

Package ‘risks’

June 13, 2023

Title Estimate Risk Ratios and Risk Differences using Regression

Version 0.4.2

Description Risk ratios and risk differences are estimated using regression models that allow for binary, categorical, and continuous exposures and confounders. Implemented are marginal standardization after fitting logistic models (g-computation) with delta-method and bootstrap standard errors, Miettinen's case-duplication approach (Schouten et al. 1993, <[doi:10.1002/sim.4780121808](https://doi.org/10.1002/sim.4780121808)>), log-binomial (Poisson) models with empirical variance (Zou 2004, <[doi:10.1093/aje/kwh090](https://doi.org/10.1093/aje/kwh090)>), binomial models with starting values from Poisson models (Spiegelman and Hertzmark 2005, <[doi:10.1093/aje/kwi188](https://doi.org/10.1093/aje/kwi188)>), and others.

License GPL-3

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BugReports <https://github.com/stopsack/risks/issues>

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breastcancer

*Breast Cancer Data***Description**

A cohort of women with breast cancer and complete follow-up, as used by Spiegelman and Hertzmark (Am J Epidemiol 2005) and Greenland (Am J Epidemiol 2004).

Usage

breastcancer

Format

breastcancer:

A tibble with 192 rows and 3 columns:

death Death, binary: 0, 1**stage** Cancer stage, 3 categories**receptor** Hormone receptor status, binary: "High", "Low" ...**Source**

Newman SC. Biostatistical methods in epidemiology. New York, NY: Wiley, 2001, table 5.3

confint.duplicate *Clustering-corrected confidence intervals for case duplication model*

Description

Estimate confidence intervals for the case duplication model with robust/sandwich/empirical covariance structure.

Usage

```
## S3 method for class 'duplicate'
confint(object, parm = NULL, level = 0.95, ...)
```

Arguments

object	Fitted model
parm	Not used
level	Confidence level, defaults to 0.95
...	Additional arguments, not used

Value

Matrix: First column, lower bound; second column, upper bound.

confint.margstd_boot *Bootstrap confidence intervals*

Description

Confidence intervals for models fit using marginal standardization based on parametric bootstrapping.

Usage

```
## S3 method for class 'margstd_boot'
confint(
  object,
  parm = NULL,
  level = 0.95,
  bootrepeats = 1000,
  bootci = c("bca", "normal", "nonpar"),
  jacksd = FALSE,
  ...
)
```

Arguments

object	Model fitted through marginal standardization
parm	Not used, for compatibility
level	Confidence level, defaults to 0.95.
bootrepeats	Bootstrap repeats. Defaults to 1000. Consider increasing.
bootci	Type of bootstrap confidence interval: <ul style="list-style-type: none"> • "bca" Default. Parametric BCa (bias-corrected accelerated) confidence intervals. • "normal" Parametric normality-based confidence intervals, which require lower repeat numbers but are less accurate and may result in invalid results for ratios. • "nonpar" Non-parametric BCa confidence intervals, which should be used with caution because of the risk of sparse-data bias with non-parametric bootstrapping.
jacksd	Return jackknife Monte-Carlo error for the confidence limits? Only functional with BCa confidence intervals. Defaults to FALSE.
...	Not used

Value

Matrix: First column, lower bound; second column, upper bound.

confint.margstd_delta *Delta method confidence intervals*

Description

Confidence intervals for models fit using marginal standardization based on the delta method.

Usage

```
## S3 method for class 'margstd_delta'
confint(object, parm = NULL, level = 0.95, ...)
```

Arguments

object	Model fitted through marginal standardization, delta method
parm	Not used, for compatibility
level	Confidence level, defaults to 0.95.
...	Not used

Value

Matrix: First column, lower bound; second column, upper bound.

confint.robpoisson *Robust confidence intervals for Poisson model*

Description

Estimate confidence intervals for the Poisson model with robust/sandwich/empirical covariance structure.

Usage

```
## S3 method for class 'robpoisson'
confint(object, parm = NULL, level = 0.95, ...)
```

Arguments

object	Fitted model
parm	Not used
level	Confidence level, defaults to 0.95
...	Additional arguments, not used

Value

Matrix: First column, lower bound; second column, upper bound.

print.risks *Print model*

Description

Print fitted risks model. The only change, compared to `print.glm()`, is the addition of the main type of model: relative risk or risk difference. If multiple models were fitted via `approach = "all"`, then the first converged model will be printed.

