# Package 'mma'

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Type Package

Title Multiple Mediation Analysis

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**Depends** R (>= 2.14.1), gbm, splines, survival, car, gplots

Imports lattice, methods

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**Description** Used for general multiple mediation analysis.

The analysis method is described in Yu and Li (2022) (ISBN: 9780367365479) ``Statistical Methods for Mediation, Confounding and Moderation Analysis Using R and SAS", published by Chapman and Hall/CRC; and Yu et al.(2017) <DOI:10.1016/j.sste.2017.02.001> ``Exploring racial disparity in obesity: a mediation analysis considering geo-coded environmental factors", published on Spatial and Spatio-temporal Epidemiology, 21, 13-23.

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# **Description**

This package is used to identify mediators and for general mediation analysis. Mediation effect refers to the effect conveyed by intervening variables to an observed relationship between an exposure and a response variable (outcome). In this package, the exposure is called the predictor, the intervening variables are called mediators. The mediation effects include the total effect, direct effect, and indirect effect, which are defined and the statistical inferences described in Yu et al.(2014).

# Details

"data.org" is used to identify potential mediators. "med", and "boot.med" are used for statistical inferences on the mediation effects when the predictor is binary or continuous. "mma" is a combined function that automatically identify potential mediators, based on which to make statistical inference on the mediation effects.

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# Examples

```
data("weight_behavior")
#binary predictor
#binary y
x=weight_behavior[,c(2,4:14)]
pred=weight_behavior[,3]
y=weight_behavior[,15]
temp.b.b.glm<-mma(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),binref=c(1,1),
                    catmed=5,catref=1,predref="M",alpha=0.4,alpha2=0.4,n=2,n2=2)
 temp.b.b.mart<-mma(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),binref=c(1,1),
             catmed=5,catref=1,predref="M",alpha=0.4,alpha2=0.4,nonlinear=TRUE,n=2,n2=5)
 #continuous y
 x=weight_behavior[,c(2,4:14)]
pred=weight_behavior[,3]
y=weight_behavior[,1]
temp.b.c.glm<-mma(x,y,pred=pred,mediator=5:12,jointm=list(n=1,j1=7:9),
                     predref="M",alpha=0.4,alpha2=0.4,n2=20)
```

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| boot.med | Statistical Inference on Mediation Analysis with Continuous or Binary<br>Predictor |
|----------|--|
|          |  |

# Description

To make inferences on the mediation effects when the predictor is continuous or binary.

# Usage

# Arguments

| data      | the list of result from data.org that organize the covariates, mediators, predictor and outcome. If data is FALSE, then need to set $x$ , $y$ , dirx, contm, catm, and jointm.  |
|-----------|---|
| margin    | the change in predictor when calculating the mediation effects, see Yu et al. (2014).   |
| n         | the time of resampling in calculating the indirect effects, default is $n=20$ , see Yu et al. (2014).   |
| nonlinear | if TURE, Multiple Additive Regression Trees (MART) will be used to fit the final full model in estimating the outcome. The default value of nonlinear is FALSE, in which case, a generalized linear model will be used to fit the final full model. |
| df1       | if nonlinear is TURE, natural cubic spline will be used to fit the relationship between the predictor and each mediator. The df is the degree of freedom in the ns() function, the default is 1.  |
| nu        | set the parameter "shrinkage" in gbm function if MART is to be used, by default, nu=0.001. See also help(gbm.fit).  |
| D         | set the parameter "interaction.depth" in gbm function if MART is to be used, by default, D=3. See also help(gbm.fit).   |
| distn     | the assumed distribution of the outcome if MART is used for final full model. The default value of distn is "gaussian" for coninuous y, and "bernoulli" for binary y.   |
| n2        | the number of times of bootstrap resampling. The default value is n2=50.  |
| W         | the weight for observations.  |

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x.new of the same format as x, with a new set of covariates and mediators on which to

calculate the mediation effects.

pred.new a new set of predictor(s).
cova.new a new set of covaraite(s).

type the type of prediction when y is class Surv. Is "risk" if not specified.

w.new the weights for new.x.

all.model save all the fitted model from bootstrap samples if TRUE. This needs to be true

to make inferences on moderation effects.

xmod If there is a moderator, xmod gives the moderator's name in cova and/or x.

custom.function

a string of customer defined final model for predicting the outcome(s). The response variable should be noted as "responseY", the dataset should be noted as "dataset123". The weights for observations should be noted as "weights123". The covariates should be in x or pred. Use "~." for all variables in x and pred is allowed. The customer defined model should be able to make prediction by using "predict(object, newdata=...)", where the object is the results of the fitted model. If a specific package needs to be called to fit the model, the user should call the package first. For example, if the gamlss package is used to fit a piecewise polynomial with "age" to be the mediator and "race" be the predictor, we

can set custom.function="gamlss(responseV~b(age,df=3)+race,data=dataset123,tace=FALSE)".

The length of custom.function should be the same as the dimension of y. If cus-

tom.function[j] is NA, the usual method will be used to fit y[j].

para It is for binary predictors. If it is true, we would like the x-m relationship be

fitted parametrically.

echo If echo is FALSE, there is no counting printed for the number of bootstrap iter-

ations.

## Details

The mediators are not tested in this function. data.org should be used for the tests and data organizing, and then the resulting list from data.org can be used directly to define the arguments in this function. boot.med considers all variables in x as mediators or covariates in the final model and all variables identified by contm, binm, catm, or jointm as mediators.

## Value

Returns an mma object.

estimation list the estimation of ie (indirect effect), te (total effect), and de (direct effect

from the predictor) separately.

bootsresults a list where the first item, ie, is a matrix of n2 rows where each column gives

the estimated indirect effect from the corresponding mediator (identified by the column name) from the n2 bootstrap samples; the second item, te, is a vector of estimated total effects from the bootstrap sample; and the 3rd item, de, is a vector of estimated direct effect of the predictor from the bootstrap sample.

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| model       | a list where the first item, MART, is TRUE if mart is fitted for the final model; the second item, Survival, is TRUE if a survival model is fitted; the third item, type, is the type of prediction when a survival model is fitted; the fourth item, model, is the fitted final full model where y is the outcome and all predictor, covariates, and mediators are the explanatory variables; and the fourth item, best.iter is the number of best iterations if MART is used to fit the final model. |
|-------------|--|
| data        | a list that contains all the used data: x=x, y=y, dirx=dirx, binm=binm, contm=contm, catm=catm, jointm=jointm, binpred=FALSE.  |
| boot.detail | a list that contains the mediation effects on each row of new.x: new.x=new.x, te1, de1, ie1.   |
| all_model   | a list with all fitted models from bootstrap samples if all.model is TRUE.   |
| all_iter    | if all.model is TRUE, a matrix with all fitted best iterations if MART is used. Each row is from one bootstrap sample.   |
| all_boot    | if all.model is TRUE, a matrix with all bootstrap samples. Each row is one bootstrap sample.   |

#### Author(s)

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#### References

Yu, Q., Fan, Y., and Wu, X. (2014) <doi:10.4172/2155-6180.1000189>. "General Multiple Mediation Analysis With an Application to Explore Racial Disparity in Breast Cancer Survival," Journal of Biometrics & Biostatistics,5(2): 189.

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## See Also

"med" just estimate the mediation effects.

