Package 'meta'

September 12, 2014

Title Meta-Analysis with R Version 3.8-0 **Depends** R (>= 2.9.1) Imports grid Suggests metafor **Date** 2014-09-12 **Description** User-friendly general package providing standard methods for meta-analysis: - fixed effect and random effects meta-analysis; - several plots (forest, funnel, Galbraith / radial, L'Abbe, Baujat, bubble); - statistical tests and trim-and-fill method to evaluate bias in meta-analysis; - import data from RevMan 5; - prediction interval, Hartung-Knapp and Paule-Mandel method for random effects model; - cumulative meta-analysis and leave-one-out meta-analysis; - meta-regression (if R package metafor is installed). **License** GPL (>= 2) **Encoding** UTF-8 Author Guido Schwarzer [cre, aut] Maintainer Guido Schwarzer <sc@imbi.uni-freiburg.de> NeedsCompilation yes Repository CRAN **Date/Publication** 2014-09-12 16:15:23

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addvar

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Additional functions for objects of class meta

Description

The as.data.frame method returns a data frame containing information on individual studies, e.g., estimated treatment effect and its standard error. The function addvar can be used to add a single variable to an object of class meta which for example is useful to add columns to a forest plot.

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Usage

```
## $3 method for class 'meta'
as.data.frame(x, row.names=NULL, optional=FALSE, ...)
addvar(x, y, varname, by.x="studlab", by.y=by.x)
```

Arguments

Χ	An object of class meta.
row.names	NULL or a character vector giving the row names for the data frame.
optional	logical. If TRUE, setting row names and converting column names (to syntactic names) is optional.
у	A data frame with an additional covariate
varname	A character specifying name of additional variable
by.x, by.y	Specifications of the common columns (see merge)
	other arguments

Value

A data frame is returned by the function as.data.frame.

A single covariate is returned by the function addvar which can be added to an object of class meta. Internally, the merge function is utilised. See help page metagen for an example on the use of R function addvar.

Author(s)

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See Also

```
metabin, metacont, metagen, forest.meta
```

Examples

```
data(Fleiss93cont)
#
# Generate additional variable with grouping information
#
Fleiss93cont$group <- c(1,2,1,1,2)
#
# Do meta-analysis without grouping information
#
meta1 <- metacont(n.e, mean.e, sd.e, n.c, mean.c, sd.c, study, data=Fleiss93cont, sm="SMD")
#
# Update meta-analysis object and do subgroup analyses
# summary(update(meta1, byvar=group))</pre>
```

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amlodipine

Amlodipine for Work Capacity

Description

Meta-analysis on the effect of amlodipine on work capacity.

This meta-analysis is used as a data example in Hartung and Knapp (2001).

Usage

```
data(amlodipine)
```

Format

A data frame with the following columns:

study Study label

n.amlo Number of observations in amlodipine group

mean.amlo Estimated mean in amlodipine group

var.amlo Variance in amlodipine group

n.plac Number of observations in placebo group

mean.plac Estimated mean in placebo group

var.plac Variance in placebo group

Source

Hartung J & Knapp G (2001), On tests of the overall treatment effect in meta-analysis with normally distributed responses. *Statistics in Medicine*, **20**, 1771–82. doi: 10.1002/sim.791.

See Also

metacont

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Examples

```
data(amlodipine)
m <- metacont(n.amlo, mean.amlo, sqrt(var.amlo),</pre>
              n.plac, mean.plac, sqrt(var.plac),
              data=amlodipine, studlab=study)
m.hakn <- metacont(n.amlo, mean.amlo, sqrt(var.amlo),</pre>
                    n.plac, mean.plac, sqrt(var.plac),
                    data=amlodipine, studlab=study,
                    hakn=TRUE)
# Results for mean difference - see Table III in Hartung and Knapp (2001)
res.md <- rbind(data.frame(summary(m)$fixed)[c("TE", "lower", "upper")],</pre>
                 data.frame(summary(m)$random)[c("TE", "lower", "upper")],
                 data.frame(summary(m.hakn)$random)[c("TE", "lower", "upper")])
res.md <- round(res.md, 5)</pre>
row.names(res.md) <- c("FE", "RE", "RE (HaKn)")</pre>
names(res.md) <- c("Absolute difference", "CI lower", "CI upper")</pre>
res.md
```

baujat

Baujat plot to explore heterogeneity in meta-analysis

Description

Draw a Baujat plot to explore heterogeneity in meta-analysis.

Usage

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Arguments

x	An object of class meta.
yscale	Scaling factor for values on y-axis.
xlim	The x limits (min,max) of the plot.
ylim	The y limits (min,max) of the plot.
xlab	A label for the x-axis.
ylab	A label for the y-axis.
pch	The plotting symbol used for individual studies.
cex	The magnification to be used for plotting symbol.
col	A vector with colour of plotting symbols.
bg	A vector with background colour of plotting symbols (only used if pch in 21:25).
studlab	A logical indicating whether study labels should be printed in the graph. A vector with study labels can also be provided (must be of same length as x TE then).
cex.studlab	The magnification for study labels.
xmin	A numeric specifying minimal value to print study labels (on x-axis).
ymin	A numeric specifying minimal value to print study labels (on y-axis).
pos	A position specifier for study labels (see text).
offset	Offset for study labels (see text).
grid	A logical indicating whether a grid is printed in the plot.
col.grid	Colour for grid lines.
lty.grid	The line type for grid lines.
lwd.grid	The line width for grid lines.
pty	A character specifying type of plot region (see par).
	Graphical arguments as in par may also be passed as arguments.

Details

Baujat et al. (2002) introduced a scatter plot to explore heterogeneity in meta-analysis. On the x-axis the contribution of each study to the overall heterogeneity statistic (see list object Q of the meta-analysis object x) is plotted. On the y-axis the standardised difference of the overall treatment effect with and without each study is plotted; this quantity describes the influence of each study on the overall treatment effect.

Internally, the metainf function is used to calculate the values on the y-axis.

Value

A data.frame with the following variables:

x Coordinate on x-axis (contribution to heterogeneity statistic).

y Coordinate on y-axis (influence on overall treatment effect).

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Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Baujat B, Mahé C, Pignon JP, Hill C (2002), A graphical method for exploring heterogeneity in meta-analyses: Application to a meta-analysis of 65 trials. *Statistics in Medicine*, **30**, 2641–2652.

See Also

```
metagen, metainf
```

Examples

```
data(Olkin95)
m1 <- metabin(event.e, n.e, event.c, n.c, data=Olkin95,</pre>
              studlab=author, sm="OR", method="I")
# Generate Baujat plot
baujat(m1)
# Do not print study labels if the x-value is smaller than 4 and the
# y-value is smaller than 1.
baujat(m1, yscale=10, xmin=4, ymin=1)
# Change position of study labels
baujat(m1, yscale=10, xmin=4, ymin=1,
       pos=1, xlim=c(0, 6.5))
# Generate Baujat plot and assign x- and y- coordinates to R object b1
b1 <- baujat(m1)</pre>
# Calculate overall heterogeneity statistic
sum(b1$x)
m1$Q
```

bubble

Bubble plot to display the result of a meta-regression

Description

Draw a bubble plot to display the result of a meta-regression.

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Usage

Arguments

•	3	
	X	An object of class metareg.
	xlim	The x limits (min,max) of the plot.
	ylim	The y limits (min,max) of the plot.
	xlab	A label for the x-axis.
	ylab	A label for the y-axis.
	cex	The magnification to be used for plotting symbols.
	min.cex	Minimal magnification for plotting symbols.
	max.cex	Maximal magnification for plotting symbols.
	pch	The plotting symbol used for individual studies.
	col	A vector with colour of plotting symbols.
	bg	A vector with background colour of plotting symbols (only used if pch in 21:25).
	lty	The line type for the meta-regression line.
	lwd	The line width for the meta-regression line.
	col.line	Colour for the meta-regression line.
	studlab	A logical indicating whether study labels should be printed in the graph. A vector with study labels can also be provided (must be of same length as the numer of studies in the meta-analysis then).
	cex.studlab	The magnification for study labels.
	pos	A position specifier for study labels (see text).
	offset	Offset for study labels (see text).
	regline	A logical indicating whether a regression line should be added to the bubble plot.
	axes	A logical indicating whether axes should be printed.
	box	A logical indicating whether a box should be printed.
		Graphical arguments as in par may also be passed as arguments.

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Details

A bubble plot can be used to display the result of a meta-regression. It is a scatter plot with the treatment effect for each study on the y-axis and the covariate used in the meta-regression on the x-axis. Typically, the size of the plotting symbol is inversely proportional to the variance of the estimated treatment effect (Thompson & Higgins, 2002).

Argument cex specifies the plotting size for each individual study. If this argument is missing the weights from the meta-regression model will be used (which typically is a random effects model). Use weight="fixed" in order to utilise weights from a fixed effect model to define the size of the plotted symbols (even for a random effects meta-regression). If a vector with individual study weights is provided, the length of this vector must be of the same length as the number of studies.

Arguments min.cex and max.cex can be used to define the size of the smallest and largest plotting symbol. The plotting size of the most precise study is set to max.cex whereas the plotting size of all studies with a plotting size smaller than min.cex will be set to min.cex.

For a meta-regression with more than one covariate. Only a scatter plot of the first covariate in the regression model is shown. In this case the effect of the first covariate adjusted for other covariates in the meta-regression model is shown.

For a factor or categorial covariate separate bubble plots for each group compared to the baseline group are plotted.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Thompson SG, Higgins JP (2002), How should meta-regression analyses be undertaken and interpreted? *Statistics in Medicine*, **21**, 1559–1573.

See Also

```
metagen, metainf
```

Examples

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```
mr2 <- metareg(meta1, age)
mr2

bubble(mr2, lwd=2, col.line="blue", xlim=c(50, 70))
bubble(mr2, lwd=2, col.line="blue", xlim=c(50, 70), cex="fixed")

# Do not print regression line
#
bubble(mr2, lwd=2, col.line="blue", xlim=c(50, 70), regline=FALSE)</pre>
```

ci

Calculation of confidence intervals (based on normal approximation or t-distribution)

Description

Calculation of confidence intervals; based on normal approximation or t-distribution.

Usage

```
ci(TE, seTE, level=0.95, df=NULL)
```

Arguments

TE Estimated treatment effect.

seTE Standard error of treatment estimate.

level The confidence level required.

df Degrees of freedom (for confidence intervals based on t-distribution).

Value

List with components

TE Estimated treatment effect.

seTE Standard error of treatment estimate.

lower Lower confidence limits.

upper Upper confidence limits.

z Test statistic (either z-score or t-score).
p P-value of test with null hypothesis TE=0.

level The confidence level required.

df Degrees of freedom (t-distribution).

Note

This function is primarily called from other functions of the library meta, e.g. forest.meta, summary.meta.

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Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

Examples

```
data.frame(ci(170, 10))
data.frame(ci(170, 10, 0.99))
data.frame(ci(1.959964, 1))
data.frame(ci(2.2621571628, 1, df=9))
```

cisapride

Cisapride in Non-Ulcer Dispepsia

Description

Meta-analysis on cisapride in non-ulcer dispepsia.

This meta-analysis is used as a data example in Hartung and Knapp (2001).

Usage

```
data(cisapride)
```

Format

A data frame with the following columns:

```
study Study label
```

event.cisa Number of events in cisapride group

n.cisa Number of observations in cisapride group

event.plac Number of events in placebo group

n.plac Number of observations in placebo group

Source

Hartung J & Knapp G (2001), A Refined Method for the Meta-analysis of Controlled Clinical Trials with Binary Outcome. *Statistics in Medicine*, **20**, 3875–89.

See Also

metabin

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Examples

```
data(cisapride)
m.or <- metabin(event.cisa, n.cisa, event.plac, n.plac,</pre>
               data=cisapride, sm="OR", method="Inverse",
               studlab=study, addincr=TRUE)
m.rr <- metabin(event.cisa, n.cisa, event.plac, n.plac,</pre>
                data=cisapride, sm="RR", method="Inverse",
                studlab=study, addincr=TRUE)
m.or.hakn <- metabin(event.cisa, n.cisa, event.plac, n.plac,</pre>
                    data=cisapride, sm="OR", method="Inverse",
                     studlab=study, addincr=TRUE,
                    hakn=TRUE)
m.rr.hakn <- metabin(event.cisa, n.cisa, event.plac, n.plac,</pre>
                    data=cisapride, sm="RR", method="Inverse",
                    studlab=study, addincr=TRUE,
                    hakn=TRUE)
# Results for log risk ratio - see Table VII in Hartung and Knapp (2001)
res.rr <- rbind(data.frame(summary(m.rr)$fixed)[c("TE", "lower", "upper")],</pre>
                data.frame(summary(m.rr)$random)[c("TE", "lower", "upper")],
                data.frame(summary(m.rr.hakn)$random)[c("TE", "lower", "upper")])
row.names(res.rr) <- c("FE", "RE", "RE (HaKn)")</pre>
names(res.rr) <- c("Log risk ratio", "CI lower", "CI upper")</pre>
res.rr
# Results for log odds ratio (Table VII in Hartung and Knapp 2001)
data.frame(summary(m.or.hakn)$random)[c("TE", "lower", "upper")])
row.names(res.or) <- c("FE", "RE", "RE (HaKn)")</pre>
names(res.or) <- c("Log odds ratio", "CI lower", "CI upper")</pre>
res.or
```

Fleiss93

Aspirin after Myocardial Infarction

Description

Meta-analysis on aspirin in preventing death after myocardial infarction.

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Data example in Fleiss (1993) for meta-analysis with binary outcomes.

Usage

```
data(Fleiss93)
```

Format

A data frame with the following columns:

```
study Study label
year Year of publication
event.e Number of deaths in aspirin group
n.e Number of observations in aspirin group
event.c Number of deaths in placebo group
n.c Number of observations in placebo group
```

Source

Fleiss JL (1993), The statistical basis of meta-analysis. *Statistical Methods in Medical Research*, **2**, 121–145.

Examples

Fleiss93cont

Mental Health Treatment

Description

Meta-analysis on the Effect of Mental Health Treatment on Medical Utilisation.

Data example in Fleiss (1993) for meta-analysis with continuous outcomes.

Usage

```
data(Fleiss93cont)
```

Format

```
A data frame with the following columns: study Study label
```

year Year of publication

n.e Number of observations in psychotherapy group

mean.e Estimated mean in psychotherapy group

sd.e Standard deviation in psychotherapy group

n.c Number of observations in control group

mean.c Estimated mean in control group

sd.c Standard deviation in control group

Source

Fleiss JL (1993), The statistical basis of meta-analysis. *Statistical Methods in Medical Research*, **2**, 121–145.

See Also

Fleiss93

Examples

forest

Forest plot to display the result of a meta-analysis

Description

Draws a forest plot in the active graphics window (using grid graphics system).

Usage

```
round(x$level.comb*100), "%-CI)", sep="")
            else "Fixed effect model",
 text.random=if (x$level!=x$level.comb) paste("Random effects model (",
              round(x$level.comb*100), "%-CI)", sep="")
            else "Random effects model",
 lty.fixed=2, lty.random=3,
 prediction=x$prediction,
 text.predict=if (!(length(x$level.predict)==0) &&
                  x$level!=x$level.predict)
               paste("Prediction interval (",
                     round(x$level.predict*100), "%)",
               sep="") else "Prediction interval",
 bylab=x$bylab, print.byvar=x$print.byvar,
 text.fixed.w=text.fixed, text.random.w=text.random,
 bysort=FALSE,
 pooled.totals=comb.fixed|comb.random, pooled.events=FALSE,
 xlab="", xlab.pos=ref,
 smlab=NULL, smlab.pos=ref, xlim="symmetric",
 allstudies=TRUE,
 weight,
 pscale=1,
 ref=ifelse(backtransf & is.relative.effect(x$sm), 1, 0),
 leftcols=NULL, rightcols=NULL,
 leftlabs=NULL, rightlabs=NULL,
 lab.e=x$label.e, lab.c=x$label.c,
 lab.e.attach.to.col=NULL, lab.c.attach.to.col=NULL,
 label.right=x$label.right, label.left=x$label.left,
 lab.NA=".",
 lwd=1,
 at=NULL, label=TRUE.
 col.i="black", col.i.inside.square="white",
 col.square="gray", col.square.lines=col.square,
 col.diamond="gray",
 col.diamond.fixed=col.diamond, col.diamond.random=col.diamond,
 col.diamond.lines="black",
 col.diamond.fixed.lines=col.diamond.lines.
 col.diamond.random.lines=col.diamond.lines,
 col.predict="red", col.predict.lines="black",
 col.by="darkgray",
 print.I2=comb.fixed|comb.random, print.tau2=comb.fixed|comb.random,
 print.Q=FALSE, print.pval.Q=comb.fixed|comb.random,
 hetstat=print.I2|print.tau2|print.Q|print.pval.Q,
 overall.hetstat=overall&hetstat,
 hetlab="Heterogeneity: ",
 fontsize=12,
 fs.heading=fontsize,
fs.fixed=fontsize, fs.random=fs.fixed, fs.predict=fs.fixed, fs.study=fontsize,
fs.fixed.labels=fs.fixed, fs.random.labels=fs.random, fs.predict.labels=fs.predict,
```