Usage

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

Arguments

x	Fitted model
...	Passed to <code>print.glm()</code>

Value

No return value, called for printing

```
print.summary.risks Print model summary
```

Description

Print summaries for "risks" models. The printout is the same as for regular summaries of generalized linear models fit via `stats::glm()`, except that the type of "risks" model is printed first (e.g., "Poisson model with robust covariance") and confidence intervals for model parameters are printed at the end.

Usage

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

Arguments

x	Model
...	Passed on

Value

No return value, called for printing

```
riskratio Fit risk ratio and risk difference models
```

Description

`riskratio` and `riskdiff` provide a flexible interface to fitting risk ratio and risk difference models.

In cohort studies with a binary outcome, risk ratios and risk differences are typically more appropriate to report than odds ratios from logistic regression, yet such models have historically been difficult to implement in standard software.

The `risks` package selects an efficient way to fit risk ratio or risk difference models successfully, which will converge whenever logistic models converge. Optionally, a specific approach to model fitting can also be requested. Implemented are Poisson models with robust covariance, binomial models, logistic models with case duplication, binomial models aided in convergence by starting values obtained through Poisson models or logistic models with case duplication, binomial models fitted via combinatorial expectation maximization (optionally also with Poisson starting values), and estimates obtained via marginal standardization after logistic regression with bootstrapped or delta method for confidence intervals.

Adjusting for covariates (e.g., confounders) in the model specification (`formula =`) is possible.

Usage

```

riskratio(
  formula,
  data,
  approach = c("auto", "all", "robpoisson", "duplicate", "glm", "glm_startp",
    "glm_startd", "glm_cem", "glm_cem_startp", "margstd_boot", "margstd_delta",
    "logistic", "legacy"),
  variable = NULL,
  at = NULL,
  ...
)

riskdiff(
  formula,
  data,
  approach = c("auto", "all", "robpoisson", "glm", "glm_startp", "glm_cem",
    "glm_cem_startp", "margstd_boot", "margstd_delta", "legacy"),
  variable = NULL,
  at = NULL,
  ...
)

```

Arguments

formula	A formula object of the form response ~ predictors.
data	A tibble or data.frame object.
approach	Optional: Method for model fitting. <ul style="list-style-type: none"> • "auto" (default) is recommended; it will return results of "margstd_delta" unless interaction terms between exposure and confounders are included. This these cases, results from "margstd_boot" are returned. • "all" will attempt to fit the model via all implemented approaches to allow for comparisons. • "legacy" selects the most efficient approach that converges and ensures that predicted probabilities are within range (< 1).

The other options allow for directly selecting a fitting approach, some of which may not converge or yield out-of-range predicted probabilities. See full documentation for details.

- "glm" Binomial model.
- "glm_startp" Binomial model with starting values from Poisson model.
- "glm_startd" Binomial model with starting values from logistic model with case duplication.
- "robpoisson" Poisson model with robust covariance.
- "duplicate" Logistic model with duplication of cases. Only available in riskratio().
- "glm_cem" Binomial model fitted with combinatorial expectation maximization.

	<ul style="list-style-type: none"> • "glm_cem_startp" As <code>glm_cem</code>, with Poisson starting values. • "margstd_boot" Marginal standardization after logistic model, bootstrap standard errors/confidence intervals. • "margstd_delta" Marginal standardization after logistic model, delta method standard errors/confidence intervals. • "logistic" For comparison only: the logistic model. Only available in <code>riskratio()</code>.
variable	Optional: exposure variable to use for marginal standardization. If <code>variable</code> is not provided and marginal standardization is attempted, then the first variable in the model is used as the exposure. Levels are determined automatically for variables types <code>logical</code> , <code>character</code> , <code>factor</code> and can optionally be supplied via <code>at =</code> .
at	Optional: Levels of exposure variable <code>variable</code> for marginal standardization. <code>at =</code> determines the levels at which contrasts of the exposure are to be assessed. The level listed first is used as the reference. Levels must exist in the data for <code>character</code> , <code>factor</code> or <code>ordered factor</code> variables. For numeric variables, levels that do not exist in the data can be interpolations or extrapolations; if levels exceed the extremes of the data (extrapolation), a warning will be displayed.
...	Optional: Further arguments passed to fitting functions (<code>glm</code> , <code>logbin</code> , or <code>addreg</code>).

Value

Fitted model. This object can be passed on to post-processing functions:

- `summary.risks`: an overview of results (risks-specific S3 methods: `summary.robpoisson`, `summary.margstd_boot`, `summary.margstd_delta`).
- `tidy.risks`: a tibble of coefficients and confidence intervals.

