# Package 'regmedint'

November 15, 2020

Title Regression-Based Causal Mediation Analysis with an Interaction

Term

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Description 'R' re-implementation of the regression-based causal mediation analysis with a treatment-mediator interaction term, as originally implemented in the 'SAS' macro by Valeri and Vander-Weele (2013) <doi:10.1037 a0031034=""> and Valeri and Vander-Weele (2015) <doi:10.1097 ede.000000000000053="">. Linear and logistic models are supported for the mediator model. Linear, logistic, loglinear, Poisson, negative binomial, Cox, and accelerated failure time (exponential and Weibull) models are supported for the outcome model.</doi:10.1097></doi:10.1037>
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Author Kazuki Yoshida [cre, aut] ( <a href="https://orcid.org/0000-0002-2030-3549">https://orcid.org/0000-0002-2030-3549</a> ), Yi Li [aut] ( <a href="https://orcid.org/0000-0002-9359-210X">https://orcid.org/0000-0002-9359-210X</a> ), Maya Mathur [ctb] ( <a href="https://orcid.org/0000-0001-6698-2607">https://orcid.org/0000-0001-6698-2607</a> )
Maintainer Kazuki Yoshida <kazukiyoshida@mail.harvard.edu></kazukiyoshida@mail.harvard.edu>
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# Description

This function extracts coef from mreg\_fit and pads with zeros appropriately to create a named vector consistently having the following elements: (Intercept) avar cvar: This part is eliminated when cvar = NULL.

# Usage

```
beta_hat(mreg, mreg_fit, avar, cvar)
```

# Arguments

mreg	A character vector of length 1. Mediator regression type: "linear" or "logistic".
mreg_fit	Model fit object for mreg (mediator model).
avar	A character vector of length 1. Treatment variable name.
cvar	A character vector of length > 0. Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.

### Value

A named numeric vector of coefficients.

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calc_myreg	Return mediation analysis functions given mediator and outcome models.

# Description

This function returns functions that can be used to calculate the causal effect measures, given the mediator model fit (mreg\_fit) and the outcome model fit (yreg\_fit).

### Usage

```
calc_myreg(mreg, mreg_fit, yreg, yreg_fit, avar, mvar, cvar, interaction)
```

# Arguments

mreg	A character vector of length 1. Mediator regression type: "linear" or "logistic".
mreg_fit	Model fit from fit_mreg
yreg	A character vector of length 1. Outcome regression type: "linear", "logistic", "loglinear", "poisson", "negbin", "survCox", "survAFT_exp", or "survAFT_weibull".
yreg_fit	Model fit from fit_yreg
avar	A character vector of length 1. Treatment variable name.
mvar	A character vector of length 1. Mediator variable name.
cvar	A character vector of length > 0. Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.
interaction	A logical vector of length 1. Default to TRUE. Whether to include a mediator-treatment interaction term in the outcome regression model.

### Value

A list containing two functions. The first is for calculating point estimates. The second is for calculating the correspoding

```
calc_myreg_mreg_linear_yreg_linear

Create calculators for effects and se (mreg linear / yreg linear)
```

### **Description**

Construct functions for the conditional effect estimates and their standard errors in the mreg linear / yreg linear setting. Internally, this function deconstruct model objects and feed parameter estiamtes to the internal worker functions calc\_myreg\_mreg\_linear\_yreg\_linear\_est and calc\_myreg\_mreg\_linear\_yreg\_linear\_se.

#### Usage

```
calc_myreg_mreg_linear_yreg_linear(
   mreg,
   mreg_fit,
   yreg,
   yreg_fit,
   avar,
   mvar,
   cvar,
   interaction
)
```

### Arguments

mreg	A character vector of length 1. Mediator regression type: "linear" or "logistic".
mreg_fit	Model fit from fit_mreg
yreg	A character vector of length 1. Outcome regression type: "linear", "logistic", "loglinear", "poisson", "negbin", "survCox", "survAFT_exp", or "survAFT_weibull".
yreg_fit	Model fit from fit_yreg
avar	A character vector of length 1. Treatment variable name.
mvar	A character vector of length 1. Mediator variable name.
cvar	A character vector of length > 0. Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.
interaction	A logical vector of length 1. Default to TRUE. Whether to include a mediator-treatment interaction term in the outcome regression model.