# Examples

```
data("weight_behavior")
##binary x
#binary y
x=weight_behavior[,c(2,4:14)]
```

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```
pred=weight_behavior[,3]
 y=weight_behavior[,15]
 data.bin < -data.org(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),binref=c(1,1),
                     catmed=5, catref=1, predref="M", alpha=0.4, alpha2=0.4)
 temp1<-boot.med(data=data.bin,n=2,n2=4)</pre>
 temp2<-boot.med(data=data.bin,n=2,n2=4,nu=0.05,nonlinear=TRUE)</pre>
#continuous y
 x=weight_behavior[,c(2,4:14)]
 pred=weight_behavior[,3]
 y=weight_behavior[,1]
 data.cont<-data.org(x,y,pred=pred,mediator=5:12,jointm=list(n=1,j1=7:9),</pre>
                      predref="M",alpha=0.4,alpha2=0.4)
 temp3<-boot.med(data=data.cont,n=2,n2=4)
 temp4<-boot.med(data=data.cont,n=2,n2=4,nu=0.05, nonlinear=TRUE)</pre>
##Surv class outcome (survival analysis)
data(cgd1)
                  #a dataset in the survival package
x=cgd1[,c(4:5,7:12)]
pred=cgd1[,6]
status<-ifelse(is.na(cgd1$etime1),0,1)</pre>
y=Surv(cgd1$futime, status)
#for continuous predictor
data.surv.contx<-data.org(x,y,pred=pred,mediator=1:ncol(x),</pre>
                           alpha=0.5,alpha2=0.5)
temp9.contx<-boot.med(data=data.surv.contx,n=1,n2=2, type="lp")</pre>
temp10.contx<-boot.med(data=data.surv.contx,nonlinear=TRUE,n=1,n2=2)</pre>
#for binary predictor
x=cgd1[,c(5:12)]
pred=cgd1[,4]
data.surv.binx<-data.org(x,y,pred=pred,mediator=1:ncol(x),</pre>
                     alpha=0.4,alpha2=0.4)
temp9.binx<-boot.med(data=data.surv.binx,n=1,n2=2, type="lp")</pre>
summary(temp9.binx)
temp10.binx<-boot.med(data=data.surv.binx,nonlinear=TRUE,n=1,n2=2)</pre>
```

boot.mod

Statistical Inference on Mediation Analysis with Continuous or Binary Predictor at different level of the moderator

## **Description**

To make inferences on the mediation effects when the predictor is continuous or binary at different level of the moderator.

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#### Usage

## **Arguments**

mma1 the mma project generated from the function mma or boot.med.

vari The name of the moderator. The moderator should be included in the original

data as a covariate for the response(in x) and/or for mediators (in cova).

continuous.resolution

The number of equally space points (for continuous moderator) or levels (for categorical moderator) at which to evaluate mediation effects separately. If the moderator is categorical, continuous.resolution=10 implies using all individual

levels.

w the weight for observations.

n the time of resampling in calculating the indirect effects, default is n=20, see Yu

et al. (2014).

x.new of the same format as x, with a new set of covariates and mediators on which to

calculate the mediation effects.

w. new the weights for new.x.

pred.new a new set of predictor(s).

cova.new a new set of covariate for mediator(s).

xj the inference of mediation effects at different levels of moderator is made on the

xjth predictor.

margin the change in predictor when calculating the mediation effects, see Yu et al.

(2014).

xmod If there is a moderator, xmod gives the moderator's name in cova and/or x.

df1 if nonlinear is TURE, natural cubic spline will be used to fit the relationship

between the predictor and each mediator. The df is the degree of freedom in the

ns() function, the default is 1.

para It is for binary predictors. If it is true, we would like the x-m relationship be

fitted parametrically.

echo If echo is FALSE, there is no counting printed for the number of bootstrap iter-

ations.

#### **Details**

calculate the mediation effects from the xjth predictor to the response variable(s) at each level of the mediator.

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#### Value

Returns an mma.mod object, which is similar to the mma object. Instead of calculating mediation effects for each predictor, the function returns the mediation effects of the xjth predictor at each level of the moderator.

## Author(s)

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## References

Yu, Q., Fan, Y., and Wu, X. (2014) <doi:10.4172/2155-6180.1000189>. "General Multiple Mediation Analysis With an Application to Explore Racial Disparity in Breast Cancer Survival," Journal of Biometrics & Biostatistics,5(2): 189.

Yu, Q., Scribner, R.A., Leonardi, C., Zhang, L., Park, C., Chen, L., and Simonsen, N.R. (2017) <doi:10.1016/j.sste.2017.02.001>. "Exploring racial disparity in obesity: a mediation analysis considering geo-coded environmental factors," Spatial and Spatio-temporal Epidemiology, 21, 13-23.

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## See Also

"med" just estimate the mediation effects.

# **Examples**

```
#binary x and categorical moderator
#linear model
data("weight_behavior")
pred=weight_behavior[,3]
x=weight_behavior[,c(2,4:14)]
inter=form.interaction(x,pred,inter.cov=c("race"),predref="M")
x=cbind(x,inter)
y=weight_behavior[,15]
head(x)
data.bin<-data.org(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),</pre>
                   binref=c(1,1), catmed=5, catref=1, predref="M", alpha=0.4, alpha2=0.4)
temp1<-boot.med(data=data.bin,n=2,n2=4,all.model=TRUE)</pre>
temp1.mode=boot.mod(temp1, vari="race", continuous.resolution=10)
plot2.mma(temp1.mode,vari="exercises",moderator="race")
plot2.mma(temp1.mode,vari="sports",moderator="race")
#nonlinear model
```

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```
x=weight_behavior[,c(2,4:14)]
pred=weight_behavior[,3]
y=weight_behavior[,15]
data.bin < -data.org(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),binref=c(1,1),
                    catmed=5,catref=1,predref="M",alpha=0.4,alpha2=0.4)
temp2<-boot.med(data=data.bin,n=2,n2=4,nu=0.05,nonlinear=TRUE,all.model=TRUE)
temp2.mode=boot.mod(temp2, vari="race", continuous.resolution=10)
plot2.mma(temp2.mode,vari="exercises",moderator="race")
plot2.mma(temp2.mode, vari="sports", moderator="race")
#binary x and continuou moderator
x=weight_behavior[,c(2,4:14)]
pred=weight_behavior[,3]
y=weight_behavior[,1]
inter=form.interaction(x,pred,inter.cov=c("age"),predref="M")
x=cbind(x,inter)
head(x)
data.cont < -data.org(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),
                   binref=c(1,1),catmed=5,catref=1,predref="M",alpha=0.4,alpha2=0.4)
temp3<-boot.med(data=data.cont,n=2,n2=4,all.model=TRUE)</pre>
temp3.mode=boot.mod(temp3,vari="age",continuous.resolution=2)
summary(temp3.mode)
plot2.mma(temp3.mode,vari="exercises",moderator="age")
plot2.mma(temp3.mode,vari="sports",moderator="age")
x=weight_behavior[,c(2,4:14)]
pred=weight_behavior[,3]
y=weight_behavior[,1]
data.cont<-data.org(x,y,pred=pred,mediator=5:12,jointm=list(n=1,j1=7:9),</pre>
                     predref="M",alpha=0.4,alpha2=0.4)
temp4<-boot.med(data=data.cont,n=2,n2=4,nu=0.05, nonlinear=TRUE,all.model=TRUE)
temp4.mode=boot.mod(temp4, vari="age", continuous.resolution=2, n=20)
summary(temp4.mode)
summary(temp4.mode,bymed=TRUE)
```

cgd1

cgd1 Data Set

#### Description

This database was obtained from the survival package containing a time-to-event data.

# Usage

```
data(weight_behavior)
```

## **Format**

The data set contains many variables.

# **Examples**

```
data(cgd1)
names(cgd1)
```

data.org

Data Organization and Identify Potential Mediators

# **Description**

Do a preliminary data analysis to identify potential mediators and covariates. Each variable listed in jointm is forced in the final estimation model as a mediator. Also organize the data into a format that can be directly used for the mediation analysis functions.