Standard post-processing functions can also be used:

- `coef`: a vector of coefficients.
- `confint`: a matrix of confidence intervals (risks-specific S3 methods: `confint.robpoisson`, `confint.margstd_boot`, `confint.margstd_delta`).
- `predict.glm`(`type = "response"`): fitted values (predictions).
- `residuals`: residuals.

If model fitting using all possible approaches was requested via `approach = "all"`, then their results can be retrieved from the list `all_models` in the returned object (e.g., `fit$all_models[[1]]`, `fit$all_models[[2]]`, etc.).

Functions

- `riskratio()`: Fit risk ratio models
- `riskdiff()`: Fit risk difference models

References

- Wacholder S. Binomial regression in GLIM: Estimating risk ratios and risk differences. *Am J Epidemiol* 1986;123:174-184. (Binomial regression models; approach = "glm")
- Spiegelman D, Hertzmark E. Easy SAS Calculations for Risk or Prevalence Ratios and Differences. *Am J Epidemiol* 2005;162:199-200. (Binomial models fitted used starting values from Poisson models; approach = "glm_start")
- Zou G. A modified Poisson regression approach to prospective studies with binary data. *Am J Epidemiol* 2004;159:702-706. (Poisson model with robust/sandwich standard errors; approach = "robpoisson")
- Schouten EG, Dekker JM, Kok FJ, Le Cessie S, Van Houwelingen HC, Pool J, Vandembroucke JP. Risk ratio and rate ratio estimation in case-cohort designs: hypertension and cardiovascular mortality. *Stat Med* 1993;12:1733-45; (Logistic model with case duplication and cluster-robust standard errors, approach = "duplicate").
- Donoghoe MW, Marschner IC. logbin: An R Package for Relative Risk Regression Using the Log-Binomial Model. *J Stat Softw* 2018;86(9). (Log-binomial models fitted via combinatorial expectation maximization; riskratio(approach = "glm_cem"))
- Donoghoe MW, Marschner IC. Stable computational methods for additive binomial models with application to adjusted risk differences. *Comput Stat Data Anal* 2014;80:184-96. (Additive binomial models fitted via combinatorial expectation maximization; riskdiff(approach = "glm_cem"))
- Localio AR, Margolis DJ, Berlin JA. Relative risks and confidence intervals were easily computed indirectly from multivariable logistic regression. *J Clin Epidemiol* 2007;60(9):874-82. (Marginal standardization after fitting a logistic model; approach = "margstd_boot")

Examples

```
data(breastcancer) # Cohort study with binary outcome
                  # See for details: help(breastcancer)

# Risk ratio model
fit_rr <- riskratio(formula = death ~ stage + receptor, data = breastcancer)
fit_rr
summary(fit_rr)

# Risk difference model
fit_rd <- riskdiff(formula = death ~ stage + receptor, data = breastcancer)
fit_rd
summary(fit_rd)
```

risks

risks: Estimate risk ratios and risk differences using regression

Description

The risks package allows for fitting risk ratio and risk difference models using regression.

risks functions

`riskratio`: Fit risk ratio models.

`riskdiff`: Fit risk difference models.

`summary.risks`: Summarize fitted model.

`tidy.risks`: Tibble (data frame) of parameters, coefficients, standard errors, confidence intervals.

`confint.robpoisson`, `confint.duplicate`, `confint.margstd_boot`, `confint.margstd_delta`: Confidence intervals. (Standard confidence intervals for generalized linear models are used for other models.)

See Also

<https://github.com/stopsack/risks>

rr_rd_mantel_haenszel *Risk Ratios and Risk Differences from Mantel-Haenszel Estimators*

Description

This function implements the Mantel-Haenszel estimators for risk ratio and risk differences for a binary or categorical exposure and one or more categorical confounder(s). Compare to estimates from regression models.

Usage

```
rr_rd_mantel_haenszel(
  data,
  exposure,
  outcome,
  confounders,
  estimand = c("rr", "rd"),
  conf.level = 0.95
)
```

Arguments

<code>data</code>	Data set.
<code>exposure</code>	Exposure variable. Must be binary or categorical. The first level is treated as unexposed.
<code>outcome</code>	Outcome variable. Must be binary.
<code>confounders</code>	Optional. Binary or categorical variable(s) to perform stratification over. Supply more than one variable using <code>confounders = c(var1, var2)</code> .
<code>estimand</code>	Optional. "rr" for risk ratio; "rd" for risk difference. Defaults to "rr".
<code>conf.level</code>	Optional. Confidence level. Defaults to 0.95.