```
fs.study.labels=fs.study, fs.hetstat=fontsize-2,
 fs.axis=fontsize,
 fs.smlab=fontsize, fs.xlab=fontsize, fs.lr=fontsize,
 ff.heading="bold",
ff.fixed="bold", ff.random=ff.fixed, ff.predict=ff.fixed, ff.study="plain",
ff.fixed.labels=ff.fixed, ff.random.labels=ff.random, ff.predict.labels=ff.predict,
 ff.study.labels=ff.study, ff.hetstat="bold.italic",
 ff.axis="plain",
 ff.smlab="bold", ff.xlab="plain", ff.lr="plain",
 squaresize=0.8,
 plotwidth=unit(6, "cm"),
 colgap=unit(2, "mm"),
 colgap.left=colgap, colgap.right=colgap,
 colgap.forest=colgap,
 colgap.forest.left=colgap.forest, colgap.forest.right=colgap.forest,
 just="center",
 just.studlab="left",
 addspace=TRUE,
 new=TRUE,
 backtransf=x$backtransf,
 digits=2, ...)
```

Arguments

X	An object of class meta.
sortvar	An optional vector used to sort the individual studies (must be of same length as x \$TE).
studlab	A logical indicating whether study labels should be printed in the graph. A vector with study labels can also be provided (must be of same length as x\$TE then).
level	The level used to calculate confidence intervals for individual studies.
level.comb	The level used to calculate confidence intervals for pooled estimates.
comb.fixed	A logical indicating whether fixed effect estimate should be plotted.
comb.random	A logical indicating whether random effects estimate should be plotted.
overall	A logical indicating whether overall summaries should be plotted. This argument is useful in combination with the argument byvar if summaries should only be plotted on group level.
text.fixed	A character string used in the plot to label the pooled fixed effect estimate.
text.random	A character string used in the plot to label the pooled random effects estimate.
lty.fixed	Line type (pooled fixed effect estimate).

lty.random	Line type (pooled random effects estimate).
prediction	A logical indicating whether a prediction interval should be printed.
level.predict	The level used to calculate prediction interval for a new study.
text.predict	A character string used in the plot to label the prediction interval.
bylab	A character string with a label for the grouping variable.
print.byvar	A logical indicating whether the name of the grouping variable should be printed in front of the group labels.
text.fixed.w	A character string to label the pooled fixed effect estimate within subgroups, or a character vector of same length as number of subgroups with corresponging labels.
text.random.w	A character string to label the pooled random effect estimate within subgroups, or a character vector of same length as number of subgroups with corresponding labels.
bysort	A logical indicating whether groups should be ordered alphabetically.
pooled.totals	A logical indicating whether total number of observations should be given in the figure.
pooled.events	A logical indicating whether total number of events should be given in the figure.
xlab	A label for the x-axis.
xlab.pos	A numeric specifying the center of the label on the x-axis.
smlab	A label for the summary measurex (printed at top of figure).
smlab.pos	A numeric specifying the center of the label for the summary measure.
xlim	The x limits (min,max) of the plot, or the character "s" to produce symmetric forest plots.
allstudies	A logical indicating whether studies with inestimable treatment effects should be plotted.
weight	A character string indicating which type of plotting symbols is to be used for individual treatment estimates. One of missing (see Details), "same", "fixed", or "random", can be abbreviated. Plot symbols have the same size for all studies or represent study weights from fixed effect or random effects model.
pscale	A numeric giving scaling factor for probabilities for objects of class metaprop.
ref	A numerical giving the reference value to be plotted as a line in the forest plot. No reference line is plotted if argument ref is equal to NA.
leftcols	A character vector specifying (additional) columns to be plotted on the left side of the forest plot or a logical value (see Details).
rightcols	A character vector specifying (additional) columns to be plotted on the right side of the forest plot or a logical value (see Details).
leftlabs	A character vector specifying labels for (additional) columns on left side of the forest plot (see Details).
rightlabs	A character vector specifying labels for (additional) columns on right side of the forest plot (see Details).
lab.e	Label to be used for experimental group in table heading.

lab.c Label to be used for control group in table heading.

lab.e.attach.to.col

A character specifying the column name where label lab.e should be attached to in table heading.

lab.c.attach.to.col

A character specifying the column name where label lab.c should be attached to in table heading.

label.left Graph label on left side of forest plot.
label.right Graph label on right side of forest plot.
lab.NA A character string to label missing values.

1wd The line width, see par.

at The points at which tick-marks are to be drawn, see grid.xaxis.

A logical value indicating whether to draw the labels on the tick marks, or an expression or character vector which specify the labels to use. See grid.xaxis.

col.i The colour for individual study results and confidence limits.

col.i.inside.square

The colour for individual study results and confidence limits if confidence limits are completely within squares.

col.square The colour for squares reflecting study's weight in the meta-analysis.

col.square.lines

The colour for the outer lines of squares reflecting study's weight in the metaanalysis.

col.diamond The colour of diamonds representing the results for fixed effect and random effects models.

col.diamond.fixed

The colour of diamonds for fixed effect estimates.

col.diamond.random

The colour of diamonds for random effects estimates.

col.diamond.lines

The colour of the outer lines of diamonds representing the results for fixed effect and random effects models.

col.diamond.fixed.lines

The colour of the outer lines of diamond for fixed effect estimate.

col.diamond.random.lines

The colour of the outer lines of diamond for random effects estimate.

col.predict Background colour of prediction interval.

col.predict.lines

Colour of outer lines of prediction interval.

col.by The colour to print information on subgroups.

print.12 A logical value indicating whether to print the value of the I-squared statistic.

print.tau2 A logical value indicating whether to print the value of the between-study variance tau-squared.

A logical value indicating whether to print the value of the heterogeneity statistic

print.Q

ff.predict.labels

A logical value indicating whether to print the p-value of the heterogeneity print.pval.Q statistic Q. hetstat A logical value indicating whether to print results for heterogeneity measures at all. overall.hetstat A logical value indicating whether to print results for heterogeneity measures for overall treatment comparisons. This argument is useful in combination with subgroup analyses (list object byvar) if heterogeneity statistics should only be printed on group level. hetlab Label printed in front of results for heterogeneity measures. fontsize The size of text (in points), see gpar. fs.heading The size of text for column headings, see gpar. fs.fixed The size of text for results of fixed effect model, see gpar. fs.random The size of text for results of random effects model, see gpar. fs.predict The size of text for results of prediction interval, see gpar. fs.study The size of text for results of individual studies, see gpar. fs.fixed.labels The size of text for label of fixed effect model, see gpar. fs.random.labels The size of text for label of random effects model, see gpar. fs.predict.labels The size of text for label of prediction interval, see gpar. fs.study.labels The size of text for labels of individual studies, see gpar. fs.hetstat The size of text for heterogeneity measures, see gpar. fs.axis The size of text on x-axis, see gpar. fs.smlab The size of text of label for summary measure, see gpar. fs.xlab The size of text of label on x-axis, see gpar. fs.lr The size of text of label on left and right side of forest plot, see gpar. ff.heading The fontface for column headings, see gpar. ff.fixed The fontface of text for results of fixed effect model, see gpar. ff.random The fontface of text for results of random effects model, see gpar. ff.predict The fontface of text for results of prediction interval, see gpar. ff.study The fontface of text for results of individual studies, see gpar. ff.fixed.labels The fontface of text for label of fixed effect model, see gpar. ff.random.labels

The fontface of text for label of random effects model, see gpar.

The fontface of text for label of prediction interval, see gpar.

ff.study.labels

The fontface of text for labels of individual studies, see gpar.

ff. hetstat The fontface of text for heterogeneity measures, see gpar.

ff.axis The fontface of text on x-axis, see gpar.

ff. smlab The fontface of text of label for summary measure, see gpar.

ff. xlab The fontface of text of label on x-axis, see gpar.

ff.lr The fontface of text of label on left and right side of forest plot, see gpar.

squaresize A numeric used to increase or decrease the size of squares in the forest plot.

plotwidth A unit object specifying width of the forest plot.

colgap A unit object specifying gap between columns printed on left and right side of

forest plot.

colgap.left A unit object specifying gap between columns printed on left side of forest plot.

colgap.right A unit object specifying gap between columns printed on right side of forest

plot.

colgap.forest A unit object specifying gap between column adjacent to forest plot and the

forest plot.

colgap.forest.left

A unit object specifying gap between column on the left side of forest plot and

the forest plot.

colgap.forest.right

A unit object specifying gap between column on the right side of forest plot and

the forest plot.

just Justification of text for additional columns (possible values: "left", "right", "cen-

ter").

just.studlab Justification of text for study labels (possible values: "left", "right", "center").

addspace A logical value indicating whether additional space (i.e. a blank row) is printed

above and below study results.

new A logical value indicating whether a new figure should be printed in an existing

graphics window.

backtransf A logical indicating whether results should be back transformed in forest plots.

If backtransf=TRUE, results for sm="OR" are presented as odds ratios rather than log odds ratios and results for sm="ZCOR" are presented as correlations

rather than Fisher's z transformed correlations, for example.

digits Minimal number of significant digits, see print. default.

... Additional graphical arguments (ignored at the moment).

Details

A forest plot, also called confidence interval plot, is drawn in the active graphics window. Subgroup analyses are conducted and displayed in the plot if byvar is not missing.

Note, in R package meta, version 3.0-0 the following arguments have been removed from R function forest.meta: byvar, level, level.comb, level.predict. This functionality is now provided by R function update.meta (or directly in R functions metabin, metacont, metagen, metacor, and metaprop).

The forest function is based on the grid graphics system. In order to print the forest plot, (i) resize the graphics window, (ii) either use dev.copy2eps or dev.copy2pdf.

Information from object x is utilised if argument weight is missing. Weights from the fixed effect model are used (weight="fixed") if argument x\$comb.fixed is TRUE; weights from the random effects model are used (weight="random") if argument x\$comb.random is TRUE and x\$comb.fixed is FALSE.

The arguments leftcols and rightcols can be used to specify columns which are plotted on the left and right side of the forest plot, respectively. If argument rightcols is FALSE, no columns will be plotted on the right side. By default, i.e. if arguments leftcols and rightcols are NULL, the following default columns will be plotted.

Argument rightcols: rightcols=c("effect", "ci"), i.e., estimated treatment effect and its level-confidence interval. In addition, weights of the fixed ("w.fixed") and/or random effects model ("w.random") will be given, if comb.fixed=TRUE and/or comb.random=TRUE. For an object of class metacum or metainf only the estimated treatment effect with level-confidence interval are plotted.

Argument leftcols: (i) leftcols=c("studlab", "event.e", "n.e", "event.c", "n.c") for an object of class metabin, (ii) leftcols=c("studlab", "n.e", "mean.e", "sd.e", "n.c", "mean.c", "sd.c") for an object of class metacont, (iii) leftcols=c("studlab", "TE", "seTE") for an object of class metagen, (iv) leftcols=c("studlab", "event", "n") for an object of class metaprop, (v) leftcols=c("studlab", "n") for an object of class metacor, (vi) leftcols=c("studlab") for an object of class metacum or metainf.

The arguments leftlabs and rightlabs can be used to specify column headings which are plotted on left and right side of the forest plot, respectively. For certain columns predefined labels exist. If the arguments leftlabs and rightlabs are NULL, the following default labels will be used: for columns c("studlab", "TE", "seTE", "n.e", "n.c", "event.e", "event.c", "mean.e", "mean.e", "sd.e", "s the labels c("Study", "TE", "seTE", "Total", "Total", "Events", "Events", "Mean", "Mean", "SD", "SD", s For additional columns the column name will be used as label. It is possible to only provide labels for new columns (see Examples). Otherwise the length of leftlabs and rightlabs must be the same as the number of printed columns. The value NA can be used to specify columns using the default labels (see Example).

If arguments lab.e and lab.c are NULL, "Experimental" and "Control" are used as labels for experimental and control group, respectively.

The arguments pscale can be used to rescale proportions for objects of class metaprop, e.g. pscale=100 means that proportions are expressed per 100 observations. This is useful in situations with (very) low proportions. For pscale=100, column heading and x-axis label are changed to "Prop (in %)" and "Proportion (in %)", respectively.

A prediction interval for treatment effect of a new study (Higgins et al., 2009) is given in the forest plot if arguments prediction and comb.random are TRUE. For graphical presentation of prediction intervals the approach by Guddat et al. (2012) is used.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Guddat C, Grouven U, Bender R, Skipka G 2012), A note on the graphical presentation of prediction intervals in random-effects meta-analyses. *Systematic Reviews*, **1**, 34.

Higgins JPT, Thompson SG, Spiegelhalter DJ (2009), A re-evaluation of random-effects meta-analysis. *Journal of the Royal Statistical Society: Series A*, **172**, 137-159.

See Also

```
metabin, metacont, metagen
```

Examples

```
data(01kin95)
meta1 <- metabin(event.e, n.e, event.c, n.c,</pre>
                 data=01kin95, subset=c(41,47,51,59),
                 sm="RR", method="I",
                 studlab=paste(author, year))
# Do (symmetric) forest plot
forest(meta1)
# Forest plot specifying argument xlim
forest(meta1, xlim=c(0.01, 10))
# Add prediction interval to forest plot
forest(meta1, prediction=TRUE)
# Forest plot with 'classic' layout used in
# R package meta, version < 1.6-0
forest(meta1, col.square="black", hetstat=FALSE)
# Change set of columns printed on left side
# of forest plot
forest(meta1, comb.random=FALSE,
       leftcols="studlab")
```

```
# Do not print columns on right side of forest plot
forest(meta1, rightcols=FALSE)
# Change study label to "Author"
forest(meta1, comb.random=FALSE,
       leftlabs=c("Author", NA, NA, NA, NA))
#
# Just give effect estimate and 95% confidence interval
# on right side of forest plot
forest(meta1, rightcols=c("effect", "ci"))
# 1. Change order of columns on left side
# 2. Attach labels to columns 'event.e' and 'event.c'
     instead of columns 'n.e' and 'n.c'
forest(meta1,
       leftcols=c("studlab", "n.e", "event.e", "n.c", "event.c"),
       lab.e.attach.to.col="event.e",
       lab.c.attach.to.col="event.c")
Olkin95$studlab <- paste(Olkin95$author, Olkin95$year)</pre>
# Add variables 'year' and 'author' to meta-analysis object
meta1$year <- addvar(meta1, Olkin95, "year")</pre>
meta1$author <- addvar(meta1, Olkin95, "author")</pre>
# Specify column labels only for newly created variables
# 'year' and 'author'
#
forest(meta1,
       leftcols=c("studlab", "event.e", "n.e", "event.c", "n.c",
                  "author", "year"),
       leftlabs=c("Author", "Year of Publ"))
# Change some fontsizes and fontfaces
forest(meta1,
```

```
fs.study=10, ff.study="italic",
      fs.study.label=11, ff.study.label="bold",
      fs.axis=5, ff.axis="italic",
      ff.smlab="bold.italic",
      ff.fixed="plain", ff.hetstat="plain")
# Change some colours
forest(meta1,
      col.diamond="green", col.diamond.lines="red",
      col.i=c("green", "blue", "red", "orange"),
      col.square="pink", col.square.lines="black")
# Sort by weight in fixed effect model
forest(meta1, sortvar=1/w.fixed, comb.random=FALSE)
#
# Sort by decreasing weight in fixed effect model
forest(meta1, sortvar=rev(1/w.fixed), comb.random=FALSE)
#
# Sort by size of treatment effect
forest(meta1, sortvar=TE, comb.random=FALSE)
# Sort by size of treatment effect
forest(meta1, sortvar=-TE, comb.random=FALSE)
# Sort by size of treatment effect
forest(meta1, sortvar=rev(TE), comb.random=FALSE)
# Sort by decreasing year of publication
forest(meta1, sortvar=rev(year), comb.random=FALSE)
#
```

funnel

Plot to assess funnel plot asymmetry

Description

Draw a funnel plot or radial plot (also called Galbraith plot) to assess funnel plot asymmetry in the active graphics window.

A contour-enhanced funnel plot can be produced for assessing causes of funnel plot asymmetry.

Usage

```
funnel(x, ...)
radial(x, ...)
## Default S3 method:
funnel(x, y,
       xlim=NULL, ylim=NULL, xlab=NULL, ylab=NULL,
       comb.fixed=FALSE, comb.random=FALSE,
       axes=TRUE,
       pch=21, text=NULL, cex=1,
       lty.fixed=2, lty.random=9,
       lwd=1, lwd.fixed=lwd, lwd.random=lwd,
       col="black", bg="darkgray",
       col.fixed="black", col.random="black",
       log="", yaxis="se", sm=NULL,
       contour.levels=NULL, col.contour,
       ref=ifelse(backtransf & is.relative.effect(sm), 1, 0),
       level=NULL,
       studlab=FALSE, cex.studlab=0.8, backtransf=TRUE, ...)
## S3 method for class 'meta'
funnel(x,
       xlim=NULL, ylim=NULL, xlab=NULL, ylab=NULL,
       comb.fixed=x$comb.fixed, comb.random=x$comb.random,
       axes=TRUE,
       pch=if (!inherits(x, "trimfill")) 21 else ifelse(x$trimfill, 1, 21),
       text=NULL, cex=1,
       lty.fixed=2, lty.random=9,
       lwd=1, lwd.fixed=lwd, lwd.random=lwd,
       col="black", bg="darkgray",
```

```
col.fixed="black", col.random="black",
       log="", yaxis="se",
       contour.levels=NULL, col.contour,
       ref=ifelse(backtransf & is.relative.effect(x$sm), 1, 0),
       level=x$level,
       studlab=FALSE, cex.studlab=0.8, backtransf=x$backtransf, ...)
## Default S3 method:
radial(x, y, xlim=NULL, ylim=NULL,
       xlab="Inverse of standard error",
      ylab="Standardised treatment effect (z-score)",
       comb.fixed=TRUE, axes=TRUE,
       pch=1, text=NULL, cex=1, col=NULL,
       level=NULL, ...)
## S3 method for class 'meta'
radial(x, xlim=NULL, ylim=NULL,
       xlab="Inverse of standard error",
       ylab="Standardised treatment effect (z-score)",
       comb.fixed=TRUE, axes=TRUE,
       pch=1, text=NULL, cex=1, col=NULL,
       level=NULL, ...)
```

Arguments

x	An object of class meta, or estimated treatment effect in individual studies.
У	Standard error of estimated treatment effect.
xlim	The x limits (min,max) of the plot.
ylim	The y limits (min,max) of the plot.
xlab	A label for the x-axis.
ylab	A label for the y-axis.
comb.fixed	A logical indicating whether the pooled fixed effect estimate should be plotted.
comb.random	A logical indicating whether the pooled random effects estimate should be plotted.
axes	A logical indicating whether axes should be drawn on the plot.
pch	The plotting symbol used for individual studies.
text	A character vector specifying the text to be used instead of plotting symbol.
cex	The magnification to be used for plotting symbol.
lty.fixed	Line type (pooled fixed effect estimate).
lty.random	Line type (pooled random effects estimate).
col	A vector with colour of plotting symbols.
bg	A vector with background colour of plotting symbols (only used if pch in 21:25).
col.fixed	Color of line representign fixed effect estimate.
col.random	Color of line representign random effects estimate.

lwd The line width for confidence intervals (if level is not NULL). lwd.fixed The line width for fixed effect estimate (if comb. fixed is not NULL). lwd.random The line width for random effects estimate (if comb.random is not NULL). A character string which contains "x" if the x-axis is to be logarithmic, "y" if the log y-axis is to be logarithmic and "xy" or "yx" if both axes are to be logarithmic (applies only to function funnel). A character string indicating which type of weights are to be used. Either "se", yaxis "invvar", "invse", or "size" (applies only to function funnel). sm A character string indicating underlying summary measure, e.g., "RD", "RR", "OR", "AS", "MD", "SMD" (applies only to function funnel). contour.levels A numeric vector specifying contour levels to produce contour-enhanced funnel plot. col.contour Colour of contours. ref Reference value (null effect) used to produce contour-enhanced funnel plot. level The confidence level utilised in the plot. For the funnel plot, confidence limits are not drawn if yaxis="size". A logical indicating whether study labels should be printed in the graph. A studlab vector with study labels can also be provided (must be of same length as x\$TE then). cex.studlab Size of study labels. backtransf A logical indicating whether results for relative summary measures (argument sm equal to "OR", "RR", "HR", or "IRR") should be back transformed in funnel plots. If backtransf=TRUE, results for sm="OR" are printed as odds ratios rather

Graphical arguments as in par may also be passed as arguments.

than log odds ratios, for example.

Details

A funnel plot or radial plot, also called Galbraith plot, is drawn in the active graphics window. If comb.fixed is TRUE, the pooled estimate of the fixed effect model is plotted. If level is not NULL, the corresponding confidence limits are drawn.

In the funnel plot, if yaxis is "se", the standard error of the treatment estimates is plotted on the y-axis which is likely to be the best choice (Sterne & Egger, 2001). Other possible choices for yaxis are "invvar" (inverse of the variance), "invse" (inverse of the standard error), and "size" (study size).

For yaxis!="size", contour-enhanced funnel plots can be produced (Peters et al., 2008) by specifying the contour levels (argument contour.levels). By default (argument col.contour missing), suitable gray levels will be used to distinguish the contours. Different colours can be chosen by argument col.contour.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>, Petra Graham <pgraham@efs.mq.edu.au>

References

Galbraith RF (1988a), Graphical display of estimates having differing standard errors. *Technometrics*, **30**, 271–281.

Galbraith RF (1988b), A note on graphical presentation of estimated odds ratios from several clinical trials. *Statistics in Medicine*, **7**, 889–894.

Light RJ & Pillemer DB (1984), Summing Up. The Science of Reviewing Research. Cambridge: Harvard University Press.

Peters JL, Sutton AJ, Jones DR, Abrams KR, Rushton L (2008), Contour-enhanced meta-analysis funnel plots help distinguish publication bias from other causes of asymmetry. *Journal of Clinical Epidemiology*, **61**, 991–996.

Sterne JAC & Egger M (2001), Funnel plots for detecting bias in meta-analysis: Guidelines on choice of axis. *Journal of Clinical Epidemiology*, **54**, 1046–1055.

See Also

metabias, metabin, metagen

Examples

```
data(01kin95)
meta1 <- metabin(event.e, n.e, event.c, n.c,</pre>
                  data=01kin95, subset=c(41,47,51,59),
                  studlab=paste(author, year),
                  sm="RR", method="I")
# Radial plot
radial(meta1, level=0.95)
oldpar <- par(mfrow=c(2, 2))</pre>
# Funnel plots
funnel(meta1)
# Same result as code above:
funnel(meta1$TE, meta1$seTE, sm="RR")
# Funnel plot with confidence intervals,
# fixed effect estimate and contours
cc <- funnel(meta1, comb.fixed=TRUE,</pre>
             level=0.95, contour=c(0.9, 0.95, 0.99))$col.contour
legend(0.05, 0.05,
```

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labbe

L'Abbe plot for meta-analysis with binary outcomes

Description

Draw a L'Abbé plot for meta-analysis with binary outcomes.

Usage