# Usage

# **Arguments**

| Х        | a data frame contains the predictor, all potential mediators and covariates. Note that x has to be a data frame. All varaibles in x that are not identified as potential mediators (by mediator, contmed, binmed, catmed, or jointm) are forced in mediation analysis as covariates.   |
|----------|--|
| У        | the vector of outcome variable. The outcome can be binary, continuous, or of "Surv" class (see survival package for help).   |
| pred     | the column or matrix of predictor(s): the predictor is the exposure variable, it can be a binary or multi-categorical factor or one/a few contiuous variable(s).   |
| mediator | the list of mediators (column numbers in x or by variable names). The mediators to be checked can be identified by "contmed", "binmed" and "catmed", or by this argument, "mediator", where binary and categorical mediators in x are identified as factors or characters, the reference group is the first level of the factor or factorized character. if a mediator has only two unique values, the mediator is identified as binary. If the reference groups need to be changed, the binary or categorical mediators can be listed in binmed or catmed, and the corresponding reference group in binref or catref. |
| contmed  | a vector of variable names or column numbers that locate the potential continuous mediators in $\boldsymbol{x}$ .  |
| binmed   | a vector of column numbers that locate the potential binary mediators in x.  |
| binref   | the defined reference groups of the binary potential mediators in binmed. The first levels of the mediators if is null.  |

catmed a vector of variable names or column numbers that locate the potential categor-

ical mediators in x. The first levels of the mediators if is null.

catref the defined reference groups of the categorical potential mediators in catmed.

jointm a list that identifies the mediators that need to be considered jointly, where the

first item indicates the number of groups of mediators to be considered jointly, and each of the following items identifies the variable names or column numbers

of the mediators in x for each group of joint mediators.

refy if y is binary, the reference group of y. The default is the first level of as.factor(y).

y for the reference group is assigned as 0.

family1 define the conditional distribution of y given x, and the linkage function that

links the mean of y with the system component in generalized linear model. The default value of family1 is binomial(link = "logit") for binary y, and gaus-

sian(link="identity") for continuous y.

predref if the predictor is categorical, identify the reference group of the predictor. The

default is the first level of the predictor. The value of the predictor is 0 for the

reference group.

alpha the significance level at which to test if the potential mediators (identified by

contmed, binmed, and catmed) can be used as a covariate or mediator in estimating y when all variables in x are included in the model. The default value is

alpha=0.1

alpha2 the significant level at which to test if a potential mediator is related with the

predictor. The default value is alpha2=0.1.

testtype if the testtype is 1 (by default), covariates/mediators are identified using full

model; if the testtype is 2, covariates/mediators are tested one by one in models

with the predictor only.

w the weight for data analysis, by default is rep(1, length(y)).

cova The data frame of covariates to be used to predict the mediator(s). The covariates

in cova cannot be potential mediators. If the covariates are for some specific potential mediators, cova\$for.m is the vector of names of potential mediators that use the covariates. Works for continuous predictors (pred) only, also the specified covariates should have no missing if only some mediators uses the

covariates.

#### Value

data.org returns a list with the organized data and identifiers of the potential mediators in the organized data set. The list has two components, bin.results if there is binary or categorical exposure(s), cont.results if there is continuous exposures.

x the organized data frame that include all potential mediators and covariates that

should be used to estimate the outcome.

dirx the vector/matrix of predictor(s)/exposure variable(s).

contm the column numbers of x that locate the potential continuous mediators.

binm when the predictor is continuous, binm gives the column numbers of x that lo-

cate the potential binary mediators.

| catm      | when the predictor is binary, catm gives the column numbers of x that locate the potential binary or categorical mediators; when the predictor is continuous, catm gives a list where the first item is the number of potential categorical mediators, and the following items give the column numbers of each binarized categorical mediator in x. |
|-----------|---|
| jointm    | a list where the first item is the number of groups of joint mediators, and each of the following items identifies the column numbers of the mediators in the newly organized x for each group of joint mediators.  |
| У         | the vector/matrix of outcome(s).  |
| y_type    | the variable type of outcome(s): 1 is continuous, 2 is binary, 3 is reserved for multi-categorical (no 3 would show in y_type, since all categorical responses are binarized), and 4 is survival.   |
| fullmodel | a list with each item the full linear model fitted with all potential mediators and covariates for each response.   |
| rela      | p-values of tests on the realtionship between the predictor(s) and each potiential mediator.  |
| P1        | If testtype=2, P1 gives the p-value of the corresponding variables in predicting the outcome(s) when only the variable and predictor are covariates in the model.   |
| binpred   | The columne numbers of binary predictors in pred.   |
| contpred  | The columne numbers of continuous predictors in pred.   |
| catpred   | A list with each item includes the columne numbers of a categorical predictor in pred.  |

# Note

All other variables in x but not identified by mediator, contmed, binmed, or catmed are forced in the final model as covariates. Compared with data.org, joint mediators are considered in this function. Every variable in the jointm should be listed in contmed, binmed, or catmed, and these variables are forced to be included as mediators for further mediation analysis. A variables can be included in more than one groups of joint mediators in jointm.

# Author(s)

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## References

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#### See Also

"data.org" that does not consider joint mediators, which can be added freely in the mediation analysis functions later.

## **Examples**

```
data("weight_behavior")
#binary predictor
#binary y
x=weight_behavior[,c(2,4:14)]
pred=weight_behavior[,3]
y=weight_behavior[,15]
data.b.b.2.1<-data.org(x,y,mediator=5:12,jointm=list(n=1,j1=c(5,7,9)),
                        pred=pred,predref="M", alpha=0.4,alpha2=0.4)
summary(data.b.b.2.1)
#Or you can specify the potential mediators and change the reference
#group for binary or categorical mediators. In the following code,
#potential continuous mediators are columns 8,9,10,12, and 13 of x,
#binary mediators are columns 7 and 11, and categorical mediator is
#column 6 of x with 1 to be the reference group for all categorical
#and binary mediators.
 data.b.b.2<-data.org(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),
  binref=c(1,1), catmed=5, catref=1, jointm=list(n=1, j1=c(5,7,9)),
  predref="M",alpha=0.4,alpha2=0.4)
 summary(data.b.b.2)
#use the mediator argument instead of contmet, binmed and catmed
#multivariate predictor
x=weight_behavior[,c(2:3,5:14)]
pred=weight_behavior[,4]
y=weight_behavior[,15]
data.b.b.2.3<-data.org(x,y,mediator=5:12,jointm=list(n=1,j1=c(5,7,9)),
                        pred=pred,predref="OTHER", alpha=0.4,alpha2=0.4)
summary(data.b.b.2.3)
#multivariate responses
x=weight_behavior[,c(2:3,5:14)]
pred=weight_behavior[,4]
y=weight_behavior[,c(1,15)]
data.b.b.2.4<-data.org(x,y,mediator=5:12,jointm=list(n=1,j1=c(5,7,9)),
```

```
pred=pred,predref="OTHER", alpha=0.4,alpha2=0.4)
summary(data.b.b.2.4)
#continuous y
x=weight_behavior[,c(2,4:14)]
pred=weight_behavior[,3]
y=weight_behavior[,1]
data.b.c.2<-data.org(x,y,pred=pred,mediator=5:12,jointm=list(n=1,j1=7:9),</pre>
  predref="M",alpha=0.4,alpha2=0.4)
summary(data.b.c.2)
#continuous predictor
#binary y
x=weight_behavior[,3:14]
pred=weight_behavior[,2]
y=weight_behavior[,15]
data.c.b.2<-data.org(x,y,pred=pred,mediator=5:12,catref=1,</pre>
          jointm=list(n=2,j1=7:9,j2=c(5,7)),alpha=0.4,alpha2=0.4)
summary(data.c.b.2)
#multivariate predictors
x=weight_behavior[,c(3:12,14)]
pred=weight_behavior[,c(2,13)]
y=weight_behavior[,15]
data.c.b.2.2<-data.org(x,y,pred=pred,mediator=5:11,catref=1,</pre>
          jointm=list(n=2, j1=7:9, j2=c(5,7)), alpha=0.4, alpha2=0.4)
summary(data.c.b.2.2)
#continuous y
x=weight_behavior[,3:14]
pred=weight_behavior[,2]
y=weight_behavior[,1]
data.c.c.2 < -data.org(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),
  binref=c(1,1), catmed=5, catref=1, jointm=list(n=2, j1=7:9, j2=c(5,7)),
  alpha=0.4,alpha2=0.4)
summary(data.c.c.2)
#multivariate responses
x=weight_behavior[,c(2:3,5:14)]
pred=weight_behavior[,4]
y=weight_behavior[,c(1,15)]
data.b.c.2.4 < -data.org(x,y,mediator=5:12,jointm=list(n=1,j1=c(5,7,9)),
                        pred=pred,predref="OTHER", alpha=0.4,alpha2=0.4)
summary(data.b.c.2.4)
#
x=weight_behavior[,c(3:12,14)]
pred=weight_behavior[,c(2,13)]
y=weight_behavior[,c(1,15)]
data.c.c.2.2<-data.org(x,y,pred=pred,mediator=5:11,catref=1,</pre>
          jointm=list(n=2,j1=7:9,j2=c(5,7)),alpha=0.4,alpha2=0.4)
summary(data.c.c.2.2)
```