### Value

A list contraining a function for effect estimates and a function for corresponding standard errors.

```
calc_myreg_mreg_linear_yreg_logistic

Create calculators for effects and se (mreg linear / yreg logistic)
```

# Description

Construct functions for the conditional effect estimates and their standard errors in the mreg linear / yreg logistic setting. Internally, this function deconstruct model objects and feed parameter estiamtes to the internal worker functions calc\_myreg\_mreg\_linear\_yreg\_logistic\_est and calc\_myreg\_mreg\_linear\_yreg\_logistic\_se.

#### Usage

```
calc_myreg_mreg_linear_yreg_logistic(
   mreg,
   mreg_fit,
   yreg,
   yreg_fit,
   avar,
   mvar,
   cvar,
   interaction
)
```

### Arguments

mreg	A character vector of length 1. Mediator regression type: "linear" or "logistic".
mreg_fit	Model fit from fit_mreg
yreg	A character vector of length 1. Outcome regression type: "linear", "logistic", "loglinear", "poisson", "negbin", "survCox", "survAFT_exp", or "survAFT_weibull".
<pre>yreg_fit</pre>	Model fit from fit_yreg
avar	A character vector of length 1. Treatment variable name.
mvar	A character vector of length 1. Mediator variable name.
cvar	A character vector of length > 0. Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.
interaction	A logical vector of length 1. Default to TRUE. Whether to include a mediator-treatment interaction term in the outcome regression model.

# Value

A list contraining a function for effect estimates and a function for corresponding standard errors.

# Description

Construct functions for the conditional effect estimates and their standard errors in the mreg logistic / yreg linear setting. Internally, this function deconstruct model objects and feed parameter estiamtes to the internal worker functions calc\_myreg\_mreg\_logistic\_yreg\_linear\_est and calc\_myreg\_mreg\_logistic\_yreg\_linear\_se.

#### Usage

```
calc_myreg_mreg_logistic_yreg_linear(
    mreg,
    mreg_fit,
    yreg,
    yreg_fit,
    avar,
    mvar,
    cvar,
    interaction
)
```

### Arguments

mreg	A character vector of length 1. Mediator regression type: "linear" or "logistic".
mreg_fit	Model fit from fit_mreg
yreg	A character vector of length 1. Outcome regression type: "linear", "logistic", "loglinear", "poisson", "negbin", "survCox", "survAFT_exp", or "survAFT_weibull".
yreg_fit	Model fit from fit_yreg
avar	A character vector of length 1. Treatment variable name.
mvar	A character vector of length 1. Mediator variable name.
cvar	A character vector of length > 0. Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.
interaction	A logical vector of length 1. Default to TRUE. Whether to include a mediator-treatment interaction term in the outcome regression model.

### Value

A list contraining a function for effect estimates and a function for corresponding standard errors.

```
calc_myreg_mreg_logistic_yreg_logistic

Create calculators for effects and se (mreg logistic / yreg logistic)
```

# Description

Construct functions for the conditional effect estimates and their standard errors in the mreg logistic / yreg logistic setting. Internally, this function deconstruct model objects and feed parameter estiamtes to the internal worker functions calc\_myreg\_mreg\_logistic\_yreg\_logistic\_est and calc\_myreg\_mreg\_logistic\_yreg\_logistic\_se.

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### Usage

```
calc_myreg_mreg_logistic_yreg_logistic(
   mreg,
   mreg_fit,
   yreg,
   yreg_fit,
   avar,
   mvar,
   cvar,
   interaction
)
```

# Arguments

mreg	A character vector of length 1. Mediator regression type: "linear" or "logistic".
mreg_fit	Model fit from fit_mreg
yreg	A character vector of length 1. Outcome regression type: "linear", "logistic", "loglinear", "poisson", "negbin", "survCox", "survAFT_exp", or "survAFT_weibull".
yreg_fit	Model fit from fit_yreg
avar	A character vector of length 1. Treatment variable name.
mvar	A character vector of length 1. Mediator variable name.
cvar	A character vector of length > 0. Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.
interaction	A logical vector of length 1. Default to TRUE. Whether to include a mediator-treatment interaction term in the outcome regression model.