```
labbe(x, ...)
## Default S3 method:
labbe(x, y,
      xlim, ylim,
      xlab=NULL, ylab=NULL,
      TE.fixed, TE.random,
      comb.fixed=FALSE, comb.random=FALSE,
      axes=TRUE,
      pch=21, text=NULL, cex=1,
      col="black", bg="lightgray",
      lwd=1, lwd.fixed=lwd, lwd.random=lwd,
      lty.fixed=2, lty.random=9,
      sm=NULL, weight,
      studlab=FALSE, cex.studlab=0.8,
      ...)
## S3 method for class 'metabin'
labbe(x,
      xlim, ylim,
      xlab=NULL, ylab=NULL,
      TE.fixed=x$TE.fixed,
      TE.random=x$TE.random,
      comb.fixed=x$comb.fixed,
      comb.random=x$comb.random,
```

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```
axes=TRUE,
pch=21, text=NULL, cex=1,
col="black", bg="lightgray",
lwd=1, lwd.fixed=lwd, lwd.random=lwd,
lty.fixed=2, lty.random=9,
sm=x$sm, weight,
studlab=FALSE, cex.studlab=0.8, ...)
```

Arguments

Х The x coordinates of points of the L'Abbé plot. Alternatively, an object of class metabin. The y coordinates of the L'Abbé plot, optional if x is an appropriate structure. У xlim The x limits (min,max) of the plot. ylim The y limits (min,max) of the plot. xlab A label for the x-axis. vlab A label for the y-axis. TE.fixed A numeric or vector specifying combined fixed effect estimate(s). TF.random A numeric or vector specifying combined random effects estimate(s). comb.fixed A logical indicating whether the pooled fixed effect estimate should be plotted. A logical indicating whether the pooled random effects estimate should be plotcomb.random A logical indicating whether axes should be drawn on the plot. axes pch The plotting symbol used for individual studies. A character vector specifying the text to be used instead of plotting symbol. text The magnification to be used for plotting symbol. cex col A vector with colour of plotting symbols. A vector with background colour of plotting symbols (only used if pch in 21:25). bg The line width. lwd lwd.fixed The line width for fixed effect estimate (if comb. fixed is not NULL or FALSE). lwd.random The line width for random effects estimate (if comb.random is not NULL or FALSE). lty.fixed Line type (pooled fixed effect estimate). lty.random Line type (pooled random effects estimate). A character string indicating underlying summary measure, i.e., "RD", "RR", "OR". weight Either a numeric vector specifying relative sizes of plotting symbols or a character string indicating which type of plotting symbols is to be used for individual treatment estimates. One of missing (see Details), "same", "fixed", or

"random", can be abbreviated. Plot symbols have the same size for all studies

or represent study weights from fixed effect or random effects model.

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studlab A logical indicating whether study labels should be printed in the graph. A vec-

tor with study labels can also be provided (must be of same length as x\$event.e

then).

cex.studlab Size of study labels.

... Graphical arguments as in par may also be passed as arguments.

Details

A L'Abbé plot is a scatter plot with the risk in the control group on the x-axis and the risk in the experimental group on the y-axis (L'Abbé et al., 1987). It can be used to evaluate heterogeneity in meta-analysis. Furthermore, this plot can aid to choose a summary measure (odds ratio, risk ratio, risk difference) that will result in more consistent results.

If comb.fixed is TRUE, the pooled estimate of the fixed effect model is plotted as a line. If comb.random is TRUE, the pooled estimate of the random effects model is plotted as a line.

Information from object x is utilised if argument weight is missing. Weights from the fixed effect model are used (weight="fixed") if argument x\$comb.fixed is TRUE; weights from the random effects model are used (weight="random") if argument x\$comb.random is TRUE and x\$comb.fixed is FALSE.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

L'Abbé KA, Detsky AS, O'Rourke K (1987), Meta-analysis in clinical research. *Annals of Internal Medicine*, **107**, 224–233.

See Also

metabin

Examples

32 metabias

metabias	Test for funnel plot asymmetry	

Description

Test for funnel plot asymmetry, based on rank correlation or linear regression method.

Usage

Arguments

X	An object of class meta or estimated treatment effect in individual studies.
seTE	Standard error of estimated treatment effect (mandatory if x not of class meta).
method.bias	A character string indicating which test is to be used. Either "rank", "linreg", "mm", "count", "score", or "peters", can be abbreviated.
plotit	A logical indicating whether a plot should be produced for method.bias "rank", "linreg", "mm", or "score".
correct	A logical indicating whether a continuity corrected statistic is used for rank correlation methods "rank" and "count".
k.min	Minimum number of studies to perform test for funnel plot asymmetry.
	Additional arguments (ignored at the moment).

Details

Following recommendations by Sterne et al. (2011), by default, a test for funnel plot asymmetry is only conducted if the number of studies is ten or larger (argument k.min=10). This behaviour can be changed by setting a smaller value for argument k.min. Note, the minimum number of studies is three.

If argument method.bias is "rank", the test statistic is based on the rank correlation between standardised treatment estimates and variance estimates of estimated treatment effects; Kendall's tau is used as correlation measure (Begg & Mazumdar, 1994). The test statistic follows a standard normal distribution. By default (if correct is FALSE), no continuity correction is utilised (Kendall & Gibbons, 1990).

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If argument method.bias is "linreg", the test statistic is based on a weighted linear regression of the treatment effect on its standard error (Egger et al., 1997). The test statistic follows at distribution with number of studies - 2 degrees of freedom.

If argument method.bias is "mm", the test statistic is based on a weighted linear regression of the treatment effect on its standard error using the method of moments estimator for the additive between-study variance component (method 3a in Thompson, Sharp, 1999). The test statistic follows at distribution with number of studies - 2 degrees of freedom.

If argument method.bias is "count", the test statistic is based on the rank correlation between a standardised cell frequency and the inverse of the variance of the cell frequency; Kendall's tau is used as correlation measure (Schwarzer et al., 2007). The test statistic follows a standard normal distribution. By default (if correct is FALSE), no continuity correction is utilised (Kendall & Gibbons, 1990).

If argument method.bias is "score", the test statistic is based on a weighted linear regression utilising efficient score and score variance (Harbord et al., 2006). The test statistic follows a t distribution with number of studies - 2 degrees of freedom.

If argument method.bias is "peters", the test statistic is based on a weighted linear regression of the treatment effect on the inverse of the total sample size using the variance of the average event rate as weights (Peters et al., 2006). The test statistic follows a t distribution with number of studies - 2 degrees of freedom.

In order to calculate an arcsine test for funnel plot asymmetry (Ruecker et al., 2008), one has to use the metabin function with argument sm="AS" as input to the metabias command. The three arcsine tests described in Ruecker et al. (2008) can be calculated by setting method.bias to "rank", "linreg" and "mm", respectively.

If argument method.bias is missing, the Harbord test (method.bias="score") is used for the odds ratio as effect measure and the Egger test (method.bias="linreg") for other effect measures (Sterne et al., 2011).

No test for funnel plot asymmetry is conducted in meta-analyses with subgroups.

Value

A list with class "htest" containing the following components if a test for funnel plot asymmetry is conducted:

estimate The estimated degree of funnel plot asymmetry, with name "ks" or "bias" cor-

responding to the method employed, i.e., rank correlation or regression method.

statistic The value of the test statistic.

parameters The degrees of freedom of the test statistic in the case that it follows a t distri-

bution.

p.value The p-value for the test.

alternative A character string describing the alternative hypothesis.

Method A character string indicating what type of test was used.

data.name A character string giving the names of the data.

title Title of Cochrane review.

complab Comparison label.

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outclab Outcome label.

version Version of R package meta used to create object.

Or a list with the following elements if test is not conducted due to the number of studies:

k Number of studies in meta-analysis.

k.min Minimum number of studies to perform test for funnel plot asymmetry.

version Version of R package meta used to create object.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Begg CB & Mazumdar M (1994), Operating characteristics of a rank correlation test for publication bias. *Biometrics*, **50**, 1088–1101.

Egger M, Smith GD, Schneider M & Minder C (1997), Bias in meta-analysis detected by a simple, graphical test. *British Medical Journal*, **315**, 629–634.

Harbord RM, Egger M & Sterne J (2006), A modified test for small-study effects in meta-analyses of controlled trials with binary endpoints. *Statistics in Medicine*, **25**, 3443–3457.

Kendall M & Gibbons JD (1990), Rank Correlation Methods. London: Edward Arnold.

Peters JL, Sutton AJ, Jones DR, Abrams KR & Rushton L (2006), Comparison of two methods to detect publication bias in meta-analysis. *Journal of the American Medical Association*, **295**, 676–680.

Ruecker G, Schwarzer G, Carpenter JR (2008) Arcsine test for publication bias in meta-analyses with binary outcomes. *Statistics in Medicine*, **27**,746–763.

Schwarzer G, Antes G & Schumacher M (2007), A test for publication bias in meta-analysis with sparse binary data. *Statistics in Medicine*, **26**, 721–733.

Sterne, JAC et al. (2011), Recommendations for Examining and Interpreting Funnel Plot Asymmetry in Meta-Analyses of Randomised Controlled Trials. *BMJ (Clinical research ed.)*, **343**, 1, doi: 10.1136/bmj.d4002.

Thompson SG & Sharp, SJ (1999), Explaining heterogeneity in meta-analysis: A comparison of methods, *Statistics in Medicine*, **18**, 2693–2708.

See Also

```
funnel, funnel.meta, metabin, metacont, metagen
```

Examples

metabin 35

```
metabias(meta1, plotit=TRUE)
metabias(meta1, method.bias="rank")
metabias(meta1, method.bias="rank", correct=TRUE)
metabias(meta1, method.bias="count")
metabias(meta1, method.bias="linreg")$p.value
# Arcsine test (based on linear regression):
meta1.as <- metabin(event.e, n.e, event.c, n.c,</pre>
                     data=Olkin95, subset=1:10,
                     sm="AS", method="I")
metabias(meta1.as)
# Same result (using function metabias.default):
metabias(meta1.as$TE, meta1.as$seTE)
# No test for funnel plot asymmetry calculated:
meta2 <- metabin(event.e, n.e, event.c, n.c,</pre>
                 data=Olkin95, subset=1:5,
                 sm="RR", method="I")
metabias(meta2)
meta3 <- metabin(event.e, n.e, event.c, n.c,</pre>
                 data=Olkin95, subset=1:2,
                 sm="RR", method="I")
metabias(meta3)
# Test for funnel plot asymmetry calculated
# (use of argument k.min):
metabias(meta2, k.min=5)
```

metabin

Meta-analysis of binary outcome data

Description

Calculation of fixed and random effects estimates (risk ratio, odds ratio, risk difference or arcsine difference) for meta-analyses with binary outcome data. Mantel-Haenszel, inverse variance and Peto method are available for pooling.

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Usage

```
metabin(event.e, n.e, event.c, n.c, studlab,
       data=NULL, subset=NULL,
       method=ifelse(tau.common, "Inverse", .settings$method),
        ifelse(!is.na(charmatch(method, c("Peto", "peto"),
                                nomatch = NA)),
               "OR", .settings$smbin),
        incr=.settings$incr, allincr=.settings$allincr,
       addincr=.settings$addincr, allstudies=.settings$allstudies,
       MH.exact=.settings$MH.exact, RR.cochrane=.settings$RR.cochrane,
       level=.settings$level, level.comb=.settings$level.comb,
       comb.fixed=.settings$comb.fixed, comb.random=.settings$comb.random,
       hakn=.settings$hakn,
       method.tau=.settings$method.tau, tau.preset=NULL, TE.tau=NULL,
       tau.common=.settings$tau.common,
       prediction=.settings$prediction, level.predict=.settings$level.predict,
       method.bias=ifelse(sm=="OR", "score", .settings$method.bias),
       backtransf=.settings$backtransf,
       title=.settings$title, complab=.settings$complab, outclab="",
       label.e=.settings$label.e, label.c=.settings$label.c,
       label.left=.settings$label.left, label.right=.settings$label.right,
       byvar, bylab, print.byvar=.settings$print.byvar,
       print.CMH=.settings$print.CMH,
       keepdata=.settings$keepdata,
       warn=.settings$warn)
```

Arguments

event.e	Number of events in experimental group.
n.e	Number of observations in experimental group.
event.c	Number of events in control group.
n.c	Number of observations in control group.
studlab	An optional vector with study labels.
data	An optional data frame containing the study information, i.e., event.e, n.e, event.c, and n.c.
subset	An optional vector specifying a subset of studies to be used.
method	A character string indicating which method is to be used for pooling of studies. One of "Inverse", "MH", or "Peto", can be abbreviated.
sm	A character string indicating which summary measure ("RR", "OR", "RD", or "AS") is to be used for pooling of studies, see Details.
incr	Could be either a numerical value which is added to each cell frequency for studies with a zero cell count or the character string "TA" which stands for treatment arm continuity correction, see Details.

allincr A logical indicating if incr is added to each cell frequency of all studies if at least one study has a zero cell count. If FALSE (default), incr is added only to each cell frequency of studies with a zero cell count. addincr A logical indicating if incr is added to each cell frequency of all studies irrespective of zero cell counts. allstudies A logical indicating if studies with zero or all events in both groups are to be included in the meta-analysis (applies only if sm is equal to "RR" or "OR"). MH.exact A logical indicating if incr is not to be added to all cell frequencies for studies with a zero cell count to calculate the pooled estimate based on the Mantel-Haenszel method. A logical indicating if 2*incr instead of 1*incr is to be added to n.e and n.c RR.cochrane in the calculation of the risk ratio (i.e., sm="RR") for studies with a zero cell. This is used in RevMan 5, the Cochrane Collaboration's program for preparing and maintaining Cochrane reviews. level The level used to calculate confidence intervals for individual studies. level.comb The level used to calculate confidence intervals for pooled estimates. comb.fixed A logical indicating whether a fixed effect meta-analysis should be conducted. comb.random A logical indicating whether a random effects meta-analysis should be conducted. prediction A logical indicating whether a prediction interval should be printed. level.predict The level used to calculate prediction interval for a new study. hakn A logical indicating whether the method by Hartung and Knapp should be used to adjust test statistics and confidence intervals. method.tau A character string indicating which method is used to estimate the betweenstudy variance τ^2 . Either "DL", "PM", "REML", "ML", "HS", "SJ", "HE", or "EB", can be abbreviated. Prespecified value for the square-root of the between-study variance τ^2 . tau.preset Overall treatment effect used to estimate the between-study variance τ^2 . TE.tau A logical indicating whether tau-squared should be the same across subgroups. tau.common method.bias A character string indicating which test for funnel plot asymmetry is to be used. Either "rank", "linreg", "mm", "count", "score", or "peters", can be abbreviated. See function metabias backtransf A logical indicating whether results for odds ratio (sm="OR") and risk ratio (sm="RR") should be back transformed in printouts and plots. If TRUE (default), results will be presented as odds ratios and risk ratios; otherwise log odds ratios and log risk ratios will be shown. Title of meta-analysis / systematic review. title complab Comparison label. outclab Outcome label. label.e Label for experimental group. label.c Label for control group.

label.left Graph label on left side of forest plot. label.right Graph label on right side of forest plot. An optional vector containing grouping information (must be of same length as byvar event.e). bylab A character string with a label for the grouping variable. A logical indicating whether the name of the grouping variable should be printed print.byvar in front of the group labels. print.CMH A logical indicating whether result of the Cochran-Mantel-Haenszel test for overall effect should be printed.

A logical indicating whether original data (set) should be kept in meta object.

keepdata A logical indicating whether warnings should be printed (e.g., if incr is added warn

to studies with zero cell frequencies).

Details

Treatment estimates and standard errors are calculated for each study. The following measures of treatment effect are available:

• Risk ratio (sm="RR")

Odds ratio (sm="OR")

• Risk difference (sm="RD")

• Arcsine difference (sm="AS")

For studies with a zero cell count, by default, 0.5 is added to all cell frequencies of these studies; if incr is "TA" a treatment arm continuity correction is used instead (Sweeting et al., 2004; Diamond et al., 2007). Treatment estimates and standard errors are only calculated for studies with zero or all events in both groups if allstudies is TRUE.

For several arguments defaults settings are utilised (assignments with .settings\$). These defaults can be changed using the settings.meta function.

Internally, both fixed effect and random effects models are calculated regardless of values choosen for arguments comb.fixed and comb.random. Accordingly, the estimate for the random effects model can be extracted from component TE.random of an object of class "meta" even if argument comb, random=FALSE. However, all functions in R package meta will adequately consider the values for comb. fixed and comb. random. E.g. function print.meta will not print results for the random effects model if comb.random=FALSE.

By default, both fixed effect and random effects models are considered (see arguments comb. fixed and comb. random). If method is "MH" (default), the Mantel-Haenszel method is used to calculate the fixed effect estimate; if method is "Inverse", inverse variance weighting is used for pooling; finally, if method is "Peto", the Peto method is used for pooling. By default, the DerSimonian-Laird estimate (1986) is used in the random effects model (method.tau="DL"). For the Peto method, Peto's log odds ratio, i.e. (0-E)/V and its standard error sqrt(1/V) with 0-E and V denoting "Observed minus Expected" and "V", are utilised in the random effects model. Accordingly, results of a random effects model using sm="Peto" can be (slightly) different to results from a random effects model using sm="MH" or sm="Inverse".

For the Mantel-Haenszel method, by default (if MH. exact is FALSE), 0.5 is added to all cell frequencies of a study with a zero cell count in the calculation of the pooled risk ratio or odds ratio

as well as the estimation of the variance of the pooled risk difference, risk ratio or odds ratio. This approach is also used in other software, e.g. RevMan 5 and the Stata procedure metan. According to Fleiss (in Cooper & Hedges, 1994), there is no need to add 0.5 to a cell frequency of zero to calculate the Mantel-Haenszel estimate and he advocates the exact method (MH. exact=TRUE). Note, the estimate based on the exact method is not defined if the number of events is zero in all studies either in the experimental or control group.

A prediction interval for treatment effect of a new study is calculated (Higgins et al., 2009) if arguments prediction and comb.random are TRUE.

R function update.meta can be used to redo the meta-analysis of an existing metabin object by only specifying arguments which should be changed.

For the random effects, the method by Hartung and Knapp (2003) is used to adjust test statistics and confidence intervals if argument hakn=TRUE.

The iterative Paule-Mandel method (1982) to estimate the between-study variance is used if argument method. tau="PM". Internally, R function paulemandel is called which is based on R function mpaule.default from R package metRology from S.L.R. Ellison <s.ellison at lgc.co.uk>.

If R package metafor (Viechtbauer 2010) is installed, the following methods to estimate the betweenstudy variance τ^2 (argument method. tau) are also available:

- Restricted maximum-likelihood estimator (method.tau="REML")
- Maximum-likelihood estimator (method.tau="ML")
- Hunter-Schmidt estimator (method.tau="HS")
- Sidik-Jonkman estimator (method.tau="SJ")
- Hedges estimator (method.tau="HE")
- Empirical Bayes estimator (method.tau="EB").

For these methods the R function rma.uni of R package metafor is called internally. See help page of R function rma.uni for more details on these methods to estimate between-study variance.

Value

An object of class c("metabin", "meta") with corresponding print, summary, plot function. The object is a list containing the following components:

```
event.e, n.e, event.c, n.c, studlab,

sm, method, incr, allincr, addincr,

allstudies, MH.exact, RR.cochrane, warn,

level, level.comb, comb.fixed, comb.random,

hakn, method.tau, tau.preset, TE.tau, method.bias,

tau.common, title, complab, outclab,

label.e, label.c, label.left, label.right,
```

byvar, bylab, print.byvar

As defined above.