16 form.interaction

```
#Surv class outcome (survival analysis)
                #a dataset in the survival package
data(cgd1)
x = cgd1[,c(4:5,7:12)]
pred=cgd1[,6]
status<-ifelse(is.na(cgd1$etime1),0,1)
y=Surv(cgd1$futime, status)
 #for continuous predictor
 #all other variables are considered as potential mediator
 data.surv.contx<-data.org(x,y,pred=pred,mediator=(1:ncol(x)),</pre>
                     alpha=0.5,alpha2=0.5)
 summary(data.surv.contx)
 #for binary predictor
x = cgd1[, c(5:12)]
pred=cgd1[,4]
 data.surv.binx<-data.org(x,y,pred=pred,mediator=(1:ncol(x)),</pre>
                     alpha=0.4,alpha2=0.4)
 summary(data.surv.binx)
```

form.interaction

*Create interaction terms of predictor(s) and potential moderator(s).* 

## **Description**

Create interaction terms of predictor(S) and potential moderator(s). Mainly for linear models.

# Usage

```
form.interaction(x,pred,inter.cov,predref=NULL,kx=NULL)
```

#### **Arguments**

x As the x in data.org.

The predictor(s) that might have interaction effect with the potential moderators.

The vector of names of potential moderators, which are included in x as covari-

ates.

predref The reference group of the predictor if the predictor is categorical.

kx The interaction is with the kx-th predictor(s). kx can be a vector. If kx is null,

the interaction is with each predictor.

#### Details

form.interaction is used to create interaction terms between predictor(s) and potential moderator(s). The created interaction terms should be used as covariates in linear mediation analysis. The function is not needed for nonlinear mediation method.

## Value

Return a matrix with the interaction terms pred\*inter.cov.

#### Author(s)

```
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```

#### See Also

```
"test.moderation", "moderate"
```

## **Examples**

```
data("weight_behavior")
pred=data.frame(weight_behavior[,3])
names(pred)="pred"
x=weight_behavior[,c(2,4:14)]
inter=form.interaction(x,pred,inter.cov=c("sports","sweat"),predref=NULL)
x=cbind(x,inter)
head(x)
```

med

Mediation Analysis with Binary or Continuous Predictor

# Description

To estimate the mediation effects when the predictor is binary or continuous.

# Usage

# **Arguments**

data

the list of result from data.org that organize the covariates, mediators, predictor and outcome. If data is NULL, then need to provide all arguments requested for data.org.

x, y, pred, mediator, contmed, binmed, binref, catmed, catref, jointm, family1, predref, alpha, alpha2, testtype, w, cova

arguments to set up the data. Need to set up only when data is NULL.

margin

the change in predictor when calculating the mediation effects, see Yu et al. (2014).

n the time of resampling in calculating the indirect effects, default is n=20, see Yu et al. (2014).

nonlinear if TURE, Multiple Additive Regression Trees (MART) will be used to fit the

final full model in estimating the outcome. The default value of nonlinear is FALSE, in which case, a generalized linear model will be used to fit the final

full model.

df1 if nonlinear is TURE, natural cubic spline will be used to fit the relationship

between the predictor and each mediator. The df is the degree of freedom in the

ns() function, the default is 1.

nu set the parameter "shrinkage" in gbm function if MART is to be used, by default,

nu=0.001. See also help(gbm.fit).

D set the parameter "interaction.depth" in gbm function if MART is to be used, by

default, D=3. See also help(gbm.fit).

distn the assumed distribution(s) of the outcome(s) if MART is used for final full

model. The default value of distn is "gaussian" for continuous y, "bernoulli" for

binary y and coxph for "Surv" class outcome.

refy if y is binary, the reference group of y.

x.new A new set of covariates, of the same format as x (after data.org), on which to

calculate the mediation effects. For continuous predictor only. If is NULL, the

mediation effects will be calculated based on the original data set.

pred.new A new set of predictor(s), of the same format as pred (after data.org), on which

to calculate the mediation effects. For continuous predictor only.

cova.new A new set of covariate(s) to estimate mediators, of the same format as cova.

type the type of prediction when y is class Surv. By default, type is "risk".

w. new the weight for each case in x.new.

xmod If there is a moderator, xmod gives the moderator's name in cova and/or x.

custom.function

a string of customer defined final model for predicting the outcome(s). The response variable should be noted as "responseY", the dataset should be noted as "dataset123". The weights for observations should be noted as "weights123". The covariates should be in x or pred. Use "~." for all variables in x and pred is allowed. The customer defined model should be able to make prediction by using "predict(object, newdata=...)", where the object is the results of the fitted model. If a specific package needs to be called to fit the model, the user should call the package first. For example, if the gamlss package is used to fit a piecewise polynomial with "age" to be the mediator and "race" be the predictor, we

can set custom.function="gamlss(responseV~b(age,df=3)+race,data=dataset123,tace=FALSE)".

The length of custom.function should be the same as the dimension of y. If cus-

tom.function[j] is NA, the usual method will be used to fit y[j].

para It is for binary predictors. If it is true, we would like the x-m relationship be

fitted parametrically.

#### **Details**

The mediators are not tested in this function. data.org should be used first for the tests and data organizing, and then the resulting list from data.org can be used directly to define the arguments in this function. med considers all variables in x as mediators or covariates in the final model and all variables identified by contm, binm, catm, or jointm as mediators.

#### Value

The result is an med object with the item a.binx for results of binary or categorical exposure(s) and the item a.contx for continuous exposure(s):

denm a matrix where each column gives the estimated direct effect not from the cor-

responding mediator (identified by the column name), see Yu et al. (2014) for the definition, and each row corresponding to the results from one resampling for binary predictor or the results on a row of x.new for continuous predictor.

ie a matrix where each column gives the estimated indirect effect from the corre-

sponding mediator (identified by the column name) and each row corresponding to the results from one resampling for binary predictor or the results on a row of

x.new for continuous predictor.

te a vector of the estimated total effect on x.new.

model a list, where the first item, MART, is TRUE if a mart is fitted as the final model;

the second item, Survial is T if a survival model is fit; the third item, type, is the type of prediction when a survival model is fitted; the fourth item, full.model, is the fitted final full model where y is the outcome and all predictor, covariates, and mediators are the explanatory variables; and the fifth item, best.iter, is the number of best iterations if MART is used to fit the final model, is NULL if the

final model is a generalized linear model.

#### Author(s)

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#### See Also

"boot.med" to make inferences on the estimated mediation effects using bootstrap method.

## **Examples**