Value

Tibble in `tidy` format with

- term the (non-reference) exposure levels
- estimate Risk ratio (on log scale) or risk difference
- `std.error`, `conf.low`, and `conf.high` Square-root of M-H variance estimate, and the corresponding confidence limits (on log scale for RR)
- model always "mh"
- estimand "rr" or "rd"

References

Greenland S, Rothman KJ. Introduction to Stratified Analysis. In: Rothman KJ, Greenland S, Lash TL. Modern Epidemiology. 3rd edn. Lippincott Williams & Wilkins: Philadelphia, PA 2008. Page 275. Risk ratios: formulae 15-18, -20, -22. Risk differences: formulae 15-18, -19, -21.

Examples

```
# Newman SC. Biostatistical methods in epidemiology. New York, NY:
# Wiley, 2001, table 5.3

library(tibble) # used to set up example data
dat <- tibble(
  death   = c(rep(1, 54), rep(0, 138)),
  stage   = c(rep("Stage I", 7), rep("Stage II", 26), rep("Stage III", 21),
              rep("Stage I", 60), rep("Stage II", 70), rep("Stage III", 8)),
  receptor = c(rep("Low", 2), rep("High", 5), rep("Low", 9), rep("High", 17),
              rep("Low", 12), rep("High", 9), rep("Low", 10), rep("High", 50),
              rep("Low", 13), rep("High", 57), rep("Low", 2), rep("High", 6)))

# Risk difference
rr_rd_mantel_haenszel(
  data = dat,
  exposure = stage,
  outcome = death,
  confounders = receptor,
  estimand = "rd")

# Risk ratio, log scale:
result <- rr_rd_mantel_haenszel(
  data = dat,
  exposure = stage,
  outcome = death,
  confounders = receptor,
  estimand = "rr")
result

# Risk ratio, exponentiated:
result %>%
  dplyr::mutate(dplyr::across(.cols = c(estimate, conf.low, conf.high),
                             .fns = exp))
```

summary.duplicate	<i>Summary for logistic model with case duplication and cluster-robust covariance</i>
-------------------	---

Description

Summarize results from fitting a logistic model with case duplication and cluster-robust covariance. The output is the same as for a regular `summary(glm(...))`, except for using cluster-robust standard errors.

Usage

```
## S3 method for class 'duplicate'
summary(
  object,
  dispersion = NULL,
  correlation = FALSE,
  symbolic.cor = FALSE,
  ...
)
```

Arguments

object	Model
dispersion	Not used
correlation	Not used
symbolic.cor	Not used
...	Other arguments, not used

Value

Model summary (list)

summary.margstd_boot	<i>Summary for models using marginal standardization</i>
----------------------	--

Description

Summary for models using marginal standardization

Usage

```
## S3 method for class 'margstd_boot'
summary(
  object,
  dispersion = NULL,
  correlation = FALSE,
  symbolic.cor = FALSE,
  level = 0.95,
  bootrepeats = 1000,
  bootci = c("bca", "normal", "nonpar"),
  ...
)
```

Arguments

object	Model
dispersion	Not used
correlation	Not used
symbolic.cor	Not used
level	Confidence level, defaults to 0.95.
bootrepeats	Bootstrap repeats for standard errors. Defaults to 1000. Consider increasing.
bootci	Type of bootstrap confidence interval: <ul style="list-style-type: none"> • "bca" Default. Parametric BCa (bias-corrected accelerated) confidence intervals. • "normal" Parametric normality-based confidence intervals, which require lower repeat numbers but are less accurate and may result in invalid results for ratios. • "nonpar" Non-parametric BCa confidence intervals, which should be used with caution because of the risk of sparse-data bias with non-parametric bootstrapping.
...	Not used

Value

Model summary (list)

summary.margstd_delta *Summary for models using marginal standardization with delta method SEs*

Description

Summary for models using marginal standardization with delta method SEs

Usage

```
## S3 method for class 'margstd_delta'
summary(
  object,
  dispersion = NULL,
  correlation = FALSE,
  symbolic.cor = FALSE,
  level = 0.95,
  ...
)
```

Arguments

object	Model
dispersion	Not used
correlation	Not used
symbolic.cor	Not used
level	Confidence level, defaults to 0.95.
...	Not used

Value

Model summary (list)

summary.risks	<i>Generate model summary</i>
---------------	-------------------------------

Description

Determine type of risks model fitted and generate appropriate summary.