TE, seTE Estimated treatment effect and standard error of individual studies.

Lower and upper confidence interval limits for individual studies.

zval, pval z-value and p-value for test of treatment effect for individual studies.

w.fixed, w.random

Weight of individual studies (in fixed and random effects model).

TE.fixed, seTE.fixed

Estimated overall treatment effect and standard error (fixed effect model).

lower.fixed, upper.fixed

Lower and upper confidence interval limits (fixed effect model).

zval.fixed, pval.fixed

z-value and p-value for test of overall treatment effect (fixed effect model).

TE.random, seTE.random

Estimated overall treatment effect and standard error (random effects model).

lower.random, upper.random

Lower and upper confidence interval limits (random effects model).

zval.random, pval.random

z-value or t-value and corresponding p-value for test of overall treatment effect (random effects model).

prediction, level.predict

As defined above.

seTE.predict Standard error utilised for prediction interval.

lower.predict, upper.predict

Lower and upper limits of prediction interval.

k Number of studies combined in meta-analysis.

Q Heterogeneity statistic Q.

df.Q Degrees of freedom for heterogeneity statistic.

tau Square-root of between-study variance.

se.tau Standard error of square-root of between-study variance.

C Scaling factor utilised internally to calculate common tau-squared across sub-

groups.

Q. CMH Cochran-Mantel-Haenszel test statistic for overall effect.

incr.e, incr.c Increment added to cells in the experimental and control group, respectively.

sparse Logical flag indicating if any study included in meta-analysis has any zero cell

frequencies.

df.hakn Degrees of freedom for test of treatment effect for Hartung-Knapp method (only

if hakn=TRUE).

keepdata As defined above.

data Original data (set) used in function call (if keepdata=TRUE).

subset Information on subset of original data used in meta-analysis (if keepdata=TRUE).

call Function call.

version Version of R package meta used to create object.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

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Sweeting MJ, Sutton AJ, Lambert PC (2004), What to add to nothing? Use and avoidance of continuity corrections in meta-analysis of sparse data. *Statistics in Medicine*, **23**, 1351–1375.

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See Also

```
update.meta, funnel, metabias, metacont, metagen, print.meta
```

Examples

```
metabin(10, 20, 15, 20, sm="OR", warn=FALSE)
```

metacont

Meta-analysis of continuous outcome data

Description

Calculation of fixed and random effects estimates for meta-analyses with continuous outcome data; inverse variance weighting is used for pooling.

Usage

```
metacont(n.e, mean.e, sd.e, n.c, mean.c, sd.c, studlab,
         data=NULL, subset=NULL,
         sm=.settings$smcont, pooledvar=.settings$pooledvar,
         level=.settings$level, level.comb=.settings$level.comb,
         comb.fixed=.settings$comb.fixed, comb.random=.settings$comb.random,
         hakn=.settings$hakn,
         method.tau=.settings$method.tau, tau.preset=NULL, TE.tau=NULL,
         tau.common=.settings$tau.common,
        prediction=.settings$prediction, level.predict=.settings$level.predict,
         method.bias=.settings$method.bias,
         title=.settings$title, complab=.settings$complab, outclab="",
         label.e=.settings$label.e, label.c=.settings$label.c,
         label.left=.settings$label.left, label.right=.settings$label.right,
         byvar, bylab, print.byvar=.settings$print.byvar,
         keepdata=.settings$keepdata,
         warn=.settings$warn)
```

Arguments

	Number of observations in experimental group.
n.e	
mean.e	Estimated mean in experimental group.
sd.e	Standard deviation in experimental group.
n.c	Number of observations in control group.
mean.c	Estimated mean in control group.
sd.c	Standard deviation in control group.
studlab	An optional vector with study labels.
data	An optional data frame containing the study information.
subset	An optional vector specifying a subset of studies to be used.
level	The level used to calculate confidence intervals for individual studies.
level.comb	The level used to calculate confidence intervals for pooled estimates.
comb.fixed	A logical indicating whether a fixed effect meta-analysis should be conducted.
comb.random	A logical indicating whether a random effects meta-analysis should be conducted.
prediction	A logical indicating whether a prediction interval should be printed.
level.predict	The level used to calculate prediction interval for a new study.
hakn	A logical indicating whether the method by Hartung and Knapp should be used to adjust test statistics and confidence intervals.
method.tau	A character string indicating which method is used to estimate the between-study variance τ^2 . Either "DL", "PM", "REML", "ML", "HS", "SJ", "HE", or "EB", can be abbreviated.
tau.preset	Prespecified value for the square-root of the between-study variance τ^2 .
TE.tau	Overall treatment effect used to estimate the between-study variance tau-squared.
tau.common	A logical indicating whether tau-squared should be the same across subgroups.
method.bias	A character string indicating which test is to be used. Either "rank", "linreg", or "mm", can be abbreviated. See function metabias
title	Title of meta-analysis / systematic review.
complab	Comparison label.
outclab	Outcome label.
label.e	Label for experimental group.
label.c	Label for control group.
label.left	Graph label on left side of forest plot.
label.right	Graph label on right side of forest plot.
sm	A character string indicating which summary measure ("MD" or "SMD") is to be used for pooling of studies.
pooledvar	A logical indicating if a pooled variance should be used for the mean difference $(sm="MD")$.

byvar An optional vector containing grouping information (must be of same length as

n.e).

bylab A character string with a label for the grouping variable.

print.byvar A logical indicating whether the name of the grouping variable should be printed

in front of the group labels.

keepdata A logical indicating whether original data (set) should be kept in meta object.

warn A logical indicating whether warnings should be printed (e.g., if studies are

excluded from meta-analysis due to zero standard deviations).

Details

Calculation of fixed and random effects estimates for meta-analyses with continuous outcome data; inverse variance weighting is used for pooling. By default, the DerSimonian-Laird estimate (1986) is used in the random effects model (method.tau="DL").

The mean difference is used as measure of treatment effect if sm="MD" – which correspond to sm="WMD" in older versions (<0.9) of the meta package. For the summary measure "SMD", Hedges' adjusted g is utilised for pooling.

For several arguments defaults settings are utilised (assignments with .settings\$). These defaults can be changed using the settings.meta function.

Internally, both fixed effect and random effects models are calculated regardless of values choosen for arguments comb.fixed and comb.random. Accordingly, the estimate for the random effects model can be extracted from component TE.random of an object of class "meta" even if argument comb.random=FALSE. However, all functions in R package meta will adequately consider the values for comb.fixed and comb.random. E.g. function print.meta will not print results for the random effects model if comb.random=FALSE.

The function metagen is called internally to calculate individual and overall treatment estimates and standard errors.

A prediction interval for treatment effect of a new study is calculated (Higgins et al., 2009) if arguments prediction and comb.random are TRUE.

R function update.meta can be used to redo the meta-analysis of an existing metacont object by only specifying arguments which should be changed.

For the random effects, the method by Hartung and Knapp (2003) is used to adjust test statistics and confidence intervals if argument hakn=TRUE.

The iterative Paule-Mandel method (1982) to estimate the between-study variance is used if argument method.tau="PM". Internally, R function paulemandel is called which is based on R function mpaule.default from R package metRology from S.L.R. Ellison <s.ellison at lgc.co.uk>.

If R package metafor (Viechtbauer 2010) is installed, the following methods to estimate the betweenstudy variance τ^2 (argument method. tau) are also available:

- Restricted maximum-likelihood estimator (method.tau="REML")
- Maximum-likelihood estimator (method.tau="ML")
- Hunter-Schmidt estimator (method.tau="HS")
- Sidik-Jonkman estimator (method.tau="SJ")
- Hedges estimator (method.tau="HE")

• Empirical Bayes estimator (method. tau="EB").

For these methods the R function rma.uni of R package metafor is called internally. See help page of R function rma.uni for more details on these methods to estimate between-study variance.

Value

An object of class c("metacont", "meta") with corresponding print, summary, plot function. The object is a list containing the following components:

```
n.e, mean.e, sd.e,
n.c, mean.c, sd.c,
studlab, sm, level, level.comb,
comb.fixed, comb.random,
hakn, method.tau, tau.preset, TE.tau, method.bias,
tau.common, title, complab, outclab,
label.e, label.c, label.left, label.right,
byvar, bylab, print.byvar, warn
                 As defined above.
TE, seTE
                 Estimated treatment effect and standard error of individual studies.
lower, upper
                 Lower and upper confidence interval limits for individual studies.
zval, pval
                 z-value and p-value for test of treatment effect for individual studies.
w.fixed, w.random
                  Weight of individual studies (in fixed and random effects model).
TE.fixed, seTE.fixed
                 Estimated overall treatment effect and standard error (fixed effect model).
lower.fixed, upper.fixed
                 Lower and upper confidence interval limits (fixed effect model).
zval.fixed, pval.fixed
                 z-value and p-value for test of overall treatment effect (fixed effect model).
TE.random, seTE.random
                 Estimated overall treatment effect and standard error (random effects model).
lower.random, upper.random
                 Lower and upper confidence interval limits (random effects model).
zval.random, pval.random
                  z-value or t-value and corresponding p-value for test of overall treatment effect
                 (random effects model).
prediction, level.predict
                 As defined above.
```

seTE.predict Standard error utilised for prediction interval.

lower.predict, upper.predict

Lower and upper limits of prediction interval.

k Number of studies combined in meta-analysis.

Q Heterogeneity statistic.

tau Square-root of between-study variance.

se.tau Standard error of square-root of between-study variance.

C Scaling factor utilised internally to calculate common tau-squared across sub-

groups.

method Pooling method: "Inverse".

df.hakn Degrees of freedom for test of treatment effect for Hartung-Knapp method (only

if hakn=TRUE).

keepdata As defined above.

data Original data (set) used in function call (if keepdata=TRUE).

subset Information on subset of original data used in meta-analysis (if keepdata=TRUE).

call Function call.

version Version of R package meta used to create object.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

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Viechtbauer W (2010), Conducting Meta-Analyses in R with the Metafor Package. *Journal of Statistical Software*, **36**, 1–48.

See Also

update.meta, metabin, metagen

Examples

```
data(Fleiss93cont)
meta1 <- metacont(n.e, mean.e, sd.e, n.c, mean.c, sd.c, data=Fleiss93cont, sm="SMD")</pre>
forest(meta1)
meta2 <- metacont(Fleiss93cont$n.e, Fleiss93cont$mean.e,</pre>
                   Fleiss93cont$sd.e,
                   Fleiss93cont$n.c, Fleiss93cont$mean.c,
                   Fleiss93cont$sd.c.
                   sm="SMD")
meta2
data(amlodipine)
meta3 <- metacont(n.amlo, mean.amlo, sqrt(var.amlo),</pre>
                   n.plac, mean.plac, sqrt(var.plac),
                   data=amlodipine, studlab=study)
summary(meta3)
# Use pooled variance
meta4 <- metacont(n.amlo, mean.amlo, sqrt(var.amlo),</pre>
                   n.plac, mean.plac, sqrt(var.plac),
                   data=amlodipine, studlab=study,
                   pooledvar=TRUE)
summary(meta4)
```

metacor

Meta-analysis of correlations

Description

Calculation of fixed and random effects estimates for meta-analyses with correlations; inverse variance weighting is used for pooling.

Usage

```
title=.settings$title, complab=.settings$complab, outclab="",
byvar, bylab, print.byvar=.settings$print.byvar,
keepdata=.settings$keepdata
)
```

Arguments

cor Correlation.

n Number of observations.

studlab An optional vector with study labels.

data An optional data frame containing the study information, i.e., cor and n.

subset An optional vector specifying a subset of studies to be used.

sm A character string indicating which summary measure ("ZCOR" or "COR") is to

be used for pooling of studies.

level The level used to calculate confidence intervals for individual studies.

The level used to calculate confidence intervals for pooled estimates.

comb.fixed A logical indicating whether a fixed effect meta-analysis should be conducted.

A logical indicating whether a random effects meta-analysis should be con-

ducted.

prediction A logical indicating whether a prediction interval should be printed.

level.predict The level used to calculate prediction interval for a new study.

hakn A logical indicating whether the method by Hartung and Knapp should be used

to adjust test statistics and confidence intervals.

method.tau A character string indicating which method is used to estimate the between-

study variance τ^2 . Either "DL", "PM", "REML", "ML", "HS", "SJ", "HE", or "EB",

can be abbreviated.

tau.preset Prespecified value for the square-root of the between-study variance τ^2 .

TE. tau Overall treatment effect used to estimate the between-study variance tau-squared. tau. common

A logical indicating whether tau-squared should be the same across subgroups.

A character string indicating which test is to be used. Either "rank", "linreg",

or "mm", can be abbreviated. See function metabias

backtransf A logical indicating whether results for Fisher's z transformed correlations (sm="ZCOR")

should be back transformed in printouts and plots. If TRUE (default), results will be presented as correlations; otherwise Fisher's z transformed correlations will

be shown.

title Title of meta-analysis / systematic review.

complab Comparison label. outclab Outcome label.

byvar An optional vector containing grouping information (must be of same length as

event.e).

bylab A character string with a label for the grouping variable.

print.byvar A logical indicating whether the name of the grouping variable should be printed

in front of the group labels.

keepdata A logical indicating whether original data (set) should be kept in meta object.

Details

Fixed effect and random effects meta-analysis of correlations based either on Fisher's z transformation of correlations (sm="ZCOR") or direct combination of correlations (sm="COR") (see Cooper et al., p264-5 and p273-4). By default, the DerSimonian-Laird estimate (1986) is used in the random effects model (method.tau="DL").

Only few statisticians would advocate the use of untransformed correlations unless sample sizes are very large (see Cooper et al., p265). The artificial example given below shows that the smallest study gets the largest weight if correlations are combined directly because the correlation is closest to 1.

For several arguments defaults settings are utilised (assignments with .settings\$). These defaults can be changed using the settings.meta function.

Internally, both fixed effect and random effects models are calculated regardless of values choosen for arguments comb.fixed and comb.random. Accordingly, the estimate for the random effects model can be extracted from component TE.random of an object of class "meta" even if argument comb.random=FALSE. However, all functions in R package meta will adequately consider the values for comb.fixed and comb.random. E.g. function print.meta will not print results for the random effects model if comb.random=FALSE.

A prediction interval for treatment effect of a new study is calculated (Higgins et al., 2009) if arguments prediction and comb.random are TRUE.

R function update.meta can be used to redo the meta-analysis of an existing metacor object by only specifying arguments which should be changed.

For the random effects, the method by Hartung and Knapp (2003) is used to adjust test statistics and confidence intervals if argument hakn=TRUE.

The iterative Paule-Mandel method (1982) to estimate the between-study variance is used if argument method. tau="PM". Internally, R function paulemandel is called which is based on R function mpaule.default from R package metRology from S.L.R. Ellison <s.ellison at lgc.co.uk>.

If R package metafor (Viechtbauer 2010) is installed, the following methods to estimate the betweenstudy variance τ^2 (argument method. tau) are also available:

- Restricted maximum-likelihood estimator (method.tau="REML")
- Maximum-likelihood estimator (method.tau="ML")
- Hunter-Schmidt estimator (method.tau="HS")
- Sidik-Jonkman estimator (method.tau="SJ")
- Hedges estimator (method.tau="HE")
- Empirical Bayes estimator (method.tau="EB").

For these methods the R function rma.uni of R package metafor is called internally. See help page of R function rma.uni for more details on these methods to estimate between-study variance.

Value

An object of class c("metacor", "meta") with corresponding print, summary, plot function. The object is a list containing the following components:

```
cor, n, studlab,
```

sm, level, level.comb, comb.fixed, comb.random, hakn, method.tau, tau.preset, TE.tau, method.bias, tau.common, title, complab, outclab, byvar, bylab, print.byvar As defined above. TE, seTE Either Fisher's z transformation of correlations (sm="ZCOR") or correlations (sm="COR") for individual studies. lower, upper Lower and upper confidence interval limits for individual studies. zval, pval z-value and p-value for test of treatment effect for individual studies. w.fixed, w.random Weight of individual studies (in fixed and random effects model). TE.fixed, seTE.fixed Estimated overall effect (Fisher's z transformation of correlation) and its standard error (fixed effect model). lower.fixed, upper.fixed Lower and upper confidence interval limits (fixed effect model). zval.fixed, pval.fixed z-value and p-value for test of overall effect (fixed effect model). TE.random, seTE.random Estimated overall effect (Fisher's z transformation of correlation or correlation) and its standard error (random effects model). lower.random, upper.random Lower and upper confidence interval limits (random effects model). zval.random, pval.random z-value or t-value and corresponding p-value for test of overall effect (random effects model). prediction, level.predict As defined above. seTE.predict Standard error utilised for prediction interval. lower.predict, upper.predict Lower and upper limits of prediction interval. k Number of studies combined in meta-analysis. Q Heterogeneity statistic Q. Square-root of between-study variance. tau se.tau Standard error of square-root of between-study variance. С Scaling factor utilised internally to calculate common tau-squared across subgroups.

A character string indicating method used for pooling: "Inverse"

method

df.hakn	Degrees of freedom	for test of treatment effect for	Hartung-Knapp method (only

if hakn=TRUE).

keepdata As defined above.

data Original data (set) used in function call (if keepdata=TRUE).

subset Information on subset of original data used in meta-analysis (if keepdata=TRUE).

call Function call.

version Version of R package meta used to create object.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Cooper H, Hedges LV, Valentine JC (2009), *The Handbook of Research Synthesis and Meta-Analysis*, 2nd Edition. New York: Russell Sage Foundation.

DerSimonian R & Laird N (1986), Meta-analysis in clinical trials. *Controlled Clinical Trials*, **7**, 177–188.

Higgins JPT, Thompson SG, Spiegelhalter DJ (2009), A re-evaluation of random-effects meta-analysis. *Journal of the Royal Statistical Society: Series A*, **172**, 137–159.

Knapp G & Hartung J (2003), Improved Tests for a Random Effects Meta-regression with a Single Covariate. *Statistics in Medicine*, **22**, 2693–710, doi: 10.1002/sim.1482.

Paule RC & Mandel J (1982), Consensus values and weighting factors. *Journal of Research of the National Bureau of Standards*, **87**, 377–385.

Viechtbauer W (2010), Conducting Meta-Analyses in R with the Metafor Package. *Journal of Statistical Software*, **36**, 1–48.

See Also

```
update.meta, metacont, metagen, print.meta
```

Examples

```
m1 <- metacor(c(0.85, 0.7, 0.95), c(20, 40, 10))
#
# Print correlations (back transformed from Fisher's z transformation)
#
m1
#
# Print Fisher's z transformed correlations
#
print(m1, backtransf=FALSE)
#
# Forest plot with back transformed correlations</pre>
```

```
#
forest(m1)

#
# Forest plot with Fisher's z transformed correlations
#
forest(m1, backtransf=FALSE)

m2 <- update(m1, sm="cor")
m2
# Identical forest plots (as back transformation is the identity transformation)
# forest(m2)
# forest(m2, backtransf=FALSE)</pre>
```

metacr

Meta-analysis of outcome data from Cochrane review

Description

Wrapper function to perform meta-analysis for a single outcome of a Cochrane Intervention review.