### Value

A list contraining a function for effect estimates and a function for corresponding standard errors.

# Description

Extract point estimates evaluated at a0, a1, m\_cde, and c\_cond.

# Usage

```
## S3 method for class 'regmedint'
coef(object, a0 = NULL, a1 = NULL, m_cde = NULL, c_cond = NULL, ...)
```

### **Arguments**

object	An object of the regmedint class.
a0	A numeric vector of length one.
a1	A numeric vector of length one.
m_cde	A numeric vector of length one. A mediator value at which the controlled direct effect (CDE) conditional on the adjustment covariates is evaluated. If not provided, the default value supplied to the call to regmedint will be used. Only the CDE is affected.
c_cond	A numeric vector as long as the number of adjustment covariates. A set of covariate values at which the conditional natural effects are evaluated.
	For compatibility with the generic. Ignored.

### Value

A numeric vector of point estimates.

# **Examples**

```
library(regmedint)
data(vv2015)
regmedint_obj <- regmedint(data = vv2015,</pre>
                           ## Variables
                           yvar = "y",
                           avar = "x",
                           mvar = "m",
                           cvar = c("c"),
                           eventvar = "event",
                           ## Values at which effects are evaluated
                           a0 = 0,
                           a1 = 1,
                           m\_cde = 1,
                           c\_cond = 0.5,
                           ## Model types
                           mreg = "logistic",
                           yreg = "survAFT_weibull",
                           ## Additional specification
                           interaction = TRUE,
                           casecontrol = FALSE)
coef(regmedint_obj)
## Evaluate at different values
coef(regmedint_obj, m_cde = 0, c_cond = 1)
```

```
coef.summary_regmedint
```

Extract the result matrix from a summary\_regmedint object.

### **Description**

Extract the result matrix from a summary\_regmedint object.

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#### Usage

```
## S3 method for class 'summary_regmedint'
coef(object, ...)
```

### **Arguments**

object An object with a class of summary\_regmedint.
... For compatibility with the generic.

#### Value

A matrix populated with results.

### **Examples**

```
library(regmedint)
data(vv2015)
regmedint_obj <- regmedint(data = vv2015,</pre>
                            ## Variables
                            yvar = "y",
                            avar = "x",
                            mvar = "m",
                            cvar = c("c"),
                            eventvar = "event",
                            ## Values at which effects are evaluated
                            a0 = 0,
                            a1 = 1,
                            m\_cde = 1,
                            c\_cond = 0.5,
                            ## Model types
                            mreg = "logistic",
                            yreg = "survAFT_weibull",
                            ## Additional specification
                            interaction = TRUE,
                            casecontrol = FALSE)
coef(summary(regmedint_obj))
```

confint.regmedint

Confidence intervals for mediation prameter estimates.

### **Description**

Construct Wald approximate confidence intervals for the quantities of interest.

#### Usage

```
## S3 method for class 'regmedint'
confint(
  object,
  parm = NULL,
  level = 0.95,
  a0 = NULL,
```

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```
a1 = NULL,

m_cde = NULL,

c_cond = NULL,

...
```

### **Arguments**

An object of the regmedint class. object For compatibility with generic. Ignored. parm level A numeric vector of length one. Requested confidence level. Defaults to 0.95. a0 A numeric vector of length one. A numeric vector of length one. a1 m\_cde A numeric vector of length one. A mediator value at which the controlled direct effect (CDE) conditional on the adjustment covariates is evaluated. If not provided, the default value supplied to the call to regmedint will be used. Only the CDE is affected. A numeric vector as long as the number of adjustment covariates. A set of c\_cond covariate values at which the conditional natural effects are evaluated. For compatibility with generic.

#### Value

A numeric matrix of the lower limit and upper limit.

# **Examples**