Usage

```
## S3 method for class 'risks'
summary(object, conf.int = TRUE, default = TRUE, ...)
```

Arguments

object	Fitted model
conf.int	Add confidence intervals to printout? Defaults to TRUE.
default	Normal confidence intervals via confint.default()? Default to TRUE. By setting default = FALSE, profiling-based confidence intervals can be calculated for binomial models.
...	Passed on

Details

If multiple models were fitted (approach = "all"), then the first converged model is displayed. Other models can be accessed via the returned list \$all_models.

Value

Model summary (list)

summary.robpoisson *Summary for Poisson model with robust covariance*

Description

Summarize results from fitting a Poisson model with robust/empirical/sandwich covariance. The output is the same as for a regular summary(glm(. . .)), except for using robust standard errors.

Usage

```
## S3 method for class 'robpoisson'
summary(
  object,
  dispersion = NULL,
  correlation = FALSE,
  symbolic.cor = FALSE,
  ...
)
```

Arguments

object	Model
dispersion	Not used
correlation	Not used
symbolic.cor	Not used
...	Other arguments, not used

Value

Model summary (list)

tidy.risks

*Tidy model summaries for risks models***Description**

Obtain a tibble (data frame) with parameters, coefficients, standard errors, confidence limits, and p-values. A column with the type of model fitted is added.

Usage

```
## S3 method for class 'risks'
tidy(
  x,
  conf.int = TRUE,
  conf.level = 0.95,
  bootrepeats = 1000,
  bootci = c("bca", "normal", "nonpar"),
  bootverbose = FALSE,
  exponentiate = FALSE,
  default = TRUE,
  ...
)
```

Arguments

x	Model
conf.int	Show confidence intervals?
conf.level	Optional. Confidence level. Defaults to 0.95.
bootrepeats	Optional. Number of bootstrap repeats. Applicable to models fitted via marginal standardization and bootstrapping (approach = "margstd_boot"). Defaults to 1000. Strongly recommended to increase repeats to »1000.
bootci	Optional and applicable for approach = "margstd_boot" only. Type of bootstrap confidence interval: <ul style="list-style-type: none"> • "bca" Default. Parametric BCa (bias-corrected accelerated) confidence intervals. • "normal" Parametric normality-based confidence intervals, which require lower repeat numbers but are less accurate and may result in invalid results for ratios. • "nonpar" Non-parametric BCa confidence intervals, which should be used with caution because of the risk of sparse-data bias with non-parametric bootstrapping.
bootverbose	Optional. Add values of bootrepeats and bootci parameters and the jackknife-based Monte-Carlo error for the confidence limits (only for type = "bca") to the returned tibble? Defaults to FALSE.

exponentiate	Optional. Exponentiate coefficients and confidence limits? Defaults to FALSE. Setting exponentiate = TRUE is useful for relative risk models (log links).
default	Use default, normality-based confidence intervals? Defaults to TRUE. With default = FALSE, for binomial models only, profile likelihood-based confidence intervals can be calculated.
...	Passed on

Details

If multiple types of models are fitted, tidy() can be used to parameters for all models at once, in one tibble. The last column of the tibble includes the name of the model. See examples.

Value

tibble

Examples

```
# Define example data
library(broom) # provides tidy() function
dat <- tibble::tibble(
  death = c(rep(1, 54), rep(0, 138)),
  stage = c(rep("Stage I", 7), rep("Stage II", 26), rep("Stage III", 21),
            rep("Stage I", 60), rep("Stage II", 70), rep("Stage III", 8)),
  receptor = c(rep("Low", 2), rep("High", 5), rep("Low", 9), rep("High", 17),
              rep("Low", 12), rep("High", 9), rep("Low", 10), rep("High", 50),
              rep("Low", 13), rep("High", 57), rep("Low", 2), rep("High", 6)))

# Fit and tidy the model
fit_rr <- riskratio(formula = death ~ stage + receptor, data = dat)
tidy(fit_rr)

# Marginal standardization,
# increase number of bootstrap repeats:

fit_rr <- riskratio(
  formula = death ~ stage + receptor, data = dat,
  approach = "margstd_boot")
tidy(fit_rr, bootrepeats = 2000)

# Multiple types of models fitted:

fit_rr <- riskratio(formula = death ~ stage + receptor, data = dat,
                  approach = "all")
tidy(fit_rr)
```

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