Usage

```
metacr(x, comp.no=1, outcome.no=1,
    method, sm,
    level=.settings$level, level.comb=.settings$level.comb,
    comb.fixed, comb.random,
    hakn=FALSE,
    method.tau="DL",
    tau.common=FALSE,
    prediction=.settings$prediction, level.predict=.settings$level.predict,
    swap.events, logscale,
    backtransf=.settings$backtransf,
    title, complab, outclab, warn=FALSE)
```

Arguments

X	An object of class rmb created by R function read.rmb.
comp.no	Comparison number.
outcome.no	Outcome number.
method	A character string indicating which method is to be used for pooling of studies. One of "Inverse", "MH", or "Peto", can be abbreviated.
sm	A character string indicating which summary measure ("RR", "OR", "RD", "AS", "HR", "MD", or "SMD") is to be used for pooling of studies.
level	The level used to calculate confidence intervals for individual studies.

level.comb	The level used to calculate confidence intervals for pooled estimates.
comb.fixed	A logical indicating whether a fixed effect meta-analysis should be conducted.
comb.random	A logical indicating whether a random effects meta-analysis should be conducted.
hakn	A logical indicating whether the method by Hartung and Knapp should be used to adjust test statistics and confidence intervals.
method.tau	A character string indicating which method is used to estimate the between-study variance τ^2 . Either "DL", "PM", "REML", "ML", "HS", "SJ", "HE", or "EB", can be abbreviated.
tau.common	A logical indicating whether tau-squared should be the same across subgroups.
prediction	A logical indicating whether a prediction interval should be printed.
level.predict	The level used to calculate prediction interval for a new study.
swap.events	A logical indicating whether events and non-events should be interchanged.
logscale	A logical indicating whether effect estimates are entered on log-scale.
backtransf	A logical indicating whether results should be back transformed in printouts and plots. If backtransf=TRUE (default), results for sm="OR" are printed as odds ratios rather than log odds ratios and results for sm="ZCOR" are printed as correlations rather than Fisher's z transformed correlations, for example.
title	Title of meta-analysis / systematic review.
complab	Comparison label.
outclab	Outcome label.
warn	A logical indicating whether warnings should be printed (e.g., if incr is added to studies with zero cell frequencies).

Details

Cochrane Intervention reviews are based on the comparison of two interventions. Each Cochrane Intervention review can have a variable number of comparisons. For each comparison, a variable number of outcomes can be define. For each outcome, a seperate meta-analysis is conducted. Review Manager 5 (RevMan 5) is the current software used for preparing and maintaining Cochrane Reviews (http://www.cc-ims.net/revman/).

This wrapper function can be used to perform meta-analysis for a single outcome of a Cochrane Intervention review. Internally, R functions metabin, metacont, and metagen are called - depending on the definition of the outcome in RevMan 5.

Value

An object of class "meta" and "metabin", "metacont", or "metagen" depending on outcome type utilised in Cochrane Intervention review for selected outcome.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

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References

Review Manager (RevMan) [Computer program]. Version 5.1. Copenhagen: The Nordic Cochrane Centre, The Cochrane Collaboration, 2011.

See Also

```
metabin, metacont, metagen, read.rm5
```

Examples

```
# Locate export data file "Fleiss93_CR.csv"
# in sub-directory of package "meta"
#
filename <- system.file("data/Fleiss93_CR.csv.gz", package = "meta")
#
Fleiss93_CR <- read.rm5(filename)
# Same result as R command example(Fleiss93):
#
metacr(Fleiss93_CR)
# Same result as R command example(Fleiss93cont):
#
metacr(Fleiss93_CR, 1, 2)
forest(metacr(Fleiss93_CR, 1, 2))
# Change summary measure to RR
#
m1 <- metacr(Fleiss93_CR)
update(m1, sm="RR")</pre>
```

metacum

Cumulative meta-analysis

Description

Performs a cumulative meta-analysis.

Usage

```
metacum(x, pooled, sortvar)
```

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Arguments

x An object of class meta.

pooled A character string indicating whether a fixed effect or random effects model is

used for pooling. Either missing (see Details), "fixed", or "random", can be

abbreviated.

sortvar An optional vector used to sort the individual studies (must be of same length as

x\$TE).

Details

A cumulative meta-analysis is performed. Studies are included sequentially as defined by sortvar.

Information from object x is utilised if argument pooled is missing. A fixed effect model is assumed (pooled="fixed") if argument x\$comb.fixed is TRUE; a random effects model is assumed (pooled="random") if argument x\$comb.random is TRUE and x\$comb.fixed is FALSE.

Value

An object of class c("metacum", "meta") with corresponding print, plot function. The object is a list containing the following components:

TE, seTE Estimated treatment effect and standard error of pooled estimate in cumulative

meta-analyses.

lower, upper Lower and upper confidence interval limits. studlab Study label describing addition of studies.

p.value P-value for test of overall effect.

w Sum of weights from fixed effect or random effects model.

I2 Heterogeneity statistic I2.

tau Square-root of between-study variance.

df. hakn Degrees of freedom for test of treatment effect for Hartung-Knapp method (only

if hakn=TRUE).

sm Summary measure.

method Method used for pooling.

k Number of studies combined in meta-analysis.

pooled As defined above.

comb.fixed A logical indicating whether analysis is based on fixed effect model.

Comb.random A logical indicating whether analysis is based on random effects model.

TE.fixed, seTE.fixed

Value is NA.

TE.random, seTE.random

Value is NA.

Q Value is NA.

level.comb The level used to calculate confidence intervals for pooled estimates.

hakn A logical indicating whether the method by Hartung and Knapp is used to adjust

test statistics and confidence intervals.

method.tau A character string indicating which method is used to estimate the between-

study variance τ^2 .

tau.preset Prespecified value for the square-root of the between-study variance τ^2 .

TE. tau Overall treatment effect used to estimate the between-study variance τ^2 .

n.harmonic.mean

Harmonic mean of number of observations (for back transformation of Freeman-

Tukey Double arcsine transformation).

version Version of R package meta used to create object.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Cooper H & Hedges LV (1994), *The Handbook of Research Synthesis*. Newbury Park, CA: Russell Sage Foundation.

See Also

```
metabin, metacont, print.meta
```

Examples

metagen

Generic inverse variance meta-analysis

Description

Fixed and random effects meta-analysis based on estimates (e.g. log hazard ratios) and their standard errors; inverse variance weighting is used for pooling.

Usage

```
metagen(TE, seTE, studlab, data=NULL, subset=NULL, sm="",
       level=.settings$level, level.comb=.settings$level.comb,
       comb.fixed=.settings$comb.fixed, comb.random=.settings$comb.random,
       hakn=.settings$hakn,
       method.tau=.settings$method.tau, tau.preset=NULL, TE.tau=NULL,
       tau.common=.settings$tau.common,
       prediction=.settings$prediction, level.predict=.settings$level.predict,
       method.bias=.settings$method.bias,
       n.e=NULL, n.c=NULL,
       backtransf=.settings$backtransf,
       title=.settings$title, complab=.settings$complab, outclab="",
       label.e=.settings$label.e, label.c=.settings$label.c,
       label.left=.settings$label.left, label.right=.settings$label.right,
       byvar, bylab, print.byvar=.settings$print.byvar,
       keepdata=.settings$keepdata,
       warn=.settings$warn)
```

Arguments

TE.tau

TE	Estimate of treatment effect.
seTE	Standard error of treatment estimate.
studlab	An optional vector with study labels.
data	An optional data frame containing the study information.
subset	An optional vector specifying a subset of studies to be used.
SM	A character string indicating underlying summary measure, e.g., "RD", "RR", "OR", "AS", "MD", "SMD".
level	The level used to calculate confidence intervals for individual studies.
level.comb	The level used to calculate confidence intervals for pooled estimates.
comb.fixed	A logical indicating whether a fixed effect meta-analysis should be conducted.
comb.random	A logical indicating whether a random effects meta-analysis should be conducted.
prediction	A logical indicating whether a prediction interval should be printed.
level.predict	The level used to calculate prediction interval for a new study.
n.e	Number of observations in experimental group.
n.c	Number of observations in control group.
hakn	A logical indicating whether method by Hartung and Knapp should be used to adjust test statistics and confidence intervals.
method.tau	A character string indicating which method is used to estimate the between-study variance τ^2 . Either "DL", "PM", "REML", "ML", "HS", "SJ", "HE", or "EB", can be abbreviated.
tau.preset	Prespecified value for the square-root of the between-study variance $ au^2$.

Overall treatment effect used to estimate the between-study variance tau-squared.

tau.common A logical indicating whether tau-squared should be the same across subgroups.

Method.bias A character string indicating which test is to be used. Either "rank", "linreg",

or "mm", can be abbreviated. See function metabias

backtransf A logical indicating whether results should be back transformed in printouts

and plots. If backtransf=TRUE (default), results for sm="OR" are printed as odds ratios rather than log odds ratios and results for sm="ZCOR" are printed as correlations rather than Fisher's z transformed correlations, for example.

title Title of meta-analysis / systematic review.

complab Comparison label. outclab Outcome label.

label.e Label for experimental group.

label.c Label for control group.

label.left Graph label on left side of forest plot.
label.right Graph label on right side of forest plot.

byvar An optional vector containing grouping information (must be of same length as

TE).

bylab A character string with a label for the grouping variable.

print.byvar A logical indicating whether the name of the grouping variable should be printed

in front of the group labels.

keepdata A logical indicating whether original data (set) should be kept in meta object.

warn A logical indicating whether warnings should be printed (e.g., if studies are

excluded from meta-analysis due to zero standard errors).

Details

Generic method for meta-analysis, only treatment estimates and their standard error are needed. The method is useful, e.g., for pooling of survival data (using log hazard ratio and standard errors as input). The inverse variance method is used for pooling. By default, the DerSimonian-Laird estimate (1986) is used in the random effects model (method.tau="DL").

For several arguments defaults settings are utilised (assignments with .settings\$). These defaults can be changed using the settings.meta function.

Internally, both fixed effect and random effects models are calculated regardless of values choosen for arguments comb.fixed and comb.random. Accordingly, the estimate for the random effects model can be extracted from component TE.random of an object of class "meta" even if argument comb.random=FALSE. However, all functions in R package meta will adequately consider the values for comb.fixed and comb.random. E.g. function print.meta will not print results for the random effects model if comb.random=FALSE.

A prediction interval for treatment effect of a new study is calculated (Higgins et al., 2009) if arguments prediction and comb.random are TRUE.

R function update.meta can be used to redo the meta-analysis of an existing metagen object by only specifying arguments which should be changed.

For the random effects, the method by Hartung and Knapp (2003) is used to adjust test statistics and confidence intervals if argument hakn=TRUE.

The iterative Paule-Mandel method (1982) to estimate the between-study variance is used if argument method. tau="PM". Internally, R function paulemandel is called which is based on R function mpaule.default from R package metRology from S.L.R. Ellison <s.ellison at lgc.co.uk>.

If R package metafor (Viechtbauer 2010) is installed, the following methods to estimate the betweenstudy variance τ^2 (argument method.tau) are also available:

- Restricted maximum-likelihood estimator (method.tau="REML")
- Maximum-likelihood estimator (method.tau="ML")
- Hunter-Schmidt estimator (method.tau="HS")
- Sidik-Jonkman estimator (method.tau="SJ")
- Hedges estimator (method.tau="HE")
- Empirical Bayes estimator (method. tau="EB").

For these methods the R function rma.uni of R package metafor is called internally. See help page of R function rma.uni for more details on these methods to estimate between-study variance.

Value

An object of class c("metagen", "meta") with corresponding print, summary, plot function. The object is a list containing the following components:

```
TE, seTE, studlab, n.e, n.c
sm, level, level.comb,
comb.fixed, comb.random,
hakn, method.tau, tau.preset, TE.tau, method.bias,
tau.common, title, complab, outclab,
label.e, label.c, label.left, label.right,
byvar, bylab, print.byvar, warn
                 As defined above.
lower, upper
                 Lower and upper confidence interval limits for individual studies.
zval, pval
                 z-value and p-value for test of treatment effect for individual studies.
w.fixed, w.random
                 Weight of individual studies (in fixed and random effects model).
TE.fixed, seTE.fixed
                 Estimated overall treatment effect and standard error (fixed effect model).
lower.fixed, upper.fixed
                 Lower and upper confidence interval limits (fixed effect model).
zval.fixed, pval.fixed
                 z-value and p-value for test of overall treatment effect (fixed effect model).
TE.random, seTE.random
```

Estimated overall treatment effect and standard error (random effects model).

lower.random, upper.random

Lower and upper confidence interval limits (random effects model).

zval.random, pval.random

z-value or t-value and corresponding p-value for test of overall treatment effect (random effects model).

prediction, level.predict

As defined above.

seTE.predict Standard error utilised for prediction interval.

lower.predict, upper.predict

Lower and upper limits of prediction interval.

k Number of studies combined in meta-analysis.

Q Heterogeneity statistic.

df.Q Degrees of freedom for heterogeneity statistic.

tau Square-root of between-study variance.

se.tau Standard error of square-root of between-study variance.

C Scaling factor utilised internally to calculate common tau-squared across sub-

groups.

method Pooling method: "Inverse".

df. hakn Degrees of freedom for test of treatment effect for Hartung-Knapp method (only

if hakn=TRUE).

keepdata As defined above.

data Original data (set) used in function call (if keepdata=TRUE).

subset Information on subset of original data used in meta-analysis (if keepdata=TRUE).

call Function call.

version Version of R package meta used to create object.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Cooper H & Hedges LV (1994), *The Handbook of Research Synthesis*. Newbury Park, CA: Russell Sage Foundation.

DerSimonian R & Laird N (1986), Meta-analysis in clinical trials. *Controlled Clinical Trials*, **7**, 177–188.

Higgins JPT, Thompson SG, Spiegelhalter DJ (2009), A re-evaluation of random-effects meta-analysis. *Journal of the Royal Statistical Society: Series A*, **172**, 137–159.

Knapp G & Hartung J (2003), Improved Tests for a Random Effects Meta-regression with a Single Covariate. *Statistics in Medicine*, 22, 2693–2710, doi: 10.1002/sim.1482.

Paule RC & Mandel J (1982), Consensus values and weighting factors. *Journal of Research of the National Bureau of Standards*, **87**, 377–385.

Viechtbauer W (2010), Conducting Meta-Analyses in R with the Metafor Package. *Journal of Statistical Software*, **36**, 1–48.

See Also

```
update.meta, metabin, metacont, print.meta
```

Examples

```
data(Fleiss93)
meta1 <- metabin(event.e, n.e, event.c, n.c, data=Fleiss93, sm="RR", method="I")</pre>
meta1
# Identical results by using the following commands:
meta1
metagen(meta1$TE, meta1$seTE, sm="RR")
forest(metagen(meta1$TE, meta1$seTE, sm="RR"))
# Meta-analysis with prespecified between-study variance
summary(metagen(meta1$TE, meta1$seTE, sm="RR", tau.preset=sqrt(0.1)))
# Meta-analysis of survival data:
logHR < - log(c(0.95, 1.5))
selogHR <- c(0.25, 0.35)
metagen(logHR, selogHR, sm="HR")
# Paule-Mandel method to estimate between-study variance
# Data from Paule & Mandel (1982)
average <- c(27.044, 26.022, 26.340, 26.787, 26.796)
variance < c(0.003, 0.076, 0.464, 0.003, 0.014)
summary(metagen(average, sqrt(variance), sm="MD", method.tau="PM"))
```

metainc

Meta-analysis of incidence rates

Description

Calculation of fixed and random effects estimates (incidence rate ratio or incidence rate difference) for meta-analyses with event counts. Mantel-Haenszel, Cochran, and inverse variance method are available for pooling.

Usage

```
metainc(event.e, time.e, event.c, time.c, studlab,
       data=NULL, subset=NULL, method="MH",
        sm=.settings$sminc,
        incr=.settings$incr, allincr=.settings$allincr,
       addincr=.settings$addincr,
        level=.settings$level, level.comb=.settings$level.comb,
       comb.fixed=.settings$comb.fixed, comb.random=.settings$comb.random,
       hakn=.settings$hakn,
       method.tau=.settings$method.tau, tau.preset=NULL, TE.tau=NULL,
        tau.common=.settings$tau.common,
       prediction=.settings$prediction, level.predict=.settings$level.predict,
       method.bias=.settings$method.bias,
       n.e=NULL, n.c=NULL,
       backtransf=.settings$backtransf,
       title=.settings$title, complab=.settings$complab, outclab="",
       label.e=.settings$label.e, label.c=.settings$label.c,
       label.left=.settings$label.left, label.right=.settings$label.right,
       byvar, bylab, print.byvar=.settings$print.byvar,
       keepdata=.settings$keepdata,
       warn=.settings$warn)
```

Arguments

event.e	Number of events in experimental group.
time.e	Person time at risk in experimental group.
event.c	Number of events in control group.
time.c	Person time at risk in control group.
studlab	An optional vector with study labels.
data	An optional data frame containing the study information, i.e., event.e, time.e, event.c, and time.c.
subset	An optional vector specifying a subset of studies to be used.
method	A character string indicating which method is to be used for pooling of studies. One of "MH", "Inverse", or "Cochran", can be abbreviated.
SM	A character string indicating which summary measure ("IRR" or "IRD") is to be used for pooling of studies, see Details.
incr	A numerical value which is added to each cell frequency for studies with a zero cell count, see Details.
allincr	A logical indicating if incr is added to each cell frequency of all studies if at least one study has a zero cell count. If FALSE (default), incr is added only to each cell frequency of studies with a zero cell count.
addincr	A logical indicating if incr is added to each cell frequency of all studies irrespective of zero cell counts.
level	The level used to calculate confidence intervals for individual studies.

level.comb	The level used to calculate confidence intervals for pooled estimates.
comb.fixed	A logical indicating whether a fixed effect meta-analysis should be conducted.
comb.random	A logical indicating whether a random effects meta-analysis should be conducted.
prediction	A logical indicating whether a prediction interval should be printed.
level.predict	The level used to calculate prediction interval for a new study.
hakn	A logical indicating whether the method by Hartung and Knapp should be used to adjust test statistics and confidence intervals.
method.tau	A character string indicating which method is used to estimate the between-study variance τ^2 . Either "DL", "PM", "REML", "ML", "HS", "SJ", "HE", or "EB", can be abbreviated.
tau.preset	Prespecified value for the square-root of the between-study variance $ au^2$.
TE.tau	Overall treatment effect used to estimate the between-study variance τ^2 .
tau.common	A logical indicating whether tau-squared should be the same across subgroups.
method.bias	A character string indicating which test for funnel plot asymmetry is to be used. Either "linreg" or "rank", can be abbreviated. See function metabias
n.e	Number of observations in experimental group (optional).
n.c	Number of observations in control group (optional).
backtransf	A logical indicating whether results for incidence rate ratio (sm="IRR") should be back transformed in printouts and plots. If TRUE (default), results will be presented as incidence rate ratios; otherwise log incidence rate ratios will be shown.
title	Title of meta-analysis / systematic review.
complab	Comparison label.
outclab	Outcome label.
label.e	Label for experimental group.
label.c	Label for control group.
label.left	Graph label on left side of forest plot.
label.right	Graph label on right side of forest plot.
byvar	An optional vector containing grouping information (must be of same length as event.e).
bylab	A character string with a label for the grouping variable.
print.byvar	A logical indicating whether the name of the grouping variable should be printed in front of the group labels.
keepdata	A logical indicating whether original data (set) should be kept in meta object.
warn	A logical indicating whether warnings should be printed (e.g., if incr is added to studies with zero cell frequencies).

Details

Treatment estimates and standard errors are calculated for each study. The following measures of treatment effect are available:

- Incidence Rate Ratio (sm="IRR")
- Incidence Rate Difference (sm="IRD")

For studies with a zero cell count, by default, 0.5 is added to all cell frequencies of these studies (argument incr).

For several arguments defaults settings are utilised (assignments with .settings\$). These defaults can be changed using the settings.meta function.

Internally, both fixed effect and random effects models are calculated regardless of values choosen for arguments comb.fixed and comb.random. Accordingly, the estimate for the random effects model can be extracted from component TE.random of an object of class "meta" even if argument comb.random=FALSE. However, all functions in R package meta will adequately consider the values for comb.fixed and comb.random. E.g. function print.meta will not print results for the random effects model if comb.random=FALSE.

By default, both fixed effect and random effects models are considered (see arguments comb.fixed and comb.random). If method is "MH" (default), the Mantel-Haenszel method is used to calculate the fixed effect estimate (Greenland & Robbins, 1985); if method is "Inverse", inverse variance weighting is used for pooling; finally, if method is "Cochran", the Cochran method is used for pooling (Bayne-Jones, 1964, Chapter 8). By default, the DerSimonian-Laird estimate (1986) is used in the random effects model (method.tau="DL").