```
library(regmedint)
data(vv2015)
regmedint_obj <- regmedint(data = vv2015,</pre>
                            ## Variables
                           yvar = "y",
                            avar = "x"
                           mvar = "m"
                           cvar = c("c"),
                           eventvar = "event",
                           ## Values at which effects are evaluated
                           a0 = 0,
                           a1 = 1,
                           m_{cde} = 1,
                            c\_cond = 0.5,
                           ## Model types
                           mreg = "logistic",
                           yreg = "survAFT_weibull",
                            ## Additional specification
                           interaction = TRUE,
                           casecontrol = FALSE)
confint(regmedint_obj)
## Evaluate at different values
confint(regmedint_obj, m_cde = 0, c_cond = 1)
## Change confidence level
confint(regmedint_obj, m_cde = 0, c_cond = 1, level = 0.99)
```

fit\_mreg

fit_mreg	Fit a model for the mediator given the treatment and covariates.

# Description

lm is called if mreg = "linear". glm is called with family = binomial() if mreg = "logistic".

# Usage

```
fit_mreg(mreg, data, avar, mvar, cvar)
```

### **Arguments**

mreg	A character vector of length 1. Mediator regression type: "linear" or "logistic".
data	Data frame containing the relevant variables.
avar	A character vector of length 1. Treatment variable name.
mvar	A character vector of length 1. Mediator variable name.
cvar	A character vector of length > 0. Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.

### Value

A regression object of class lm (linear) or glm (logistic)

fit_yreg	Fit a model for the outcome given the treatment, mediator, and covari-
	ates.

# **Description**

The outcome model type yreg can be one of the following "linear", "logistic", "loglinear" (implemented as modified Poisson), "poisson", "negbin", "survCox", "survAFT\_exp", or "survAFT\_weibull".

# Usage

```
fit_yreg(yreg, data, yvar, avar, mvar, cvar, eventvar, interaction)
```

yreg	A character vector of length 1. Outcome regression type: "linear", "logistic", "loglinear", "poisson", "negbin", "survCox", "survAFT_exp", or "survAFT_weibull".
data	Data frame containing the relevant variables.
yvar	A character vector of length 1. Outcome variable name. It should be the time variable for survival outcomes.
avar	A character vector of length 1. Treatment variable name.

mvar A character vector of length 1. Mediator variable name.

cvar A character vector of length > 0. Covariate names. Use NULL if there is no

covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.

eventvar An character vector of length 1. Only required for survival outcome regression

models. Note that the coding is 1 for event and 0 for censoring, following the R

survival package convention.

interaction A logical vector of length 1. Default to TRUE. Whether to include a mediator-

treatment interaction term in the outcome regression model.

#### **Details**

The outcome regression functions to be called are the following:

- "linear" lm
- "logistic" glm
- "loglinear" glm (modified Poisson)
- "poisson" glm
- "negbin" glm.nb
- "survCox" coxph
- "survAFT\_exp" survreg
- "survAFT\_weibull" survreg

### Value

Model fit object from on of the above regression functions.

```
grad_prop_med_yreg_linear
```

Calculate the gradient of the proportion mediated for yreg linear.

### Description

Calculate the gradient of the proportion mediated for yreg linear case.

#### Usage

```
grad_prop_med_yreg_linear(pnde, tnie)
```

### **Arguments**

pnde Pure natural direct effect.
tnie Total natural indirect effect.

#### Value

Proportion mediated value.

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new_regmedint Low level constructor for a regmedint S3 class object.	
--	--

# Description

This is not a user function and meant to be executed within the regmedint function after validating the arguments.

# Usage

```
new_regmedint(
  data,
  yvar,
  avar,
  mvar,
  cvar,
  eventvar,
  a0,
  a1,
  m_cde,
  c_cond,
  yreg,
  mreg,
  interaction,
  {\tt case control}
)
```

data	Data frame containing the relevant variables.
yvar	A character vector of length 1. Outcome variable name. It should be the time variable for survival outcomes.
avar	A character vector of length 1. Treatment variable name.
mvar	A character vector of length 1. Mediator variable name.
cvar	A character vector of length $> 0$ . Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.
eventvar	An character vector of length 1. Only required for survival outcome regression models. Note that the coding is 1 for event and 0 for censoring, following the R survival package convention.
a0	A numeric vector of length 1. Reference level of treatment variable that is considered "untreated" or "unexposed".
a1	A numeric vector of length 1.
m_cde	A numeric vector of length 1. Mediator level at which controlled direct effect is evaluated at.
c_cond	A numeric vector of the same length as cvar. Covariate vector at which conditional effects are evaluated at.

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yreg	A character vector of length 1. Outcome regression type: "linear", "logistic", "loglinear", "poisson", "negbin", "survCox", "survAFT_exp", or "survAFT_weibull".
mreg	A character vector of length 1. Mediator regression type: "linear" or "logistic".
interaction	A logical vector of length 1. Default to TRUE. Whether to include a mediator-treatment interaction term in the outcome regression model.
casecontrol	A logical vector of length 1. Default to FALSE. Whether data comes from a case-control study.

### Value

A regmedint object.

print.regmedint print method for regmedint object

# Description

Print the mreg\_fit, yreg\_fit, and the mediation analysis effect estimates.

# Usage