```
data("weight_behavior")
##binary x
#binary y
x=weight_behavior[,c(2,4:14)]
pred=weight_behavior[,3]
y=weight_behavior[,15]
data.bin<-data.org(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),
 binref=c(1,1), catmed=5, catref=1, predref="M", alpha=0.4, alpha2=0.4)
temp1<-med(data=data.bin,n=2)
#med(data=NULL,x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),
     binref=c(1,1),catmed=5,catref=1,predref="M",alpha=0.4,alpha2=0.4,n=2)
#or use self-defined final function
temp1<-med(data=data.bin,n=2,custom.function =</pre>
           'glm(responseY~.,data=dataset123,family="quasibinomial",
           weights=weights123)')
temp2<-med(data=data.bin,n=2,nonlinear=TRUE)
#multiple predictors (categorical and continuous predictors)
x=weight_behavior[,c(3,5:14)]
pred=weight_behavior[,c(2,4)]
y=weight_behavior[,15]
data.b.b.2.3<-data.org(x,y,mediator=4:11,jointm=list(n=1,j1=c(5,7,9)),
                        pred=pred,predref="OTHER", alpha=0.4,alpha2=0.4)
temp1.2<-med(data.b.b.2.3,n=2)
temp2.2<-med(data.b.b.2.3,n=2,nonlinear=TRUE)</pre>
#multivariate responses
x=weight_behavior[,c(2:3,5:14)]
pred=weight_behavior[,4]
y=weight_behavior[,c(1,15)]
data.b.b.2.4 < -data.org(x,y,mediator=5:12,jointm=list(n=1,j1=c(5,7,9)),
                        pred=pred,predref="OTHER", alpha=0.4,alpha2=0.4)
temp1.3<-med(data.b.b.2.4, n=2)
#or use the self defined function
temp1.3<-med(data.b.b.2.4, n=2, custom.function =c('glm(responseY~.,</pre>
              data=dataset123, family="gaussian", weights=weights123)'
              'glm(responseY~.,data=dataset123,family="quasibinomial",
              weights=weights123)'))
```

```
temp2.3<-med(data.b.b.2.4,n=2,nonlinear=TRUE)</pre>
#continuous y
x=weight_behavior[,c(2,4:14)]
 pred=weight_behavior[,3]
 y=weight_behavior[,1]
 data.cont<-data.org(x,y,pred=pred,mediator=5:12,jointm=list(n=1,j1=7:9),</pre>
                      predref="M",alpha=0.4,alpha2=0.4)
 temp3<-med(data=data.cont,n=2)</pre>
 temp4<-med(data=data.cont,n=2,nonlinear=TRUE)</pre>
##continuous x
#binary y
x=weight_behavior[,3:14]
 pred=weight_behavior[,2]
 y=weight_behavior[,15]
 data.contx<-data.org(x,y,pred=pred,mediator=4:10,alpha=0.4,alpha2=0.4)
 temp5<-med(data=data.contx,n=2)</pre>
#or use the self defined function
 temp5<-med(data=data.contx,n=2,custom.function ='glm(responseY~.,</pre>
            data=dataset123, family="quasibinomial", weights=weights123)')
 temp6<-med(data=data.contx,n=2,nonlinear=TRUE,nu=0.05)</pre>
#continuous y
x=weight_behavior[,3:14]
y=weight_behavior[,1]
pred=weight_behavior[,2]
data.contx<-data.org(x,y,pred=pred,contmed=c(11:12),binmed=c(6,10),</pre>
                      binref=c(1,1), catmed=5, catref=1,
                      alpha=0.4,alpha2=0.4)
temp7<-med(data=data.contx,n=2)</pre>
temp8<-med(data=data.contx,n=2,nonlinear=TRUE,nu=0.05)</pre>
##Surv class outcome (survival analysis)
data(cgd1)
                  #a dataset in the survival package
x=cgd1[,c(4:5,7:12)]
pred=cgd1[,6]
status<-ifelse(is.na(cgd1$etime1),0,1)</pre>
y=Surv(cgd1$futime,status)
#for continuous predictor
data.surv.contx<-data.org(x,y,pred=pred,mediator=1:ncol(x),</pre>
                           alpha=0.5,alpha2=0.5)
temp9.contx<-med(data=data.surv.contx,n=2,type="lp")</pre>
#close to mart results when use type="lp"
temp10.contx<-med(data=data.surv.contx,n=2,nonlinear=TRUE)</pre>
#results in the linear part unit
temp10.contx
#for binary predictor
x = cgd1[, c(5:12)]
```

mma

Multiple Mediation Analysis

# **Description**

Test for mediators and do statistical inferences on the identified mediation effects.

## Usage

## **Arguments**

| x        | a data frame contains the predictor, all potential mediators and covariates. All variables in x that are not identified as potential mediators (by mediator, contmed, binmed, catmed, or jointm) are forced in mediation analysis as covariates.   |
|----------|--|
| у        | the vector of outcome variable.  |
| pred     | the vector/matrix of the predictor(s).   |
| mediator | the list of mediators (column numbers in x or by variable names). The mediators to be checked can be identified by "contmed", "binmed" and "catmed", or by this argument, "mediator", where binary and categorical mediators in x are identified by factors, the reference group is the first level of the factor. |
| contmed  | a vector of column numbers that locate the potential continuous mediators in x.  |
| binmed   | a vector of column numbers that locate the potential binary mediators in x.  |
| binref   | the defined reference groups of the binary potential mediators in binmed.  |
| catmed   | a vector of column numbers that locate the potential categorical mediators in x.   |
| catref   | the defined reference groups of the categorical potential mediators in catmed.   |
| jointm   | a list that identifies the mediators that need to be considered jointly, where the first item indicates the number of groups of mediators to be considered jointly, and each of the following items identifies the column numbers of the mediators in x for each group of joint mediators.                         |

| cova      | The data frame of covariates to be used to predict the mediator(s). The covariates in cova cannot be potential mediators. If the covariates are for some specific potential mediators, cova\$for.m is the vector of names of potential mediators that use the covariates. Works for continuous predictors (pred) only, also the specified covariates should have no missing if only some mediators uses the covariates. Works mainly for continuous predictors (pred), also the specified covariates should have no missing if only some mediators uses the covariates. |
|-----------|---|
| refy      | if y is binary, the reference group of y.By default, the reference group will be the first level of $as.factor(y)$ .  |
| predref   | if predictor is binary, identify the reference group of the binary predictor. By default, the reference group will be the first level of the predictor.   |
| alpha     | the significance level at which to test if the potential mediators (identified by contmed, binmed, and catmed) can be used as a covariate or mediator in estimating y when all variables in x are included in the model. The default value is alpha=0.1   |
| alpha2    | the significant level at which to test if a potential mediator is related with the predictor. The default value is alpha2=0.1.  |
| margin    | if binpred is FALSE, define the change in predictor when calculating the mediation effects, see Yu et al. (2014).   |
| n         | the time of resampling in calculating the indirect effects, default is $n=20$ , see Yu et al. (2014).   |
| nonlinear | if TURE, Multiple Additive Regression Trees (MART) will be used to fit the final full model in estimating the outcome. The default value of nonlinear is FALSE, in which case, a generalized linear model will be used to fit the final full model.   |
| df1       | if nonlinear is TURE, natural cubic spline will be used to fit the relationship between the countinuous predictor and each mediator. The df is the degree of freedom in the ns() function, the default is 1.  |
| nu        | set the parameter "interaction.depth" in gbm function if MART is to be used, by default, nu=0.001. See also help(gbm.fit).  |
| D         | set the parameter "shrinkage" in gbm function if MART is to be used, by default, D=3. See also help(gbm.fit).   |
| distn     | the assumed distribution of the outcome if MART is used for final full model. The default value of distn is "bernoulli" for binary y, and "gaussian" for continuous y.  |
| family1   | define the conditional distribution of y given x, and the linkage function that links the mean of y with the system component if generalized linear model is used as the final full model. The default value of family1 is binomial(link = "logit") for binary y, gaussian(link="identity") for continuous y.   |
| n2        | the number of times of bootstrap resampling. The default value is n2=50.  |
| W         | the weight for each observation.  |
| testtype  | if the testtype is 1 (by default), covariates/mediators are identified using full model; if the testtype is 2, covariates/mediators are tested one by one in models with the predictor only.  |

A new set of predictor and corresponding covariates, of the same format as x x.new

> (after data.org), on which to calculate the mediation effects. For continuous predictor only. If is NULL, the mediation effects will be calculated based on the

original data set.

pred.new A new set of predictor(s), of the same format as x (after data.org), on which to

calculate the mediation effects. For continuous predictor only.

a new set of covaraite(s). cova.new

type the type of prediction when y is class Surv. Is "risk" if not specified.

w.new the weights for new.x.

all.model save all the fitted model from bootstrap samples if TRUE. This needs to be true

to make inferences on moderation effects.

xmod If there is a moderator, xmod gives the moderator's name in cova and/or x.

custom.function

a string of customer defined final model for predicting the outcome(s). The response variable should be noted as "responseY", the dataset should be noted as "dataset123". The weights for observations should be noted as "weights123". The covariates should be in x or pred. Use "~." for all varaibles in x and pred is allowed. The customer defined model should be able to make prediction by using "predict(object, newdata=...)", where the object is the results of the fitted model. If a specific package needs to be called to fit the model, the user should call the package first. For example, if the gamlss package is used to fit a piecewise polynomial with "age" to be the mediator and "race" be the predictor, we

can set custom.function="gamlss(responseV~b(age,df=3)+race,data=dataset123,tace=FALSE)". The length of custom.function should be the same as the dimension of y. If cus-

tom.function[i] is NA, the usual method will be used to fit y[i].

It is for binary predictors. If it is true, we would like the x-m relationship be

fitted parametrically.

If echo is FALSE, there is no counting printed for the number of bootstrap iterecho

ations.

## **Details**

para

mma first tests if the potential mediators defined by binm, contm, and catm should be treated as mediators or covariates (if none, the variable will be deleted from further analysis). All variables identified by jointm are treated as mediators. All other variables in x that are not tested are treated as covariates. Then mma does the mediation effects estimation and inference on the selected variables.

#### Value

Returns an mma object.

estimation list the estimation of ie (indirect effect), te (total effect), and de (direct effect

from the predictor) separately.

bootsresults a list where the first item, ie, is a matrix of n2 rows where each column gives

> the estimated indirect effect from the corresponding mediator (identified by the column name) from the n2 bootstrap samples; the second item, te, is a vector

|             | of estimated total effects from the bootstrap sample; and the 3rd item, de, is a vector of estimated direct effect of the predictor from the bootstrap sample.   |
|-------------|--|
| model       | a list where the first item, MART, is T if mart is fitted for the final model; the second item, Survival, is T if a survival model is fitted; the third item, type, is the type of prediction when a survival model is fitted; the fourth item, model, is the fitted final full model where y is the outcome and all predictor, covariates, and mediators are the explanatory variables; and the fourth item, best.iter is the number of best iterations if MART is used to fit the final model. |
| data        | a list that contains all the used data: x=x, y=y, dirx=dirx, binm=binm, contm=contm, catm=catm, jointm=jointm, binpred=FALSE.  |
| boot.detail | for continuous predictor only: a list that contains the mediation effects on each row of new.pred: new.pred=new.pred, te1, de1, ie1.   |
| all_model   | a list with all fitted models from bootstrap samples if all.model is TRUE.   |
| all_iter    | if all.model is TRUE, a matrix with all fitted best iterations if MART is used. Each row is from one bootstrap sample.   |
| all_boot    | if all.model is TRUE, a matrix with all bootstrap samples. Each row is one bootstrap sample.   |

## Author(s)

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## References

Baron, R.M., and Kenny, D.A. (1986) <doi:10.1037/0022-3514.51.6.1173>. "The moderator-mediator variable distinction in social psychological research: conceptual, strategic, and statistical considerations," J. Pers Soc Psychol, 51(6), 1173-1182.

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J.H. Friedman (2001) <doi:10.1214/aos/1013203451>. "Greedy Function Approximation: A Gradient Boosting Machine," Annals of Statistics 29(5):1189-1232.

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Yu, Q., Scribner, R.A., Leonardi, C., Zhang, L., Park, C., Chen, L., and Simonsen, N.R. (2017) <doi:10.1016/j.sste.2017.02.001>. "Exploring racial disparity in obesity: a mediation analysis considering geo-coded environmental factors," Spatial and Spatio-temporal Epidemiology, 21, 13-23.

Yu, Q., and Li, B. (2017) <doi:10.5334/hors.160>. "mma: An r package for multiple mediation analysis," Journal of Open Research Software, 5(1), 11.

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## See Also

"data.org" is for mediator tests, and "med", and "boot.med" for mediation analysis and inferences.

## **Examples**