For Mantel-Haenszel and Cochran method, nothing is added to zero cell counts. Accordingly, Mantel-Haenszel and Cochran estimate are not defined if the number of events is zero in all studies either in the experimental or control group.

A prediction interval for treatment effect of a new study is calculated (Higgins et al., 2009) if arguments prediction and comb.random are TRUE.

R function update.meta can be used to redo the meta-analysis of an existing metainc object by only specifying arguments which should be changed.

For the random effects, the method by Hartung and Knapp (2003) is used to adjust test statistics and confidence intervals if argument hakn=TRUE.

The iterative Paule-Mandel method (1982) to estimate the between-study variance is used if argument method. tau="PM". Internally, R function paulemandel is called which is based on R function mpaule.default from R package metRology from S.L.R. Ellison <s.ellison at lgc.co.uk>.

If R package metafor (Viechtbauer 2010) is installed, the following methods to estimate the between-study variance τ^2 (argument method. tau) are also available:

- Restricted maximum-likelihood estimator (method.tau="REML")
- Maximum-likelihood estimator (method.tau="ML")
- Hunter-Schmidt estimator (method.tau="HS")
- Sidik-Jonkman estimator (method.tau="SJ")
- Hedges estimator (method.tau="HE")
- Empirical Bayes estimator (method. tau="EB").

For these methods the R function rma.uni of R package metafor is called internally. See help page of R function rma.uni for more details on these methods to estimate between-study variance.

Value

An object of class c("metainc", "meta") with corresponding print, summary, plot function. The object is a list containing the following components:

event.e, time.e, event.c, time.c, studlab,

sm, method, incr, allincr, addincr, warn,

level, level.comb, comb.fixed, comb.random,

hakn, method.tau, tau.preset, TE.tau, method.bias,

tau.common, title, complab, outclab,

label.e, label.c, label.left, label.right,

byvar, bylab, print.byvar

As defined above.

TE, seTE Estimated treatment effect and standard error of individual studies.

lower, upper Lower and upper confidence interval limits for individual studies.

zval, pval z-value and p-value for test of treatment effect for individual studies.

w.fixed, w.random

Weight of individual studies (in fixed and random effects model).

TE.fixed, seTE.fixed

Estimated overall treatment effect and standard error (fixed effect model).

lower.fixed, upper.fixed

Lower and upper confidence interval limits (fixed effect model).

zval.fixed, pval.fixed

z-value and p-value for test of overall treatment effect (fixed effect model).

TE.random, seTE.random

Estimated overall treatment effect and standard error (random effects model).

lower.random, upper.random

Lower and upper confidence interval limits (random effects model).

zval.random, pval.random

z-value or t-value and corresponding p-value for test of overall treatment effect (random effects model).

prediction, level.predict

As defined above.

seTE.predict Standard error utilised for prediction interval.

lower.predict, upper.predict

Lower and upper limits of prediction interval.

k Number of studies combined in meta-analysis.

Q	Heterogeneity	statistic Q.

df.Q Degrees of freedom for heterogeneity statistic.

tau Square-root of between-study variance.

se. tau Standard error of square-root of between-study variance.

C Scaling factor utilised internally to calculate common tau-squared across sub-

groups.

sparse Logical flag indicating if any study included in meta-analysis has any zero cell

frequencies.

incr.event Increment added to number of events.

df. hakn Degrees of freedom for test of treatment effect for Hartung-Knapp method (only

if hakn=TRUE).

keepdata As defined above.

data Original data (set) used in function call (if keepdata=TRUE).

subset Information on subset of original data used in meta-analysis (if keepdata=TRUE).

call Function call.

version Version of R package meta used to create object.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

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Viechtbauer W (2010), Conducting Meta-Analyses in R with the Metafor Package. *Journal of Statistical Software*, **36**, 1–48.

metainf 67

See Also

```
metabin, update.meta, print.meta
```

Examples

```
data(smoking)
m1 <- metainc(d.smokers, py.smokers,</pre>
              d.nonsmokers, py.nonsmokers,
              data=smoking, studlab=study)
print(m1, digits=2)
m2 <- metainc(d.smokers, py.smokers,</pre>
              d.nonsmokers, py.nonsmokers,
              data=smoking, studlab=study,
              method="Cochran")
print(m2, digits=2)
data(lungcancer)
m3 <- metainc(d.smokers, py.smokers,</pre>
              d.nonsmokers, py.nonsmokers,
              data=lungcancer, studlab=study)
print(m3, digits=2)
# Redo Cochran meta-analysis with inflated standard errors
# All cause mortality
TEa <- log( (smoking$d.smokers/smoking$py.smokers) /</pre>
            (smoking$d.nonsmokers/smoking$py.nonsmokers)
seTEa <- sqrt(1/smoking$d.smokers +</pre>
              1/smoking$d.nonsmokers + 2.5/smoking$d.nonsmokers)
metagen(TEa, seTEa, sm="IRR", studlab=smoking$study)
# Lung cancer mortality
TEl <- log( (lungcancer$d.smokers/lungcancer$py.smokers) /
            (lungcancer$d.nonsmokers/lungcancer$py.nonsmokers)
seTEl <- sqrt(1/lungcancer$d.smokers +</pre>
              1/lungcancer$d.nonsmokers + 2.25/lungcancer$d.nonsmokers)
metagen(TE1, seTE1, sm="IRR", studlab=lungcancer$study)
```

metainf

Influence analysis in meta-analysis

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Description

Performs a influence analysis. Pooled estimates are calculated omitting one study at a time.

Usage

```
metainf(x, pooled, sortvar)
```

Arguments

x An object of class meta.

pooled A character string indicating whether a fixed effect or random effects model is

used for pooling. Either missing (see Details), "fixed" or "random", can be

abbreviated.

sortvar An optional vector used to sort the individual studies (must be of same length as

x\$TE).

Details

Performs a influence analysis; pooled estimates are calculated omitting one study at a time. Studies are sorted according to sortvar.

Information from object x is utilised if argument pooled is missing. A fixed effect model is assumed (pooled="fixed") if argument x\$comb.fixed is TRUE; a random effects model is assumed (pooled="random") if argument x\$comb.random is TRUE and x\$comb.fixed is FALSE.

Value

An object of class c("metainf", "meta") with corresponding print, plot function. The object is a list containing the following components:

TE, seTE Estimated treatment effect and standard error of pooled estimate in influence

analysis.

lower, upper Lower and upper confidence interval limits. studlab Study label describing omission of studies.

p. value P-value for test of overall effect.

w Sum of weights from fixed effect or random effects model.

I2 Heterogeneity statistic I2.

tau Square-root of between-study variance.

df. hakn Degrees of freedom for test of treatment effect for Hartung-Knapp method (only

if hakn=TRUE).

sm Summary measure.

method Method used for pooling.

k Number of studies combined in meta-analysis.

pooled As defined above.

comb.fixed A logical indicating whether analysis is based on fixed effect model.

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comb.random A logical indicating whether analysis is based on random effects model.

TE.fixed, seTE.fixed

Value is NA.

TE.random, seTE.random

Value is NA.

Q Value is NA.

level.comb The level used to calculate confidence intervals for pooled estimates.

hakn A logical indicating whether the method by Hartung and Knapp is used to adjust

test statistics and confidence intervals.

method.tau A character string indicating which method is used to estimate the between-

study variance τ^2 .

tau.preset Prespecified value for the square-root of the between-study variance τ^2 .

TE. tau Overall treatment effect used to estimate the between-study variance τ^2 .

n.harmonic.mean

Harmonic mean of number of observations (for back transformation of Freeman-

Tukey Double arcsine transformation).

version Version of R package meta used to create object.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Cooper H & Hedges LV (1994), *The Handbook of Research Synthesis*. Newbury Park, CA: Russell Sage Foundation.

See Also

```
metabin, metacont, print.meta
```

Examples

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metaprop	Meta-analysis of single proportions	

Description

Calculation of an overall proportion from studies reporting a single proportion.

Usage

```
metaprop(event, n, studlab,
         data=NULL, subset=NULL,
         sm=.settings$smprop,
         incr=.settings$incr, allincr=.settings$allincr,
         addincr=.settings$addincr,
         method.ci=.settings$method.ci,
         level=.settings$level, level.comb=.settings$level.comb,
         comb.fixed=.settings$comb.fixed, comb.random=.settings$comb.random,
         hakn=.settings$hakn,
         method.tau=.settings$method.tau, tau.preset=NULL, TE.tau=NULL,
         tau.common=.settings$tau.common,
        prediction=.settings$prediction, level.predict=.settings$level.predict,
         method.bias=.settings$method.bias,
         backtransf=.settings$backtransf,
         title=.settings$title, complab=.settings$complab, outclab="",
         byvar, bylab, print.byvar=.settings$print.byvar,
         keepdata=.settings$keepdata,
         warn=.settings$warn)
```

Arguments

event	Number of events.
n	Number of observations.
studlab	An optional vector with study labels.
data	An optional data frame containing the study information, i.e., event and n.
subset	An optional vector specifying a subset of studies to be used.
sm	A character string indicating which summary measure ("PFT", "PAS", "PRAW", "PLN", or "PLOGIT") is to be used for pooling of studies, see Details.
incr	A numeric which is added to each cell frequency for studies with a zero cell count.
allincr	A logical indicating if incr is added to each cell frequency of all studies if at least one study has a zero cell count. If FALSE (default), incr is added only to each cell frequency of studies with a zero cell count.
addincr	A logical indicating if incr is added to each cell frequency of all studies irrespective of zero cell counts.

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method.ci	A character string indicating which method is used to calculate confidence intervals for individual studies, see Details.
level	The level used to calculate confidence intervals for individual studies.
level.comb	The level used to calculate confidence intervals for pooled estimates.
comb.fixed	A logical indicating whether a fixed effect meta-analysis should be conducted.
comb.random	A logical indicating whether a random effects meta-analysis should be conducted.
prediction	A logical indicating whether a prediction interval should be printed.
level.predict	The level used to calculate prediction interval for a new study.
hakn	A logical indicating whether the method by Hartung and Knapp should be used to adjust test statistics and confidence intervals.
method.tau	A character string indicating which method is used to estimate the between- study variance $ au^2$, see Details.
tau.preset	Prespecified value for the square-root of the between-study variance τ^2 .
TE.tau	Overall treatment effect used to estimate the between-study variance tau-squared.
tau.common	A logical indicating whether tau-squared should be the same across subgroups.
method.bias	A character string indicating which test is to be used. Either "rank", "linreg", or "mm", can be abbreviated. See function metabias.
backtransf	A logical indicating whether results for transformed proportions (argument sm!="PRAW") should be back transformed in printouts and plots. If TRUE (default), results will be presented as proportions; otherwise transformed proportions will be shown. See Details for presentation of confidence intervals.
title	Title of meta-analysis / systematic review.
complab	Comparison label.
outclab	Outcome label.
byvar	An optional vector containing grouping information (must be of same length as event.e).
bylab	A character string with a label for the grouping variable.
print.byvar	A logical indicating whether the name of the grouping variable should be printed in front of the group labels.
keepdata	A logical indicating whether original data (set) should be kept in meta object.
warn	A logical indicating whether the addition of incr to studies with zero cell frequencies should result in a warning.

Details

Fixed effect and random effects meta-analysis of single proportions to calculate an overall proportion. By default, the DerSimonian-Laird estimate (1986) is used in the random effects model (method.tau="DL").

The following transformations of proportions are implemented to calculate an overall proportion:

• Logit transformation (sm="PLOGIT", default)

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- Log transformation (sm="PLN")
- Freeman-Tukey Double arcsine transformation (sm="PFT")
- Arcsine transformation (sm="PAS")
- Raw, i.e. untransformed, proportions (sm="PRAW")

If the summary measure is equal to "PRAW", "PLN", or "PLOGIT", a continuity correction is applied if any studies has a zero cell count. By default, 0.5 is added to all cell frequencies of studies with a zero cell count (argument incr).

Various methods are available to calculate confidence intervals for individual study results (see Agresti & Coull 1998; Newcombe 1988):

- Clopper-Pearson interval also called 'exact' binomial interval (method.ci="CP", default)
- Wilson Score interval (method.ci="WS")
- Wilson Score interval with continuity correction (method.ci="WSCC")
- Agresti-Coull interval (method.ci="AC")
- Simple approximation interval (method.ci="SA")
- Simple approximation interval with continuity correction (method.ci="SACC")
- Normal approximation interval based on summary measure, i.e. defined by argument sm (method.ci="NAsm")

Note, with exception of the normal approximation based on the summary measure, i.e. method.ci="NAsm", the same confidence interval is calculated for any summary measure (argument sm) as only number of events and observations are used in the calculation disregarding the chosen summary measure. Results will be presented for transformed proportions if argument backtransf=FALSE in the print.meta, print.summary.meta, or forest.meta function. In this case, argument method.ci="NAsm" is used, i.e. confidence intervals based on the normal approximation based on the summary measure.

For several arguments defaults settings are utilised (assignments with .settings\$). These defaults can be changed using the settings.meta function.

Internally, both fixed effect and random effects models are calculated regardless of values choosen for arguments comb.fixed and comb.random. Accordingly, the estimate for the random effects model can be extracted from component TE.random of an object of class "meta" even if argument comb.random=FALSE. However, all functions in R package meta will adequately consider the values for comb.fixed and comb.random. E.g. function print.meta will not print results for the random effects model if comb.random=FALSE.

A prediction interval for treatment effect of a new study is calculated (Higgins et al., 2009) if arguments prediction and comb.random are TRUE.

R function update.meta can be used to redo the meta-analysis of an existing metaprop object by only specifying arguments which should be changed.

For the random effects, the method by Hartung and Knapp (2003) is used to adjust test statistics and confidence intervals if argument hakn=TRUE.

The iterative Paule-Mandel method (1982) to estimate the between-study variance is used if argument method. tau="PM". Internally, R function paulemandel is called which is based on R function mpaule.default from R package metRology from S.L.R. Ellison <s.ellison at lgc.co.uk>.

If R package metafor (Viechtbauer 2010) is installed, the following methods to estimate the betweenstudy variance τ^2 (argument method. tau) are also available:

- Restricted maximum-likelihood estimator (method.tau="REML")
- Maximum-likelihood estimator (method.tau="ML")
- Hunter-Schmidt estimator (method.tau="HS")
- Sidik-Jonkman estimator (method.tau="SJ")
- Hedges estimator (method.tau="HE")
- Empirical Bayes estimator (method.tau="EB").

For these methods the R function rma.uni of R package metafor is called internally. See help page of R function rma.uni for more details on these methods to estimate between-study variance.

Value

An object of class c("metaprop", "meta") with corresponding print, summary, plot function. The object is a list containing the following components:

```
event, n, studlab,
sm, incr, allincr, addincr, method.ci,
level, level.comb,
                  As defined above.
comb.fixed, comb.random,
hakn, method.tau, tau.preset, TE.tau, method.bias,
tau.common, title, complab, outclab,
byvar, bylab, print.byvar, warn
TE, seTE
                 Estimated (un)transformed proportion and its standard error for individual stud-
lower, upper
                 Lower and upper confidence interval limits for individual studies.
zval, pval
                 z-value and p-value for test of treatment effect for individual studies.
w.fixed, w.random
                 Weight of individual studies (in fixed and random effects model).
TE.fixed, seTE.fixed
                 Estimated overall (un)transformed proportion and standard error (fixed effect
                 model).
lower.fixed, upper.fixed
                 Lower and upper confidence interval limits (fixed effect model).
zval.fixed, pval.fixed
                  z-value and p-value for test of overall effect (fixed effect model).
```

TE.random, seTE.random

Estimated overall (un)transformed proportion and standard error (random effects model).

lower.random, upper.random

Lower and upper confidence interval limits (random effects model).

zval.random, pval.random

z-value or t-value and corresponding p-value for test of overall effect (random

effects model).

prediction, level.predict

As defined above.

seTE.predict Standard error utilised for prediction interval.

lower.predict, upper.predict

Lower and upper limits of prediction interval.

k Number of studies combined in meta-analysis.

Q Heterogeneity statistic Q.

tau Square-root of between-study variance.

se. tau Standard error of square-root of between-study variance.

C Scaling factor utilised internally to calculate common tau-squared across sub-

groups.

sm A character string: "proportion"

method A character string indicating method used for pooling: "Inverse"

df.hakn Degrees of freedom for test of treatment effect for Hartung-Knapp method (only

if hakn=TRUE).

incr.event Increment added to number of events.

keepdata As defined above.

data Original data (set) used in function call (if keepdata=TRUE).

subset Information on subset of original data used in meta-analysis (if keepdata=TRUE).

call Function call.

version Version of R package meta used to create object.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

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Freeman MF & Tukey JW (1950), Transformations related to the angular and the square root. *Annals of Mathematical Statistics*, **21**, 607–611.

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Miller JJ (1978), The inverse of the Freeman-Tukey double arcsine transformation. *The American Statistician*, **32**, 138.

Newcombe RG (1998), Two-sided confidence intervals for the single proportion: Comparison of seven methods. *Statistics in Medicine*, **17**, 857–872.

Paule RC & Mandel J (1982), Consensus values and weighting factors. *Journal of Research of the National Bureau of Standards*, **87**, 377–385.

Pettigrew HM, Gart JJ, Thomas DG (1986), The bias and higher cumulants of the logarithm of a binomial variate. *Biometrika*, **73**, 425–435.

Viechtbauer W (2010), Conducting Meta-Analyses in R with the Metafor Package. *Journal of Statistical Software*, **36**, 1–48.