```
## S3 method for class 'regmedint'
print(
    x,
    a0 = NULL,
    a1 = NULL,
    m_cde = NULL,
    c_cond = NULL,
    args_mreg_fit = list(),
    args_yreg_fit = list(),
    ...
)
```

x	An object of the regmedint class.
a0	A numeric vector of length one.
a1	A numeric vector of length one.
m_cde	A numeric vector of length one. A mediator value at which the controlled direct effect (CDE) conditional on the adjustment covariates is evaluated. If not provided, the default value supplied to the call to regmedint will be used. Only the CDE is affected.
c_cond	A numeric vector as long as the number of adjustment covariates. A set of covariate values at which the conditional natural effects are evaluated.
args_mreg_fit	A named list of argument to be passed to the method for the mreg_fit object.
args_yreg_fit	A named list of argument to be passed to the method for the mreg_fit object.
• • •	For compatibility with the generic. Ignored.

#### Value

Invisibly return the regmedint class object as is.

### **Examples**

```
library(regmedint)
data(vv2015)
regmedint_obj <- regmedint(data = vv2015,</pre>
                           ## Variables
                           yvar = "y",
                           avar = "x",
                           mvar = "m"
                           cvar = c("c"),
                           eventvar = "event",
                           ## Values at which effects are evaluated
                           a0 = 0,
                           a1 = 1,
                           m\_cde = 1,
                           c\_cond = 0.5,
                           ## Model types
                           mreg = "logistic",
                           yreg = "survAFT_weibull",
                           ## Additional specification
                           interaction = TRUE,
                           casecontrol = FALSE)
## Implicit printing
regmedint_obj
## Explicit printing
print(regmedint_obj)
## Evaluate at different values
print(regmedint_obj, m_cde = 0, c_cond = 1)
```

```
print.summary_regmedint
```

Print method for summary objects from summary.regmedint

### **Description**

Print results contained in a summary\_regmedint object with additional explanation regarding the evaluation settings.

#### Usage

```
## S3 method for class 'summary_regmedint'
print(x, ...)
```

- x An object of the class summary\_regmedint.
- ... For compatibility with the generic function.

#### Value

Invisibly return the first argument.

# **Examples**

```
library(regmedint)
data(vv2015)
regmedint_obj <- regmedint(data = vv2015,</pre>
                            ## Variables
                            yvar = "y",
                            avar = "x",
                            mvar = "m",
                            cvar = c("c"),
                            eventvar = "event",
                            ## Values at which effects are evaluated
                            a0 = 0,
                            a1 = 1,
                            m_{cde} = 1,
                            c\_cond = 0.5,
                            ## Model types
                            mreg = "logistic",
                            yreg = "survAFT_weibull",
                            ## Additional specification
                            interaction = TRUE,
                            casecontrol = FALSE)
## Implicit printing
summary(regmedint_obj)
## Explicit printing
print(summary(regmedint_obj))
```

prop\_med\_yreg\_linear Calculate the proportion mediated for yreg linear.

### **Description**

Calculate the proportion mediated on the mean difference scale.

### Usage

```
prop_med_yreg_linear(pnde, tnie)
```

# Arguments

pnde Pure natural direct effect.
tnie Total natural indirect effect.

### Value

Proportion mediated value.