```
data("weight_behavior")
#binary predictor
#binary y
x=weight_behavior[,c(2,4:14)]
 pred=weight_behavior[,3]
y=weight_behavior[,15]
 temp.b.b.glm<-mma(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),binref=c(1,1),
                    catmed=5,catref=1,predref="M",alpha=0.4,alpha2=0.4,n=2,n2=2)
 temp.b.b.mart<-mma(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),binref=c(1,1),
             catmed=5, catref=1, predref="M", alpha=0.4, alpha2=0.4, nonlinear=TRUE, n=2, n2=5)
 #continuous y
 x=weight_behavior[,c(2,4:14)]
 pred=weight_behavior[,3]
 y=data.frame(weight_behavior[,1])
 colnames(y)="bmi"
 temp.b.c.glm<-mma(x,y,pred=pred,mediator=5:12,jointm=list(n=1,j1=7:9),</pre>
                     predref="M",alpha=0.4,alpha2=0.4,n2=20)
 temp.b.c.mart<-mma(x,y,pred=pred,mediator=5:12,jointm=list(n=1,j1=7:9),</pre>
                     predref="M", alpha=0.4, alpha2=0.4,
                     n=2,nonlinear=TRUE,n2=20)
##Surv class outcome (survival analysis)
                 #a dataset in the survival package
data(cgd1)
x=cgd1[,c(4:5,7:12)]
pred=cgd1[,6]
status<-ifelse(is.na(cgd1$etime1),0,1)
y=Surv(cgd1$futime, status)
#for continuous predictor
temp.cox.contx<-mma(x,y,pred=pred,mediator=1:ncol(x),</pre>
                    alpha=0.5,alpha2=0.5,n=1,n2=2,type="lp")
summary(temp.cox.contx)
temp.surv.mart.contx<-mma(x,y,pred=pred,mediator=1:ncol(x),
                    alpha=0.5,alpha2=0.5,nonlinear=TRUE,n2=2)
#for binary predictor
x = cgd1[, c(5:12)]
pred=cgd1[,4]
temp.cox.binx<-mma(x,y,pred=pred,mediator=1:ncol(x),</pre>
                    alpha=0.4,alpha2=0.4,n=1,n2=2,type="lp")
summary(temp.cox.binx)
```

moderate 27

| moderate | Calculate and plot the direct effect of the selected exposure variable at each level of the moderator. |
|----------|--|
|          | ar each sever of the meachain.   |

#### **Description**

Calculate and plot the direct effect of the selected exposure variable at each level of the moderator.

# Usage

```
moderate(med1,vari,j=1,kx=1,continuous.resolution=100,plot=TRUE)
```

# **Arguments**

med1 The med object from the med function.

vari The name of the moderator.

j The jth response if the response is multiple.

kx The moderate effect is with the kx-th predictor(s).

continuous.resolution

The number of equally space points at which to evaluate continuous predictors.

Plot the direct effect at each level of the moderator if ture.

#### Details

Calculate and plot the direct effect of the selected exposure variable at each level of the moderator base on the result from the med function.

## Value

The moderate returns a list where the item result is a data frame with two or three elements

moderator the moderator levels.

x the level of the exposure variable – available only for continuous exposure and

moderate with nonlinear method.

de the direct effect at the corresonding moderator (and exposure) level(s).

## Author(s)