See Also

```
update.meta, metacont, metagen, print.meta
```

```
#
# Apply various meta-analysis methods to estimate proportions
#
m1 \leftarrow metaprop(4:1, c(10, 20, 30, 40))
m2 <- update(m1, sm="PAS")</pre>
m3 <- update(m1, sm="PRAW")
m4 <- update(m1, sm="PLN")
m5 <- update(m1, sm="PFT")</pre>
m1
m2
m3
m4
m5
#
forest(m1)
# forest(m2)
# forest(m3)
# forest(m3, pscale=100)
# forest(m4)
# forest(m5)
# Do not back transform results, e.g. print logit transformed
# proportions if sm="PLOGIT"
oldset <- settings.meta(backtransf=FALSE)</pre>
m6 < -metaprop(4:1, c(10, 20, 30, 40))
m7 <- update(m6, sm="PAS")
```

```
m8 <- update(m6, sm="PRAW")</pre>
m9 <- update(m6, sm="PLN")</pre>
m10 <- update(m6, sm="PFT")</pre>
forest(m6)
# forest(m7)
# forest(m8)
# forest(m8, pscale=100)
# forest(m9)
# forest(m10)
# Reset settings
settings.meta(oldset)
#
# Examples with zero events
m1 \leftarrow metaprop(c(0, 0, 10, 10), rep(100, 4))
m2 \leftarrow metaprop(c(0, 0, 10, 10), rep(100, 4), incr=0.1)
summary(m1)
summary(m2)
# forest(m1)
# forest(m2)
# Example from Miller (1978):
death <- c(3, 6, 10, 1)
animals <- c(11, 17, 21, 6)
m3 <- metaprop(death, animals, sm="PFT")</pre>
forest(m3)
# Data examples from Newcombe (1998)
# - apply various methods to estimate confidence intervals for
#
    individual studies
event <- c(81, 15, 0, 1)
n < -c(263, 148, 20, 29)
m1 <- metaprop(event, n, sm="PLOGIT", method.ci="SA")</pre>
m2 <- update(m1, method.ci="SACC")</pre>
m3 <- update(m1, method.ci="WS")</pre>
m4 <- update(m1, method.ci="WSCC")</pre>
m5 <- update(m1, method.ci="CP")</pre>
#
```

```
lower <- round(rbind(NA, m1$lower, m2$lower, NA, m3$lower, m4$lower, NA, m5$lower), 4)</pre>
upper <- round(rbind(NA, m1$upper, m2$upper, NA, m3$upper, m4$upper, NA, m5$upper), 4)
tab1 <- data.frame(</pre>
  scen1=meta:::p.ci(lower[,1], upper[,1]),
  scen2=meta:::p.ci(lower[,2], upper[,2]),
  scen3=meta:::p.ci(lower[,3], upper[,3]),
  scen4=meta:::p.ci(lower[,4], upper[,4]),
  stringsAsFactors=FALSE
  )
names(tab1) <- c("r=81, n=263", "r=15, n=148", "r=0, n=20", "r=1, n=29")
"Binomial", "- CP")
tab1[is.na(tab1)] <- ""
# Newcombe (1998), Table I, methods 1-5:
#
tab1
# Same confidence interval, i.e. unaffected by choice of summary measure
print(metaprop(event, n, sm="PLOGIT", method.ci="WS"), ma=FALSE)
print(metaprop(event, n, sm="PLN", method.ci="WS"), ma=FALSE)
print(metaprop(event, n, sm="PFT", method.ci="WS"), ma=FALSE)
print(metaprop(event, n, sm="PAS", method.ci="WS"), ma=FALSE)
print(metaprop(event, n, sm="PRAW", method.ci="WS"), ma=FALSE)
# Different confidence intervals as argument sm="NAsm"
print(metaprop(event, n, sm="PLOGIT", method.ci="NAsm"), ma=FALSE)
print(metaprop(event, n, sm="PLN", method.ci="NAsm"), ma=FALSE)
print(metaprop(event, n, sm="PFT", method.ci="NAsm"), ma=FALSE)
print(metaprop(event, n, sm="PAS", method.ci="NAsm"), ma=FALSE)
print(metaprop(event, n, sm="PRAW", method.ci="NAsm"), ma=FALSE)
# Different confidence intervals as argument backtransf=FALSE.
# Accordingly, method.ci="NAsm" used internally.
print(metaprop(event, n, sm="PLOGIT", method.ci="WS"), ma=FALSE, backtransf=FALSE)
\label{eq:print}  \texttt{print}(\texttt{metaprop}(\texttt{event}, \ \mathsf{n}, \ \texttt{sm="PLN"}, \ \texttt{method.ci="WS"}), \ \texttt{ma=FALSE}, \ \texttt{backtransf=FALSE}) 
print(metaprop(event, n, sm="PFT", method.ci="WS"), ma=FALSE, backtransf=FALSE)
print(metaprop(event, n, sm="PAS", method.ci="WS"), ma=FALSE, backtransf=FALSE)
print(metaprop(event, n, sm="PRAW", method.ci="WS"), ma=FALSE, backtransf=FALSE)
# Same results (printed on original and log scale, respectively)
print(metaprop(event, n, sm="PLN", method.ci="NAsm"), ma=FALSE)
print(metaprop(event, n, sm="PLN"), ma=FALSE, backtransf=FALSE)
# Results for first study (on log scale)
round(log(c(0.3079848, 0.2569522, 0.3691529)), 4)
```

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metareg	Meta-regression	

Description

Meta-regression for objects of class meta. This is a wrapper function for the R function rma. uni in the R package metafor (Viechtbauer 2010).

Usage

```
metareg(x, formula, method.tau=x$method.tau, ...)
```

Arguments

x An object of class meta.

formula Either a character string or a formula object.

method.tau A character string indicating which method is used to estimate the between-study variance tau-squared. Either "FE", "DL", "REML", "ML", "HS", "SJ", "HE", or "EB", can be abbreviated.

... Additional arguments (ignored at the moment).

Details

This R function is a wrapper function for R function rma. uni in the R package metafor (Viechtbauer 2010), i.e. the function metareg can only be used if the R package metafor is installed.

Value

An object of class c("metareg", "rma.uni","rma"). Please look at the help page of R function rma.uni for more details on the output from this function.

In addition, a list .meta is added to the output containing the following components:

x, formula, method.tau

As definied above.

call Function call.

version Version of R package meta used to create object.

version.metafor

Version of R package metafor used to create object.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Viechtbauer W (2010), Conducting Meta-Analyses in R with the Metafor Package. *Journal of Statistical Software*, **36**, 1–48.

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See Also

```
bubble, summary.meta, metagen
```

```
data(Fleiss93cont)
# Add some (fictious) grouping variables:
Fleiss93cont$age <- c(55, 65, 55, 65, 55)
Fleiss93cont$region <- c("Europe", "Europe", "Asia", "Asia", "Europe")
meta1 <- metacont(n.e, mean.e, sd.e,</pre>
                  n.c, mean.c, sd.c,
                  data=Fleiss93cont, sm="MD")
mu1 <- update(meta1, byvar=region)</pre>
mu2 <- update(meta1, byvar=region,</pre>
              tau.common=TRUE, comb.fixed=FALSE)
## Not run:
# Warnings due to wrong ordering of arguments (order has changed with
# version 3.0-0 of R package meta)
metareg(~region, meta1)
metareg(~region, data=meta1)
# Warning as no information on covariate is available
metareg(meta1)
## End(Not run)
# Do meta-regression for covariate region
# (see R code to create object mu2)
metareg(mu2)
# Same result for
# - tau-squared
# - test of heterogeneity
# - test for subgroup differences
# (as argument 'tau.common' was used to create mu2)
mu2
metareg(mu2)
metareg(meta1, region)
# Different result for
# - tau-squared
# - test of heterogeneity
```

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```
# - test for subgroup differences
# (as argument 'tau.common' is - by default - FALSE)
#
mu1

# Generate bubble plot
#
bubble(metareg(mu2))

# Do meta-regression with two covariates
#
metareg(mu1, region + age)

# Do same meta-regressions using 'official' formula notation
#
metareg(meta1, ~region)
metareg(mu1, ~region + age)
```

Olkin95

Thrombolytic Therapy after Acute Myocardial Infarction

Description

Meta-analysis on Thrombolytic Therapy after Acute Myocardial Infarction

Usage

```
data(Olkin95)
```

Format

A data frame with the following columns:

```
author First author
year Year of publication
event.e Number of events in experimental group
n.e Number of observations in experimental group
event.c Number of events in control group
n.c Number of observations in control group
```

Source

Olkin I (1995), Statistical and theoretical considerations in meta-analysis. *Journal of Clinical Epidemiology*, **48**, 133–146.

```
data(01kin95)
summary(metabin(event.e, n.e, event.c, n.c, data=01kin95))
```

print.meta

Print and summary method for objects of class meta

Description

Print and summary method for objects of class meta.

Usage

```
## S3 method for class 'meta'
print(x, sortvar,
        comb.fixed=x$comb.fixed,
        comb.random=x$comb.random,
        prediction=x$prediction,
        details=FALSE, ma=TRUE, backtransf=x$backtransf,
        digits=max(4, .Options$digits - 3), ...)
## S3 method for class 'metabias'
print(x, ...)
## S3 method for class 'meta'
summary(object,
        comb.fixed=object$comb.fixed, comb.random=object$comb.random,
        prediction=object$prediction,
        backtransf=object$backtransf,
        bylab=object$bylab, print.byvar=object$print.byvar,
        bystud=FALSE, print.CMH=object$print.CMH,
        warn=object$warn, ...)
## S3 method for class 'summary.meta'
print(x, digits = max(3, .0ptions$digits - 3),
        comb.fixed=x$comb.fixed, comb.random=x$comb.random,
        prediction=x$prediction,
        print.byvar=x$print.byvar, print.CMH=x$print.CMH,
        header=TRUE, backtransf=x$backtransf,
        bylab.nchar=35, ...)
cilayout(bracket="[", separator="; ")
```

Arguments

comb.fixed

X	An object of class meta, metabias, or summary.meta.
object	An object of class meta.
sortvar	An optional vector used to sort the individual studies (must be of same length as x \$TE).

A logical indicating whether a fixed effect meta-analysis should be conducted.

comb.random	A logical indicating whether a random effects meta-analysis should be conducted.
prediction	A logical indicating whether a prediction interval should be printed.
bylab	A character string with a label for the grouping variable.
print.byvar	A logical indicating whether the name of the grouping variable should be printed in front of the group labels.
header	A logical indicating whether information on title of meta-analysis, comparison and outcome should be printed at the beginning of the printout.
details	A logical indicating whether further details of individual studies should be printed.
ma	A logical indicating whether the summary results of the meta-analysis should be printed.
backtransf	A logical indicating whether printed results should be back transformed. If backtransf=TRUE, results for sm="OR" are printed as odds ratios rather than log odds ratios and results for sm="ZCOR" are printed as correlations rather than Fisher's z transformed correlations, for example.
bylab.nchar	A numeric specifying the number of characters to print from label for the grouping variable.
bystud	A logical indicating whether results of individual studies should be printed by grouping variable.
print.CMH	A logical indicating whether result of the Cochran-Mantel-Haenszel test for overall effect should be printed.
digits	Minimal number of significant digits, see print.default.
warn	A logical indicating whether the use of summary. meta in connection with metacum or metainf should result in a warning.
bracket	A character with bracket symbol to print lower confidence interval: "[", "(", " $\{$ ", "".
separator	A character string with information on separator between lower and upper confidence interval.
	Additional arguments.

Details

Note, in R package meta, version 3.0-0 some arguments have been removed from R functions summary.meta (arguments: byvar, level, level.comb, level.prediction) and print.summary.meta (arguments: level, level.comb, level.prediction). This functionality is now provided by R function update.meta (or directly in R functions metabin, metacont, metagen, metacor, and metaprop).

Review Manager 5 (RevMan 5) is the current software used for preparing and maintaining Cochrane Reviews (http://www.cc-ims.net/revman/). In RevMan 5, subgroup analyses can be defined and data from a Cochrane review can be imported to R using the function read.rm5. If a meta-analysis is then conducted using function metacr, information on subgroups is available in R (components byvar, bylab, and print.byvar, byvar in an object of class "meta"). Accordingly, by using function metacr there is no need to define subgroups in order to redo the statistical analysis conducted in the Cochrane review.

Note, for an object of type metaprop, starting with version 3.7-0 of meta, list elements TE, lower and upper in element study correspond to transformed proportions and confidence limits (regardless whether exact confidence limits are calculated; argument ciexact=TRUE in metaprop function). Accordingly, the following results are based on the same transformation defined by argument sm: list elements TE, lower and upper in elements study, fixed, random, within.fixed and within.random.

R function cilayout can be utilised to change the layout to print confidence intervals (both in printout from print.meta and print.summary.meta function as well as in forest plots). The default layout is "[lower; upper]". Another popular layout is "(lower - upper)" which is used throughout an R session by using R command cilayout("(", " - ").

Value

within.fixed

A list is returned by the function summary.meta with the following elements:

study	Results for individual studies (a list with elements TE, seTE, lower, upper, z, p, level, df).
fixed	Results for fixed effect model (a list with elements TE, seTE, lower, upper, z, p, level, df).
random	Results for random effects model (a list with elements TE, seTE, lower, upper, z, p, level, df).
k	Number of studies combined in meta-analysis.
Q	Heterogeneity statistic Q.
tau	Square-root of between-study variance.
se.tau	Standard error of square-root of between-study variance.
С	Scaling factor utilised internally to calculate common tau-squared across subgroups.
Н	Heterogeneity statistic H (a list with elements TE, lower, upper).
I2	Heterogeneity statistic I2 (a list with elements TE, lower, upper), see Higgins & Thompson (2002).
k.all	Total number of studies.
Q.CMH	Cochran-Mantel-Haenszel test statistic for overall effect.
sm	A character string indicating underlying summary measure.
method	A character string with the pooling method.
call	Function call.
ci.lab	Label for confidence interval.
hakn	A logical indicating whether method by Hartung and Knapp was used.
method.tau	A character string indicating which method is used to estimate the between-study variance tau-squared.
tau.common	A logical indicating whether tau-squared is assumed to be the same across subgroups.
	D. 14 C. C. 1 CC 4 1.1 Cd (114 TE TE 1

Result for fixed effect model within groups (a list with elements TE, seTE, lower,

upper, z, p, level, df, harmonic.mean) - if byvar is not missing.

within.random	Result for random effects model within groups (a list with elements TE, seTE, lower, upper, z, p, level, df, harmonic.mean) - if byvar is not missing.	
k.w	Number of studies combined within groups - if byvar is not missing.	
Q.w	Heterogeneity statistic Q within groups - if byvar is not missing.	
Q.b.fixed	Heterogeneity statistic Q between groups (based on fixed effect model) - if byvar is not missing.	
Q.b.random	Heterogeneity statistic \boldsymbol{Q} between groups (based on random effects model) - if byvar is not missing.	
tau.w	Square-root of between-study variance within subgroups - if byvar is not missing.	
C.w	Scaling factor utilised internally to calculate common tau-squared across subgroups.	
H.w	Heterogeneity statistic H within subgroups (a list with elements TE, lower, upper) - if byvar is not missing.	
I2.w	Heterogeneity statistic I2 within subgroups (a list with elements TE, lower, upper) - if byvar is not missing.	
by.levs	Levels of grouping variable - if byvar is not missing.	
title	Title of meta-analysis / systematic review.	
complab	Comparison label.	
outclab	Outcome label.	
data	Original data (set) used to create meta object.	
subset	Information on subset of original data used in meta-analysis.	
prediction, level.predict		
comb.fixed, com	mb.random, print.CMH As defined above.	
version	Version of R package meta used to create object.	

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Cooper H & Hedges LV (1994), *The Handbook of Research Synthesis*. Newbury Park, CA: Russell Sage Foundation.

Higgins JPT & Thompson SG (2002), Quantifying heterogeneity in a meta-analysis. *Statistics in Medicine*, **21**, 1539–1558.

See Also

 ${\tt update.meta}, {\tt metabin}, {\tt metacont}, {\tt metagen}$

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Examples

```
data(Fleiss93cont)
meta1 <- metacont(n.e, mean.e, sd.e, n.c, mean.c, sd.c, data=Fleiss93cont, sm="SMD")
summary(meta1)
summary(update(meta1, byvar=c(1,2,1,1,2), bylab="group"))
forest(update(meta1, byvar=c(1,2,1,1,2), bylab="group"))</pre>
```

print.rm5

Print and summary methods for objects of class rm5

Description

Print and summary methods for objects of class rm5.

Usage

Arguments

x An object of class rm5.
object An object of class rm5.
comp.no Comparison number.
outcome.no Outcome number.
method.bias A character string indicating which test for small-study effects is to be used for all outcomes. Either "rank", "linreg", or "mm", can be abbreviated. See function metabias
method.bias.binary

A character string indicating which test is to be used for binary outcomes. Either "rank", "linreg", "mm", "count", "score", or "peters", can be abbreviated. See function metabias

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```
method.bias.or A character string indicating which test is to be used for binary outcomes with odds ratio as summary measure. Either "rank", "linreg", "mm", "count", "score", or "peters", can be abbreviated. See function metabias k.min Minimum number of studies to perform test for small-study effects.

Additional arguments (ignored at the moment)
```

Details

Review Manager 5 (RevMan 5) is the current software used for preparing and maintaining Cochrane Reviews (http://www.cc-ims.net/revman/). In RevMan 5, subgroup analyses can be defined and data from a Cochrane review can be imported to R using the function read.rm5.

The R function summary.rm5 can be used to redo all meta-analyses of the imported Cochrane Review.

The R function metabias.rm5 can be used to conduct a test for funnel plot asymmetry for all meta-analyses of the imported Cochrane Review.

The R function metacr is called internally.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Higgins, J.P.T and S. Green (2011), Cochrane Handbook for Systematic Reviews of Interventions Version 5.1.0 [Updated March 2011]. The Cochrane Library: http://www.cochrane-handbook.org

See Also

```
metabias.meta, summary.meta, read.rm5
```

```
# Locate export data file "Fleiss93_CR.csv"
# in sub-directory of package "meta"
#
filename <- system.file("data/Fleiss93_CR.csv.gz", package = "meta")
#
Fleiss93_CR <- read.rm5(filename)
#
# Print summary results for all meta-analysis:
# summary(Fleiss93_CR)
#
# Print results for tests of small-study effects:
# metabias(Fleiss93_CR, k.min=5)</pre>
```

read.mtv 87

read.mtv	Import RevMan 4 data files (.mtv)

Description

Reads a file created with RevMan 4 and creates a data frame from it.

Usage

```
read.mtv(file)
```

Arguments

file The name of a file to read data values from.

Details

Reads a file created with RevMan 4 (Menu: "File" - "Export" - "Analysis data file...") and creates a data frame from it.

Value

A data frame containing the following components:

comp.no	Comparison number.
outcome.no	Outcome number.
group.no	Group number.
studlab	Study label.
year	Year of publication.
event.e	Number of events in experimental group.
n.e	Number of observations in experimental group.
event.c	Number of events in control group.
n.c	Number of observations in control group.
mean.e	Estimated mean in experimental group.
sd.e	Standard deviation in experimental group.
mean.c	Estimated mean in control group.
sd.c	Standard deviation in control group.
0.E	Observed minus expected (IPD analysis).
V	Variance of 0.E (IPD analysis).
order	Ordering of studies.
conceal	Concealment of treatment allocation.
grplab	Group label.

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type	Type of outcome. $D = dichotomous$, $C = continuous$, $P = IPD$.
outclab	Outcome label.
graph.exp	Graph label for experimental group.
graph.cont	Graph label for control group.
label.exp	Label for experimental group.
label.cont	Label for control group.
complab	Comparison label.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Review Manager (RevMan) [Computer program]. Version 4.2 for Windows. Copenhagen: The Nordic Cochrane Centre, The Cochrane Collaboration, 2003.

See Also

```
metabin, metacont, metagen
```

read.rm5

|--|

Description

Reads data file from Cochrane Intervention review created with RevMan 5 and creates a data frame from it.

Usage

Arguments

file The name of a file to read data values from.

sep The field separator character. Values on each line of the file are separated by this

character. The comma is the default field separator character in RevMan 5.

quote The set of quoting characters. In RevMan 5 a "\"" is the default quoting charac-

ter.

title Title of Cochrane review.

numbers.in.labels

A logical indicating whether comparision number and outcome number should be printed at the beginning of the comparison (argument complab) and outcome label (argument outclab); this is the default in RevMan 5.

Details

Review Manager 5 (RevMan 5) is the current software used for preparing and maintaining Cochrane Reviews (http://www.cc-ims.net/revman/). RevMan 5 includes the ability to write Systematic reviews of interventions, Diagnostic test accuracy reviews, Methodology reviews and Overviews of reviews.

This function provides the ability to read a data file from a Cochrane Intervention review created with RevMan 5; a data frame is created from it. Cochrane Intervention reviews are based on the comparison of two interventions.

In order to generate a data analysis file in RevMan 5 use the following Menu points: "File" - "Export" - "Data and analyses". It is mandatory to include the following fields in the exported data file by selecting them with the mouse cursor in the Export Analysis Data Wizard: (i) Comparison Number, (ii) Outcome Number, (iii) Subgroup Number. When these fields are not selected a corresponding error message will be printed in R. It is recommended to include all fields in the exported data file except for the last field "Risk of bias tables". For example, in order to redo the meta-analysis in R for the RevMan 5 data type "0-E and Variance" the fields "0-E" and "Variance" have to be selected in the Export Analysis Data Wizard. If the last field "Risk of bias tables" is selected the import in R fails with an error message "line X did not have Y elements".

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By default in RevMan 5, the name of the exported data file is the title of the Cochrane Review. Accordingly, information on the title is extracted from the name of the exported data file (argument: file) if argument title is missing (default).

Each respective meta-analysis for arguments event.e.pooled – df.pooled is defined by values for "comp.no" and "outcome.no", and "grp.no".

Value

A data frame containing the following components:

comp.no Comparison number.
outcome.no Outcome number.
group.no Group number.
studlab Study label.

year Year of publication.

event.e Number of events in experimental group.

n.e Number of observations in experimental group.

event.c Number of events in control group.

n.c Number of observations in control group.

mean.e Estimated mean in experimental group.

sd.e Standard deviation in experimental group.

mean.c Estimated mean in control group.

sd.c Standard deviation in control group.

