+0.5)
r1 <- sample(classes, 30, replace=TRUE)
r2 <- sample(classes, 30, replace=TRUE)
icc.est <- intervalICC(r1,r2, predefined.classes=TRUE, classes, class.limits)
summary(icc.est)</pre>
```

summary.ntestMeans Summary for ntestMeans Objects

## Description

The function summarizes the results of normality check for means.

## Usage

```
## S3 method for class 'ntestMeans'
summary(object, ...)
```

#### Arguments

object	object of the class ntestMeans (output of the ntest.means function)
	additional arguments passed to the function (they do not affect the summary produced)

## Details

For more details about normality check and output values shortly described below, please refer to the documentation for ntest.means.

#### Value

An object of class "summary.ntestMeans". The object is a list with the components:

test.res	a data frame containing the chi-squared statistic and p-value for normality test
mu	mean of the expected normal distribution for means
stdev	standard deviation of the expected normal distribution for means
bins	number of categories in chi-squared normality test
df	number of degrees of freedom in chi-squared normality test

## Author(s)

Jelena Kovacic < jkovacic@imi.hr>

#### References

Kovacic J, Varnai VM. Reproducibility of grouped data assessed by intraclass correlation coefficient. Accepted for publication in *Epidemiology*.

#### See Also

ntest.means

## Examples

```
# Example with 6 predefined classes (grouped data)
classes <- 1:6
class.limits <- cbind(classes-0.5, classes+0.5)
r1 <- sample(classes, 30, replace=TRUE)
r2 <- sample(classes, 30, replace=TRUE)
nm <- ntest.means(r1,r2, predefined.classes=TRUE, classes, class.limits, bins=10)
summary(nm)</pre>
```

summary.ntestRes Summary for ntestRes Objects

## Description

The function summarizes the results of normality check for residuals.

## Usage

## S3 method for class 'ntestRes'
summary(object, ...)

#### Arguments

object	object of the class ntestRes (output of the ntest.res function)
	additional arguments passed to the function (they do not affect the summary produced)

## Details

For more details about normality check and output values shortly described below, please refer to the documentation for ntest.res.

#### Value

An object of class "summary.ntestRes". The object is a list with the components:

test.res a	a data frame containing the chi-squared statistics and p-values for normality tests
mu a	mean of the expected normal distribution for residuals
stdev	standard deviation of the expected normal distribution for residuals
bins i	number of categories in chi-squared normality test
df	number of degrees of freedom in chi-squared normality test

## Author(s)

Jelena Kovacic <jkovacic@imi.hr>

### References

Kovacic J, Varnai VM. Reproducibility of grouped data assessed by intraclass correlation coefficient. Accepted for publication in *Epidemiology*.

## See Also

ntest.res

## summary.ntestRes

## Examples

```
# Example with 6 predefined classes (grouped data)
classes <- 1:6
class.limits <- cbind(classes-0.5,classes+0.5)
r1 <- sample(classes,30,replace=TRUE)
r2 <- sample(classes,30,replace=TRUE)
nr <- ntest.res(r1,r2,predefined.classes=TRUE,classes,class.limits,bins=10)
summary(nr)</pre>
```

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