```
Qingzhao Yu <qyu@lsuhsc.edu>
```

#### See Also

```
"form.interaction", "test.moderation"
```

28 plot.med

## **Examples**

plot.med

Plot the mediation effect on the fitted med object

## **Description**

Plot the marginal effect of the selected variable on the outcome, and the marginal effect of the predictor on the selected variable.

## Usage

```
## S3 method for class 'med'
plot(x,...,vari,xlim=NULL)
```

# **Arguments**

x a med object created initially call to med, med.binx, or med.contx.
 vari an indices or the name of the variable to plot.
 xlim the range of the variable to be plotted.
 other arguments passed to the plot function.

# **Details**

plot.med plots the marginal effect of the selected variable on the outcome, and the marginal effect of the predictor on the selected variable. If the predictor is binary, draw a histogram or boxplot of the marginal density of the variable at each different value of the predictor.

#### Value

No return value, called for graphs

## Author(s)

Qingzhao Yu <qyu@lsuhsc.edu>

plot.mma 29

## References

Yu, Q., Fan, Y., and Wu, X. (2014) <doi:10.4172/2155-6180.1000189>. "General Multiple Mediation Analysis With an Application to Explore Racial Disparity in Breast Cancer Survival," Journal of Biometrics & Biostatistics,5(2): 189.

#### See Also

```
"med"
```

# **Examples**

```
data("weight_behavior")
  x=weight_behavior[,c(2,4:14)]
  pred=weight_behavior[,3]
  y=weight_behavior[,15]
  data.bin<-data.org(x,y,pred=pred,contmed=c(7:9,11:12),binmed=c(6,10),
    binref=c(1,1),catmed=5,catref=1,predref="M",alpha=0.4,alpha2=0.4)
  temp1<-med(data=data.bin,n=2)
  temp2<-med(data=data.bin,n=2,nonlinear=TRUE)
  plot(temp1,data.bin,vari="exercises",xlim=c(0,50))
  plot(temp2,data.bin,vari="sports")</pre>
```

plot.mma

Relative effects plot of the fitted mma object

## **Description**

Plot the marginal effect of the selected variable on the outcome, and the marginal effect of the predictor on the selected variable.

# Usage

```
## $3 method for class 'mma'
plot(x,...,vari,xlim=NULL,alpha=0.95,
quantile=FALSE)
```

## **Arguments**

| x        | a mma object created initially call to mma, boot.met.binx, or boot.met.contx.  |
|----------|--|
| vari     | an indices or the name of the variable to plot.  |
| xlim     | the range of the variable to be plotted.   |
| alpha    | for continuous predictor only, to draw the alpha confidence interval of the indirect effect.   |
| quantile | for continuous predictor only, if true to draw the alpha confidence interval of the indirect effect based on quantile, otherwise, based on the normal approximation. |
|          | other arguments passed to the plot function.   |

30 plot.mma

#### **Details**

plot.mma plots the marginal effect of the selected variable on the outcome, and the marginal effect of the predictor on the selected variable. If the predictor is binary, draw a histogram or boxplot of the marginal density of the variable at each different value of the predictor.

#### Value

No return value, called for drawing graphs

#### Author(s)

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#### References

Yu, Q., Fan, Y., and Wu, X. (2014) <doi:10.4172/2155-6180.1000189>. "General Multiple Mediation Analysis With an Application to Explore Racial Disparity in Breast Cancer Survival," Journal of Biometrics & Biostatistics,5(2): 189.

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Yu, Q., Wu, X., Li, B., and Scribner, R. (2018). <doi:10.1002/sim.7977>. "Multiple Mediation Analysis with Survival Outcomes – With an Application to Explore Racial Disparity in Breast Cancer Survival," Statistics in Medicine.

Yu, Q., Medeiros, KL, Wu, X., and Jensen, R. (2018). <doi:10.1007/s11336-018-9612-2>. "Explore Ethnic Disparities in Anxiety and Depression Among Cancer Survivors Using Nonlinear Mediation Analysis," Psychometrika, 83(4), 991-1006.

# See Also

```
"mma","boot.med"
```

## **Examples**

plot2.mma 31

| plot2.mma | Relative effects plot of the fitted mma object with moderator |
|-----------|---|
|           |   |

# Description

Plot the marginal effect of the selected variable on the outcome, and the marginal effect of the predictor on the selected variable, at each level of the moderator.

# Usage

```
plot2.mma(x,...,vari,xlim=NULL,alpha=0.95,quantile=FALSE,moderator,xj=1)
```

# **Arguments**

| x         | a mma object created initially call to boot.mod.   |
|-----------|--|
| vari      | an indices or the name of the variable to plot.  |
| xlim      | the range of the variable to be plotted.   |
| alpha     | for continuous predictor only, to draw the alpha confidence interval of the indirect effect.   |
| quantile  | for continuous predictor only, if true to draw the alpha confidence interval of the indirect effect based on quantile, otherwise, based on the normal approximation. |
|           | other arguments passed to the plot function.   |
| moderator | the name of the moderator.   |
| хj        | the moderation effect on the xjth predictor.   |
|           |  |

## **Details**

plot2.mma plots the marginal effect of the selected variable on the outcome, and the marginal effect of the predictor on the selected variable, at each level of the moderator. If the predictor is binary, draw a histogram or boxplot of the marginal density of the variable at each different value of the predictor.

# Value

No return value, called for graphs

# Author(s)

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32 print.med

#### References

Yu, Q., Fan, Y., and Wu, X. (2014) <doi:10.4172/2155-6180.1000189>. "General Multiple Mediation Analysis With an Application to Explore Racial Disparity in Breast Cancer Survival," Journal of Biometrics & Biostatistics,5(2): 189.

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#### See Also

```
"boot.mod"
```

## **Examples**

#see boot.mod menu.

print.med

Print an med object

#### **Description**

Print the estimation of mediation effects from an med object: from functions med.

## Usage

```
## S3 method for class 'med'
print(x,...,digit=4)
```

# **Arguments**

x a med object created initially call to med.... other arguments passed to the print function.digit the number of digits to keep at printing.

## Value

No return value, called for graphs

print.mma 33

## Author(s)

Qingzhao Yu <qyu@lsuhsc.edu>

#### References

Yu, Q., Fan, Y., and Wu, X. (2014) <doi:10.4172/2155-6180.1000189>. "General Multiple Mediation Analysis With an Application to Explore Racial Disparity in Breast Cancer Survival," Journal of Biometrics & Biostatistics,5(2): 189.

Yu, Q., Scribner, R.A., Leonardi, C., Zhang, L., Park, C., Chen, L., and Simonsen, N.R. (2017) <doi:10.1016/j.sste.2017.02.001>. "Exploring racial disparity in obesity: a mediation analysis considering geo-coded environmental factors," Spatial and Spatio-temporal Epidemiology, 21, 13-23.

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Yu, Q., Medeiros, KL, Wu, X., and Jensen, R. (2018). <doi:10.1007/s11336-018-9612-2>. "Explore Ethnic Disparities in Anxiety and Depression Among Cancer Survivors Using Nonlinear Mediation Analysis," Psychometrika, 83(4), 991-1006.

## See Also

"med"

## **Examples**

print.mma

Print a mma object

# Description

Print the estimation of mediation effects from an mma object.

34 print.mma

## Usage

```
## S3 method for class 'mma'
print(x,...,digit=3)
```

# Arguments

x a mma object created initially call to mma, boot.med.binx, or boot.med.contx.

... other arguments passed to the print function.

digit the number of decimal digits to keep.

#### Value

#### Return a LIST

te estimation of the indirect effects from the mma object.
te estimation of the total effect from the mma object.
de estimation of the direct effect from the mma object.

## Author(s)

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#### References

Yu, Q., Fan, Y., and Wu, X. (2014) <doi:10.4172/2155-6180.1000189>. "General Multiple Mediation Analysis With an Application to Explore Racial Disparity in Breast Cancer Survival," Journal of Biometrics & Biostatistics,5(2): 189.

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#### See Also