```
prop_med_yreg_logistic
```

Calculate the proportion mediated for yreg logistic.

#### **Description**

Calculate the approximate proportion mediated on the risk difference scale.

### Usage

```
prop_med_yreg_logistic(pnde, tnie)
```

### **Arguments**

pnde Pure natural direct effect on the log scale.
tnie Total natural indirect effect on the log scale.

### Value

Proportion mediated value.

regmedint

regmedint: A package for regression-based causal mediation analysis

### **Description**

The package is a simple R implementation of the SAS macro as described in Valeri & VanderWeele 2013 and Valeri & VanderWeele 2015 https://www.hsph.harvard.edu/tyler-vanderweele/tools-and-tutorials/.

This is a user-interface for regression-based causal mediation analysis as described in Valeri & VanderWeele 2013 and Valeri & VanderWeele 2015.

### Usage

```
regmedint(
 data,
 yvar,
 avar,
 mvar,
 cvar,
 eventvar = NULL,
 a0,
 a1,
 m_cde,
 c_cond,
 mreg,
 yreg,
 na_omit = FALSE,
 interaction = TRUE,
  casecontrol = FALSE
)
```

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# Arguments

data	Data frame containing the relevant variables.
yvar	A character vector of length 1. Outcome variable name. It should be the time variable for survival outcomes.
avar	A character vector of length 1. Treatment variable name.
mvar	A character vector of length 1. Mediator variable name.
cvar	A character vector of length > 0. Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.
eventvar	An character vector of length 1. Only required for survival outcome regression models. Note that the coding is 1 for event and 0 for censoring, following the R survival package convention.
a0	A numeric vector of length 1. Reference level of treatment variable that is considered "untreated" or "unexposed".
a1	A numeric vector of length 1.
m_cde	A numeric vector of length 1. Mediator level at which controlled direct effect is evaluated at.
c_cond	A numeric vector of the same length as cvar. Covariate vector at which conditional effects are evaluated at.
mreg	A character vector of length 1. Mediator regression type: "linear" or "logistic".
yreg	A character vector of length 1. Outcome regression type: "linear", "logistic", "loglinear", "poisson", "negbin", "survCox", "survAFT_exp", or "survAFT_weibull".
na_omit	A logical vector of length 1. Default to FALSE. Whether to use na.omit() function in stats package to remove NAs in columns of interest before fitting the models.
interaction	A logical vector of length 1. Default to TRUE. Whether to include a mediator-treatment interaction term in the outcome regression model.
casecontrol	A logical vector of length 1. Default to FALSE. Whether data comes from a case-control study.

# Value

regmedint object, which is a list containing the mediator regression object, the outcome regression object, and the regression-based mediation results.

# Fitting models

Use the regmedint function to fit models and set up regression-based causal mediation analysis.

# **Examining results**

Several methods are available to examine the regmedint object. print summary coef confint FIXME: Document once implemented.

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#### **Examples**

```
library(regmedint)
data(vv2015)
regmedint_obj <- regmedint(data = vv2015,</pre>
                            ## Variables
                            yvar = "y",
                            avar = "x",
                            mvar = "m",
                            cvar = c("c"),
                            eventvar = "event",
                            ## Values at which effects are evaluated
                            a0 = 0,
                            a1 = 1,
                            m\_cde = 1,
                            c\_cond = 0.5,
                            ## Model types
                            mreg = "logistic",
                            yreg = "survAFT_weibull",
                            ## Additional specification
                            interaction = TRUE,
                            casecontrol = FALSE)
summary(regmedint_obj)
```

summary.regmedint

summary method for regmedint object

### Description

Summarize the mreg\_fit, yreg\_fit, and the mediation analysis effect estimates.

### Usage