0.E Observed minus expected (IPD analysis).

V Variance of 0.E (IPD analysis).

TE, seTE Estimated treatment effect and standard error of individual studies.

lower, upper Lower and upper limit of 95% confidence interval for treatment effect in indi-

vidual studies.

weight Weight of individual studies (according to meta-analytical method used in re-

spective meta-analysis - see below for details).

order Ordering of studies.

grplab Group label.

type Type of outcome. D = dichotomous, C = continuous, P = IPD.

method A character string indicating which method has been used for pooling of studies.

One of "Inverse", "MH", or "Peto".

sm A character string indicating which summary measure has been used for pooling

of studies.

model A character string indicating which meta-analytical model has been used (either

"Fixed" or "Random").

comb. fixed A logical indicating whether fixed effect meta-analysis has been used in respec-

tive meta-analysis (see below for details).

read.rm5

comb.random	A logical indicating whether random effects meta-analysis has been used in respective meta-analysis (see below for details).
outclab	Outcome label.
k	Total number of studies combined in respective meta-analysis).
event.e.pooled	Number of events in experimental group in respective meta-analysis (see below for details).
n.e.pooled	Number of observations in experimental group in respective meta-analysis (see below for details).
event.c.pooled	Number of events in control group in respective meta-analysis (see below for details).
n.c.pooled	Number of observations in control group in respective meta-analysis (see below for details).
TE.pooled	Estimated treatment effect in respective meta-analysis (see below for details).
lower, upper	Lower and upper limit of 95% confidence interval for treatment effect in respective meta-analysis (see below for details).
weight.pooled	Total weight in respective meta-analysis (see below for details).
Z.pooled	Z-score for test of overall treatment effect in respective meta-analysis (see below for details).
pval.pooled	P-value for test of overall treatment effect in respective meta-analysis (see below for details).
Q	Heterogeneity statistic Q in respective meta-analysis (see below for details).
pval.Q	P-value of heterogeneity statistic Q in respective meta-analysis (see below for details).
I2	Heterogeneity statistic I2 in respective meta-analysis (see below for details).
tau2	Between-study variance (moment estimator of DerSimonian-Laird) in respective meta-analysis.
Q.w	Heterogeneity statistic Q within groups in respective meta-analysis (see below for details).
pval.Q.w	P-value of heterogeneity statistic Q within groups in respective meta-analysis (see below for details).
I2.w	Heterogeneity statistic I2 within groups in respective meta-analysis (see below for details).
label.e	Label for experimental group.
label.c	Label for control group.
label.left	Graph label on left side of forest plot.
label.right	Graph label on right side of forest plot.
RR.cochrane	A logical indicating if 2*incr instead of 1*incr is to be added to n.e and n.c in the calculation of the risk ratio (i.e., sm="RR") for studies with a zero cell. This is used in RevMan 5.
complab	Comparison label.

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Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

References

Review Manager (RevMan) [Computer program]. Version 5.1. Copenhagen: The Nordic Cochrane Centre, The Cochrane Collaboration, 2011.

See Also

```
metabin, metacont, metagen, metacr
```

Examples

```
# Locate export data file "Fleiss93_CR.csv"
# in sub-directory of package "meta"
#
filename <- system.file("data/Fleiss93_CR.csv.gz", package = "meta")
#
Fleiss93_CR <- read.rm5(filename)
# Same result as R command example(Fleiss93):
#
metacr(Fleiss93_CR)
# Same result as R command example(Fleiss93cont):
#
metacr(Fleiss93_CR, 1, 2)</pre>
```

settings.meta

Print and change default settings for meta-analyses in R package meta.

Description

Print and change default settings for meta-analyses in R package meta.

Usage

```
settings.meta(...)
```

Arguments

... Arguments to change default settings.

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Details

This function can be used to define defaults for several arguments of the following R functions: metabin, metacont, metacor, metacor, metagen, metainc, metaprop.

A list of all arguments with current settings is printed using the command settings.meta(print=TRUE).

In order to reset all settings to the default settings of R package meta the command settings.meta(reset=TRUE) can be used.

Author(s)

Guido Schwarzer <sc@imbi.uni-freiburg.de>

```
# Get list of settings
settings.meta(print=TRUE)
# Meta-analyses using default settings of R pacakge meta
metabin(10, 20, 15, 20, warn=FALSE)
metaprop(4, 20)
metabin(10, 20, 15, 20, sm="RD", warn=FALSE)
metaprop(4, 20, sm="PLN")
# Change default summary measure for R functions metabin and metaprop
oldset <- settings.meta(smbin="RD", smprop="PLN")</pre>
metabin(10, 20, 15, 20, warn=FALSE)
metaprop(4, 20)
# Reset settings
settings.meta(oldset)
# Change level for confidence intervals
metagen(1:3, (2:4)/10, sm="MD")
settings.meta(level=0.99, level.comb=0.999)
metagen(1:3, (2:4)/10, sm="MD")
# Always print a prediction interval
settings.meta(prediction=TRUE)
metagen(1:3, (2:4)/10, sm="MD")
#
```

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```
# Try to set unknown argument
settings.meta(unknownarg=TRUE)
# Reset to default settings of R package meta
#
settings.meta(reset=TRUE)
metabin(10, 20, 15, 20, warn=FALSE)
metaprop(4, 20)
metagen(1:3, (2:4)/10, sm="MD")
#
# Never back transform results (e.g. print log odds ratios instead of
# odds ratios, print transformed correlations/proportions instead of
# correlations/proportions)
#
oldset <- settings.meta(backtransf=FALSE)</pre>
metabin(10, 20, 15, 20, warn=FALSE)
metaprop(4, 20)
metacor(c(0.85, 0.7, 0.95), c(20, 40, 10))
settings.meta(oldset)
```

smoking

Smoking example

Description

Meta-analyses on the effect of smoking on mortality risk.

Data have been reconstructed based on the famous Smoking and Health Report to the Surgeon General (Bayne-Jones S et al., 1964). Data sets can be used to evaluate the risk of smoking on overall mortality and lung-cancer deaths, respectively. The person time is attributed such that the rate ratios are equal to the reported mortality ratios implicitly assuming that the data have arisen from a homogeneous age group; more detailed information by age is not available from the report. Note, the group of "non-smokers" actually consists of all participants except those who are smokers of cigarettes only. Information on real non-smokers is not available from the published Smoking and Health Report.

Usage

```
data(smoking)
data(lungcancer)
```

Format

A data frame with the following columns:

study Study label

```
participants Total number of participants
d.smokers Number of deaths in smokers' group
py.smokers Person years at risk in smokers' group
d.nonsmokers Number of deaths in non-smokers' group
py.nonsmokers Person years at risk in non-smokers' group
```

Source

Bayne-Jones S et al. (1964), Smoking and Health: Report of the Advisory Committee to the Surgeon General of the United States. U-23 Department of Health, Education, and Welfare. Public Health Service Publication No. 1103. http://profiles.nlm.nih.gov/ps/retrieve/ResourceMetadata/NNBBMQ

See Also

metainc

Examples

trimfill

Trim-and-fill method to adjust for bias in meta-analysis

Description

Trim-and-fill method for estimating and adjusting for the number and outcomes of missing studies in a meta-analysis.

Usage

```
trimfill(x, ...)
## Default S3 method:
trimfill(x, seTE, left=NULL, ma.fixed=TRUE, type="L", n.iter.max=50,
```

```
sm=NULL, studlab=NULL, level=0.95, level.comb=level,
    comb.fixed=FALSE, comb.random=TRUE,
    hakn=FALSE, method.tau="DL",
    prediction=FALSE, level.predict=level,
    backtransf=TRUE,
    silent=TRUE, ...)

## S3 method for class 'meta'
trimfill(x, left=NULL, ma.fixed=TRUE, type="L", n.iter.max=50,
    level=x$level, level.comb=x$level.comb,
    comb.fixed=FALSE, comb.random=TRUE,
    hakn=x$hakn, method.tau=x$method.tau,
    prediction=x$prediction, level.predict=x$level.predict,
    backtransf=x$backtransf,
    silent=TRUE, ...)
```

Arguments

prediction

X	An object of class meta, or estimated treatment effect in individual studies.
seTE	Standard error of estimated treatment effect.
left	A logical indicating whether studies are supposed to be missing on the left or right side of the funnel plot. If NULL, the linear regression test for funnel plot symmetry (i.e., function metabias(, method="linreg")) is used to determine whether studies are missing on the left or right side.
ma.fixed	A logical indicating whether a fixed effect or random effects model is used to estimate the number of missing studies.
type	A character indicating which method is used to estimate the number of missing studies. Either "L" or "R".
n.iter.max	Maximum number of iterations to estimate number of missing studies.
sm	An optional character string indicating underlying summary measure, e.g., "RD", "RR", "OR", "AS", "MD", "SMD"; ignored if x is of class meta.
studlab	An optional vector with study labels; ignored if x is of class meta.
level	The level used to calculate confidence intervals for individual studies. If existing, x\$level is used as value for level; otherwise 0.95 is used.
level.comb	The level used to calculate confidence interval for the pooled estimate. If existing, x\$level.comb is used as value for level.comb; otherwise 0.95 is used.
comb.fixed	A logical indicating whether a fixed effect meta-analysis should be conducted.
comb.random	A logical indicating whether a random effects meta-analysis should be conducted.
hakn	A logical indicating whether the method by Hartung and Knapp should be used to adjust test statistics and confidence intervals.
method.tau	A character string indicating which method is used to estimate the between-study variance τ^2 . Either "DL", "PM", "REML", "ML", "HS", "SJ", "HE", or "EB", can be abbreviated.

A logical indicating whether a prediction interval should be printed.

level.predict The level used to calculate prediction interval for a new study.

backtransf A logical indicating whether results should be back transformed in printouts and

plots. If backtransf=TRUE, results for sm="OR" are printed as odds ratios rather than log odds ratios and results for sm="ZCOR" are printed as correlations rather

than Fisher's z transformed correlations, for example.

silent A logical indicating whether basic information on iterations shown.

... other arguments

Details

The trim-and-fill method (Duval, Tweedie 2000a, 2000b) can be used for estimating and adjusting for the number and outcomes of missing studies in a meta-analysis. The method relies on scrutiny of one side of a funnel plot for asymmetry assumed due to publication bias.

Three different methods have been proposed originally to estimate the number of missing studies. Two of these methods (L- and R-estimator) have been shown to perform better in simulations, and are available in this R function (argument type).

A fixed effect or random effects model can be used to estimate the number of missing studies (argument ma.fixed). Furthermore, a fixed effect and/or random effects model can be used to summaries study results (arguments comb.fixed and comb.random). Simulation results (Peters et al. 2007) indicate that the fixed-random model, i.e. using a fixed effect model to estimate the number of missing studies and a random effects model to summaries results, (i) performs better than the fixed-fixed model, and (ii) performs no worse than and marginally better in certain situations than the random-random model. Accordingly, the fixed-random model is the default.

An empirical comparison of the trim-and-fill method and the Copas selection model (Schwarzer et al. 2010) indicates that the trim-and-fill method leads to excessively conservative inference in practice. The Copas selection model is available in R package copas.

The function metagen is called internally.

Value

An object of class c("metagen", "meta", "trimfill"). The object is a list containing the following components:

```
studlab, sm, left, ma.fixed, type,
```

n.iter.max, level, level.comb, level.predict,

As defined above.

comb.fixed, comb.random, prediction, hakn, method.tau

TE, seTE Estimated treatment effect and standard error of individual studies.

lower, upper Lower and upper confidence interval limits for individual studies.

zval, pval z-value and p-value for test of treatment effect for individual studies.

w.fixed, w.random

Weight of individual studies (in fixed and random effects model).

TE.fixed, seTE.fixed

Estimated overall treatment effect and standard error (fixed effect model).

TE.random, seTE.random

Estimated overall treatment effect and standard error (random effects model).

seTE.predict Standard error utilised for prediction interval.

lower.predict, upper.predict

Lower and upper limits of prediction interval.

k Number of studies combined in meta-analysis.

Q Heterogeneity statistic Q.

tau Square-root of between-study variance.

method Pooling method: "Inverse".

call Function call.

n.iter Actual number of iterations to estimate number of missing studies.

trimfill A logical vector indicating studies that have been added by trim-and-fill method.

df. hakn Degrees of freedom for test of treatment effect for Hartung-Knapp method (only

if hakn=TRUE).

title Title of meta-analysis / systematic review.

complab Comparison label.
outclab Outcome label.

label.e Label for experimental group.

label.c Label for control group.

label.left Graph label on left side of forest plot.label.right Graph label on right side of forest plot.k0 Number of studies added by trim-and-fill.

n.e Number of observations in experimental group (only for object x of class metabin

or metacont).

n.c Number of observations in control group (only for object x of class metabin or

metacont).

event.e Number of events in experimental group (only for object x of class metabin).

event.c Number of events in control group (only for object x of class metabin).

mean.e Estimated mean in experimental group (only for object x of class metacont).

sd.e Standard deviation in experimental group (only for object x of class metacont).

mean.c Estimated mean in control group (only for object x of class metacont).

sd.c Standard deviation in control group (only for object x of class metacont).

n Number of observations (only for object x of class metaprop).

event Number of events (only for object x of class metaprop).

cor Corelation (only for object x of class metacor).

class.x Main class of object x (e.g. 'metabin' or 'metacont').

version Version of R package meta used to create object.

Author(s)

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References

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Peters JL, Sutton AJ, Jones DR, Abrams KR, Rushton L (2007), Performance of the trim and fill method in the presence of publication bias and between-study heterogeneity. *Statistics in Medicine*, **10**, 4544–62.

Schwarzer G, Carpenter J, Rücker G (2010), Empirical evaluation suggests Copas selection model preferable to trim-and-fill method for selection bias in meta-analysis. *Journal of Clinical Epidemiology*, **63**, 282–8.

See Also

metagen, metabias, funnel

Examples

update.meta

Update a meta-analysis object

Description

Update an existing meta-analysis object.

Usage

```
## S3 method for class 'meta'
update(object,
       data=object$data, subset=object$subset,
       studlab=object$data$.studlab,
      method=object$method, sm=object$sm,
       incr=object$incr, allincr=object$allincr,
       addincr=object$addincr, allstudies=object$allstudies,
       MH.exact=object$MH.exact, RR.cochrane=object$RR.cochrane,
       level=object$level, level.comb=object$level.comb,
       comb.fixed=object$comb.fixed, comb.random=object$comb.random,
       hakn=object$hakn, method.tau=object$method.tau,
       tau.preset=object$tau.preset,
       TE.tau=object$TE.tau, tau.common=object$tau.common,
       prediction=object$prediction, level.predict=object$level.predict,
      method.bias=object$method.bias, backtransf = object$backtransf,
       title=object$title, complab=object$complab, outclab=object$outclab,
       label.e=object$label.e, label.c=object$label.c,
       label.left=object$label.left, label.right=object$label.right,
       n.e=object$n.e, n.c=object$n.c,
       pooledvar=object$pooledvar, method.ci=object$method.ci,
       byvar=object$byvar, bylab=object$bylab, print.byvar=object$print.byvar,
       print.CMH=object$print.CMH, keepdata=TRUE,
       left=object$left, ma.fixed=object$ma.fixed,
       type=object$type, n.iter.max=object$n.iter.max,
       warn=object$warn, ...)
```

Arguments

obiect	An object of class meta.

data Dataset.
subset Subset.
studlab Study label.

method A character string indicating which method is to be used for pooling of studies.

One of "Inverse", "MH", or "Peto", can be abbreviated. (only for metabin

object)

sm A character string indicating which summary measure is used for pooling.

incr Could be either a numerical value which is added to each cell frequency for stud-

ies with a zero cell count or the character string "TA" which stands for treatment

arm continuity correction.

allincr A logical indicating if incr is added to each cell frequency of all studies if at

least one study has a zero cell count. If FALSE (default), incr is added only to

each cell frequency of studies with a zero cell count.

addincr A logical indicating if incr is added to each cell frequency of all studies irre-

spective of zero cell counts.

A logical indicating if studies with zero or all events in both groups are to be allstudies included in the meta-analysis (applies only if sm is equal to "RR" or "OR"). MH.exact A logical indicating if incr is not to be added to all cell frequencies for studies with a zero cell count to calculate the pooled estimate based on the Mantel-Haenszel method. RR.cochrane A logical indicating if 2*incr instead of 1*incr is to be added to n.e and n.c in the calculation of the risk ratio (i.e., sm="RR") for studies with a zero cell. This is used in RevMan 5, the Cochrane Collaboration's program for preparing and maintaining Cochrane reviews. level The level used to calculate confidence intervals for individual studies. level.comb The level used to calculate confidence intervals for pooled estimates. comb.fixed A logical indicating whether a fixed effect meta-analysis should be conducted. comb.random A logical indicating whether a random effects meta-analysis should be conducted. hakn A logical indicating whether the method by Hartung and Knapp should be used to adjust test statistics and confidence intervals. method.tau A character string indicating which method is used to estimate the betweenstudy variance τ^2 . Either "DL", "PM", "REML", "ML", "HS", "SJ", "HE", or "EB", can be abbreviated. See function metagen. Prespecified value for the square-root of the between-study variance τ^2 . tau.preset TE.tau Overall treatment effect used to estimate the between-study variance τ^2 . A logical indicating whether tau-squared should be the same across subgroups. tau.common prediction A logical indicating whether a prediction interval should be printed. level.predict The level used to calculate prediction interval for a new study. method.bias A character string indicating which test for funnel plot asymmetry is to be used. Either "rank", "linreg", "mm", "count", "score", or "peters", can be abbreviated. See function metabias backtransf A logical indicating whether results should be back transformed in printouts and plots. If backtransf=TRUE, results for sm="OR" are printed as odds ratios rather than log odds ratios and results for sm="ZCOR" are printed as correlations rather than Fisher's z transformed correlations, for example. title Title of meta-analysis / systematic review. Comparison label. complab outclab Outcome label. label.e Label for experimental group. label.c Label for control group. label.left Graph label on left side of forest plot. label.right Graph label on right side of forest plot. Number of observations in experimental group. (only for metagen object) n.e Number of observations in control group. (only for metagen object) n.c

pooledvar	A logical indicating if a pooled variance should be used for the mean difference (only for metacont object with sm="MD").
method.ci	A character string indicating which method is used to calculate confidence intervals for individual studies. Either "CP", "WS", "WSCC", "AC", "SA",, "SACC", or "NAsm", can be abbreviated. See function metaprop.
byvar	An optional vector containing grouping information (must be of same length as $event.e$).
bylab	A character string with a label for the grouping variable.
print.byvar	A logical indicating whether the name of the grouping variable should be printed in front of the group labels.
print.CMH	A logical indicating whether result of the Cochran-Mantel-Haenszel test for overall effect should be printed.
keepdata	A logical indicating whether original data (set) should be kept in meta object.
left	A logical indicating whether studies are supposed to be missing on the left or right side of the funnel plot. If NULL, the linear regression test for funnel plot symmetry (i.e., function metabias(, method="linreg")) is used to determine whether studies are missing on the left or right side.
ma.fixed	A logical indicating whether a fixed effect or random effects model is used to estimate the number of missing studies.
type	A character indicating which method is used to estimate the number of missing studies. Either "L" or "R".
n.iter.max	Maximum number of iterations to estimate number of missing studies.
warn	A logical indicating whether warnings should be printed (e.g., if incr is added to studies with zero cell frequencies).
	Additional arguments (ignored at the moment).

Details

Wrapper function to update an existing meta-analysis object which was created with R function metabin, metacont, metagen, metacor, metaprop.

This function can also be used for objects of class 'trimfill', 'metacum', and 'metainf'.

More details on function arguments are available in help files of respective R functions, i.e. metabin, metacont, \dots

Value

An object of class "meta" and "metabin", "metacont", "metagen", "metaprop", or "metacor".

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See Also

metabin, metacont, metagen, metaprop, metacor

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