```
"mma", "boot.med"
```

summary.med\_iden 35

## **Examples**

summary.med\_iden

Summary method for class "med\_iden".

## **Description**

Compute the estimations, standard deviations and confidence intervals of the mediation effects.

## Usage

```
## S3 method for class 'med_iden'
summary(object,...,only=FALSE)
## S3 method for class 'summary.med_iden'
print(x,...)
```

# **Arguments**

object a med\_iden object created initially call to data.org.

x a summary.med\_iden object created initially call to summary.med\_iden

other arguments passed to the print function.

only if only=T, show test results for selected covariates and mediators only.

# **Details**

summary.med\_iden gives a list of identified mediators, covariates and the test results.

## Value

The function summary.med\_iden return a list of covariates and mediators as identified by a series tests.

mediator variable names of the identified mediators, either signicant in both full model

and in relate to the predictor, or being a member of the pre-identified joint me-

diators.

covariate variable names of covariates: being significant in the full model but not signifi-

cantly relate with the predictor.

tests statistical test results.
results the original object.

36 summary.mma

## Author(s)

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#### References

Yu, Q., Fan, Y., and Wu, X. (2014) <doi:10.4172/2155-6180.1000189>. "General Multiple Mediation Analysis With an Application to Explore Racial Disparity in Breast Cancer Survival," Journal of Biometrics & Biostatistics,5(2): 189.

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Yu, Q., Wu, X., Li, B., and Scribner, R. (2018). <doi:10.1002/sim.7977>. "Multiple Mediation Analysis with Survival Outcomes – With an Application to Explore Racial Disparity in Breast Cancer Survival," Statistics in Medicine.

Yu, Q., Medeiros, KL, Wu, X., and Jensen, R. (2018). <doi:10.1007/s11336-018-9612-2>. "Explore Ethnic Disparities in Anxiety and Depression Among Cancer Survivors Using Nonlinear Mediation Analysis," Psychometrika, 83(4), 991-1006.

#### See Also

```
"mma", "boot.med"
```

# **Examples**

summary.mma

Summary of an mma project

# **Description**

Compute the estimations, standard deviations and confidence intervals of the mediation effects.

# Usage

```
## S3 method for class 'mma'
summary(object,..., alpha=0.05, plot=TRUE, RE=FALSE,
quant=FALSE,ball.use=FALSE,bymed=FALSE,win1=0.2)
## S3 method for class 'summary.mma'
print(x,...,digit=3)
```

summary.mma 37

## Arguments

object a mma object created initially call to mma, boot.met.binx, or boot.met.contx.

x a summary.mma object created initially call to summary.mma.

... other arguments passed to the print function.

alpha the alpha level for confidence interval.

plot default is TRUE, if ture, draw a barplot of the mediation effects with confidence

intervals.

RE default is FALSE, if ture, show the inferences on relative mediation effects.

quant default is TRUE, if ture and ball.use is FALSE, draw the confidence intervals of

relative effects using quantile.

ball.use default is TRUE, if ture, draw the confidence intervals of relative effects using

the confidence ball. If both quant and ball.use are false, draw the confidence

intervals based on the standard deviaitons from bootstrap estimates.

digit the number of decimal digits to keep.

bymed if true, show results by each mediator.

win1 The rate of winsarize.

#### **Details**

summary.mma gives a list of the estimations and summary statistics based on the bootstrap results. If plot=T, draw a barplot of the relative effects of the direct effect of the predictor and indirect effects of the mediators. Relative effects is defined as the (in)direct effect divided by the total effect. The plot is arranged in order from the largest to the smallest relative effect.

## Value

The function summary.mma return a list. The first item, results, is the list for mediation effects, and the second item, re, is the list for relative effects. Under them, results have the items ie, te and de; re has the items ie and de. In each of the items, est is the estimation of the corresponding (relative) mediation effects based on the whole data, mean is the average estimated (relative) effects from the bootstrap samples, sd is the standard deviation of the estimates from the bootstrap sample. upbd and lwbd are the upper and lower bound of the confidence interval of the estimation using parametric method from the bootstrap sample, upbd\_q and lwbd\_q are the corresponding quantiles of the estimation from the bootstrap sample.

ie a matrix of statistics inference on the (relative) indirect effects from the mma

object. est is the estimate using the full sample. mean is the estimate that average over the bootstrap estimates. sd is the standard deviation of the bootstrap estimates. upbd and lwbd are the confidence bounds based on sd. upbd\_q and lwbd\_q are the confidence bounds based on quantiles of the bootstrap estimates. upbd b and lwbd b are condidence ball bounds based on bootstap estimates.

te statistics inference on the total effects from the mma object.

de statistics inference on the (relative) direct effects from the mma object.

If plot=T, draw a barplot of the relative mediation effects.

38 test.moderation

## Author(s)

Qingzhao Yu <qyu@lsuhsc.edu>

#### References

Yu, Q., Fan, Y., and Wu, X. (2014) <doi:10.4172/2155-6180.1000189>. "General Multiple Mediation Analysis With an Application to Explore Racial Disparity in Breast Cancer Survival," Journal of Biometrics & Biostatistics,5(2): 189.

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#### See Also

```
"mma", "boot.med"
```

## **Examples**

test.moderation

Test for moderation effects.

# Description

Used on a med object, test if there are moderation effects.

# Usage

```
test.moderation(med1,vari,j=1,kx=NULL)
```

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## Arguments

med1 a med object created initially call to med.

vari If med1 was from generalized linear method, vari is the vector of names of po-

tential moderation effects – the interaction terms formed by the potential moderator(s) times the predictor(s). Those interaction terms have to be included in

mediation analysis as covariates.

If med1 was formed by nonlinear method, vari is the vector of names of potential moderators, which have been included in mediation analysis as covariates.

j To check the moderation effect on the jth response.

kx The moderation effect is with the kx-th predictor(s). kx can be a vector. If kx is

null, the moderation effect is with each predictor.

#### **Details**

test.moderation is used to test whether direct moderation effect is significant based on the med object from med function (mediation analysis). If geralized linear models were used for mediation anlysis, the moderation effects (formed by the interactions of potential moderators and predictor) should be included in the mediation analysis as covariates. The function test.modeartion will give the significance levels of these interaction terms. If nonlinear models were used for mediation analysis, the potential moderators (but not the interaction terms) should have been included in mediation analysis as covarites. The function test.moderation will give not only significance levels of their interaction with the predictor(s) in generalized linear model, but also the H-statistics by Friedman and Popescue (2008).

#### Value

No return value, called for test results.

## Author(s)

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#### See Also

```
"med",, "moderate"
```

## **Examples**

weight\_behavior

Weight\_Behavior Data Set

## **Description**

This database was obtained from the Louisiana State University Health Sciences Center, New Orleans, by Dr. Richard Scribner. He explored the relationship between BMI and kids behavior through a survey at children, teachers and parents in Grenada in 2014. This data set includes 691 observations and 15 variables.

## Usage

```
data(weight_behavior)
```

#### **Format**

The data set contains the following variables:

```
bmi - body mass index, calculated by weight(kg)/height(cm)^2, numeric
```

age - children's age in years at the time of survey, numeric

sex - sex of the children, factor

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race - race of the children, factor
numpeople - number of people in family, numeric
car - the number of cars in family, numeric
gotosch - the method used to go to school, factor
snack - eat snack or not in a day, binary
tvhours - number of hours watching TV per week, numeric
cmpthours - number of hours using computer per week, numeric
cellhours - number of hours playing with cell phones per week, numeric
sports - join in a sport team or not, 1: yes; and 2: no
exercises - number of hours of exercises per week, numeric
sweat - number of hours of sweating activities per week, numeric
overweigh - the child is overweighed or not, binary

# **Examples**

data(weight\_behavior)
names(weight\_behavior)

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