```
## S3 method for class 'regmedint'
summary(
   object,
   a0 = NULL,
   a1 = NULL,
   m_cde = NULL,
   c_cond = NULL,
   args_mreg_fit = list(),
   args_yreg_fit = list(),
   exponentiate = FALSE,
   level = 0.95,
   ...
)
```

# Arguments

a0 An object of the regmedint class.
 a0 A numeric vector of length one.
 a1 A numeric vector of length one.

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m_cde	A numeric vector of length one. A mediator value at which the controlled direct effect (CDE) conditional on the adjustment covariates is evaluated. If not provided, the default value supplied to the call to regmedint will be used. Only the CDE is affected.
c_cond	A numeric vector as long as the number of adjustment covariates. A set of covariate values at which the conditional natural effects are evaluated.
args_mreg_fit	A named list of argument to be passed to the method for the mreg_fit object.
args_yreg_fit	A named list of argument to be passed to the method for the mreg_fit object.
exponentiate	Whether to add exponentiated point and confidence limit estimates. When yreg = "linear", it is ignored.
level	Confidence level for the confidence intervals.
	For compatibility with the generic. Ignored.

#### Value

A summary\_regmedint object, which is a list containing the summary objects of the mreg\_fit and the yreg\_fit as well as the mediation analysis results.

# **Examples**

```
library(regmedint)
data(vv2015)
regmedint_obj <- regmedint(data = vv2015,</pre>
                           ## Variables
                           yvar = "y",
                           avar = "x",
                           mvar = "m",
                           cvar = c("c"),
                           eventvar = "event",
                           ## Values at which effects are evaluated
                           a0 = 0,
                           a1 = 1,
                           m\_cde = 1,
                           c\_cond = 0.5,
                           ## Model types
                           mreg = "logistic",
                           yreg = "survAFT_weibull",
                           ## Additional specification
                           interaction = TRUE,
                           casecontrol = FALSE)
## Detailed result with summary
summary(regmedint_obj)
## Add exponentiate results for non-linear outcome models
summary(regmedint_obj, exponentiate = TRUE)
## Evaluate at different values
summary(regmedint_obj, m_cde = 0, c_cond = 1)
## Change confidence level
summary(regmedint_obj, m_cde = 0, c_cond = 1, level = 0.99)
```

```
summary.regmedint_mod_poisson
```

Summary with robust sandwich variance estimator for modified Poisson

#### **Description**

This is a version of summary, glm modified to use the robust variance estimator sandwich.

#### Usage

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

### **Arguments**

object A model object of the class regmedint\_mod\_poisson

... For compatibility with the generic.

# Value

An object of the class summary.glm

theta\_hat

Create a vector of coefficients from the outcome model (yreg)

# **Description**

This function extracts coef from yreg\_fit and pads with zeros appropriately to create a named vector consistently having the following elements: (Intercept): A zero element is added for yreg = "survCox" for which no intercept is estimated (the baseline hazard is left unspecified). avar mvar avar:mvar: A zero element is added when interaction = FALSE. cvar: This part is eliminated when cvar = NULL.

### Usage

```
theta_hat(yreg, yreg_fit, avar, mvar, cvar, interaction)
```

### **Arguments**

yreg	A character vector of length 1. Outcome regression type: "linear", "logistic", "loglinear", "poisson", "negbin", "survCox", "survAFT_exp", or "survAFT_weibull".
yreg_fit	Model fit object for yreg (outcome model).
avar	A character vector of length 1. Treatment variable name.
mvar	A character vector of length 1. Mediator variable name.
cvar	A character vector of length > 0. Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, myar is not. Thus, there should usually be some confounder(s) to

covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.

interaction A logical vector of length 1. Default to TRUE. Whether to include a mediator-

treatment interaction term in the outcome regression model.

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### Value

A named numeric vector of coefficients.

validate\_args

Validate arguments to regmedint before passing to other functions

# Description

Internal functions (usually) do not validate arguments, thus, we need to make sure informative errors are raised when the arguments are not safe for subsequent computation.

# Usage

```
validate_args(
  data,
  yvar,
  avar,
  mvar,
  cvar,
  eventvar,
  a0,
  a1,
  m_cde,
  c_cond,
  mreg,
  yreg,
  interaction,
  {\tt case control}
)
```

data	Data frame containing the relevant variables.
yvar	A character vector of length 1. Outcome variable name. It should be the time variable for survival outcomes.
avar	A character vector of length 1. Treatment variable name.
mvar	A character vector of length 1. Mediator variable name.
cvar	A character vector of length $> 0$ . Covariate names. Use NULL if there is no covariate. However, this is a highly suspicious situation. Even if avar is randomized, mvar is not. Thus, there should usually be some confounder(s) to account for the common cause structure (confounding) between avar and yvar.
eventvar	An character vector of length 1. Only required for survival outcome regression models. Note that the coding is 1 for event and 0 for censoring, following the R survival package convention.
a0	A numeric vector of length 1. Reference level of treatment variable that is considered "untreated" or "unexposed".
a1	A numeric vector of length 1.

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m_cde	A numeric vector of length 1. Mediator level at which controlled direct effect is evaluated at.
c_cond	A numeric vector of the same length as cvar. Covariate vector at which conditional effects are evaluated at.
mreg	A character vector of length 1. Mediator regression type: "linear" or "logistic".
yreg	A character vector of length 1. Outcome regression type: "linear", "logistic", "loglinear", "poisson", "negbin", "survCox", "survAFT_exp", or "survAFT_weibull".
interaction	A logical vector of length 1. Default to TRUE. Whether to include a mediator-treatment interaction term in the outcome regression model.
casecontrol	A logical vector of length 1. Default to FALSE. Whether data comes from a case-control study.

# Value

No return value, called for side effects.

	validate_regmedint	Validate soundness of a regmedint object.	
--	--------------------	---	--

# Description

Check the structure of a proposed regmedint object for soundness.

# Usage

```
validate_regmedint(x)
```

# Arguments

x A regmedint object.

### Value

No return value, called for side effects.

vcov.regmedint	Extract variance estimates in the vcov form.

# Description

Extract variance estimates evaluated at a0, a1, m\_cde, and c\_cond.

# Usage

```
## S3 method for class 'regmedint'
vcov(object, a0 = NULL, a1 = NULL, m_cde = NULL, c_cond = NULL, ...)
```

### **Arguments**

object	An object of the regmedint class.
a0	A numeric vector of length one.
a1	A numeric vector of length one.
m_cde	A numeric vector of length one. A mediator value at which the controlled direct effect (CDE) conditional on the adjustment covariates is evaluated. If not provided, the default value supplied to the call to regmedint will be used. Only the CDE is affected.
c_cond	A numeric vector as long as the number of adjustment covariates. A set of covariate values at which the conditional natural effects are evaluated.
	For compatibility with the generic. Ignored.

### Value

A numeric matrix with the diagonals populated with variance estimates. Off-diagnonals are NA since these are not estimated.

### **Examples**

```
library(regmedint)
data(vv2015)
regmedint_obj <- regmedint(data = vv2015,</pre>
                            ## Variables
                            yvar = "y",
avar = "x",
                            mvar = "m",
                            cvar = c("c"),
                            eventvar = "event",
                            ## Values at which effects are evaluated
                            a0 = 0,
                            a1 = 1,
                            m\_cde = 1,
                            c\_cond = 0.5,
                            ## Model types
                            mreg = "logistic",
                            yreg = "survAFT_weibull",
                            ## Additional specification
                            interaction = TRUE,
                            casecontrol = FALSE)
vcov(regmedint_obj)
## Evaluate at different values
vcov(regmedint_obj, m_cde = 0, c_cond = 1)
```

```
vcov.regmedint_mod_poisson
```

Robust sandwich variance estimator for modified Poisson

### **Description**

Provide robust sandwich variance-covariance estimate using sandwich.

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#### Usage

```
## S3 method for class 'regmedint_mod_poisson'
vcov(object, ...)
```

# Arguments

object A model object of the class regmedint\_mod\_poisson ... For compatibility with the generic.

### Value

A variance-covariance matrix using the sandwich.

vv2015

Example dataset from Valeri and VanderWeele 2015.

# Description

An example dataset from Valeri and VanderWeele (2015) <doi:10.1097/EDE.00000000000000253>.

# Usage

vv2015

#### **Format**

A tibble with 100 rows and 7 variables:

- id Positive integer id.
- x Binary treatment assignment variable.
- m Binary mediator variable.
- y Time to event outcome variable.

cens Binary censoring indicator. Censored is 1.

c Continuous confounder variable.

event Binary event indicator. Event is 1.

### **Source**

https://www.hsph.harvard.edu/tyler-vanderweele/tools-and-tutorials/

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